

SCIENTIFIC REPORT OF EFSA AND ECDC The European Union Summary Report on antimicrobial resistance in zoonotic and indicator bacteria from humans, animals and food in 2012¹ European Food Safety Authority^{2,3} European Centre for Disease Prevention and Control^{2,3}

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ABSTRACT

The antimicrobial resistance data among zoonotic and indicator bacteria in 2012, submitted by 26 European Union Member States, were jointly analysed by the EFSA and the ECDC. Resistance in zoonotic Salmonella and Campylobacter isolates from humans, animals and food and resistance in indicator Escherichia coli, as well as data on methicillin-resistant Staphylococcus aureus, in animals and food were addressed. Resistance in human isolates was mainly interpreted using clinical breakpoints, while microbiological resistance in animal and food isolates was assessed using epidemiological cut-off values. Resistance was commonly found in isolates from humans, animals and food, although marked disparities in resistance were frequently observed between Member States. In Salmonella from humans, high resistance levels were recorded to ampicillin, sulfonamides and tetracyclines, while resistance to thirdgeneration cephalosporins and fluoroquinolones remained low. In Salmonella and Escherichia coli isolates from fowl, pigs, cattle and meat thereof, microbiological resistance to ampicillin, tetracyclines and sulfonamides was commonly detected, while microbiological resistance to third-generation cephalosporins was generally low. High to very high microbiological resistance to (fluoro)quinolones was observed in Salmonella isolates from turkeys, fowl and broiler meat. In *Campylobacter* from humans, resistance to ampicillin, ciprofloxacin, nalidixic acid and tetracyclines was high, while resistance to erythromycin was low to moderate. High to extremely high microbiological resistance to ciprofloxacin, nalidixic acid and tetracyclines was observed in Campylobacter isolates from fowl, broiler meat, pigs and cattle, whereas much lower levels were observed for erythromycin and gentamicin. Increasing trends for ciprofloxacin resistance was observed in Campylobacter isolates from humans, broilers and/or pigs in several Member States. Multiresistance and co-resistance to critically important antimicrobials in both human and animal isolates were presented, and for the first time, multi-resistance patterns in Salmonella serovars. Very few isolates from animals were co-resistant to critically important antimicrobials. A minority of isolates from animals belonging to a few Salmonella serovars (notably Kentucky and Infantis) were resistant to high levels of ciprofloxacin.

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KEY WORDS

antimicrobial resistance, zoonotic bacteria, indicator bacteria

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^{*} Changes have been made to human data in the *Campylobacter* chapter where resistance levels to ciprofloxacin in Iceland have been modified in Tables CA2 and CA4. In addition, text revision has been done in text boxes related to the revision of epidemiological cut-off values presented in the Introduction part on page 12, in the *E. coli* chapter on page 201, and in Materials and methods chapter on page 248. The changes do not affect the main findings and the overall discussion of the report. To avoid any confusion the original version of the output has been removed from the website but is available on request.



EUROPEAN UNION SUMMARY REPORT

Antimicrobial resistance in zoonotic and indicator bacteria from humans, animals and food in the European Union in 2012

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About EFSA

The European Food Safety Authority (EFSA), located in Parma, Italy, was established and funded by the European Union (EU) as an independent agency in 2002 following a series of food scares that caused the European public to voice concerns about food safety and the ability of regulatory authorities to protect consumers. EFSA provides objective scientific advice on all matters, in close collaboration with national authorities and in open consultation with its stakeholders, with a direct or indirect impact on food and feed safety, including animal health and welfare and plant protection. EFSA is also consulted on nutrition in relation to EU legislation. EFSA's work falls into two areas: risk assessment and risk communication. In particular, EFSA's risk assessments provide risk managers (EU institutions with political accountability, i.e. the European Commission, the European Parliament and the Council) with a sound scientific basis for defining policy-driven legislative or regulatory measures required to ensure a high level of consumer protection with regard to food and feed safety. EFSA communicates to the public in an open and transparent way on all matters within its remit. Collection and analysis of scientific data, identification of emerging risks and scientific support to the European Commission, particularly in the case of a food crisis, are also part of EFSA's mandate, as laid down in the founding Regulation (EC) No 178/2002⁴ of 28 January 2002.

About ECDC

The European Centre for Disease Prevention and Control (ECDC), an EU agency based in Stockholm, Sweden, was established in 2005. The objective of ECDC is to strengthen Europe's defences against infectious diseases. According to Article 3 of the founding Regulation (EC) No 851/2004⁵ of 21 April 2004, ECDC's mission is to identify, assess and communicate current and emerging threats to human health posed by infectious diseases. In order to achieve this mission, ECDC works in partnership with national public health bodies across Europe to strengthen and develop EU-wide disease surveillance and early warning systems. By working with experts throughout Europe, ECDC pools Europe's knowledge in health so as to develop authoritative scientific opinions about the risks posed by current and emerging infectious diseases.

About the report

Based on Article 33 in the Regulation (EC) 178/2002, EFSA's Zoonoses Unit is responsible for examining data on zoonoses, antimicrobial resistance and food-borne outbreaks collected from the Member States in accordance with Directive 2003/99/EC⁶ and for preparing the European Union Summary Report from the results. Regarding antimicrobial resistance data from 2012, this European Union Summary Report was produced in collaboration with ECDC and the Animal Health and Veterinary Laboratories Agency (AHVLA), the United Kingdom and the University of Hasselt in Belgium, contracted by EFSA.

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⁴ Regulation (EC) No 178/2002 of the European Parliament and of the Council of 28 January 2002 laying down the general principles and requirements of food law, establishing the European Food Safety Authority and laying down procedures in matters of food safety. OJ L 31, 1.2.2002, p. 1-24.

⁵ Regulation (EC) No 851/2004 of the European Parliament and of the Council of 21 April 2004 establishing a European centre for disease prevention and control. OJ L 142, 30.4.2004, p. 1-11.

⁶ Directive 2003/99/EC of the European Parliament and of the Council of 17 November 2003 on the monitoring of zoonoses and zoonotic agents, amending Council Decision 90/424/EEC and repealing Council Directive 92/117/EEC. OJ L 325, 12.12.2003, p. 31-40.



Summary

Zoonoses are infections and diseases that are transmissible between animals and humans. Infection can be acquired directly from animals, or through the ingestion of contaminated foodstuffs. The severity of these diseases in humans can vary from mild symptoms to life-threatening conditions. The zoonotic bacteria that are resistant to antimicrobials are of special concern, since they might compromise the effective treatment of infections in humans. In order to follow the occurrence of antimicrobial resistance in zoonotic bacteria isolated from humans, animals and food in the European Union, information is collected and analysed from the European Union Member States.

In 2012, 26 Member States reported data on antimicrobial resistance in zoonotic bacteria to the European Commission and the European Food Safety Authority, and 19 Member States submitted data to the European Centre for Disease Prevention and Control. In addition, three other European countries provided information. Assisted by its contractors, the Animal Health and Veterinary Laboratories Agency in the United Kingdom and the University of Hasselt in Belgium, the European Food Safety Authority and the European Centre for Disease Prevention and Control analysed the data, the results of which are published in this European Union Summary Report on antimicrobial resistance. Information on resistance was reported regarding *Salmonella* and *Campylobacter* isolates from human cases, food and animals, whereas data on indicator *Escherichia coli* and indicator enterococci isolates related only to animals and food. Information was reported by some Member States on the occurrence of methicillin-resistant *Staphylococcus aureus* isolates was additionally reported by two countries.

Data on antimicrobial resistance in isolates from human cases were mainly interpreted by using clinical breakpoints, while the quantitative data on antimicrobial resistance in isolates from food and animals were assessed using harmonised epidemiological cut-off values that detect microbiological resistance, i.e. reduced susceptibility to the antimicrobials tested, as well as using clinical breakpoints where considered appropriate. Direct comparisons should only be made between isolates from different sources using the same measure of determining resistance (i.e. by applying the same breakpoint).

The reporting of antimicrobial resistance data at isolate-based level by a significant number of Member States allowed the second analysis at the European Union level of multi-resistance and co-resistance patterns to critically important antimicrobials in both human and animal isolates. Detailed analyses of multi-drug resistance in certain *Salmonella* serovars, including analysis of high-level resistance to ciprofloxacin and pentavalent resistance, were possible for Member States reporting isolate-based data and included for the first time in the report. In addition, for certain bacterial species, antimicrobial resistance data could be analysed at the production-type level, such as broilers, laying hens and breeders of *Gallus gallus*, which allows the analysis of the data to be fine-tuned.

Antimicrobial resistance was commonly detected in isolates of *Salmonella* and *Campylobacter* from human cases as well as from food-producing animals and food in the European Union. This was also the case for indicator (commensal) *Escherichia coli* isolated from animals and food. For many of the antimicrobials, the levels of resistance varied greatly between different Member States.

In the European Union, the occurrence of resistance in *Salmonella* isolates from cases of salmonellosis in humans was high for ampicillin, streptomycin, sulfonamides and tetracyclines and moderate for nalidixic acid, with high levels of multi-drug resistance observed in some countries. Resistance to the critically important antimicrobials for human medicine, cefotaxime (a third-generation cephalosporin) and ciprofloxacin (a fluoroquinolone), was relatively low, although the resistance levels for ciprofloxacin were generally higher in countries using more sensitive interpretive criteria, such as epidemiological cut-off values. Co-resistance to ciprofloxacin and cefotaxime among *Salmonella* isolates was very low. The resistance levels also differed substantially between the three most commonly reported serovars, with higher resistance to ciprofloxacin, gentamicin and nalidixic acid observed in *Salmonella* Enteritidis than in *Salmonella* Typhimurium and the opposite for the other antimicrobials.

There was a high level of resistance to ampicillin, ciprofloxacin, nalidixic acid and tetracyclines among *Campylobacter* isolates from human cases, with high and very high levels of multi-drug resistance observed in some countries. The levels of resistance to erythromycin in human *Campylobacter jejuni* isolates was overall low, but moderately high in *Campylobacter coli*. Very high resistance levels to ciprofloxacin were reported in human *Campylobacter* isolates, with increasing trends observed in several Member States.



Almost one in six human *Campylobacter coli* isolates were also resistant to both erythromycin and ciprofloxacin, which is worrying as these two antimicrobials are the clinically most important for treatment of campylobacteriosis in humans.

The high proportions of Salmonella, Campylobacter and indicator Escherichia coli isolates exhibiting 'microbiological resistance' or reduced susceptibility to fluoroquinolones (ciprofloxacin) remain of concern. In Salmonella spp. isolates of food and animal origin, the highest occurrence of resistance to ciprofloxacin was noted in fattening turkeys, broiler meat, turkeys and broilers of Gallus gallus, where the proportion of such isolates varied between 46.0 % and 86.2 % in the reporting Member State group. Ciprofloxacin resistance was recorded more often in broilers than in breeders and laying hens. Two Member States demonstrated a significant increasing trend for ciprofloxacin and nalidixic acid resistance observed in 2006 to 2012. Considering the indicator Escherichia coli isolates, the levels of ciprofloxacin resistance observed in isolates from broilers and pigs were 52.7 % and 7.5 %, respectively. Furthermore, high to extremely high resistance to fluoroquinolones was commonly observed in *Campylobacter* isolates from *Gallus gallus* and broiler meat, as well as from pigs and cattle, at levels ranging from 32.0 % (*Campylobacter coli* from pigs) to 82.7 % (*Campylobacter coli* from meat from broilers).

Resistance to the third-generation cephalosporin cefotaxime was observed in *Salmonella* spp. isolates from *Gallus gallus*, turkeys, pigs, cattle and meat derived from broilers, pigs and bovine at very low or low levels varying between 0.4 % and 4.5 %, as well as in indicator *Escherichia coli* isolates from *Gallus gallus*, pigs and cattle at low or moderate levels ranging from 1.4 % to 10.2 %. Resistance to erythromycin was detected in *Campylobacter* isolates from *Gallus gallus*, pigs, cattle and broiler meat at levels of 0.4 % (*Campylobacter jejuni* from *Gallus gallus*) to 23.9 % (*Campylobacter coli* from pigs).

Among *Salmonella* isolates from meat and animals, microbiological resistance to tetracyclines, ampicillin and sulfonamides was reported at levels of 9.5 % to 66.7 % and it was higher in isolates from pigs and turkeys than in those from broilers, laying hens, breeding hens and cattle. Resistance to ciprofloxacin and nalidixic acid was higher in *Salmonella* isolates from fattening turkeys and broilers (41.5-86.2 %) than it was in isolates from breeding hens, pigs or cattle (5.8-25.5 %). In isolates of *Campylobacter* from meat and animals, resistance was commonly detected to tetracyclines at levels up to 76.8 %, whereas much lower resistance was reported to gentamicin (levels lower than 4.1 %).

Among indicator *Escherichia coli* from broilers and pigs, microbiological resistance to tetracyclines, ampicillin and sulfonamides was commonly reported at levels of 29.5 % to 54.7 %, resistance levels being lower in laying hens (18.3 % to 25.2 %). In the case of cattle, levels of resistance to these antimicrobials fell within the range 34.7 % to 46.7 % in younger age groups, mainly fattening veal calves, but values were lower in older cattle, mainly adult cows. In general, resistance levels were lower among isolates from cattle and layers than in isolates from broilers and pigs.

Multi-resistance (reduced susceptibility to at least three antimicrobial classes according to epidemiological cut-off values) was generally high in *Salmonella* isolates from broilers, pigs and cattle in those countries reporting isolate-based data. However, co-resistance/reduced susceptibility to the clinically important antimicrobials ciprofloxacin and cefotaxime in the same isolate was detected in very few isolates of *Salmonella* species. Multi-resistance was either not detected or reported at very low or low levels in *Campylobacter jejuni* isolates from broilers, and co-resistance to ciprofloxacin and erythromycin at the same time was reported at low levels. High-level ciprofloxacin resistance was observed in a limited number of *Salmonella* isolates, notably belonging to the serovars Kentucky and Infantis, from broilers, laying hens and turkeys, but not in isolates from pigs or cattle, although it was detected in isolates from pig meat. A small number of serovars, including notably the serovar Infantis, displayed pentavalent resistance, which is potentially significant because certain *Salmonella* serovars which have shown epidemic spread have shown such pentavalent resistance in the past.

Several statistically significant national trends in resistance levels in isolates from animals were observed. Among *Salmonella* isolates, more decreasing than increasing trends were found, whereas in the case of *Campylobacter*, the statistically significant national trends were mostly increasing.



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1. INTRODUCTION

The antimicrobial agents used in food-producing animals in Europe are frequently the same, or belong to the same classes, as those used in human medicine. Antimicrobial resistance is the main undesirable side effect of antimicrobial use in both humans and animals and results from the continuous positive selection of resistant bacterial clones, whether these are pathogenic, commensal or even environmental bacteria. This will modify the population structure of microbial communities, leading to accelerated evolutionary trends with unpredictable consequences for human health. The use of antimicrobials can differ in humans and food-producing animals, in terms of both the methods of administration and the quantities administered; there are important variations between and within food-producing animal species, as well as between countries.

Bacterial resistance to antimicrobials occurring in food-producing animals can spread to people not only via food-borne routes, but also by routes such as water or environmental contamination as well as through direct animal contact. *Campylobacter, Salmonella* and some strains of *Escherichia coli* are examples of zoonotic bacteria which can infect people by the food-borne route. Infections with bacteria which are resistant to antimicrobials may result in treatment failures or necessitate the use of second-line antimicrobials for therapy. The commensal bacterial flora can also form a reservoir of resistance genes which may transfer between bacterial species, including transference to organisms capable of causing disease in both humans and animals (EFSA, 2008).

The monitoring of antimicrobial resistance in zoonotic and commensal bacteria in food-producing animals and food thereof is a prerequisite for understanding the development and diffusion of resistance, providing relevant risk assessment data, and evaluating targeted interventions. Resistance monitoring entails specific and continuous data collection, analysis and reporting that quantitatively follow temporal trends in the occurrence and distribution of resistance to antimicrobials, and should also allow the identification of emerging or specific patterns of resistance.

1.1. Antimicrobial resistance monitoring and reporting at the European Union level

According to Directive 2003/99/EC on the monitoring of zoonoses and zoonotic agents, Member States (MSs) are obliged to monitor and report antimicrobial resistance in *Salmonella* and *Campylobacter* isolates obtained from healthy food-producing animals and from food. In addition, Commission Decision 2007/407/EC⁷ lays down detailed requirements on the harmonised monitoring and reporting of antimicrobial resistance of *Salmonella* isolates from various poultry populations and pigs, sampled under the corresponding national control and monitoring programmes of *Salmonella*. The monitoring and reporting of antimicrobial resistance data from the indicator organisms *Escherichia coli* and enterococci is voluntary.

Decision 1082/2013⁸ on serious cross-border threats to health and repealing Decision No 2119/98/EC, as complemented by Decision 2000/96/EC⁹ with amendment Decision 2003/542/EC¹⁰ on the diseases to be progressively covered by the network, provides the basis for data collection on human diseases in MSs and reporting to the European Centre for Disease Prevention and Control (ECDC). ECDC has provided data on zoonotic infections in humans, as well as their analyses, for the Community Summary Reports since 2005. Starting in 2007, data on human cases have been reported from The European Surveillance System (TESSy), maintained by ECDC.

This EU Summary Report 2012 includes data related to the occurrence of antimicrobial resistance both in isolates from animals and foodstuffs, collected in the framework of Directive 2003/99/EC, and in isolates from human cases, derived from the networks under Decision 2119/98/EC. This report is a joint collaboration between the European Food Safety Authority (EFSA) and ECDC with the assistance of EFSA's contractors, the Animal Health and Veterinary Laboratories Agency (AHVLA) in the United Kingdom and the University of

⁷ Commission Decision 2007/407/EC of 12 June 2007 on a harmonised monitoring of antimicrobial resistance in *Salmonella* in poultry and pigs. OJ L 153, 14.6.2007, p. 26–29.

⁸ Decision No 1082/2013/EU of the European Parliament and of the Council of 22 October 2013 on serious cross-border threats to health and repealing Decision No 2119/98/EC. OJ L 293, 5.11.2013, p. 1–15.

⁹ Commission Decision 2009/539/EC of 10 July 2009 amending Decision 2000/96/EC on communicable diseases to be progressively covered by the Community network under Decision No 2119/98/EC of the European Parliament and of the Council. OJ L 180, 11.7.2009, p. 22–23.

¹⁰ Commission Decision 2003/542/EC of 17 July 2003 amending Decision 2000/96/EC as regards the operation of dedicated surveillance networks. OJ L 185, 24.7.2003, p. 55–58.



Hasselt in Belgium. MSs, other reporting countries, the European Commission (EC) and the relevant European Union Reference Laboratories (EU-RL) were consulted while preparing the report. The efforts made by MSs, the reporting non-MSs and the EC in the reporting of zoonoses data and in the preparation of this report are gratefully acknowledged.

The main issues when comparing antimicrobial resistance data originating from different countries are the use of different laboratory methods and different interpretive criteria of resistance. These issues have been addressed by the development of EFSA's guidelines for harmonised monitoring and reporting of resistance in food-producing animals and food thereof. The resistance monitoring performed under these guidelines utilises epidemiological cut-off values (ECOFFs) which separate the naïve, susceptible wild-type bacterial populations from isolates that have developed reduced susceptibility to a given antimicrobial agent (Kahlmeter et al., 2003). The ECOFFs may differ from breakpoints used for clinical purposes, which are defined against a background of clinically relevant data, including therapeutic indication, clinical response data, dosing schedules, pharmacokinetics and pharmacodynamics. In the EU Summary Reports on antimicrobial resistance from 2004 to 2011, ECOFFs were applied to minimum inhibitory concentration (MIC) data to define resistant *Salmonella, Campylobacter*, indicator *E. coli* and indicator enterococci isolates from animals and food. The use of harmonised methods and ECOFFs ensured the comparability of data over time at the country level and also facilitated the comparison of the occurrence of resistance between MSs. The same methods and principles have been applied in this 2012 Summary Report on antimicrobial resistance.

The antimicrobial susceptibility data reported to EFSA for the year 2012 for *Campylobacter, Salmonella* and indicator *E. coli* isolates from animals and food were analysed and all quantitative data were interpreted using ECOFFs. This report also includes results of phenotypic monitoring of resistance caused by extended-spectrum beta-lactamases (ESBLs) in *Salmonella* and indicator *E. coli*, conferring resistance to third-generation cephalosporins, as well as the second investigation at the EU level of the occurrence of complete susceptibility and multi-resistance in data reported at the isolate level. A list of the antimicrobials included in this evaluation of multi-resistance can be found in Chapter 8 'Materials and methods'. The majority of antimicrobial resistance data reported to EFSA by MSs comprised data collected in accordance with EFSA's monitoring guidelines; quantitative disc diffusion data constituted only a small percentage of the total data and were analysed in the report as qualitative data only. This has circumvented the problem that ECOFFs are not available for the different disc diffusion methods used by MSs.

The report also encompasses resistance in Salmonella and Campylobacter isolates from human cases of salmonellosis and campylobacteriosis, respectively. These data were reported as qualitative data (with one exception), mostly interpreted using clinical breakpoints (CBPs), by MSs to TESSy. An important general feature of this report is that human data are largely based on susceptibility testing of clinical isolates, whereas animal data are based mainly on the testing of isolates from healthy animals, where testing has been performed in accordance with EFSA's recommendations. The data on zoonotic bacteria from humans have largely been collated and collected using CBPs. Such data are therefore not always directly comparable with data from food-producing animals and food, which have been analysed using ECOFFs. Indeed, the use of ECOFFs in animal and food isolates generally conveys the picture of 'microbiological resistance' levels in these isolates higher than 'clinical resistance' levels recorded in human isolates, where CBPs have been used. These issues are discussed further in the chapters on Campylobacter and Salmonella. Universal adoption and understanding of the distinction between CBPs and ECOFFs would enable clinicians to choose the appropriate treatment based on information relevant to the individual patient, yet would recognise that epidemiologists need to be aware of small changes in bacterial susceptibility, which may indicate emerging resistance and allow for appropriate control measures to be considered. ECOFFs, CBPs and related concepts regarding antimicrobial resistance/susceptibility are presented in detail hereafter.



1.2. Epidemiological cut-off values and clinical breakpoints

The European Committee on Antimicrobial Susceptibility Testing (EUCAST) has defined clinical breakpoints (CBPs) and epidemiological cut-off values (ECOFFs). A microorganism is defined as clinically resistant when the degree of resistance shown is associated with a high likelihood of therapeutic failure. The microorganism is categorised as resistant by applying the appropriate CBP in a defined phenotypic test system, and this breakpoint may alter with legitimate changes in circumstances (for example alterations in dosing regime, drug formulation, patient factors).

A microorganism is defined as wild-type for a bacterial species when no acquired or mutational resistance mechanisms are present to the antimicrobial in question. A microorganism is categorised as wild-type for a given bacterial species presenting a lower minimum inhibitory concentration (MIC) to the antimicrobial in question than the appropriate ECOFF in a defined phenotypic test system. This cut-off value will not be altered by changing circumstances (such as alterations in frequency of antimicrobial administration). Wild-type microorganisms may or may not respond clinically to antimicrobial treatment. A microorganism is defined as non-wild-type for a given bacterial species by the presence of an acquired or mutational resistance mechanism to the antimicrobial in question. A microorganism is categorised as non-wild-type for a given bacterial species by the presence of an acquired or mutational resistance mechanism to the antimicrobial in question. A microorganism is categorised as non-wild-type for a given bacterial species by the presence of an acquired or mutational resistance mechanism to the antimicrobial in question. A microorganism is categorised as non-wild-type for a given bacterial species by the presence of an acquired or mutational resistance mechanism to the antimicrobial in question. A microorganism is categorised as non-wild-type for a given bacterial species by applying the appropriate ECOFF value in a defined phenotypic test system; non-wild-type organisms are considered to show 'microbiological resistance' (as opposed to 'clinical resistance'). CBPs and ECOFFs may be the same, although it is often the case that the ECOFF is lower than the CBP.

Comparative advantages and disadvantages of the use of CBPs versus ECOFFs (see box hereafter) have been taken into account in the detailed specifications for harmonised monitoring schemes on antimicrobial resistance in animals and food devised by EFSA. These guidelines have been published (EFSA, 2007, 2008) and the terminology used is that devised by EUCAST (Kahlmeter et al., 2003). As far as possible, ECOFFs have been used in this report, as recommended in the guidelines, to determine non-wild-type organisms also termed 'microbiologically resistant' organisms, and to ensure that results from different MSs are comparable. Hereafter in this report, 'microbiologically antimicrobial-resistant' organisms are referred to as 'resistant' for brevity.

CLINICAL BREAKPOINTS (CLINICAL RESISTANCE)

The clinician, or veterinarian, choosing an antimicrobial agent to treat humans or animals with a bacterial infection requires information that the antimicrobial selected is effective against the bacterial pathogen. Such information will be used, together with clinical details such as the site of infection, ability of the antimicrobial to reach the site of infection, formulations available and dosage regimes, when determining an appropriate therapeutic course of action. The in vitro susceptibility of the bacterial pathogen can be determined and clinical breakpoints used to ascertain whether the organism is likely to respond to treatment. Clinical breakpoints (CBPs) will take into account the clinical behaviour of the drug following administration and assume that a clinical response will be obtained if the drug is given as recommended and there are no other adverse factors which affect the outcome. Conversely, if the CBP indicates resistance, then it is likely that treatment will be unsuccessful. Frequency of dosing is one factor that can affect the antimicrobial concentration achieved at the site of infection. Therefore, different dosing regimes can lead to the development of different CBPs, as occurs in some countries for certain antimicrobials where different therapeutic regimes are in place. Although the rationale for the selection of different CBPs may be clear, their use makes the interpretation of results from different countries in reports of this type problematic, as the results are not directly comparable between those different countries.



EPIDEMIOLOGICAL CUT-OFF VALUES (MICROBIOLOGICAL RESISTANCE)

For a given bacterial species, the pattern of the minimum inhibitory concentration (MIC) distribution or the inhibition zone diameter distribution (i.e. the frequency of occurrence of each given MIC or zone diameter plotted against the MIC value or zone diameter obtained) can enable the separation of the wild-type population of microorganisms from those populations which show a degree of resistance. The wild-type susceptible population is assumed to have no acquired or mutational resistance and commonly shows a normal distribution.

When bacteria acquire resistance by a clearly defined and efficacious mechanism, such as the acquisition of a plasmid bearing a gene which produces an enzyme capable of destroying the antimicrobial, then the MIC or zone diameter distribution commonly shows two major sub-populations, one a fully susceptible normal distribution of isolates and the other a fully resistant population which has acquired the resistance mechanism. Resistance may be achieved by a series of small steps, such as changes in the permeability of the bacterial cell wall to the antimicrobial or other mechanisms which occur lying between the fully susceptible population and more resistant populations. The epidemiological cut-off value (ECOFF) indicates the MIC or zone diameter above which the pathogen has some detectable reduction in susceptibility. ECOFFs are derived by testing an adequate number of isolates to ensure that the wild-type population can be confidently identified for a given antimicrobial. The clinical breakpoint, which is set to determine the therapeutic effectiveness of the antimicrobial, may fail to detect emergent resistance. Conversely, the ECOFF detects any deviation in susceptibility from the wild-type population, although it may not be appropriate for determining the likelihood of success or failure for clinical treatment.

The EUCAST ECOFFs which should be applied to interpret the results obtained by MSs are quoted in Commission Decision 2007/407/EC. However, since this Decision was adopted, there have been some minor changes to a few of the ECOFFs for some antimicrobials. This occurs because, as more data are collected relating to more bacterial isolates, the normal distribution of the wild-type population can in some cases be better defined. This 2012 EU Summary Report interprets the antimicrobial resistance data in accordance with the current Decision. The Decision is currently undergoing review by the EC, notably on the basis of the technical specifications proposed for harmonised monitoring of antimicrobial resistance in animals and food recently issued by EFSA (EFSA, 2012a, b, c), and the expected revision in the future will update a number of the ECOFFs to be used.

REVISION OF EPIDEMIOLOGICAL CUT-OFF VALUES

The epidemiological cut-off value (ECOFF) for E. coli versus ciprofloxacin has been recently revised by the European Committee on Antimicrobial Susceptibility Testing (EUCAST). Wild-type isolates are now considered to have a ciprofloxacin minimum inhibitory concentration lower than or equal to 0.06 mg/L (which is a change from the original tentative ECOFF of 0.03 mg/L and which now corresponds to the ECOFF for Salmonella spp.). The proportion of isolates showing microbiological resistance according to this breakpoint will alter when the new breakpoint is adopted and in fact will be reduced. For reasons of continuity and to comply with the current legislation where applicable, the ECOFFs used in this report have been those adopted in EFSA's recommendations (EFSA, 2007, 2008) and quoted in Commission Decision 2007/407/EC. For these reasons, the most recent revisions by EUCAST have not been included in this report. The report for 2013 will incorporate all of these changes in a comprehensive revision, which will also re-evaluate the historical data using the revised ECOFFs.



1.3. Developments in the harmonised monitoring of antimicrobial resistance

The ECDC has, during 2012 and 2013, arranged several expert workshops together with its Food- and Waterborne Diseases and Zoonoses (FWD) network in order to develop an EU protocol for harmonised monitoring of antimicrobial resistance in human *Salmonella* and *Campylobacter* isolates (ECDC, 2014). Consultation was also sought from EFSA, EUCAST and the EU Reference Laboratory for Antimicrobial Resistance to facilitate comparison of data between countries and with results from the antimicrobial resistance monitoring performed in isolates from animals and food products. The protocol is effective from 2014 and supports the implementation of the Commission Action Plan on antimicrobial resistance.

In 2012, EFSA, at the request of the EC, reviewed and revised the detailed specifications for the harmonised monitoring of antimicrobial resistance in food-producing animals (EFSA, 2007, 2008). Three reports have been produced describing proposals to improve (1) the harmonisation, analysis and reporting of data on antimicrobial resistance in animals and food collected from the MSs (EFSA, 2012a), (2) the harmonised monitoring and reporting of antimicrobial resistance in *Salmonella*, *Campylobacter* and indicator *E. coli* and *Enterococcus* spp. bacteria transmitted through food (EFSA, 2012b) and (3) the harmonised monitoring and reporting of antimicrobial resistance in methicillin-resistant *Staphylococcus aureus* (EFSA, 2012c).

A NEW LEGISLATION ON HARMONISED MONITORING OF ANTIMICROBIAL RESISTANCE IN ANIMALS AND FOOD

In 2013, based on the proposals issued by EFSA, the European Commission put forward and discussed with the MSs a new legislation on the harmonised monitoring of antimicrobial resistance in Salmonella, Campylobacter and indicator bacteria in food-producing animals and food. The Commission Decision 2013/652/EU¹¹ of 12 November 2013 establishes a list of combinations of bacterial species, food-producing animal populations and food products and sets up priorities for the monitoring of antimicrobial resistance from a public health perspective.

Monitoring of antimicrobial resistance in E. coli become mandatory, as it is for Salmonella and C. jejuni in the major food producing animal population and their derived meat. Sampling should be performed at the level of domestically produced animal populations, corresponding to different production types, and not at the animal species level, with the aim of collecting data that, in the future, could be combined with those on exposure to antimicrobials. The concept of a threshold is introduced for some animal populations and their derived meat to determine whether monitoring of antimicrobial resistance should be mandatory. Provisions have been taken where possible to exploit samples that would be collected under other existing control programmes.

Microdilution methods for testing are confirmed and this should be accompanied by the application of European Committee on Antimicrobial Susceptibility Testing epidemiological cut-off values (ECOFFs) for the interpretation of microbiological resistance. The harmonised panel of antimicrobials used for Salmonella, Campylobacter, E. coli and Enterococcus spp. is broadened with the inclusion of substances that either are important for human health or can provide clearer insight into the resistance mechanisms involved. The concentration ranges to be used ensure that both the ECOFF and the clinical breakpoint are included so that comparability of results with human data is made possible.

The specific monitoring of extended-spectrum beta-lactamase-, AmpC- and carbapenemase-producing Salmonella and indicator commensal E. coli is also foreseen. The collection and reporting of data is to be performed at the isolate level, in order to enable more in-depth analyses to be conducted, in particular on the occurrence of multi-resistance. The Commission Implementing Decision 2013/652/EU will enter into force in 2014, as well as the Commission Implementing Decision 2013/653/EU¹² of 12 November 2013 as regards a Union financial aid towards a coordinated control plan for antimicrobial resistance monitoring in zoonotic agents in 2014.

¹¹ Commission Implementing Decision 2013/652/EU of 12 November 2013 on the monitoring and reporting of antimicrobial resistance in zoonotic and commensal bacteria. OJ L 303, 14.11.2013, p. 26–39.

¹² Commission Implementing Decision 2013/653/EU of 12 November 2013 as regards a Union financial aid towards a coordinated control plan for antimicrobial resistance monitoring in zoonotic agents in 2014. OJ L 303, 14.11.2013, p. 40–47.



2. MAIN FINDINGS

2.1. Main findings of the European Union Summary Report on antimicrobial resistance 2012

CLINICAL RESISTANCE AND MICROBIOLOGICAL RESISTANCE: USE OF CLINICAL BREAKPOINTS AND EPIDEMIOLOGICAL CUT-OFF VALUES

Development of resistance in bacteria is a major threat to public health. It is therefore important to detect any occurrence of resistance and increases in resistance levels as early as possible. In this report, acquired resistance in bacteria is denoted 'microbiological resistance' and harmonised epidemiological cut-off values (ECOFFs) are used to interpret the results of susceptibility testing in isolates from animals and food. In contrast, results of susceptibility testing of clinical isolates from humans are interpreted using clinical breakpoints (CBPs) to guide medical treatment of the patient. The CBP is in many cases less sensitive than the ECOFF for a specific bacteria–drug combination, resulting in that isolates interpreted with ECOFFs more often will be classified as (microbiologically) resistant than isolates interpreted with CBPs. Direct comparisons between isolates from different sources should therefore only be made when the criteria for interpretation is at the same level (see breakpoint figures SA1 and CA1). The harmonisation efforts on antimicrobial resistance monitoring in humans led by ECDC include introduction of quantitative reporting. This will enable use of ECOFFs for interpretation of susceptibility testing results from human isolates.

- In 2012, MSs reported qualitative data (and one non-MS reported quantitative data) on antimicrobial
 resistance in *Salmonella* and *Campylobacter* isolates from human cases mostly interpreted by using
 clinical breakpoints (CBPs) to define the resistant isolates. In contrast, quantitative data (minimum
 inhibitory concentrations (MICs) and/or inhibition zone diameter (IZD) results) on antimicrobial
 resistance, reported for isolates from food and animals, were interpreted by using epidemiological cutoff values (ECOFFs). ECOFFs are often lower than CBPs, and this can result in more isolates being
 classified as resistant, although that is dependent on the distribution of MICs obtained.
- Antimicrobial resistance was regularly observed in isolates of *Salmonella* and *Campylobacter* from human cases as well as from food-producing animals and food in the EU. For many of the antimicrobials, the levels of resistance varied greatly between different MSs and animal production types.
- Fluoroquinolones, such as ciprofloxacin, and third-generation cephalosporins, such as cefotaxime, are considered critically important antimicrobials in the treatment of severe or invasive salmonellosis in humans. Likewise, fluoroquinolones and macrolides, such as erythromycin, are considered critically important for treating severe *Campylobacter* infections. Therefore, special attention was paid to resistance against these substances in the analyses of the data.
- 'Clinical resistance' at the EU level in *Salmonella* spp. isolates from human cases was high (between 23.6 % and 30.0 %) to ampicillin, streptomycin, sulfonamides and tetracyclines. In contrast, resistance to ciprofloxacin and cefotaxime was relatively low (on average <6 % and <2 %, respectively); however, even low levels of resistance to these critically important antimicrobials are important. The determined resistance levels to ciprofloxacin were influenced by the interpretive criteria used in each country, resulting in higher resistance levels in countries using more sensitive criteria. The introduction of harmonised EUCAST methods and interpretive criteria in more and more MSs is therefore much welcomed and supported by ECDC.
- Multi-drug resistance (MDR, defined as reduced susceptibility to at least three antimicrobial classes) was high to very high in human *Salmonella* isolates in eight out of 12 reporting countries; however, there were very low levels of co-resistance to ciprofloxacin and cefotaxime. Furthermore, more than half of all *Salmonella* isolates were susceptible to the complete range of antimicrobials tested.
- In food and animal isolates, the highest occurrence of 'microbiological resistance' to ciprofloxacin was noted in *Salmonella* spp. isolates from fattening turkeys, broiler meat and fowl (*Gallus gallus*) (from 37.3 % to 86.2 % at the MS group level). The further sub-division of the *Gallus gallus* species into production types revealed higher overall resistance to ciprofloxacin in *Salmonella* spp. isolates from



broilers (46.0 %) than in those from breeding hens (25.5 %) or laying hens (19.4 %). In cattle, pigs and pig meat, low resistance levels were observed (7.6 %–9.1 %).

- 'Microbiological resistance' to cefotaxime (a third-generation cephalosporin) was observed in Salmonella spp. isolates from Gallus gallus, turkeys, pigs and cattle and in meat derived from broilers, pigs and cattle, but at low or very low levels (0.4 %–4.5 %), when all reporting MSs were considered. However, even low levels of resistance to this critically important antimicrobial are important. Resistance to cefotaxime was not detected in Salmonella strains isolated from cattle in reporting countries in 2011, but was detected in a single MS in 2012.
- 'Microbiological resistance' to tetracyclines, ampicillin and sulfonamides was frequently reported among Salmonella spp. isolates from meat and animals (9.5 %–66.7 % at MS group level). Resistance to these antimicrobials was higher in isolates from pigs, turkeys and cattle (34.5 %–66.7 %) than in isolates from Gallus gallus (21.2 %–28.3 %).
- Multi-drug resistance (MDR, reduced susceptibility to at least three antimicrobial classes according to ECOFFs) was high in *Salmonella* spp. isolates from animals in some countries reporting isolate-based data; however, co-resistance/reduced susceptibility to the clinically important antimicrobials, ciprofloxacin and cefotaxime, was at very low to low levels.
- Detailed analyses of MDR in certain Salmonella serovars, including analysis of high-level resistance to ciprofloxacin and pentavalent resistance, was possible for MSs reporting isolate-based data from animals and food. High-level ciprofloxacin resistance was observed in a limited number of Salmonella isolates, notably belonging to the serovars Kentucky and Infantis, from broilers, laying hens and turkeys, but not in isolates from pigs or cattle, although it was detected in isolates from pig meat. A small number of serovars, in particular S. Infantis, displayed pentavalent resistance, which is potentially significant because certain Salmonella serovars which have shown epidemic spread have shown such pentavalent resistance in the past.
- MDR data on *Salmonella* isolates from animals and meat also provided evidence relating to (1) the nondetection of a serovar (*S.* Enteritidis) with a particular resistance pattern which has shown increased virulence for humans and (2) the spread of particular strains of *Salmonella* which have a typical antimicrobial resistance pattern (well-illustrated by monophasic *S.* Typhimurium isolates, where examination of the antimicrobial resistance patterns shown allows presumptive identification of the main clones which have been described).
- The 'clinical resistance' among *Campylobacter* spp. isolates from human cases was high (between 32.4 % and 48.8 %) for ampicillin, ciprofloxacin, nalidixic acid and tetracyclines. Low resistance levels (average 3.1 %) were observed to the clinically important antimicrobial erythromycin. Multi-resistance in human *Campylobacter* isolates was moderate or high in some countries. Levels of co-resistance to the clinically important antimicrobials, ciprofloxacin and erythromycin, were on average low among *Campylobacter jejuni* (*C. jejuni*) isolates and moderate among *Campylobacter coli* (*C. coli*) isolates. However, even low levels of co-resistance to both of these critically important antimicrobials are of significance.
- Extremely high 'microbiological resistance' levels to ciprofloxacin (a fluoroquinolone) was commonly observed in *C. coli* isolates from broiler meat and broilers (*Gallus gallus*) (82.7 % and 78.4 %, respectively), with somewhat lower levels in *C. jejuni* (59.5 % and 44.1 %, respectively). High levels were also reported for isolates from pigs and cattle (32.0 % to 32.9 %). Important differences were observed between animal species and MSs.
- 'Microbiological resistance' to erythromycin was detected at very low to moderate levels in *Campylobacter* isolates from broilers (*Gallus gallus*) and broiler meat (0.4 %–16.5 %). The highest level of resistance to erythromycin at the reporting MS group level was observed in *C. coli* isolates from pigs (23.9 %), while the level of erythromycin resistance in isolates of *C. jejuni* from cattle across reporting MSs was very low (0.6 %).
- Considering all reporting MSs, 'microbiological resistance' to nalidixic acid and tetracyclines was common among *Campylobacter* isolates from meat and animals (31.6 %–81.0 %), whereas resistance



to gentamicin was low (0.2 %–4.1 %). As for *Salmonella*, levels of resistance to nalidixic acid followed closely those observed for ciprofloxacin.

- Multi-resistance (reduced susceptibility to at least three antimicrobial classes according to ECOFFs) was generally low in *C. jejuni* isolates from broilers and co-resistance to the clinically important antimicrobials, ciprofloxacin and erythromycin, was, in the same isolates, either not detected or recorded at low levels. Multi-resistance and co-resistance levels were generally higher in *C. coli* isolates from broilers and fattening pigs.
- Analysis of MDR data for Campylobacter can also provide indications of certain mechanisms of
 resistance. Resistance to ciprofloxacin and erythromycin in Campylobacter is usually the result of
 mutation with or without the additional action of efflux pumps. Additionally, the efflux pump CmeABC
 has been shown to confer a degree of resistance to erythromycin, ciprofloxacin and tetracyclines.
 Isolates of both C. coli and C. jejuni, from animals and humans were detected which showed resistance
 to erythromycin, ciprofloxacin and tetracyclines, raising the possibility that CmeABC may have been
 responsible for, or contributed to, the observed pattern of 'microbiological resistance'.
- Several statistically significant national trends in resistance levels in isolates from animals were observed. Among *Salmonella* spp. isolates, more decreasing than increasing trends were found, whereas in *Campylobacter* isolates the statistically significant national trends were mostly increasing. Increasing (decreasing) trends in resistance to (fluoro)quinolones in *C. jejuni*, *C. coli* and *E. coli* in broilers and pigs were observed in a number of countries having consistently reported AMR data through active monitoring programmes over the last years.
- Among indicator (commensal) *E. coli* isolates from animals, 'microbiological resistance' to ampicillin, streptomycin sulfonamides and tetracyclines, was commonly reported in *Gallus gallus* and pigs (29.5 %–54.7 %), lower levels being reported in cattle (24.5 %–30.6 %). Resistance to ciprofloxacin and nalidixic acid was highest among *E. coli* isolates from *Gallus gallus* (57.6 % and 47.4 %, respectively), while levels were lower in pigs and cattle (4.9 %–12.2 %). Cefotaxime resistance was low in pigs and cattle (1.4 % and 2.4 %, respectively), and moderate in isolates from *Gallus gallus* (10.2 %), considering all reporting MSs. At the MS level, resistance to cefotaxime in indicator *E. coli* showed wider variation in some species or production types, for example ranging between 0 % and 13.5 % in broilers.
- Indicator *E. coli* from meat from *Gallus gallus*, pigs and cattle showed moderate or high levels of resistance to ampicillin, sulfonamides and tetracyclines, considering all reporting MSs. Resistance to cefotaxime was 3 % or less in *E. coli* from meat from all three animal species and for all reporting MSs.
- In general, resistance to third-generation cephalosporins in *E. coli* was higher than that observed in *Salmonella* spp. (in which resistance was sometimes not detected) for the same species of animals, which is consistent with the hypothesis that *E. coli* may provide a reservoir of cephalosporin resistance genes for organisms such as *Salmonella*.
- Multi-resistance was high in indicator (commensal) *E. coli* isolates from animals in some countries reporting isolate-based data; however, co-resistance to the clinically important antimicrobials, ciprofloxacin and cefotaxime, was generally reported at very low to low levels. Considering the *E. coli* for which isolate-based data was available, ciprofloxacin 'microbiological resistance' was a frequent component of MDR patterns in *E. coli* from broilers occurring in 72.3 % of isolates, whereas it occurred in 16.0 % of MDR patterns in isolates from pigs.

2.2. Zoonotic and indicator agent-specific summaries

2.2.1. Salmonella

The Salmonella spp. data presented in this report comprise results for all reported Salmonella serovars which have been amalgamated to represent the overall occurrence of antimicrobial resistance in Salmonella within humans and the various animal and food categories. The differences in the distribution and prevalence of particular serovars and phage types of Salmonella in different countries and in different animal species, and their associated patterns of resistance, may explain some of the differences in the levels of



antimicrobial resistance observed as well as in those of multi-resistance. The spread of particularly resistant clones, and the occurrence of resistance genes within these clones, can be exacerbated by the use of antimicrobials in the human and animal populations and the selective pressure this exerts. Other factors, such as foreign travel by humans, international food trade, animal movements, farming systems, animal husbandry and the pyramidal structure of some types of animal primary production can also influence the spread of resistant clones.

In addition to the amalgamated data for *Salmonella* spp., resistance data for the most numerous *Salmonella* serovars in humans, *S.* Enteritidis, *S.* Typhimurium and monophasic *S.* Typhimurium, were analysed separately.

2.2.1.1. In humans

In 2012, 19 MSs and two non-MSs provided information on antimicrobial resistance in *Salmonella* isolates from cases of salmonellosis in humans.

The reported data represented 25.6 % of the confirmed salmonellosis cases reported in the EU/European Economic Area (EEA) in 2012. 'Clinical resistance' in human *Salmonella* isolates was high for ampicillin (27.6 %), streptomycin (23.6 %), sulfonamides (28.9 %) and tetracyclines (30.0 %), and moderate for nalidixic acid (14.4 %), and high levels of multi-resistance were observed in some countries (28.9 % overall). For these first four antimicrobials this was largely a result of the high to extremely high resistance levels observed among *S*. Typhimurium and monophasic *S*. Typhimurium isolates. However, more than half of all isolates tested were susceptible to the complete range of antimicrobials in the human data collection. In addition, the resistance to the clinically important antimicrobials, ciprofloxacin and cefotaxime, was on average relatively low (5.1 % and 1.1 %, respectively), although in the case of ciprofloxacin, markedly higher in countries using more sensitive interpretive criteria such as EUCAST ECOFFs. Co-resistance to ciprofloxacin and cefotaxime among isolates was very low (0.2 %). Resistance to quinolones (ciprofloxacin and nalidixic acid) was generally higher in *S*. Enteritidis isolates than in *S*. Typhimurium isolates.

When assessed by geographical region, *Salmonella* spp. isolates acquired within the EU/EEA countries exhibited greater resistance to ampicillin, sulfonamides, streptomycin and tetracyclines, while the highest levels of resistance to ciprofloxacin were observed in isolates from cases that had travelled in Asia, Africa or European countries outside of the EU/EEA.

2.2.1.2. In animals and food

In 2012, information on antimicrobial resistance in *Salmonella* isolates from animals and food was reported by 19 MSs and two non-MSs.

Among *Salmonella* spp. isolates from *Gallus gallus*, the resistance level to tetracyclines, ampicillin and sulfonamides in all reporting MSs was at a high level, 25.9 %, 21.2 % and 28.3 %, respectively. Resistance to ciprofloxacin and nalidixic acid was higher (37.3 % and 34.3 %, respectively), for all reporting MSs. In general, there were large variations in the levels of resistance to these antimicrobials between different reporting MSs. The occurrence of resistance to cefotaxime in all reporting MSs was low, at 4.5 %.

For the second year, data were presented at the production-type level, where possible, throughout the report. In 2012, 13 MSs reported quantitative data from broilers, and 12 MSs reported quantitative data from laying hens. In general, the levels of resistance in this production type were slightly higher in isolates from broilers than those reported when all *Gallus gallus* production types were considered for isolates from layers or breeding flocks. Twelve MSs reported quantitative data from laying hens in 2012, and in contrast to the data from broilers, the levels of resistance in this production type were lower than those reported when all *Gallus gallus* and in contrast to the data from broilers, the levels of resistance in this production type were lower than those reported when all *Gallus gallus* were considered. Quantitative data on isolates of *Salmonella* spp. from breeding flocks of *Gallus gallus* were reported by four MSs in 2012 and the levels of resistance among isolates from breeding flocks of *Gallus gallus* were generally higher than those observed in laying hens but lower than in isolates from broilers.

Multi-resistance levels (reduced susceptibility to at least three different antimicrobial classes using ECOFFs) were generally high in *Salmonella* spp. isolates from broilers and moderate in those from laying hens. In general, co-resistance to ciprofloxacin and cefotaxime was low, and not detected when using CBPs.

Some MSs showed statistically significant increasing trends in resistance among *Salmonella* spp. isolates from *Gallus gallus* over the years 2006 to 2012, whereas other MSs exhibited decreasing trends. Statistically significant decreasing trends were more frequently observed than significant increasing trends. Two MSs demonstrated a significant increasing trend for ciprofloxacin and nalidixic acid 'microbiological resistance' and one a decreasing trend for both antimicrobials. In particular, resistance to cefotaxime remained generally low, very low or absent in reporting MSs between 2006 and 2012.

Resistance in *S*. Enteritidis was lower than in *Salmonella* spp. isolates from *Gallus gallus*. In *S*. Enteritidis, the occurrence of resistance for all reporting MSs was 3.3 % for tetracyclines, 5.5 % for ampicillin and 4.3 % for sulfonamides, whereas the level of 'microbiological resistance' to ciprofloxacin and nalidixic acid was 23.9 % and 22.2 %, respectively.

In *Salmonella* spp. isolates from broiler meat, resistance levels for all reporting MSs for tetracyclines and sulfonamides were high and very high at 48.9 % and 53.0 %, respectively. 'Microbiological resistance' to ciprofloxacin and nalidixic acid resistance was also very high, with overall resistance levels of 63.1 % and 57.3 %, respectively. The resistance level for cefotaxime was low, at 4.3 %.

Among *Salmonella* spp. isolates from turkeys, the level of resistance to tetracyclines, ampicillin and sulfonamides in all reporting MSs was very high at 66.7 %, 56.5 % and 58.9 %, respectively. The levels of resistance to ciprofloxacin and nalidixic acid were also very high and high, at 59.9 % and 42.0 %, respectively, for all reporting MSs. Often, there were large variations in the levels of resistance to these antimicrobials among the different reporting MSs. The occurrence of resistance to cefotaxime in all reporting MSs was very low, at 0.8 %. Five MSs reported quantitative data from fattening turkeys and, in general, the levels of resistance in this production type were slightly higher than those reported when all turkey production types were considered. Multi-resistance was generally very high in *Salmonella* spp. isolates from turkeys; however, co-resistance to ciprofloxacin and cefotaxime (interpreted using CBPs) was not detected.

For *Salmonella* spp. isolates from pigs, resistance levels in the reporting group of MSs were very high: 63.3 % for tetracyclines and sulfonamides, and 60.2 % for ampicillin. Ciprofloxacin and nalidixic acid 'microbiological resistance' levels remained low, at 7.6 % and 5.8 %, respectively, and the level of resistance to cefotaxime was also low, at 2.3 % overall. Five MSs reported quantitative data from fattening pigs and the levels of resistance in this production type were lower than those reported when all pig production types were considered. Resistance to tetracyclines, ampicillin and sulfonamides was common in *Salmonella* spp. from pig meat, 49.2 %, 47.5 % and 53.5 %, respectively, considering all reporting MSs. 'Microbiological resistance' to ciprofloxacin and nalidixic acid was at a low level (7.6 % and 4.2 %, respectively) and cefotaxime resistance was very low, at 0.9 %. The trends in resistance observed in *Salmonella* spp. isolates from pigs over the years 2006 to 2012 remained stable in some countries, while fluctuation was observed in others. Among the few statistically significant national trends, slightly more decreasing trends were observed than increasing ones. However, it is noteworthy that 'microbiological resistance' to cefotaxime remained generally low, very low or absent in reporting MSs over the period 2006 to 2012. Multi-resistance was generally very high in *Salmonella* spp. isolates from fattening pigs; however, co-resistance to ciprofloxacin and cefotaxime was not detected when using CBPs.

Among Salmonella spp. isolates from cattle, the occurrence of resistance to tetracyclines, ampicillin and sulfonamides in all reporting MSs was high, at 36.0 %, 34.5 % and 42.4 %, respectively. The level of resistance to ciprofloxacin and nalidixic acid was low, 9.1 % for both, for all reporting MSs, while cefotaxime resistance was observed among the reporting MSs at a very low level (0.4 %). Although variation was observed between MSs in the level of resistance to some antimicrobials, overall trends in resistance between 2006 and 2012 were mainly decreasing ones among Salmonella spp. from cattle. Variability between MSs was observed in multi-resistance levels in Salmonella spp. isolates from cattle; however, corresistance to ciprofloxacin and cefotaxime was not detected when using CBPs. The few statistically significant trends observed in resistance levels among Salmonella isolates from cattle were all decreasing ones.

With respect to detailed analyses of MDR in *Salmonella* serovars, high-level resistance to ciprofloxacin in *S*. Kentucky was identified in isolates from broilers, laying hens and turkeys. *S*. Kentucky with high-level ciprofloxacin resistance was also detected in meat from turkeys and in meat from broilers. *S*. Kentucky isolates showing high-level resistance to ciprofloxacin are likely to belong to the clone of *S*. Kentucky sequence type 198, which recently emerged in North Africa and the Middle East exhibiting such resistance and which has subsequently been detected in poultry in some European countries.



S. Infantis showing high-level ciprofloxacin resistance, together with resistance to a core antimicrobial pairing of sulfonamides and tetracyclines (but with other resistances which were not invariably present in all high-level ciprofloxacin S. Infantis isolates) was detected in meat from broilers, pig meat and broilers. The occurrence of S. Infantis with this resistance pattern in several MSs and in different types of animal or meat probably indicates that either a clone of S. Infantis showing such resistance has spread within Europe or that several clones have gained such high-level ciprofloxacin resistance independently. In addition to showing high-level ciprofloxacin resistance, S. Infantis also featured as a serovar displaying 'pentavalent' resistance that is resistance to ampicillin, chloramphenicol, streptomycin, sulfonamides and tetracyclines. A small number of serovars displayed this resistance phenotype which is potentially significant because certain *Salmonella* serovars which have shown epidemic spread have shown such pentavalent resistance in the past. Although a relatively low number of serovars were detected showing such pentavalent resistance, these included several important serovars relevant to public health, including, for example, *S.* Saintpaul, as well as *S*. Typhimurium.

Resistance to ampicillin, chloramphenicol, streptomycin, sulfonamides, tetracyclines and trimethoprim was not detected in isolates of *S*. Enteritidis from animals or meat for which isolate-based data was available. This pattern of resistance has previously been reported in human *S*. Enteritidis isolates which possess a combined virulence-resistance plasmid; such strains cause human infections of increased severity. The absence of certain resistance phenotypes can therefore be just as significant as the detection of resistance in evaluating the current situation in Europe.

Considering monophasic *S*. Typhimurium, the MDR results assist in tracking the increasing spread of this *Salmonella* serovar. Different clones have been identified and using the MDR information it is possible to presumptively identify each clone. The so-called 'Spanish clone' is resistant to ampicillin, chloramphenicol, gentamicin, streptomycin, sulfonamides, tetracyclines and trimethoprim, whereas the 'US clone' is fully susceptible and the 'European clone' is resistant to ampicillin, streptomycin, sulfonamides. The pattern associated with the Spanish clone was only seen in 1.8 % (5 out of 279) isolates from fattening pigs, where the pattern associated with the European clone was much more frequently detected.

2.2.2. Campylobacter

2.2.2.1. In humans

Overall, 14 MSs and one non-MS provided information on antimicrobial resistance in isolates from campylobacteriosis cases in humans for the year 2012.

Data from antimicrobial susceptibility testing represented 17.9 % of the total confirmed campylobacteriosis cases reported in the EU/EEA in 2012. Fewer countries reported results for *Campylobacter* than for *Salmonella*. The variety of methods and interpretive criteria used by MSs in antimicrobial susceptibility testing for *Campylobacter* was still large, even though some harmonisation towards the use of EUCAST CBPs could be observed. The launch of CBPs for disc diffusion by EUCAST in 2012 will most likely facilitate this harmonisation further, as many countries use disc diffusion for testing of human isolates.

The 'clinical resistance' levels in human *Campylobacter* isolates were highest for nalidixic acid (48.8 %) and ciprofloxacin (47.4 %) followed by ampicillin (36.4 %) and tetracyclines (32.4 %), with high levels of multi-resistance observed in some countries. Resistance to the clinically important antimicrobial erythromycin was low overall (3.1 %), but moderately high in *C. coli* (15.1 %), although the number of isolates of this species tested was small.

Sufficient data were available for levels of resistance to be compared by geographical region for ciprofloxacin, erythromycin, nalidixic acid and tetracyclines. Isolates acquired in EU/EEA countries had the lowest frequency of resistance to all these antimicrobials, with resistance to both ciprofloxacin and erythromycin notably lower than in isolates acquired in Asia and Africa. However, the number of isolates tested that originated from infections acquired outside of the EU/EEA was very low.

2.2.2.2. In animals and food

In 2012, 15 MSs and one non-MS reported quantitative MIC data for *Campylobacter* isolates from food and animals. Five MSs additionally reported qualitative data where the method of testing was not specified; however, these data are not presented in the report. When considering all host species, the highest levels of resistance were seen for the (fluoro)quinolones (ciprofloxacin and nalidixic acid) and tetracyclines.

Resistance to erythromycin and gentamicin was comparatively low among *Campylobacter* isolates from food and animals. Resistance was generally higher in *C. coli* than in *C. jejuni* from the same host species (*Gallus gallus*).

For *C. jejuni* isolates from *Gallus gallus*, resistance was high for ciprofloxacin (44.1 %), nalidixic acid (41.4 %) and tetracyclines (34.1 %), while levels of resistance to erythromycin and gentamicin were very low, at 0.4 % and 0.7 %, respectively. A similar pattern was seen for *C. coli* isolates from *Gallus gallus*; however, levels of resistance were higher overall. Levels of resistance to ciprofloxacin, nalidixic acid and tetracyclines were extremely high at 78.4 %, 75.7 % and 73.1 %, respectively, while levels of resistance to erythromycin and gentamicin were moderate (11.2 %) and low (4.1 %), respectively.

Multi-resistance (reduced susceptibility to at least three antimicrobial classes according to ECOFFs) was very low, low or not detected in *C. jejuni* isolates from broilers, and co-resistance to the clinically important antimicrobials ciprofloxacin and erythromycin in the same isolates was either not detected or recorded at low levels in the reporting MSs. The situation was different for *C. coli* from broilers where multi-resistance as a percentage of all isolates received by the individual MSs ranged from 3.0 % to 55.6 % and co-resistance to ciprofloxacin and erythromycin ranged from 0 % to 22.2 %.

Although resistance to ciprofloxacin and nalidixic acid in *Gallus gallus* varied greatly among reporting MSs over the period 2006 to 2012, some statistically increasing trends in resistance to these antimicrobials were observed for several MSs, both for *C. jejuni* and *C. coli*.

For *C. jejuni* isolates from broiler meat, resistance, considering all reporting MSs, ranged from high to very high for ciprofloxacin (59.5 %), nalidixic acid (57.9 %) and tetracyclines (47.5 %), while levels of resistance to erythromycin and gentamicin ranged from low to very low at 1.8 % and 0.7 %, respectively. A similar pattern was seen for *C. coli* isolates from broiler meat; however, levels of resistance were higher overall. Levels of resistance to ciprofloxacin and nalidixic acid were extremely high at 82.7 % and 81.0 %, respectively, very high for tetracyclines at 57.3 %, moderate for erythromycin at 16.5 % and low for gentamicin at 1.7 %.

C. coli isolates from pigs were derived from fattening pigs (one MS did not specify the production level). Resistance to ciprofloxacin, nalidixic acid and tetracyclines ranged from high to extremely high at 32.0 %, 31.6 % and 76.8 %, respectively. Resistance was high to erythromycin (23.9 %) and low to gentamicin (2.9 %). Resistance to ciprofloxacin and/or nalidixic acid in *C. coli* from pigs showed a significantly increasing trend in three reporting MSs over the period 2006 to 2012.

Multi-resistance (reduced susceptibility to at least three antimicrobial classes according to ECOFFs) varied greatly in occurrence with *C. coli* isolates from pigs from the different MSs, ranging from 3.9 % to 97.3 %. Co-resistance to the clinically important antimicrobials, ciprofloxacin and erythromycin, was low for three reporting countries, ranging from 2.9 % to 9.4 %, but extremely high in one MS at 76.7 %.

C. jejuni isolates from cattle were also considered. Overall, resistance was high for ciprofloxacin (32.9 %), nalidixic acid (32.5 %) and tetracyclines (43.5 %), while resistance to erythromycin and gentamicin was very low at 0.6 % and 0.2 %, respectively. No statistically significant trends in ciprofloxacin and nalidixic acid resistance were observed in any of the reporting countries, but erythromycin resistance significantly decreased statistically in the Netherlands over the period 2006 to 2012.

Multi-resistance (reduced susceptibility to at least three antimicrobial classes according to ECOFFs) ranged from 0 % to 12.3 % in *C. jejuni* isolates from cattle from the reporting MSs. Co-resistance to the clinically important antimicrobials, ciprofloxacin and erythromycin, was low or not detected.

2.2.3. Indicator (commensal) Escherichia coli

Eleven MSs and two non-MSs reported quantitative data on antimicrobial resistance in indicator *E. coli* isolates from animals and food in 2012. Most of the data related to isolates from *Gallus gallus*, pigs and cattle; four MSs reported results for meat derived from each of those species.

Indicator *E. coli* from meat from *Gallus gallus*, pigs and cattle showed high or very high levels of 'microbiological resistance' to ampicillin, sulfonamides and tetracyclines, considering all reporting MSs. Resistance was less than 4 % to gentamicin, less than 3 % to cefotaxime and less than 7 % to



chloramphenicol for all reporting MSs and for meat from all three animal species. The levels of resistance were therefore broadly similar for meat from broilers, pigs and bovine animals for all reporting MSs for these antimicrobials. The situation was different for ciprofloxacin and nalidixic acid, where 'microbiological resistance' was high in meat from broilers considering all reporting MSs at 29.1 % and 24.1 %, respectively, but low in meat from pigs and cattle at less than 7 %.

Most data on *Gallus gallus* referred to broilers, although two MSs provided data on *E. coli* from laying hens. Regarding broilers, the highest overall 'microbiological resistance' levels observed at the reporting MS group level were to ciprofloxacin (52.7 %), ampicillin (50.4 %), sulfonamides (45.4 %), tetracyclines (43.3 %), nalidixic acid (43.2 %) and streptomycin (38.4 %). The isolates from laying hens also most commonly showed reduced susceptibility to these antimicrobials, but resistance levels were lower, ranging between 10.1 % and 43.3 %. Resistance to cefotaxime was low in both broilers (6.2 %) and layers (6.0 %). There was substantial variation in the level of resistance to these antimicrobials between reporting MSs. Countries mostly reported relatively stable resistance in *E. coli* isolates from *Gallus gallus* between 2006 and 2012. However, statistically significant trends in resistance to all of these antimicrobials, except tetracyclines, have been identified: these trends have more commonly been increasing ones than decreasing ones.

Concerning indicator *E. coli* isolates from pigs, the highest overall 'microbiological resistance' levels in the reporting group of MSs were observed for tetracyclines (54.7 %), streptomycin (50.3 %), sulfonamides (41.7 %) and ampicillin (29.5 %). Resistance to both ciprofloxacin and nalidixic acid was low at 7.5 % and 4.9 %, respectively. Overall, only 1.4 % of isolates were resistant to cefotaxime. There were large differences in the occurrence of resistance between MSs. There were fewer statistically significant trends than in isolates from *Gallus gallus*. No significant trends were observed for cefotaxime.

Multi-resistance levels (reduced susceptibility to at least three antimicrobial classes according to ECOFFs) were generally high in indicator *E. coli* isolates from broilers and pigs, and in a number of reporting countries. Co-resistance/reduced susceptibility to the clinically important antimicrobials, ciprofloxacin and cefotaxime, was also detected in very few isolates from these species.

In the reporting group of MSs, resistance levels in indicator *E. coli* isolates from cattle were generally lower than among isolates from *Gallus galllus* and pigs. The highest resistance levels observed were to tetracyclines (30.6 %), sulfonamides (28.3 %), streptomycin (25.1 %) and ampicillin (24.5 %). 'Microbiological resistance' to ciprofloxacin was moderate, at 12.2 %, and resistance to nalidixic acid was low, at 8.6 %. Overall, only a few isolates (2.4 %) expressed resistance to cefotaxime. The occurrence of resistance was variable between MSs for most of the antimicrobials. As for *Salmonella*, some MSs presented data at the production-type level for cattle, although only three MSs did so. One of these MSs reported much higher resistance among younger animals, mainly fattening veal calves, compared with older cattle, mainly adult cows, but this was not observed in the other countries. There have been numerous statistically significant trends in resistance since 2006, mainly of a decreasing nature.

Strains of *E. coli* are not separated on phenotypic characteristics (e.g. serotype) in the current monitoring programme and a less detailed analysis is therefore possible than for Salmonella where isolates can be subdivided by serovar. The common core patterns of 'microbiological resistance' to ampicillin, streptomycin, sulfonamides, tetracyclines and trimethoprim (and combinations thereof) frequently observed in the monitoring of E. coli isolates are probably related to the presence of class 1 or class 2 integrons, which generally carry genes conferring resistance to these antimicrobials. A common core of 'microbiological resistance' to ampicillin, sulfonamides and tetracyclines, generally with 'microbiological resistance' to ciprofloxacin and frequently with such resistance to streptomycin and trimethoprim, was discernible in broilers. However, no single pattern or patterns of 'microbiological resistance' occurred at a high frequency in broilers. In fattening pigs, two MDR patterns were predominant (streptomycin, sulfonamides, tetracyclines and streptomycin, sulfonamides, tetracyclines, ampicillin and trimethoprim) and each accounted for more than 5 % of the total number of E. coli isolates from fattening pigs for which isolate-based data were available. Ciprofloxacin resistance (microbiological) frequently occurred as a component of MDR in E. coli from broilers and was observed in 72.3 % of MDR isolates (120 out of 166), whereas 'microbiological resistance' to ciprofloxacin occurred infrequently as a component of MDR in pigs and was present in 16.0 % (32 out of 200) of porcine MDR in E. coli isolates.



2.2.4. Methicillin-resistant Staphylococcus aureus

A low number of MSs reported the results of monitoring food for methicillin-resistant *Staphylococcus aureus* (MRSA). MRSA was detected in meat from broilers, turkeys, pigs and bovine animals. Some MSs also examined products such as fish and fruit and detected MRSA in a low number of such products. The occurrence of MRSA in meat and products derived from animals may reflect colonisation of those animals with MRSA; however, the occurrence of MRSA in products such as fish and fruit may also reflect contamination from human personnel who may be colonised with MRSA involved in the preparation of such foods. Strain typing would probably assist in elucidating the likely origin of MRSA isolates in these samples. Some MSs undertook comprehensive monitoring at various stages along the food chain revealing interesting differences in the occurrence of MRSA at different stages of food production.

In relation to healthy food-producing animals, MRSA was detected in meat-producing turkey flocks, but not in breeding flocks in one MS. There was a large degree of variation between MSs in the occurrence of MRSA in pigs, with one MS not detecting MRSA in a farm-based national survey, whilst another MS reported that 99 % of animals were positive in slaughterhouse monitoring. Three MSs examined cattle for MRSA; the number of animals which were positive in sampling on farms for one MS was considerably lower than when calves were sampled at slaughter. The occurrence of MRSA in dairy cows in this MS (9.9 %) was similar to the occurrence in calves under one year old (10.2 %), both types of animals being sampled on farms. In calves under one year old monitored at slaughterhouse through sampling of nasal swabs, the occurrence of MRSA ranged from 45.0 % to 47.1 %. Sheep and goats were investigated by one MS and no MRSA was detected. Molecular typing data was reported by one MS in relation to isolates from cattle; the majority of isolates were *spa*-type t011 belonging to MRSA clonal complex (CC) 398, the common livestock-associated type of MRSA occurring in Europe.

Several MSs reported results of clinical investigations which yielded MRSA in food-producing animals and companion animals. MRSA was detected in cats, dogs and horses in one MS as well as in clinical diagnostic samples from pigs.

Temporal trends in the occurrence of MRSA in animals could be assessed for two MSs and one non-MS. Monitoring of sheep and goats on farms in 2011 and 2012 in one MS did not reveal the presence of MRSA in these animals in either year. A further MS monitored calves under one year old on the farms in 2010 and 2012 and reported similar numbers of animals positive for MRSA in each year (19.6 % and 19.2 %, respectively). The same MS also reported results on the occurrence of MRSA in meat production turkeys, at the farm in 2010 and 2012. The prevalence reported was moderate in both years (19.6 % in 2010 and 12.8 % in 2012). One non-MS reported data on the occurrence of MRSA in fattening pigs at slaughter through the monitoring of nasal swabs in consecutive years from 2009 to 2012. The numbers of animals positive for MRSA showed a slow increase over this period from 2.2 % in 2009 to 18.1 % in 2012. Molecular typing data were also available for these MRSA isolates, the majority of which belonged to *spa*-type t034, CC398, while much lower numbers of MRSA sequence type ST 49 were reported.



3. ANTIMICROBIAL RESISTANCE IN SALMONELLA

3.1. Introduction

Salmonella is an important zoonotic pathogen of economic significance in both humans and animals. The genus Salmonella is divided into two species: *S. enterica* and *S. bongori*. There are six sub-species of *S. enterica* and most Salmonella belong to the sub-species *S. enterica* subsp. *enterica*. Salmonella are further sub-divided into serovars based on the serological reactions of their somatic O-antigens and flagellar H-antigens. Different serovars have often been named based on the location where the serovar was first isolated. In this report, the organisms are identified by genus followed by serovar, e.g. *S*. Typhimurium. There are more than 2,500 serovars of zoonotic Salmonella which have been recognised, and the prevalence of these different serovars can change over time.

Human salmonellosis is usually characterised by the acute onset of fever, abdominal pain, nausea and sometimes vomiting. The majority of *Salmonella* infections result in mild, self-limiting, gastrointestinal illness and usually do not require antimicrobial treatment. In some patients the infection may be more serious and the associated dehydration can be life threatening. Invasive disease, such as *Salmonella* bacteraemia or meningitis, can occur in a smaller sub-set of patients, with a higher risk in patients who are immuno-compromised. In cases of severe enteric disease, or when *Salmonella* invades and causes a bloodstream infection, effective antimicrobials are essential for treatment and can be life saving. The treatment of choice for *Salmonella* infection is fluoroquinolones for adults and third-generation cephalosporins for children. Resistance in *Salmonella* to these first-line treatments, resulting in infections with antimicrobial-resistant strains, may cause treatment failure, which in turn can lead to more severe outcomes in patients. Salmonellosis has also been associated with long-term or chronic sequelae, e.g. reactive arthritis.

The common reservoir of non-typhoidal *Salmonella* strains is the intestinal tract of a wide range of domestic and wild animals. A wide variety of food stuffs of both animal and plant origin can be contaminated with *Salmonella*, which may cause infection in humans. Transmission usually occurs when the bacteria are introduced during food preparation or are allowed to multiply in food (for example because of inadequate storage temperature, inadequate cooking or cross-contamination of ready-to-eat food and uncooked food). *Salmonella* may also be transmitted through direct contact with infected animals or humans, or by contact with contaminated environments.

Overall, considering all *Salmonella* infections in the EU, *S*. Enteritidis and *S*. Typhimurium are the serovars most frequently associated with human illness. *S*. Enteritidis cases in humans are most commonly associated with the consumption of contaminated eggs and poultry meat, while *S*. Typhimurium cases are mostly associated with the consumption of contaminated pig, bovine and poultry meat.

In animals, particularly in certain species, sub-clinical infections or healthy carriage can be common. The organism may spread rapidly and easily between animals in a herd or flock without the animals showing any clinical signs in some cases and animals may become intermittent or persistent carriers. In other species, clinical disease may occur following *Salmonella* infection and, in particular, cattle may succumb to fever, diarrhoea and abortion following infection, particularly with some serovars such as *S*. Dublin. In calves, *Salmonella* can cause outbreaks of diarrhoea with high mortality. Fever and diarrhoea are less common in pigs than in cattle and sheep and poultry may also show no signs of infection.

Salmonella spp. comprise the amalgamated results for all Salmonella serovars reported by a MS. In the case of sampling in animals performed in accordance with EFSA's recommendations (EFSA, 2007) and related to National Salmonella Control Programmes (NCP), there is a defined method of selecting isolates for inclusion in the monitoring. The relative contribution of different serovars possessing a particular resistance should ideally be considered when interpreting the results, in order to evaluate the influence of clonal dissemination of serovars. If a MS has reported the susceptibility of particular serovars and excluded others, then this would introduce a source of bias in the susceptibility figures relating to Salmonella spp.

3.2. Overview of reported resistance data in Salmonella from humans, animals and food

Nineteen MSs, as well as Iceland and Norway, provided data for 2012 on human *Salmonella* isolates. Countries reported qualitative data (i.e. interpreted antibiotic susceptibility testing (AST) results for tested isolates; susceptible (S), intermediate (I) or resistant (R)) with the exception of Norway, which used the isolate-based reporting under piloting at ECDC to report measured IZDs. Twenty MSs and two non-MSs (Norway and Switzerland) reported quantitative MIC and disc inhibition zones data on the antimicrobial resistance of *Salmonella* isolates recovered from animals and food in 2012. Table SA1 presents an overview of the MSs reporting on antimicrobial resistance, either MIC or IZD data, on *Salmonella* spp. from humans and various animal and food categories in 2012.

Table SA1. Overview of countries reporting antimicrobial resistance data using MICs and disc diffusion inhibition zones on Salmonella spp. (all serovars) from humans and various animal and food categories in 2012

Method	Origin	Total number of MSs reporting	Countries
	Human	12	MSs: AT, EE, ES, FR, GR, HU, IT, LU, LT, RO, SI, SK
		2	Non-MSs: IS, NO
Diffusion	Gallus gallus (fowl)	1	MS: GR
	Meat from broilers (<i>Gallus gallus</i>)	1	MS: GR
	Meat from pigs	1	MS: GR
	Human	10	MSs: DE, DK, FR, HU, IE, MT, NL, RO, SK, UK
	Gallus gallus (fowl)	18	MSs: AT, BE, CZ, DE, DK, ES, FI, HU, IE, IT, LV, NL, PL, PT, RO, SK, SE, UK
	3 ()		Non-MS: CH
	Turkeys	13	MSs: AT, BE, CZ, DE, ES, FI, HU, IE, IT ,PL, PT, SK, UK
	Pigs	16	MSs: BE, CZ, DE, DK, EE, ES, FI, HU, IE, IT, LV, NL, PL, RO, SK, SE
Dilution	C		Non-MS: CH
	Cattle (bevine enimele)	12	MSs: BE, CZ, DK, EE, ES, FI, DE, IE, IT, LV, NL, SE
	Cattle (bovine animals)	12	Non-MSs: NO, CH
	Meat from broilers (<i>Gallus gallus</i>)	14	MSs: BE, CZ, DE, EE, ES, HU, IE, IT, LV, NL, PL, PT, RO, SK
	Meat from turkeys	10	MSs: CZ, DE, EE, HU, IE, IT, LV, PL, PT, SK
	Meat from pigs	14	MSs: BE, CZ, DE, DK, EE, ES, HU, IE, IT, LV, NL, PL, RO, SK
	Meat from bovine animals	9	MSs: CZ, DE, EE, ES, FI, HU, IE, IT, NL

MIC: minimum inhibitory concentration.

Note: for abbreviations of Member States (MS) and other reporting countries see Appendix 7.

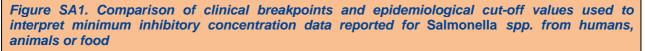


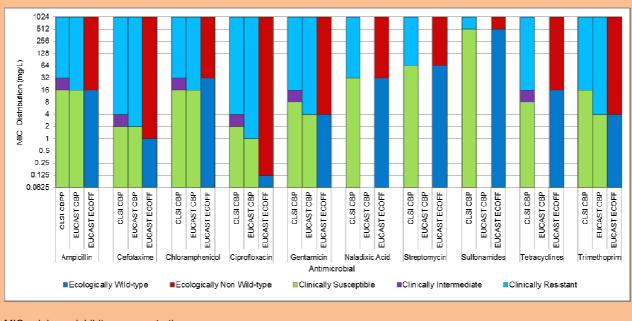
3.3. Antimicrobial resistance in *Salmonella* isolates from humans

METHODS AND INTERPRETIVE CRITERIA USED FOR ANTIMICROBIAL SUSCEPTIBILITY TESTING OF SALMONELLA ISOLATES FROM HUMANS

The method of testing for antimicrobial susceptibility and the selection of the isolates to be tested varied between countries. The methods and interpretive criteria used for antimicrobial susceptibility testing (AST) of Salmonella are presented in Table MM1 Chapter 8. Materials and methods. The European Centre for Disease Prevention and Control (ECDC) is working together with the national public health reference laboratories to harmonise the guidelines and interpretive criteria being used in AST for Salmonella (and Campylobacter). While nine of the reporting countries in 2012 were using criteria from the Clinical and Laboratory Standards Institute (CLSI), two countries were only using criteria from the European Committee on Antimicrobial Susceptibility Testing (EUCAST) and an additional five were using a combination of EUCAST and CLSI (since EUCAST do not provide clinical breakpoints for all antimicrobials included in the AST for Salmonella). Five countries used other criteria. Two of the Member States applied epidemiological cut-off values (ECOFFs) from EUCAST, while the other applied clinical breakpoints.

Of the 10 antimicrobials reported from both human and animal/food isolates, the minimum inhibitory concentration values or zone diameters differ markedly between the clinical breakpoints and the ECOFFs for four: cefotaxime, ciprofloxacin, gentamicin and trimethoprim. In particular, the ECOFF for ciprofloxacin is three dilution steps lower than the EUCAST clinical breakpoint and five dilution steps lower than the CLSI clinical breakpoint (Figure SA1). The results for these four antimicrobials must therefore be interpreted with caution and no direct comparison between countries should be made. Where countries have used the same method over the time period covered by the report, the trends in occurrence of resistance are likely to be valid, although sensitivity may vary depending on the specific thresholds used.





MIC: minimum inhibitory concentration.

Note: CLSI (M100-S22 2012), EUCAST clinical breakpoints (2011), EUCAST ECOFFS (as by EFSA, 2007).



3.3.1. Antimicrobial resistance in *Salmonella* spp. in humans

In total, 23,625 *Salmonella* spp. isolates were tested by the 19 reporting MSs, as well as Iceland and Norway, for resistance to one or more antimicrobials, representing 25.6 % (N=92,443) of all confirmed human salmonellosis cases reported in the EU/EEA in 2012. Levels of resistance are only shown when at least 20 isolates are tested. The number of antimicrobials tested per isolate varied by country. Ampicillin and ciprofloxacin were tested by all 19 MSs and 2 non-MSs, while sulfonamides were only tested by 15 MSs and 1 non-MS. Please note that the Danish data were excluded from Table SA2 since Denmark only reported data for *S*. Typhimurium and inclusion of these data would have skewed the EU average for *Salmonella* spp.

In order to better assess the impact from food consumed within each reporting country on the antimicrobial resistance levels found in human *Salmonella* isolates, the analysis focused on domestically-acquired cases. Several countries, however, did not provide information on travel (or non-travel) of their cases. Cases with unknown travel status were therefore also included in the analysis. A separate analysis was made on travel-associated cases by geographical regions.

3.3.1.1. Resistance levels in *Salmonella* spp. isolates from human cases

The highest average levels of resistance in human *Salmonella spp.* isolates in 2012 were reported for tetracyclines (30.0 %; N=15,233), sulfonamides (28.9 %; N=14,582), ampicillin (27.6 %, N=18,972) and streptomycin (23.6 %; N=16,643) (Table SA2). However, as in previous years, wide variability in frequencies of resistance to different antimicrobials was observed among the reporting countries. The variability may reflect the differences in the *Salmonella* population that people are exposed to in different countries, but may also arise due to differences in testing or sampling methods applied by individual countries as well as the use of different interpretive criteria. Sampling bias could also arise by only including isolates from hospitalised cases. This may explain some of the extreme observations.

Resistance levels can also differ substantially between *Salmonella* serovars and therefore an in-depth analysis is presented separately below for the three most common serovars, *S.* Enteritidis, *S.* Typhimurium and monophasic *S.* Typhimurium <u>1</u>,4,[5],12:i-,.

3.3.1.2. Comparison of resistance levels in *Salmonella* spp. isolates acquired within EU/EEA and in other geographical regions

To compare resistance levels in isolates acquired across the world, isolates from travel-associated cases were classified into seven different geographical regions¹³ EU/EEA, non-EU/EEA, Africa, Asia, North and Central America, South America and Oceania, based on the probable country of infection. Isolates from non-travel-associated cases were combined with those from cases infected in another EU/EEA country. Please note that the number of isolates tested per region does not necessarily reflect the number of travels to that region. Moreover, only 13 isolates were tested for infections acquired in Oceania and the results for this region should therefore be interpreted with caution.

Varying levels of resistance were observed among *Salmonella* spp. infections acquired in the different geographical regions (Table SA3). Isolates acquired within EU/EEA countries had the highest level of resistance to ampicillin (27.8 %; N=20,124), streptomycin (23.9 %; N=17,759), sulfonamides (29.2 %; N=15,686) and tetracyclines (30.2 %; N=16,340) of all regions. Resistance levels to ciprofloxacin was however higher in isolates from all other regions and particularly high in isolates originating in Asia (22.7 %; N=1,160), Africa (20.5 %; N=774) and Europe (non-EU/EEA countries) (19.5 %, N=41). Resistance levels to cefotaxime was also higher in isolates acquired in South America (4.8 %; N=21), Africa (2.1 %; N=767) and Asia (1.5 %; N=1,145) compared to the EU/EEA (1.1 %; N=18,949), though only a few isolates were tested from South America (Table SA3).

¹³ Regional classification from United Nations Statistical Division. Available online <u>http://unstats.un.org/unsd/methods/m49/m49regin.htm</u>



Country	Ampi	cillin	Cefota	axime	Chloram	phenicol	Ciprof	loxacin	Genta	micin	Kanai	nycin
Country	N	% Res	Ν	% Res	N	% Res	N	% Res	N	% Res	N	% Res
Austria	1,815	17.4	1,815	0.6	1,815	3.5	1,815	1.1	1,815	2.0	1,815	0.9
Denmark ²	_	_	-	_	_	_	Ι	-	_	_	Ι	_
Estonia	218	15.1	216	0.9	175	0	214	3.3	175	4.0	171	0.6
France	1,278	28.2	1,278	2.6	1,278	6.3	1,278	12.8	1,278	9.4	1,278	3.8
Germany	1,994	36.4	1,994	0.9	-	-	1,994	1.5	1,994	2.4	1,994	1.5
Greece	106	20.8	34	5.9	72	5.6	106	0	34	100	72	4.2
Hungary	588	55.1	588	2.4	588	12.9	588	0.3	588	2.6	588	1.0
Ireland	217	39.2	217	0.9	217	13.8	217	1.4	217	0.9	217	0.9
Italy	133	61.7	133	0	133	6.8	133	1.5	133	2.3	131	1.5
Latvia	54	11.1	17	NA	-	-	48	0	3	NA	_	_
Lithuania	1,734	13.7	1,478	0.7	812	2.0	1,362	0.3	673	0.3	624	0
Luxembourg	135	40.7	135	0	135	8.1	135	3.0	135	1.5	135	0.7
Malta	88	21.6	_	_	_	_	88	18.2	88	84.1	_	_
Netherlands ³	1,028	35.6	1,028	0.7	1,028	5.4	1,028	6.3	1,028	1.5	_	-
Romania	137	41.6	137	1.5	137	7.3	137	1.5	137	1.5	137	0
Slovakia	965	14.1	338	10.4	67	7.5	356	1.7	341	96.2	1	NA
Slovenia	392	15.3	392	0	392	8.2	392	0.8	392	6.9	392	5.6
Spain	1,874	47.9	1,875	1.5	1,873	8.1	1,874	0.9	1,874	2.3	1,874	2.1
United Kingdom	6,216	23.5	6,161	0.6	6,177	4.9	6,233	9.1	6,186	1.6	6,158	1.5
Total (18 MSs)	18,972	27.6	17,836	1.1	14,899	5.7	17,998	5.1	17,091	5.0	15,587	1.7
Iceland	22	0.1	-	_	22	0.1	22	0	_	_	Ι	_
Norway	359	20.1	_	_	359	7.2	359	0.3	_	_	_	_

Table SA2. Antimicrobial resistance in Salmonella spp. (all non-typhoidal serovars) from humans per country in 2012, using clinical breakpoints¹

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated; MS: Member State.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM2.

2. Data from Denmark not included in this table as only available for S. Typhimurium (see Table SA6) and monophasic S. Typhimurium (see Table SA7).

3. Epidemiological cut-off values were used for interpretation.

Table continued overleaf.



Table SA2 (continued). Antimicrobial resistance in Salmonella spp. (all non-typhoidal serovars) from humans per country in 2012, using clinical breakpoints¹

Country	Nalidixi	c acid	Strepto	omycin	Sulfonar	nides	Tetracy	/clines	Trimeth	oprim
Country	Ν	% Res	N	% Res	N	% Res	N	% Res	Ν	% Res
Austria	1,815	16.4	1,815	18.5	1,815	17.9	1,815	19.4	1,815	3.3
Denmark ²	_	_	_	_	_	_	_	_	_	_
Estonia	174	21.3	169	10.7	171	9.9	172	11.0	214	1.4
France	1,278	23.5	1,278	33.2	1,278	36.0	1,278	37.7	1,278	6.6
Germany	1,993	9.9	1,994	43.5	-	_	-	_	1,994	8.4
Greece	73	6.8	72	18.1	-	_	72	22.2	30	6.7
Hungary	588	33.0	588	46.9	588	57.1	588	55.3	588	8.0
Ireland	217	7.4	217	39.2	217	43.3	217	42.9	217	8.3
Italy	133	5.3	133	62.4	133	61.7	132	64.4	133	6.0
Latvia	-	-	-	-	-	-	-	-	53	0
Lithuania	643	11.7	623	6.6	622	6.4	621	9.0	1,736	2.2
Luxembourg	135	10.4	135	39.3	135	40.0	135	36.3	135	4.4
Malta	-	-	_	-	-	-	-	-	88	46.6
Netherlands ³	1,028	5.5	1,028	35.3	1,028	34.8	1,028	37.6	_	_
Romania	137	23.4	136	27.2	136	62.5	137	27.7	137	15.3
Slovakia	7	NA	11	NA	38	5.3	613	11.9	_	_
Slovenia	392	8.9	392	15.6	390	16.9	392	19.9	392	2.6
Spain	1,875	22.6	1,874	39.2	1,873	44.2	1,875	47.0	1,874	6.4
United Kingdom	6,201	11.5	6,178	8.7	6,158	23.9	6,158	26.6	6,228	8.8
Total (18 MSs)	16,689	14.4	16,643	23.6	14,582	28.9	15,233	30.0	16,912	6.9
Iceland	3	NA	_	_	_	_	_	_	22	0
Norway	359	10.3	47	91.5	47	95.7	359	21.4	359	3.9

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated; MS: Member State.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.

2. Data from Denmark not included in this table as only available for S. Typhimurium (see Table SA6) and monophasic S. Typhimurium (see Table SA7).

3. Epidemiological cut-off values were used for interpretation.



Table SA3. Antimicrobial resistance in Salmonella spp. (all non-typhoidal serovars) from humans acquired in the EU/EEA and other geographical regions in 2012, using clinical breakpoints¹

Country	Ampicillin		Cefotaxime		Chloramphenicol		Ciprofloxacin		Gentamicin		Kanamycin	
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Europe (EU/EEA Countries)	20,124	27.8	18,949	1.1	16,038	5.7	19,157	5.4	18,209	4.8	16,671	1.7
Europe (non-EU/EEA Countries)	41	4.9	40	0	36	2.8	41	19.5	40	7.5	38	2.6
Africa	773	20.2	767	2.1	756	7.8	774	20.5	770	7.1	734	2.7
Asia	1,158	24.3	1,145	1.5	1,132	6.9	1,160	22.7	1,147	4.6	1,075	4.0
Northern and Central America	178	3.4	175	0	176	2.3	181	10.5	178	2.8	173	0.6
South America	21	9.5	21	4.8	19	10.5	21	9.5	21	4.8	18	11.1
Oceania	13	15.4	13	0	13	15.4	13	15.4	13	7.7	13	0

Country	Nalidixic acid		Strept	Streptomycin		Sulfonamides		Tetracyclines		Trimethoprim	
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	
Europe (EU/EEA Countries)	17,814	14.2	17,759	23.9	15,686	29.2	16,340	30.2	18,041	6.8	
Europe (non-EU/EEA Countries)	38	26.3	38	5.3	35	5.7	35	11.4	40	5.0	
Africa	769	22.9	768	10.7	754	21.6	755	26.2	739	14.3	
Asia	1,147	22.9	1,146	10.9	1,126	24.6	1,128	25.7	1,086	10.7	
Northern and Central America	177	10.2	177	5.1	174	10.9	174	11.5	177	5.6	
South America	21	19.0	21	14.3	19	15.8	19	10.5	18	11.1	
Oceania	13	15.4	13	7.7	13	15.4	13	15.4	13	7.7	

N: number of isolates tested; % Res: percentage of resistant isolates; NA: not applicable - if less than 10 isolates were tested resistance was not calculated.

1. Epidemiological cut-off values were used for interpretation in the Netherlands and Denmark.



3.3.1.3. Multi-drug resistance among Salmonella spp. isolates from human cases

Thirteen MSs tested isolates for the full range of antimicrobials included in the human data collection for *Salmonella* spp., however as Denmark only provided data for serovars *Salmonella* Typhimurium and monophasic *S*. Typhimurium, which are known to be multi-drug resistant, the Danish data were excluded from the analysis to avoid bias.

About half of the human *Salmonella* spp. isolates in the 12 MSs were susceptible to all 10 antimicrobials (51.3 %; N=13,496), varying from 13.2 % (N=136) in Romania to 74.8 % (N=612) in Lithuania (Table SA4). Multi-drug resistance was high (28.9 %; N=13,496; country average 33.5 %) at the EU level, with the highest levels reported from Italy (63.1 %; N=130) and Hungary (55.8 %; N=588). The proportions of isolates susceptible to all and resistant (or non-susceptible) to any one up to 10 antimicrobials are presented by MSs in Figure SA2. The proportions differed substantially between countries. Isolates resistant to five antimicrobials were reported from all 12 MSs, and seven MSs (Austria, France, Hungary, Ireland, Lithuania, Spain and the United Kingdom) even reported a few isolates resistant to 9 or all 10 antimicrobials. The serotypes of those isolates resistant to 9 or 10 antimicrobials included monophasic S. Typhimurium 1,4,[5],12:i- (six isolates), S. Brandenburg (four), S. Typhimurium (four), S. Infantis (three), S. Montevideo (three), S. Agona (one), S. Albany (one), S. Concord (one), S. Derby (one), S. Haifa (one), S. Havana (one), S. Kentucky (one), and S. Panama (one).

Few isolates exhibited co-resistance to both ciprofloxacin and cefotaxime at the EU level (0.2 %; N=13,496) (Table SA4).

Country	Susceptible to all (%)	Multi-resistant (%)	Co-resistant to Cip and Ctx (%)		
Austria (N=1,815)	57.6	19.0	0.1		
Estonia (N=167)	64.1	11.4	0		
France (N=1,278)	49.1	37.0	0.4		
Hungary (N=588)	16.5	55.8	0.2		
Ireland (N=217)	52.5	41.9	0		
Italy (N=130)	33.1	63.1	0		
Lithuania (N=612)	74.8	7.2	0		
Luxembourg (N=135)	43.7	40.7	0		
Romania (N=136)	13.2	38.2	0		
Slovenia (N=390)	64.4	16.4	0		
Spain (N=1,870)	18.7	47.4	0.1		
United Kingdom (N=6,158)	60.9	23.7	0.3		
Total (12 MSs) (N=13,496)	51.3	28.9	0.2		

Table SA4. Complete susceptibility, multi-resistance and co-resistance (non-susceptibility) to ciprofloxacin and cefotaxime as determined by clinical breakpoints in Salmonella spp. from humans by MS, 2012¹

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; Cip: ciprofloxacin; Ctx: cefotaxime; MS: Member State.

Susceptible to all: proportion of isolates clinically susceptible to all antimicrobial substances of the European Centre for Disease Prevention and Control (ECDC) common set for *Salmonella*.

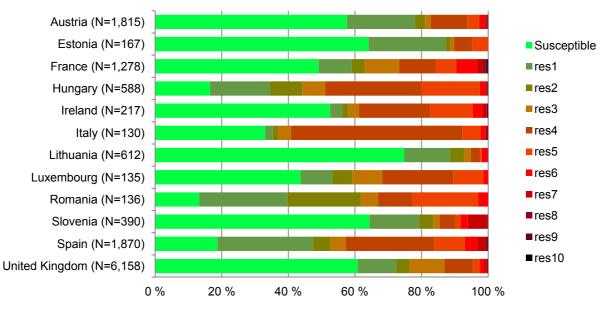
Multi-resistant: proportion of isolates clinically non-susceptible (resistant and intermediate) to at least three different antimicrobial substances belonging to any three antimicrobial families from the ECDC common antimicrobial set for *Salmonella*.

Co-resistant to Cip and Ctx: proportion of isolates clinically non-susceptible to both ciprofloxacin and cefotaxime.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.







N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella

Susceptible: total number of isolates susceptible to all antimicrobial substances of the common set for Salmonella.

res1-res10: total number of isolates non-susceptible (resistant and intermediate) to between 1 and 10 antimicrobial substances of the common set for *Salmonella*.

1. Isolates from cases reported as related to travel outside the country were excluded from this graph. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.

3.3.2. Antimicrobial resistance in Salmonella Enteritidis in humans

As in previous years, S. Enteritidis was the most common Salmonella serovar identified in 2012, with 34,019 cases reported in the EU/EEA. Seventeen MSs and Norway reported data on antimicrobial resistance in S. Enteritidis isolates for \geq 20 isolates (ranging from 13 MSs for sulfonamides to all 17 MSs and Norway for ciprofloxacin and ampicillin (Table SA5)).

3.3.2.1. Resistance levels in *Salmonella* Enteritidis isolates from human cases

The highest level of resistance among *S*. Enteritidis isolates in 2012 was observed for nalidixic acid (18.8 %; N=4,628). By country, the highest level was observed in Spain (60.3 %; N=527), followed by Romania (37.0 %; N=46), France (31.7 %; N=101) and Estonia (26.3 %; N=137) (Table SA5).

Compared with the data presented in this report for 2011 (EFSA and ECDC, 2013), the average level of resistance to ciprofloxacin was significantly lower in 2012 (4.9 %; N=5,598 in 2012 vs 12.7 %; N=7,965 in 2011). This can be explained by the exclusion in 2012 of isolates acquired during travelling as ciprofloxacin resistance levels were markedly higher in all other parts of the world than in the EU/EEA (see further Table SA3). Another reason for the lower resistance level is also that the countries reporting the highest ciprofloxacin resistance levels in 2011 (Denmark and Italy) did not test or only tested very few isolates of *S*. Enteritidis in 2012, thus having less effect on the total average than in 2011. By country, the highest resistance to ciprofloxacin in 2012 was reported in the United Kingdom (14.2 %; N=1,636), followed by the Netherlands (9.6 %; N=281) and Malta (7.4 %; N=27), all three of which used sensitive interpretive criteria for this antimicrobial.

As in previous years, resistance to cefotaxime was generally not detected or very low in the reporting MSs in 2012, total average 0.7 % (N=5,588), with the exception of Slovakia (11.3 %; N=240) (Table SA5). In Slovakia, cefotaxime resistance is not tested at the National Public Health Reference Laboratory and the methods and breakpoints applied in the testing laboratories are not known. Other noteworthy observations were the extremely high resistance levels to gentamicin among *S*. Entertitidis in Slovakia (97.1 %; N=244) and Malta (96.3 %; N=27), and the high resistance level to sulfonamides in Romania (41.3 %; N=46) and to trimethoprim/sulfamethoxazole in Malta (33.3 %; N=27) (Table SA5).



Table SA5. Antimicrobial resistance in S. Enteritidis from humans per country in 2012, using clinical breakpoints¹

Country	Ampicillin		Cefotaxime		Chloram	phenicol	Ciprof	loxacin	Genta	micin	Kana	mycin
Country	Ν	% Res	N	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	890	2.4	890	0.1	890	0	890	0	890	0	890	0
Estonia	172	7.6	170	0.6	138	0	169	3.6	139	0	135	0
France	101	3.0	101	0	101	1.0	101	0	101	0	101	0
Germany	207	1.4	207	0.5	-	-	207	0	207	0	207	0
Greece	30	6.7	5	NA	25	0	30	0	5	NA	25	4.0
Hungary	43	27.9	43	0	43	0	43	0	43	0	43	0
Ireland	30	3.3	30	0	30	0	30	0	30	0	30	0
Italy	7	NA	7	NA	7	NA	7	NA	7	NA	7	NA
Latvia	40	0	13	NA	_	-	39	0	_	-	-	-
Lithuania	1,405	8.5	1,200	0.3	659	0.2	1,106	0.2	531	0	499	0
Luxembourg	34	8.8	34	0	34	0	34	0	34	0	34	0
Malta	27	7.4	_	-	-	-	27	7.4	27	96.3	-	-
Netherlands ²	281	2.8	281	0.4	281	0.4	281	9.6	281	0	-	-
Romania	46	2.2	46	0	46	0	46	0	46	0	46	0
Slovakia	752	4.0	240	11.3	40	5.0	256	1.6	244	97.1	1	NA
Slovenia	169	2.4	169	0	169	0.6	169	0	169	0	169	0
Spain	527	12.9	527	0.4	526	0.6	527	0.4	527	0	527	0.2
United Kingdom	1,634	4.7	1,625	0.1	1,627	0.5	1,636	14.2	1,628	0.1	1,624	0.1
Total (18 MSs)	6,395	5.7	5,588	0.7	4,616	0.4	5,598	4.9	4,909	5.5	4,338	0.1
Iceland	3	NA	_	_	3	NA	3	NA	_	-	-	_
Norway	79	6.3	_	_	79	2.5	79	0	_	_	-	_

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated; MS: Member State.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.

2. Epidemiological cut-off values were used for interpretation.

Table continued overleaf.



Country	Nalidix	tic acid	Strepto	omycin	Sulfon	amides	Tetrac	yclines	Trimet	hoprim ³
Country	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res
Austria	890	5.5	890	0.6	890	0.4	890	1.3	890	0.2
Estonia	137	26.3	134	1.5	135	1.5	136	2.2	169	0.6
France	101	31.7	101	1.0	101	2.0	101	3.0	101	1.0
Germany	207	1.0	207	0.5	_	_	_	_	207	0.5
Greece	25	0	25	4.0	_	_	25	4.0	5	NA
Hungary	43	2.3	43	4.7	43	2.3	43	4.7	43	0
Ireland	30	16.7	30	3.3	30	3.3	30	3.3	30	0
Italy	7	NA	7	NA	7	NA	7	NA	7	NA
Latvia	_	-	_	_	_	_	-	_	42	0
Lithuania	499	9.8	498	0.2	498	0	496	3.6	1,405	1.1
Luxembourg	34	14.7	34	2.9	34	2.9	34	0	34	0
Malta	_	_	_	_	_	_	_	_	27	33.3
Netherlands ²	281	9.3	281	0.4	281	0.7	281	1.4	-	-
Romania	46	37.0	46	4.3	46	41.3	46	2.2	46	0
Slovakia	2	NA	-	-	31	6.5	468	1.9	-	_
Slovenia	169	4.1	169	1.2	167	3.0	169	3.6	169	0.6
Spain	527	60.3	527	1.9	527	2.8	527	2.7	527	1.1
United Kingdom	1,630	19.8	1,627	0.6	1,624	1.8	1,624	3.0	1,636	2.8
Total (18 MSs)	4,628	18.8	4,619	0.9	4,414	1.9	4,877	2.5	5,338	1.5
Iceland	-	_	_	_	_	_	_	_	3	NA
Norway	79	13.9	_	_	_	_	79	3.8	79	1.3

Table SA5 (continued). Antimicrobial resistance in S. Enteritidis from humans per country in 2012, using clinical breakpoints¹

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated; MS: Member State.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.

2. Epidemiological cut-off values were used for interpretation.

3. For at least the following countries, the results are for the combination trimethoprim/sulfamethoxazole: France, Germany, Hungary, Luxembourg, Malta, Spain and Norway.



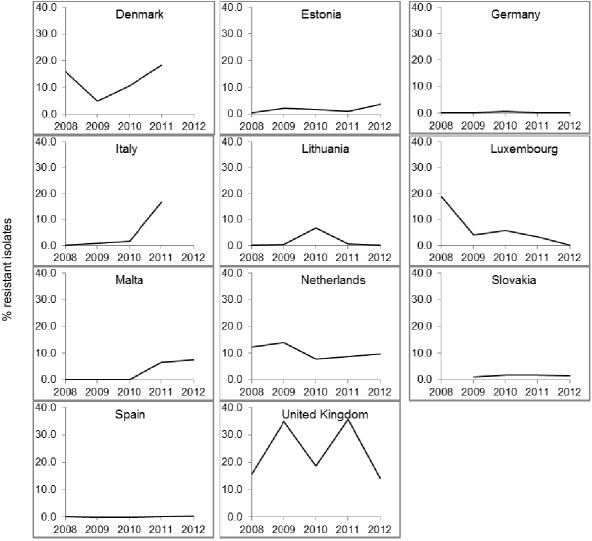
3.3.2.2. Trends in resistance levels in *Salmonella* Enteritidis isolates from human cases

Country-specific trends in resistance to ciprofloxacin and cefotaxime over the years 2008–2012 are presented in Figures SA3 and SA4. Only countries reporting data for at least three consecutive years and 10 isolates per year were included.

In eight MSs (Austria, France, Greece, Hungary, Ireland, Latvia, Romania and Slovenia) and Iceland, no isolates resistant to ciprofloxacin were reported in the five year period. There were few similarities in the trends of ciprofloxacin resistance levels across countries (Figure SA3). A sharp increase was observed in Denmark in 2010, continuing in 2011, and in Italy and Malta in 2011. To our knowledge, none of these could be explained by changes made in the interpretive criteria. No information was available to explain the resistance levels observed in the United Kingdom, which fluctuated greatly over the period (e.g. 35.6 %; N=1,751 in 2011 to 14.2 %; N=1,636 in 2012). In Luxembourg, a decreasing trend could be observed over the whole period.

Figure SA3. Resistance to ciprofloxacin in S. Enteritidis in humans in reporting MSs, 2008–2012¹

Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²



Year

Note: Countries reporting zero resistant isolates in the period were not shown as graphs: Austria, France, Greece, Hungary, Iceland, Ireland, Latvia, Romania and Slovenia; MS: Member State.

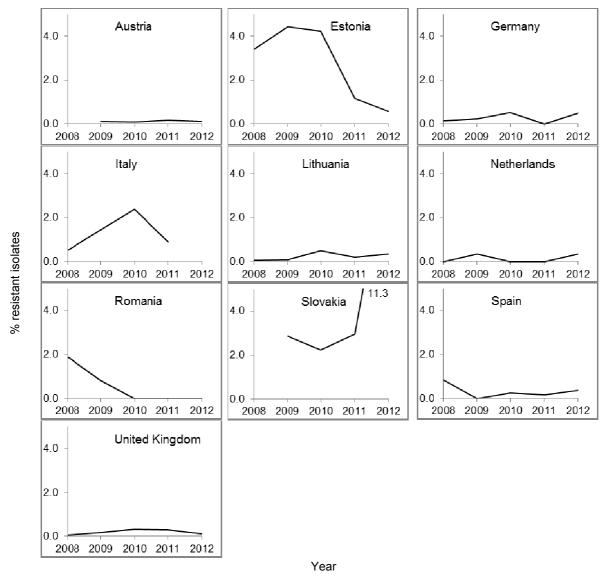
- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM2.
- Guidelines for interpretive criteria: Denmark (epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST)), Estonia (Clinical breakpoints (CBP) from Technical University of Denmark (DTU)/Clinical and Laboratory Standards Institute (CLSI)), Germany (CBP from Deutsches Institut für Normung (DIN)), Italy (CBP from CLSI), Lithuania (CB from CLSI), Luxembourg (CPB from CLSI), Malta (CBP from EUCAST), the Netherlands (ECOFFs from EUCAST), Slovakia (CBP from EUCAST), Spain (CB from CLSI), the United Kingdom (CBP from HPA). See also Table MM1.



The cefotaxime resistance were generally at very low levels in 2008 to 2012 in reporting MSs (Figure SA4). Denmark, Hungary, Ireland, Latvia, Luxembourg and Slovenia did not report any resistant isolates during the period. A sharp decrease in resistance levels was observed in Estonia and Italy in 2011. No visible increases in cefotaxime resistance levels could be observed in 2010 or 2011 in the countries using CLSI breakpoints although the breakpoint was changed from ≥ 64 mg/L to ≥ 4 mg/L in 2010. One exception was Slovakia where a slight increase in the resistance level was observed in 2011 and a drastic increase in 2012 (when Slovakia changed to EUCAST breakpoints which, however, are the same as CLSI regarding cefotaxime).

Figure SA4. Resistance to cefotaxime in S. Enteritidis in humans in reporting MSs, 2008–2012¹

Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²



MS: Member State.

Note: Countries reporting zero resistant isolates in the period were not shown as graphs: Denmark, Hungary, Ireland, Latvia, Luxembourg and Slovenia.

- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM2.
- 2. Guidelines for interpretive criteria: Austria (Clinical breakpoints (CBP) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST)), Estonia (CBP from Technical University of Denmark (DTU)/Clinical and Laboratory Standards Institute (CLSI)), Germany (CBP from Deutsches Institut für Normung (DIN)), Italy (CBP from CLSI), Lithuania (CBP from CLSI), the Netherlands (ECOFFs from EUCAST), Romania (CBP from CLSI), Slovakia (CBP from EUCAST), Spain (CBP from CLSI), the United Kingdom (CBP from the Health Protection Agency (HPA)). See also Table MM1.



3.3.3. Antimicrobial resistance in *Salmonella* Typhimurium in humans

As in previous years, S. Typhimurium was the second most common Salmonella serovar identified in 2012, with 18,248 cases reported in the EU/EEA. Sixteen MSs and Norway reported data on antimicrobial resistance in S. Typhimurium isolates for \geq 20 isolates (ranging from 13 MSs for kanamycin and sulfonamides to all 16 MSs and Norway for ciprofloxacin and ampicillin) (Table SA6).

3.3.3.1. Resistance levels in *Salmonella* Typhimurium isolates from human cases

The highest level of resistance in *S*. Typhimurium was observed for ampicillin (66.6 %; N=4,183), tetracyclines (63.7 %; N=3,272), sulfonamides (62.4 %; N=3,182) and streptomycin (46.2 %; N=3,951) (Table SA6). The levels of resistance to these antimicrobials were high to extremely high in the reporting MSs. Resistance levels observed in *S*. Typhimurium isolates to the two clinically most important antimicrobials were 2.2 % (N=4,114) for ciprofloxacin and 0.9 % (N=4,057) for cefotaxime. The highest resistance levels to ciprofloxacin were observed in the United Kingdom (4.3 %; N=1,686), the Netherlands (4.0 %; N=224) and Denmark (2.6 %; N=117), all three of which used more sensitive interpretive criteria than other countries. The highest level of resistance to cefotaxime was observed in France (3.6 %; N=110) (Table SA6).

Other noteworthy observations were the extremely high resistance in *S*. Typhimurium to gentamicin in Slovakia (92.9 %; N=56) and to streptomycin in Germany (82.3 %; N=744) (Table SA6).

3.3.3.2. Trends in resistance levels in *Salmonella* Typhimurium isolates from human cases

Country-specific trends in S. Typhimurium resistance to ciprofloxacin and cefotaxime over the years 2008 to 2012 are presented in Figures SA5 and SA6, respectively. Only countries reporting data for at least three consecutive years and 10 isolates per year were included.

The five-year trend (2008–2012) in resistance to ciprofloxacin by country showed that the countries using more sensitive interpretive criteria throughout the period (Denmark, the Netherlands and the United Kingdom) reported consistently higher levels of resistance compared with other countries (Figure SA5). A slight increasing trend over the period was observed in Denmark and decreasing trends were observed in Luxembourg and the Netherlands. A peak (smaller or larger) in resistance levels could be noted in 2010 in Denmark, Estonia, Lithuania, Luxembourg, the Netherlands and the United Kingdom, while in Italy, Malta and Slovakia, a peak was observed in 2011. Austria, France, Greece and Slovenia did not report any resistant isolates during the period.

There were few common trends between countries regarding cefotaxime resistance in *S*. Typhimurium during the period 2008 to 2012 (Figure SA6). An exception was a proportionally large increase in resistance levels observed in 2012 compared with previous years in Denmark, Hungary, Ireland and the Netherlands, and compared with 2011 in Lithuania, Romania and Spain. Please note, however, that the number of cefotaxime resistant isolates overall was very low and therefore even minor increases or decreases in the number of resistant isolates may influence the proportion of resistant isolates. A peak (smaller or larger) in resistance levels could also be noted in 2010 in Germany, Italy, Lithuania, the Netherlands and Spain, while in Austria, Slovakia and the United Kingdom, a peak was observed in 2011. Luxembourg did not report any resistant isolates during the period.



Table SA6. Antimicrobial resistance in S. Typhimurium from humans per country in 2012, using clinical breakpoints¹

Country	Amp	icillin	Cefota	axime	Chloram	phenicol	Ciprofl	oxacin	Genta	micin	Kanar	nycin
Country	N	% Res	N	% Res	N	% Res	N	% Res	N	% Res	N	% Res
Austria	236	56.8	236	0.4	236	21.2	236	0	236	1.3	236	1.7
Denmark ²	117	48.7	117	1.7	117	13.7	117	2.6	117	2.6	117	2.6
Estonia	19	NA	19	NA	10	NA	19	NA	10	NA	10	NA
France	110	60.9	110	3.6	110	43.6	110	0	110	0	110	0.9
Germany	744	81.2	744	0.9	_	_	744	0	744	0.9	744	1.9
Greece	29	37.9	15	NA	14	NA	29	0	15	NA	14	NA
Hungary	240	72.1	240	2.1	240	26.7	240	0	240	0.8	240	0.8
Ireland	59	59.3	59	1.7	59	45.8	59	1.7	59	1.7	59	1.7
Italy	33	69.7	33	0	33	12.1	33	0	33	0	33	0
Latvia	11	NA	4	NA	-	-	6	NA	3	NA	_	_
Lithuania	132	59.8	116	0.9	87	16.1	119	0	70	0	66	0
Luxembourg	27	70.4	27	0	27	25.9	27	0	27	0	27	0
Malta	7	NA	-	-	-	-	7	NA	7	NA	—	-
Netherlands ²	224	60.7	224	0.9	224	18.3	224	4.0	224	0.9	-	-
Romania	54	75.9	54	1.9	54	18.5	54	1.9	54	1.9	54	0
Slovakia	114	71.1	56	1.8	20	15.0	56	0	56	92.9	-	-
Slovenia	32	28.1	32	0	32	12.5	32	0	32	0	32	0
Spain	316	76.9	316	1.3	316	30.1	316	0	316	0.9	316	3.2
United Kingdom	1,679	63.0	1,655	0.4	1,662	12.3	1,686	4.3	1,666	1.6	1,654	1.6
Total (19 MSs)	4,183	66.6	4,057	0.9	3,241	18.3	4,114	2.2	4,019	3.0	3,712	1.7
Iceland	8	NA	_	-	8	NA	8	NA	_	-	_	_
Norway	76	28.9	_	-	76	21.1	76	0	_	-	_	_

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated; MS: Member State.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.

2. Epidemiological cut-off values were used for interpretation.

Table continued overleaf.



Country	Nalidix	ic acid	Strept	omycin	Sulfon	amides	Tetrac	yclines	Trimet	hoprim ³
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res
Austria	236	2.5	236	55.9	236	54.2	236	58.5	236	4.7
Denmark ²	117	1.7	117	52.1	117	53.0	117	31.6	117	7.7
Estonia	10	NA	9	NA	10	NA	10	NA	19	NA
France	110	6.4	110	61.8	110	66.4	110	64.5	110	5.5
Germany	744	9.9	744	82.3	_	_	_	_	744	10.2
Greece	14	NA	14	NA	_	_	14	NA	12	NA
Hungary	240	2.9	240	57.1	240	64.2	240	59.6	240	12.1
Ireland	59	8.5	59	55.9	59	61.0	59	61.0	59	16.9
Italy	33	3.0	33	69.7	33	72.7	33	72.7	33	12.1
Latvia	_	_	_	_	_	_	_	_	8	NA
Lithuania	66	12.1	66	48.5	66	48.5	66	43.9	135	5.9
Luxembourg	27	3.7	27	55.6	27	63.0	27	59.3	27	0
Malta	-	-	-	_	-	-	-	-	7	NA
Netherlands ²	224	3.6	224	50.0	224	55.8	224	57.1	-	-
Romania	54	3.7	54	51.9	54	83.3	54	51.9	54	27.8
Slovakia	3	NA	8	NA	4	NA	80	53.8	-	-
Slovenia	32	9.4	32	31.3	32	28.1	32	31.3	32	6.3
Spain	316	15.2	316	57.6	316	74.1	316	76.3	316	12.0
United Kingdom	1,671	4.5	1,662	22.6	1,654	63.1	1,654	68.5	1,685	14.7
Total (19 MSs)	3,956	6.4	3,951	46.2	3,182	62.4	3,272	63.7	3,834	12.0
Iceland	2	NA	_	_	I	-	_	_	8	NA
Norway	76	7.9	14	NA	14	NA	76	27.6	76	1.3

Table SA6 (continued). Antimicrobial resistance in S. Typhimurium from humans per country in 2012, using clinical breakpoints¹

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated; MS: Member State.

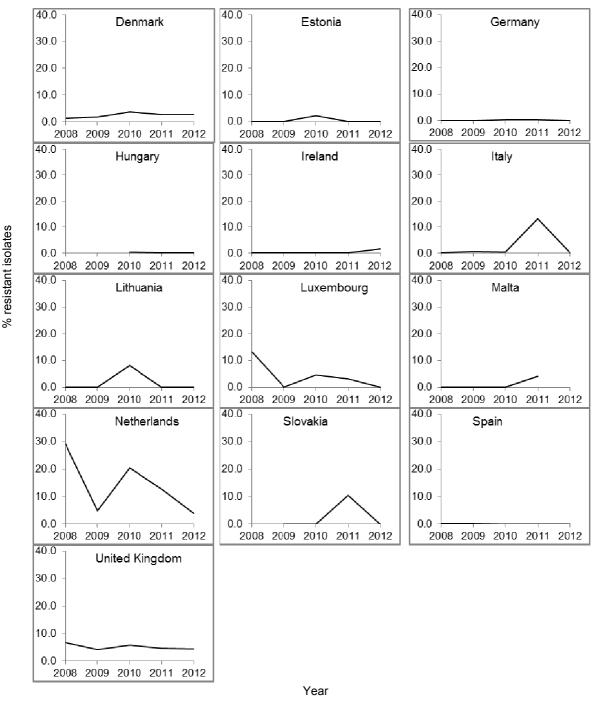
1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.

2. Epidemiological cut-off values were used for interpretation.

3. For at least the following countries, the results are for the combination trimethoprim/sulfamethoxazole: France, Germany, Hungary, Luxembourg, Malta, Spain and Norway.

Figure SA5. Resistance to ciprofloxacin in S. Typhimurium in humans in reporting MSs, 2008–2012¹

Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²



MS: Member State.

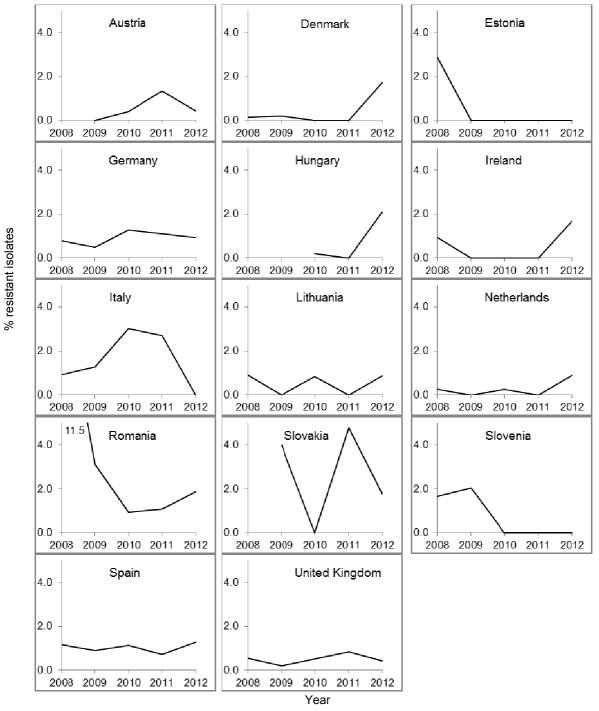
Note: Countries reporting zero resistant isolates in the period were not shown as graphs: Austria, France, Greece and Slovenia.

- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM2.
- 2. Guidelines for interpretive criteria: Denmark (epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST)), Estonia (Clinical breakpoints (CBP) from Technical University of Denmark (DTU)/Clinical and Laboratory Standards Institute (CLSI)), Germany (CBP from Deutsches Institut für Normung (DIN)), Hungary (CBP from EUCAST), Ireland (CBP from EUCAST), Italy (CBP from CLSI), Lithuania (CBP from CLSI), Luxembourg (CBP from CLSI), Malta (CB from EUCAST), the Netherlands (ECOFFs from EUCAST), Slovakia (CBP from EUCAST), Spain (CBP from CLSI), the United Kingdom (CBP from the Health Protection Agency (HPA)). See also Table MM1.



Figure SA6. Resistance to cefotaxime in S. Typhimurium in humans in reporting MSs, 2008-2012¹

Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²



MS: Member State.

Note: Countries reporting zero resistant isolates in the period were not shown as graphs: Luxembourg.

- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM2.
- 2. Guidelines for interpretive criteria: Austria (Clinical breakpoints (CBP) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST)), Denmark (epidemiological cut-off values (ECOFFs) from EUCAST), Estonia (Clinical breakpoints (CBP) from Technical University of Denmark (DTU)/Clinical and Laboratory Standards Institute (CLSI)), Germany (CBP from Deutsches Institut für Normung (DIN)), Hungary (CBP from EUCAST), Ireland (CBP from EUCAST), Italy (CBP from CLSI), Lithuania (CBP from CLSI), the Netherlands (ECOFFs from EUCAST), Romania (CBP from CLSI), Slovakia (CBP from EUCAST), Slovenia (CBP from CLSI), Spain (CBP from CLSI), the United Kingdom (CBP from the Health Protection Agency (HPA)). See also Table MM1.



3.3.4. Antimicrobial resistance in monophasic *Salmonella* Typhimurium <u>1</u>,4,[5],12:i:- in humans

Since the introduction of a separate serovar code in the reporting in 2010, monophasic *S*. Typhimurium <u>1</u>,4,[5],12:i:- has become the third most common *Salmonella* serovar in humans. In 2012, 5,932 cases were reported by the EU/EEA countries. Eight MSs reported data on antimicrobial resistance in monophasic *S*. Typhimurium <u>1</u>,4,[5],12:i:- isolates for \geq 20 isolates for all 11 antimicrobials except for kanamycin and trimethoprim which seven MSs reported data for (Table SA7).

3.3.4.1. Resistance levels in monophasic *Salmonella* Typhimurium <u>1</u>,4,[5],12:i:- isolates human cases

Extremely high levels of resistance were observed for tetracyclines (92.7 %; N=1,263), ampicillin (89.5 %; N=1,263), sulfonamides (89.1 %; N=1,262) and streptomycin (86.9 %; N=1,263) in monophasic S. Typhimurium <u>1</u>,4,[5],12:i- (Table SA7). This resistance pattern, ASSuT, is a well-known character for monophasic S. Typhimurium <u>1</u>,4,[5],12:i- and was observed at similar levels in all nine reporting MSs.

Resistance levels to the two clinically most important antimicrobials were 0.7 % (N=1,263) for ciprofloxacin and 2.1 % (N=1,263) for cefotaxime, which regarding ciprofloxacin was lower than in other S. Typhimurium but regarding cefotaxime was higher. The highest resistance levels to ciprofloxacin was observed in Italy (3.2 %; N=62) and Denmark (1.9 %; N=107). It should be noted that Denmark used ECOFFs for interpretation which are more sensitive than the clinical breakpoints for ciprofloxacin. The highest levels of resistance to cefotaxime were observed in France (5.2 %; N=115), Spain (2.8 %; N=571) and Austria (2.1 %; N=95) (Table SA7).



Country	Ampi	cillin	Cefotaxime		Chloramphenicol		Ciprof	loxacin	Genta	micin	Kanamycin		
Country	N	% Res	Ν	% Res	N	% Res	N	% Res	N	% Res	N	% Res	
Austria	95	93.7	95	2.1	95	5.3	95	0	95	3.2	95	2.1	
Denmark ²	107	83.2	107	0	107	0	107	1.9	107	0.9	107	0	
Estonia	12	NA	12	NA	12	NA	12	NA	12	NA	12	NA	
France	115	90.4	115	5.2	115	2.6	115	0	115	0	115	0	
Ireland	45	86.7	45	0	45	0	45	0	45	0	45	0	
Italy	62	91.9	62	0	62	6.5	62	3.2	62	3.2	60	0	
Luxembourg	30	96.7	30	0	30	6.7	30	0	30	3.3	30	3.3	
Netherlands ²	226	85.4	226	0.9	226	2.7	226	0.9	226	2.2	-	-	
Spain	571	90.9	571	2.8	571	5.3	571	0.5	571	3.7	571	3.2	
Total (9 MSs)	1,263	89.5	1,263	2.1	1,263	4.0	1,263	0.7	1,263	3.2	1,035	2.0	

Table SA7. Antimicrobial resistance in monophasic S. Typhimurium <u>1</u>,4,[5],12:i:- from humans per country in 2012, using clinical breakpoints¹

Country	Nalidix	ic acid	Strepto	omycin	Sulfon	amides	Tetracyclines		Trimethoprim	
Country	N	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res
Austria	95	0	95	92.6	95	93.7	95	91.6	95	6.3
Denmark ²	107	0	107	87.9	107	89.7	107	96.3	107	0
Estonia	12	NA	12	NA	12	NA	12	NA	12	NA
France	115	2.6	115	89.6	115	92.2	115	93.0	115	6.1
Ireland	45	2.2	45	97.8	45	97.8	45	97.8	45	0
Italy	62	3.2	62	90.3	62	90.3	62	90.3	62	3.2
Luxembourg	30	3.3	30	90.0	30	93.3	30	80.0	30	10.0
Netherlands ²	226	0.4	226	88.1	226	85.4	226	94.2	-	-
Spain	571	1.8	571	83.0	570	87.7	571	91.9	571	6.8
Total (9 MSs)	1,263	1.4	1,263	86.9	1,262	89.1	1,263	92.7	1,037	5.5

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated; MS: Member States.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM2.

2. Epidemiological cut-off values were used for interpretation.

3.4. Antimicrobial resistance in *Salmonella* isolates from animals and food

Nineteen MSs and two non-MSs (Norway and Switzerland) reported quantitative MIC data and one MS reported IZD data on antimicrobial resistance in *Salmonella* isolates recovered from animals and food in 2012. The MSs reporting either MIC or IZD data, for each animal or food category, are listed in Tables SA1, SA8 and SA9. As quantitative IZD data constitute a relatively small percentage of the total data available, these data have, therefore, been analysed as qualitative data only. The susceptibility test results for *Salmonella* isolates reported as qualitative data are presented in Appendix 2.

In this report, antimicrobial resistance data for all reported *Salmonella* isolates were collated to generate a figure for *Salmonella* spp. (covering all reported serovars) for each country, year and animal/food category. In addition, the *Salmonella* serovars which are most prevalent and significant for public health, *S*. Enteritidis, *S*. Typhimurium and monophasic *S*. Typhimurium, were reported separately where sufficient quantitative data were available from the various animal/food categories. Frequency distributions of serovars per animal and meat categories are presented in Appendix 1.

Table SA8. Overview of countries reporting quantitative antimicrobial resistance data using MIC and disc inhibition zones on S. Typhimurium from various animal and food categories in 2012

Method	Origin	Total number of MSs reporting	Countries
Diffusion	Gallus gallus (fowl)	1	MS : GR
Dillusion	Meat from pig	1	MS: GR
		12	MSs: AT, BE, DE, DK, ES, FI, HU, IT, PL, RO, SE, UK
	Gallus gallus (fowl)	12	Non-MS: CH
	Turkeys	7	MSs: AT, BE, DE, ES, FI, IT, UK
			MSs: BE, CZ, DE, DK, EE, ES, FI, HU, IE, LV, NL, RO,
	Pigs	14	SE, SK
			Non-MS: CH
Dilution	Cattle (bovine animals)	11	MSs: BE, CZ, DE, DK, EE, ES, FI, IE, IT, NL, SE
		11	Non-MS: CH
	Meat from broilers (Gallus gallus)	3	MSs: IE, IT, LV
	Meat from turkeys	4	MSs: DE, HU, IT, PT
	Meat from pigs	12	MSs: BE, CZ, DE, DK, EE, HU, IE, IT, LV, NL, PL, RO
	Meat from bovine animals	7	MSs: CZ, DE, EE, FI, IE, IT, NL

Note: For abbreviations of Member States (MS) and other reporting countries, see Appendix 7; MIC: minimum inhibitory concentration.

Table SA9. Overview of countries reporting quantitative antimicrobial resistance data using MIC and disc inhibition zones on S. Enteritidis from various animal and food categories in 2012

Method	Origin	Total number of MSs reporting	Countries
Diffusion	Gallus gallus (fowl)	1	MS: GR
	Gallus gallus (fowl)	15	MSs: AT, BE, CZ, DE, DK, ES, HU, IT, LV, NL, PL, PT, RO, SK, UK
	0 ()		Non-MS: CH
	Turkeys	5	MSs: AT, CZ, DE, HU, SK
	Pigs	5	MSs: BE, DE, DK, EE, HU
Dilution	Cattle (bovine animals)	4	MSs: BE, CZ, DK, EE
	Calle (Dovine animals)	4	Non-MS: CH
	Meat from broilers (Gallus gallus)	7	MSs: BE, CZ, DE, LV, NL, PL, SK
	Meat from pigs	2	MSs: CZ, RO
	Meat from turkeys	3	MSs: CZ, DE, HU

Note: For abbreviations of Member States (MS) and other reporting countries, see Appendix 7; MIC: minimum inhibitory concentration.

In this chapter, resistance to ampicillin, cefotaxime, ciprofloxacin, chloramphenicol, gentamicin, nalidixic acid, sulfonamides and tetracyclines has been analysed in detail in line with the antimicrobials listed in the EFSA monitoring and reporting specifications for antimicrobial resistance in *Salmonella* (EFSA, 2007). The occurrence of resistance to these substances was tabulated, temporal trends and spatial distribution of resistance was drawn and the multi-resistance and a specific co-resistance pattern was analysed. The results of these analyses are displayed in this report according to the following inclusion criteria: whenever a country subjected more than 10 isolates to susceptibility testing for a given animal or food category then these data were not included in any further analyses. In addition, tables were generated and analysis was performed only if four or more countries tested and reported quantitative data for a given *Salmonella* category and sampling origin.

- Temporal trend graphs of resistance were generated showing resistance to ampicillin, cefotaxime, ciprofloxacin and nalidixic acid for *Salmonella* isolates from animals and food over the 2006–2012 period. Only countries which had reported data for four or more years during 2006–2012 were included. Data from 2004 and 2005 were excluded from the temporal trends graphs because of the relative scarcity of data compared with the 2006–2012 period. Statistical analysis of the trend within individual countries was performed using logistic regression when data were available for five or more years.
- The spatial distributions of ampicillin and nalidixic acid resistance rates in *Salmonella* spp. from *Gallus gallus*, turkeys, pigs and cattle were presented. For countries where resistance level figures for 2012 were not available, 2011 figures were used instead.
- Multi-resistance was analysed in isolate-based data on *Salmonella* isolates tested for the full harmonised set of antimicrobials (nine substances) belonging to different classes. Multi-resistance was defined as the non-susceptibility to at least three different antimicrobial classes. The proportions of isolates susceptible to all antimicrobials and resistant (non-susceptible) from one to nine antimicrobials are presented. Co-resistance to cefotaxime (Ctx) and ciprofloxacin (Cip) was estimated as these two antimicrobials are of particular interest in human medicine in the case of treatment of severe salmonellosis. Co-resistance was addressed using both ECOFFs (Ctx>0.5 mg/L and Cip>0.06 mg/L) and clinical breakpoints (Ctx>2 mg/L and Cip>1 mg/L). Complete susceptibility and multi-resistance were analysed for isolate-based resistance data reported by the MSs and the results are shown at Appendix 4.

The antimicrobials selected by the different MSs and non-MSs for susceptibility testing of *Salmonella* are shown in Chapter 8 Materials and methods, Table MM6. For further information on reported MIC distributions and number of resistant isolates for apramycin, ceftazidime, ceftiofur, colistin, florfenicol, kanamycin, neomycin, spectinomycin, streptomycin and trimethoprim, refer to the Level 3 tables published on the EFSA website.

3.4.1. Antimicrobial resistance in *Salmonella* isolates from food

This section describes the MIC data for isolates of *Salmonella* spp. and *S.* Enteritidis from meat from broilers, *Salmonella* spp., *S.* Typhimurium and monophasic *S.* Typhimurium from meat from pigs and *Salmonella* spp. from meat from bovine animals.

3.4.1.1. Meat from broilers and spent hens (Gallus gallus)

Quantitative MIC susceptibility data for isolates of *Salmonella* spp. from broiler meat from 12 MSs in 2012 are included in the following analysis. Data for *S.* Typhimurium isolates are not presented separately for meat from broilers as no MS tested more than 10 isolates.

Representative sampling and monitoring

In reporting MSs, data on antimicrobial resistance in *Salmonella* isolates from meat from *Gallus gallus* derived from active monitoring carried out within the framework of either official sampling or Hazard Analysis and Critical Point Control (HACCP) and own-check programmes. Samples of meat from broilers and spent hens were collected randomly at either the slaughterhouse, the cutting/processing plant, or at retail outlets. In Belgium, the Czech Republic, Ireland and Romania, representative random sampling of meat from



broilers, whether carcase swabs or samples of neck skin, fresh meat, minced meat, meat products or meat preparations, was carried out entirely or primarily at the slaughterhouse or at the processing plant. In Portugal and Romania, sampling of fresh meat and meat preparations from broilers was also performed at retail. Moreover, Belgium was the only reporting MS to monitor resistance in *Salmonella* isolates derived from swabbing of both carcases of spent hens and chilled broilers at the slaughterhouse. Germany, Hungary, the Netherlands, Poland and Spain did not provide any details on the sampling scheme used.

Resistance levels in *Salmonella* spp. isolates from meat from broilers and spent hens

Table SA10 describes the occurrence of resistance to selected antimicrobials in *Salmonella* spp. isolated mostly from broiler meat in MSs in 2012.

In 2012, resistance to ampicillin, sulfonamides and tetracyclines was generally high to extremely high in the reporting MSs, although levels of resistance were highly variable across countries, ranging from 0 % to 48.6 % for ampicillin and from 0 % to 87.3 % for both sulfonamides and tetracyclines. Resistance levels to chloramphenicol and gentamicin were typically lower, a number of MSs observing no resistance to one or both of these antimicrobials. Variation in the resistance recorded was important, between 0 % and 18.1 % for chloramphenicol and from 0 % to 45.5 % for gentamicin.

As in previous years, the occurrence of resistance to ciprofloxacin and nalidixic acid was typically similar within MSs, while, between countries, resistance levels ranged from 0 % to 100 %. The overall level of resistance to cefotaxime across the reporting MSs remained low in 2012 at 4.3 %, as generally resistance was either not detected or observed at low levels. The Netherlands reported a high level of resistance to cefotaxime of 31.1 %, which was similar to that (31.9 %) reported in 2011.

Considering the figures at reporting MS group level for *Salmonella* spp. isolated from broiler meat, there were increases in the levels of resistance to sulfonamides and tetracyclines which stood at 44.8 % and 43.7 % in 2011 and were 53.0 % and 48.9 % in 2012. However, there are a number of factors which can influence changes in the occurrence of resistance in *Salmonella* spp. in different years: (1) differences in the contributing MSs and their relative contribution to the total, since levels of resistance vary greatly between MSs, (2) differences in the relative contribution of different serovars (which can vary greatly in the resistances shown), (3) changes in the level of resistance.

Salmonella spp. comprise the amalgamated results for all Salmonella serovars reported by a reporting MS for a different animal or food category. The relative contribution of different serovars possessing a particular resistance should ideally be considered when interpreting the results, in order to evaluate the influence of clonal dissemination of serovars. The chapter has also presented data at the individual serovar level for a number of *Salmonella* serovars.

Resistance levels in *Salmonella* Enteritidis isolates from meat from broilers and spent hens

Resistance among *S*. Enteritidis isolates from broiler meat in reporting MSs was generally lower than that reported in *Salmonella* spp. As low numbers of isolates of *S*. Enteritidis (fewer than 10) were recovered from meat from broilers in the Czech Republic, Germany, the Netherlands and Slovakia, these countries have been excluded from the detailed analysis, leaving only Belgium, Latvia and Poland contributing to the analysis; thus, there are insufficient data to present a specific table.

Data from Belgium showed marked differences in the occurrence of ciprofloxacin resistance in *Salmonella* spp. from meat from broilers and spent hens. While no resistance was detected in isolates from spent hens there was 39.5 % ciprofloxacin resistance in isolates from broilers.

One isolate (4.3 %) was resistant to ampicillin, sulfonamides and tetracyclines in Latvia, while four isolates (15.4 %) were resistant to ampicillin and sulfonamides and one isolate (3.8 %) was resistant to tetracyclines in Poland.

In both Latvia and Poland, no resistance was detected to cefotaxime, chloramphenicol and gentamicin, while resistance to ciprofloxacin and nalidixic acid was recorded in both countries. In Latvia, 16 isolates (69.6 %) were resistant to both antimicrobials, while 14 isolates and 13 isolates (53.8 % and 50.0 %) were resistant to ciprofloxacin and nalidixic acid, respectively.



Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidia	kic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Belgium ¹	43	34.9	43	0	43	0	43	39.5	43	0	43	39.5	43	11.6	43	0
Belgium ²	66	0	66	0	66	0	66	0	66	0	66	0	66	0	66	0
Czech Republic	47	4.3	47	0	47	0	47	34.0	47	0	47	34.0	47	34.0	_	-
Germany	94	39.4	94	12.8	94	18.1	94	70.2	94	3.2	94	58.5	94	58.5	94	48.9
Hungary	164	4.3	168	0.6	168	1.8	168	97.6	168	1.2	168	95.8	168	79.8	168	76.8
Ireland	70	22.9	70	2.9	70	7.1	70	11.4	70	4.3	70	11.4	70	21.4	70	7.1
Latvia	32	9.4	32	0	32	3.1	32	53.1	32	0	32	53.1	32	9.4	32	6.3
Netherlands	74	48.6	74	31.1	74	9.5	74	51.4	72	1.4	74	47.3	74	52.7	74	36.5
Poland	93	23.7	93	0	93	2.2	93	67.7	93	1.1	93	52.7	93	35.5	93	31.2
Portugal	37	24.3	37	0	37	5.4	37	13.5	37	13.5	37	13.5	37	43.2	37	56.8
Romania	189	18.0	189	1.1	189	10.1	189	89.4	189	5.3	189	89.9	189	87.3	189	87.3
Slovakia	14	0	14	0	14	0	14	71.4	14	0	14	71.4	14	71.4	14	71.4
Spain	33	27.3	-	_	15	0	33	100	33	45.5	33	21.2	-	_	15	26.7
Total (12 MSs)	956	19.9	927	4.3	942	5.9	960	63.1	958	4.2	960	57.3	927	53.0	895	48.9

Table SA10. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella spp.</u> from <u>meat from broilers and spent hens</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested, % Res: percentage of resistant isolates; -: no data reported.

1. Broilers

2. Spent hens



Multi-resistance among Salmonella isolates from meat from broilers

In 2012, four MSs provided isolate-based data concerning resistance in *Salmonella* spp. in meat from broilers. Important variability was observed in both rates of complete susceptibility and multi-resistance of the isolates tested in the reporting MSs (Table SA11). The frequency distributions (Figure SA7) showed that reduced susceptibility to six or up to eight different substances might be observed in some tested isolates, while the Czech Republic recorded multi-resistance to three classes at a maximum. Very few isolates were resistant to both ciprofloxacin and cefotaxime (Table SA11).

Table SA11. Complete susceptibility, multi-resistance and index of diversity in <u>Salmonella spp.</u> from <u>meat from broilers</u> in MSs reporting isolate-based data, 2012

Country	Suscep	tible to all	Multi-r	esistant	Index of	Co-resistant to Cip and Ctx			
	n	%	n	%	diversity	n	%		
Czech Republic (N=47)	30	63.8	16	34.0	0.289	0 (0)	0 (0)		
Germany (N=94)	23	24.5	61	64.9	0.582	9 (1)	9.6 (1.1)		
Ireland (N=70)	51	72.9	16	22.9	0.395	2 (0)	2.9 (0)		
Romania (N=188)	13	6.9	163	86.7	0.542	2 (0)	1.1 (0)		

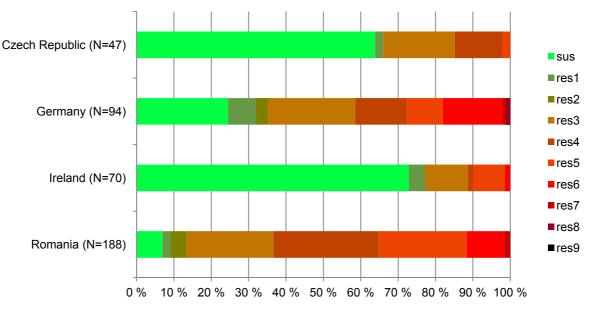
MS:Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella; n: number of isolates.

Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set. Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of Salmonella isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip:>1 mg/L).

Figure SA7. Frequency distribution of <u>Salmonella spp.</u> in <u>meat from broilers</u> completely susceptible or resistant to one or up to nine antimicrobials, in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*.



3.4.1.2. Meat from pigs

Twelve MSs reported quantitative MIC data for *Salmonella* spp. from pig meat in 2012. Data for *S*. Enteritidis isolates are not presented separately for meat from pigs as none of the MSs reporting data tested more than 10 isolates. Tables SA12 and SA13 present the level of resistance to selected antimicrobials for *Salmonella* spp. and *S*. Typhimurium.

Representative sampling and monitoring

Pig meat sample types collected at the slaughterhouse by Belgium, Estonia, Ireland and Poland consisted mainly of carcase swabs, while different kinds of meat samples (fresh meat, minced meat and meat preparations) were collected by the Czech Republic, Denmark, Estonia, Ireland, Italy and Romania either at meat cutting/processing plants, at retail outlets or at catering outlets. Whether based on official sampling or HACCP and own-check programmes, objective sampling was primarily carried out in MSs reporting information on sampling design. Conversely, Germany, Hungary, Poland and Spain did not report detailed information on the sampling design of meat from pigs used.

Resistance levels in Salmonella spp. isolates from meat from pigs

Salmonella spp. isolated from pig meat mostly displayed high to extremely high occurrence of resistance to ampicillin, tetracyclines and sulfonamides, although resistance varied from not detected (in Latvia on 13 isolates) to very/extremely high levels across the reporting MSs, varying from 0 % to nearly 70 % for ampicillin and sulfonamides and to more than 90 % for tetracyclines. Chloramphenicol resistance remained moderate, at 12.6 %, at MS group level, and ranged from 0 % to 44.4 % among the reporting MSs. Overall, gentamicin resistance was 2.4 % in the reporting group of MSs; it was not detected in seven MSs and ranged between 0.4 % and 31.3 % in the remaining five reporting MSs. The proportion of *Salmonella* spp. isolates resistant to ciprofloxacin and nalidixic acid was generally low to moderate among the reporting MSs. Once more in 2012, Denmark and Estonia did not detect resistance to either ciprofloxacin or nalidixic acid, and neither did Latvia on the limited number of 13 isolates tested. Resistance to cefotaxime in reporting MSs was either not detected (in eight MSs) or reported at low levels, ranging from 0.6 % to 3.0 %, between the four MSs reporting resistance.

Resistance levels in Salmonella Typhimurium isolates from meat from pigs

Ten MSs reported quantitative MIC data for *S*. Typhimurium isolates from pig meat in 2012. Although the features of the resistance in *S*. Typhimurium parallel those observed in *Salmonella* spp., resistance levels to most of the antimicrobials in *S*. Typhimurium isolates were typically higher than the levels reported in *Salmonella* spp. isolates in the vast majority of reporting MSs. Contrastingly, in Denmark, *S*. Typhimurium isolates presented lower levels of resistance to ampicillin, sulfonamides and tetracyclines than the overall group of *Salmonella* spp. isolates. The 13 *S*. Typhimurium isolates tested in Latvia did not exhibit resistance to any of the antimicrobials tested.

Resistance levels in monophasic Salmonella Typhimurium isolates from meat from pigs

As low numbers of isolates of monophasic *S*. Typhimurium (fewer than 10) were recovered from meat from pigs in the Czech Republic, Greece, Hungary, Italy, the Netherlands and Slovakia, these countries have been excluded from the detailed analysis, leaving only Denmark, Germany and Ireland contributing to the analysis; thus, there are insufficient data to present a specific table.

Denmark detected no resistance to cefotaxime, gentamicin, ciprofloxacin and nalidixic acid. In both Germany and Ireland, no resistance was detected to cefotaxime. Resistance was detected against ampicillin, sulfonamides and tetracyclines at extremely high levels ranging from 73.9 % to 100 %. One isolate (2.7 %) was resistant to both ciprofloxacin and nalidixic acid in Germany, whilst Ireland reported no resistance for these antimicrobials. One isolate (4.5 %) was resistant to chloramphenicol in Denmark, nine (24.3 %) in Germany and two (8.7 %) in Ireland. Denmark detected no resistance to gentamicin whilst one isolate and two isolates (2.7 % and 8.7 %, respectively) were resistant to gentamicin in Germany and Ireland, respectively.



Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Gent	amicin	Nalidia	kic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Belgium	262	60.7	262	1.5	262	11.5	262	1.1	262	0.4	262	1.1	262	60.3	262	45.4
Czech Republic	33	42.4	33	3.0	33	12.1	33	6.1	33	0	33	6.1	33	51.5	_	_
Denmark ¹	41	56.1	41	0	41	2.4	41	0	41	0	41	0	41	61.0	41	51.2
Estonia	22	4.5	22	0	22	4.5	22	0	22	0	22	0	22	13.6	22	9.1
Germany	163	49.1	163	0.6	163	14.1	163	6.1	163	0.6	163	5.5	163	50.3	163	49.1
Hungary	16	68.8	16	0	16	31.3	16	12.5	16	0	16	12.5	16	56.3	16	93.8
Ireland	69	53.6	69	0	69	17.4	69	1.4	69	4.3	69	1.4	69	69.6	69	66.7
Italy	85	41.2	85	0	85	10.6	85	1.2	85	2.4	85	1.2	85	50.6	85	58.8
Latvia	13	0	13	0	13	0	13	0	13	0	13	0	13	0	13	0
Netherlands	52	28.8	52	0	52	1.9	52	3.8	51	0	52	3.8	52	42.3	52	34.6
Poland	31	22.6	22	0	31	9.7	31	16.1	31	0	11	9.1	31	16.1	31	32.3
Romania	125	37.6	125	1.6	125	16.0	125	10.4	125	4.8	125	11.2	125	60.8	125	54.4
Spain	34	58.8	_	_	18	44.4	34	97.1	32	31.3	34	11.8	_	_	18	66.7
Total (13 MSs)	946	47.5	903	0.9	930	12.6	946	7.6	943	2.4	926	4.2	912	53.5	897	49.2

Table SA12. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella spp</u>. isolates from <u>meat from pigs</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

MS: Member State; MIC: minimum inhibitory concentration; N: total number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

1. Denmark reported only monophasic S. Typhimurium and S. Typhimurium isolates.



Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidix	kic acid	Sulfon	amides	Tetrac	yclines
Country	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Belgium	105	73.3	105	1.0	105	13.3	105	1.0	105	0	105	1.0	105	69.5	105	52.4
Denmark	18	27.8	18	0	18	0	18	0	18	0	18	0	18	38.9	18	11.1
Germany	58	51.7	58	0	58	19.0	58	8.6	58	0	58	8.6	58	56.9	58	56.9
Hungary	10	80.0	10	0	10	40.0	10	10.0	10	0	10	10.0	10	60.0	10	100
Ireland	22	63.6	22	0	22	36.4	22	4.5	22	0	22	4.5	22	72.7	22	63.6
Italy	18	100	18	0	18	27.8	18	0	18	0	18	0	18	100	18	100
Latvia	13	0	13	0	13	0	13	0	13	0	13	0	13	0	13	0
Netherlands	16	50.0	16	0	16	6.3	16	6.3	16	0	16	6.3	16	50.0	16	50.0
Poland	11	45.5	11	0	11	18.2	11	9.1	11	0	11	9.1	11	45.5	11	63.6
Romania	43	62.8	43	2.3	43	34.9	43	18.6	43	2.3	43	14.0	42	66.7	42	71.4
Total (10 MSs)	314	61.1	314	0.6	314	19.1	314	5.7	314	0.3	314	5.1	313	62.0	313	56.5

Table SA13. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>S. Typhimurium</u> isolates from <u>meat from pigs</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values



Multi-resistance among Salmonella spp. isolates from meat from pigs

In 2012, seven MSs provided isolate-based data concerning resistance in *Salmonella* spp. in meat from pigs. In most cases, complete susceptibility in the isolates tested ranged between 20 % and 40 % and multiresistance was typically high to very high. Contrastingly, complete susceptibility reached above 85 % and multiresistance was lower than 15 % in Estonia, although these figures were assessed on an isolate sample of small size (Table SA14). The frequency distributions (Figure SA8) showed similarities among the multiresistance recorded in three reporting MSs, with some isolates showing reduced susceptibility to up to six or seven different substances, while Estonia recorded multi-resistance to four classes at a maximum. Very few isolates were resistant to both ciprofloxacin and cefotaxime in a single MS (Table SA14).

Table SA14. Complete susceptibility, multi-resistance and index of diversity in <u>Salmonella spp.</u> from <u>meat from pigs</u> in MSs reporting isolate-based data, 2012

Country	Suscep	otible to all	Multi-r	esistant	Index of	Co-resistant to Cip and Ctx			
	n	%	n	%	diversity	n	%		
Czech Republic (N=33)	12	36.4	14	42.4	0.464	0 (0)	0 (0)		
Denmark ¹ (N=41)	12	29.3	24	58.5	0.315	0 (0)	0 (0)		
Estonia (N=22)	19	86.4	3	13.6	0.189	0 (0)	0 (0)		
Germany (N=163)	62	38.0	79	48.5	0.46	0 (0)	0 (0)		
Ireland (N=69)	15	21.7	41	59.4	0.545	0 (0)	0 (0)		
Italy (N=85)	32	37.6	43	50.6	0.5	0 (0)	0 (0)		
Romania (N=125)	30	24.0	70	56.0	0.427	1 (0)	0.8 (0)		

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; n: number of isolates.

Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

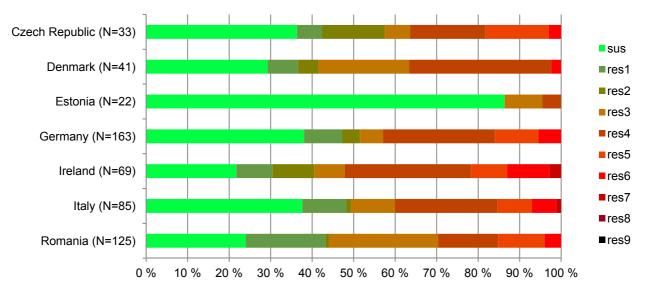
Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of *Salmonella* isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip:>1 mg/L).

1. Denmark reported only monophasic S. Typhimurium and S. Typhimurium isolates.

Figure SA8. Frequency distribution of <u>Salmonella spp.</u> in <u>meat from pigs</u> completely susceptible or resistant to one or up to nine antimicrobials, in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*. Note: Denmark reported only monophasic *S*. Typhimurium and *S*. Typhimurium isolates.



3.4.1.3. Meat from bovine animals

Five MSs reported quantitative MIC data for *Salmonella* spp. from bovine meat in 2012. Table SA15 presents the level of resistance to selected antimicrobials for *Salmonella* spp. isolates.

Representative sampling and monitoring

Monitoring and surveillance programmes for *Salmonella* spp. in meat from bovine animals are in place in Germany, Ireland, Italy, the Netherlands and Spain, while surveillance of HACCP and own checks submissions takes place in Ireland. Sample types collected by MSs at slaughterhouses consisted of carcase swabs. Germany, Ireland, the Netherlands and Spain tested meat, without details about sampling stage. Italy tested meat preparations at retail outlets.

Resistance levels in Salmonella spp. isolates from meat from bovine animals

Among the five reporting MSs, *Salmonella* spp. isolated from meat from bovine animals generally displayed high levels of resistance to ampicillin, sulfonamides and tetracyclines, varying from 20.8 % to 78.6 % for ampicillin, 25.0 % to 55.6 % for sulfonamides and 10.0 % to 61.5 % for tetracyclines. Chloramphenicol resistance was low, at 9.9 %, at MS group level, and ranged from 0 % to 25.0 % across the reporting MSs, with the Netherlands and Spain detecting no resistance. Gentamicin resistance was only detected in Spain (57.1 %). Resistance to either ciprofloxacin or nalidixic acid was generally similar among the reporting MSs, ranging from no resistance detected to low/moderate levels, with the notable exception of Spain reporting high resistance to nalidixic acid (28.6 %) and extremely high resistance to ciprofloxacin (100 %). The occurrence of resistance to cefotaxime was generally not detected in reporting MSs, as only the Netherlands reported 5.6 % resistance.

Multi-resistance among Salmonella spp. isolates from meat from bovine animals

As only three MSs reported resistance for isolate-based data on 10 or more isolates of *Salmonella* spp. from meat from bovine animals, the corresponding multi-resistance analysis is not presented in this report.



 Table SA15. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among

 Salmonella spp. isolates from meat from bovine animals
 in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	Ampicillin Co		Cefotaxime		Chloramphenicol		Ciprofloxacin		Gentamicin		kic acid	Sulfonamides		Tetrac	yclines
Country	Ν	% Res	Ν	% Res	N	% Res	N	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res
Germany	16	50.0	16	0	16	25.0	16	0	16	0	16	0	16	50.0	16	43.8
Ireland	24	20.8	24	0	24	12.5	24	0	24	0	24	0	24	25.0	24	33.3
Italy	13	30.8	13	0	13	7.7	13	7.7	13	0	13	7.7	13	38.5	13	61.5
Netherlands	18	33.3	18	5.6	18	0	18	11.1	18	0	18	11.1	18	55.6	18	44.4
Spain	14	78.6	_	_	10	0	14	100	14	57.1	14	28.6	_	_	10	10.0
Total (5 MSs)	85	40.0	71	1.4	81	9.9	85	20.0	85	9.4	85	8.2	71	40.8	81	39.5



3.4.2. Antimicrobial resistance in *Salmonella* isolates from animals

3.4.2.1. Domestic fowl (Gallus gallus)

In 2012, 16 MSs submitted quantitative antimicrobial susceptibility data for *Salmonella* spp. from *Gallus gallus*. As in previous years, an overview of all data relating to *Gallus gallus* including breeding and laying hens and broiler flocks, as well as unspecified flocks of *Gallus gallus*, is presented. However, as in 2011, data from flocks of broilers, laying hens and breeders are also presented separately.

Representative sampling and monitoring

In the vast majority of MSs, isolates for antimicrobial resistance testing are obtained from national surveillance and control programmes carried out according to the EU legislation. In Latvia, isolates were obtained from faecal samples from broilers before slaughter and from laying hens on the farm.

Resistance levels in Salmonella spp. isolates from Gallus gallus

Table SA16 shows the level of resistance to antimicrobials among isolates of *Salmonella* spp. from *Gallus gallus* in 2012. There was generally high resistance reported to ampicillin, sulfonamides and tetracyclines in the reporting MS group, although the levels of resistance varied between 0 % and 44.4 % for ampicillin, between 0 % and 57.5 % for sulfonamides and between 6.3 % and 55.9 % for tetracyclines across the 16 reporting countries. Resistance to chloramphenicol and gentamicin was mostly either not detected or reported at low levels among the reporting MSs. The levels of resistance to ciprofloxacin and nalidixic acid were mostly moderate to very high within individual MSs, although four MSs did not detect resistance or reported low resistance to these compounds. This considerable disparity in resistance to ciprofloxacin and nalidixic acid among *Salmonella* isolates, from different MSs, may reflect the variability of serovars of *Salmonella* spp. included in the analyses of the different MSs. Cefotaxime resistance was reported by 11 of the 16 reporting MSs at low levels, with the exception of Belgium which reported for the first time in 2012 susceptibility results at a level of 18.1 %.

Resistance levels in Salmonella spp. isolates from broilers

Thirteen MSs reported quantitative data on resistance among isolates of *Salmonella* spp. from broiler flocks in 2012 (Table SA17). Resistance to ampicillin and tetracyclines varied from low to high levels among the reporting MSs, while resistance to sulfonamides was moderate to high in all reporting MSs. For chloramphenicol and gentamicin, absence or low levels of resistance were mostly reported, as only Romania reported moderate resistance to these substances. Resistance to ciprofloxacin and nalidixic acid was generally high in the reporting MSs, although Denmark, Ireland and the United Kingdom reported either no resistance or low levels of resistance to both compounds. Cefotaxime resistance was either not detected or observed at very low and low levels, with the exception of Italy which reported a moderate resistance at 13.3 %. In general, the levels of resistance in broiler flocks were slightly higher than those reported when all *Gallus gallus* were considered, as this category includes data from broilers, laying hens and breeders.

Resistance levels in Salmonella spp. isolates from laying hens

Twelve MSs reported quantitative data on resistance among isolates of *Salmonella* spp. from laying hens in 2012 (Table SA18). Resistance to ampicillin, sulfonamides and tetracyclines was generally reported at low to moderate levels across the reporting MSs, with the exception of Romania reporting high resistance to these substances. Resistance to chloramphenicol and gentamicin was either not detected or recorded at low levels, ranging from 0 % to 3.7 % and from 0 % to 6.2 %, respectively. Portugal reported resistance only to ciprofloxacin (9.4 %) and nalidixic acid (12.5 %). Resistance to ciprofloxacin and nalidixic acid was generally reported at moderate to high levels, while Germany, the Netherlands and the United Kingdom recorded low resistance to these substances. Latvia observed no resistance to both compounds. Cefotaxime resistance was observed only by Austria and Italy at low levels of 1.6 % and 1.2 %. Generally, resistance in laying hens was similar to or lower than that reported in broilers.



Resistance levels in Salmonella spp. isolates from breeding flocks

Quantitative data on isolates of *Salmonella* spp. from breeding flocks of *Gallus gallus* were reported by four MSs in 2012 (Table SA19). The breeder categories reported were: unspecified breeding flocks, unspecified grandparent breeding flocks, parent breeding flocks for the broiler production line and elite breeding flocks for egg production line. The levels of resistance among isolates from breeding flocks of *Gallus gallus* were generally higher than those observed in laying hens. The occurrence of resistance to ampicillin, sulfonamides and tetracyclines was high across the reporting MSs (22.7 %, 20.9 % and 32.5 %, respectively). High levels of resistance to ciprofloxacin and nalidixic acid were observed at the MS group level (25.5 % and 22.7 % respectively), but these levels were at almost half of the level reported for broilers. For these antimicrobials, reported resistance levels ranged from 7.4 % to 53.1 %. The Czech Republic and Poland reported no resistance to cefotaxime, chloramphenicol and gentamicin. The overall occurrence of gentamicin resistance, considering all reporting MSs, was 5.5 %. Cefotaxime resistance was reported by two MSs with an overall resistance at MS group level of 1.8 %.



Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Gent	amicin	Nalidix	kic acid	Sulfor	namides	Tetrac	cyclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	176	4.0	176	0.6	176	1.1	176	18.2	176	0.6	176	16.5	176	17.0	176	22.7
Belgium	664	44.4	664	18.1	664	7.1	664	39.0	664	3.5	664	37.0	664	37.0	664	24.4
Czech Republic	386	4.7	386	0.3	386	0	386	19.4	386	1.3	386	19.2	386	18.1	-	-
Denmark	28	32.1	28	0	28	0	28	0	28	0	28	0	28	28.6	28	32.1
Germany	238	6.7	238	0.8	238	2.1	238	14.3	238	0	238	22.7	238	15.5	238	13.0
Hungary	261	9.2	261	0.4	261	1.1	261	69.0	261	1.9	261	67.4	261	57.5	261	55.9
Ireland	38	10.5	38	0	38	0	38	2.6	38	0	38	2.6	38	10.5	38	7.9
Italy	328	21.3	328	5.2	328	4.0	328	23.8	328	2.4	328	23.5	328	18.0	328	25.3
Latvia	14	7.1	14	0	14	0	14	0	14	0	14	0	14	0	14	7.1
Netherlands	192	22.9	192	4.2	192	3.1	192	18.2	192	5.2	192	16.1	192	15.1	192	9.4
Poland	739	7.8	739	0.7	739	2.2	738	39.2	739	0	739	35.0	734	12.1	739	8.4
Portugal	79	6.3	174	2.3	174	2.9	174	23.6	174	0	174	21.8	174	10.9	174	6.3
Romania	964	41.2	964	5.3	964	10.9	964	68.4	964	15.7	964	58.6	964	52.9	964	47.4
Slovakia	85	0	85	0	85	0	85	30.6	85	0	85	30.6	85	25.9	85	25.9
Spain	179	9.5	179	0.6	179	1.1	179	22.3	179	3.4	179	19.0	179	8.4	179	12.3
United Kingdom	236	4.7	236	0	236	2.1	236	2.1	236	4.2	236	2.1	236	17.4	236	21.6
Total (16 MSs)	4,607	21.2	4,702	4.5	4,702	4.4	4,701	37.3	4,702	4.7	4,702	34.3	4,697	28.3	4,316	25.9

 Table SA16. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among

 Salmonella spp.
 isolates from Gallus gallus in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values



Country	Amp	icillin	Cefo	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidix	ic acid	Sulfor	amides	Tetrac	yclines
Country	N	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res
Austria	113	3.5	113	0	113	0	113	22.1	113	0	113	22.1	113	18.6	113	23.9
Czech Republic	351	4.0	351	0.3	351	0	351	20.8	351	1.4	351	20.5	351	19.4	_	_
Denmark	24	29.2	24	0	24	0	24	0	24	0	24	0	24	29.2	24	29.2
Hungary	175	6.9	175	0.6	175	1.1	175	90.3	175	1.1	175	90.3	175	77.7	175	73.7
Ireland	38	10.5	38	0	38	0	38	2.6	38	0	38	2.6	38	10.5	38	7.9
Italy	105	37.1	105	13.3	105	4.8	105	32.4	105	6.7	105	30.5	105	31.4	105	38.1
Netherlands	130	27.7	130	6.2	130	3.1	130	24.6	130	7.7	130	21.5	130	16.9	130	6.2
Poland	189	6.3	189	0.5	189	0.5	189	40.7	189	0	189	36.5	189	14.8	189	10.1
Portugal	27	18.5	122	3.3	122	4.1	122	27.9	122	0	122	26.2	122	15.6	122	9.0
Romania	784	43.9	784	6.4	784	12.2	784	73.0	784	17.5	784	62.0	784	57.4	784	48.7
Slovakia	55	0	55	0	55	0	55	41.8	55	0	55	41.8	55	34.5	55	34.5
Spain	29	34.5	29	3.4	29	3.4	29	65.5	29	13.8	29	62.1	29	20.7	29	31.0
United Kingdom	170	5.3	170	0	170	2.9	170	2.4	170	5.9	170	2.4	170	21.8	170	25.3
Total (13 MSs)	2,190	22.6	2,285	3.5	2,285	5.2	2,285	46.0	2,285	7.7	2,285	41.5	2,285	37.2	1,934	36.0

 Table SA17. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among

 Salmonella spp.
 isolates from broilers in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values



Country	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidia	kic acid	Sulfor	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	63	4.8	63	1.6	63	3.2	63	11.1	63	1.6	63	6.3	63	14.3	63	20.6
Germany	51	3.9	51	0	51	2.0	51	2.0	51	0	51	2.0	51	9.8	51	2.0
Hungary	86	14.0	86	0	86	1.2	86	25.6	86	3.5	86	20.9	86	16.3	86	19.8
Italy	161	15.5	161	1.2	161	3.1	161	18.0	161	0	161	18.0	161	10.6	161	16.8
Latvia	14	7.1	14	0	14	0	14	0	14	0	14	0	14	0	14	7.1
Netherlands	54	3.7	54	0	54	3.7	54	5.6	54	0	54	5.6	54	1.9	54	5.6
Poland	132	5.3	132	0	132	0	132	25.8	132	0	132	22.0	132	5.3	132	0.8
Portugal	32	0	32	0	32	0	32	9.4	32	0	32	12.5	32	0	32	0
Romania	145	22.1	145	0	145	2.8	145	46.2	145	6.2	145	43.4	145	30.3	145	37.2
Slovakia	29	0	29	0	29	0	29	10.3	29	0	29	10.3	29	10.3	29	10.3
Spain	150	4.7	150	0	150	0.7	150	14.0	150	1.3	150	10.7	150	6.0	150	8.7
United Kingdom	66	3.0	66	0	66	0	66	1.5	66	0	66	1.5	66	6.1	66	12.1
Total (12 MSs)	983	9.5	983	0.3	983	1.6	983	19.4	983	1.5	983	17.4	983	11.5	983	14.3

Table SA18. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella spp.</u> isolates from <u>laying hens</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

Table SA19. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella spp.</u> isolates from <u>breeders</u> of Gallus gallus in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	oicillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidi	xic acid	Sulfor	namides	Tetrac	yclines
Country	N	% Res	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Czech Republic	27	11.1	27	0	27	0	27	7.4	27	0	27	7.4	27	7.4	-	-
Italy	36	8.3	36	2.8	36	2.8	36	16.7	36	2.8	36	19.4	36	13.9	36	22.2
Poland	15	0	15	0	15	0	15	20.0	15	0	15	20.0	15	0	15	0
Romania	32	59.4	32	3.1	32	15.6	32	53.1	32	15.6	32	40.6	32	50.0	32	59.4
Total (4 MSs)	110	22.7	110	1.8	110	5.5	110	25.5	110	5.5	110	22.7	110	20.9	83	32.5



Resistance levels in Salmonella Enteritidis isolates from Gallus gallus

Susceptibility data on *S*. Enteritidis isolates from *Gallus gallus* were reported by 13 MSs in 2012 (Table SA20). Resistance to ampicillin, sulfonamides and tetracyclines was generally not detected or recorded at low levels in most MSs, with the exceptions of Belgium and Romania which reported moderate to high levels of resistance to these substances. Chloramphenicol and gentamicin resistance was often rarely detected or recorded at low levels; resistance to these substances in the reporting MS group equalled 0.8 % and 0.4 %, respectively. In contrast to the other antimicrobials tested, the occurrence of ciprofloxacin and nalidixic acid resistance varied substantially, from 0 % to 88.9 % for ciprofloxacin, and, from 0 % to 92.6 % for nalidixic acid, among three distinct groups of reporting MSs. While seven MSs either did not detect any resistance or recorded low resistance. Once more, the levels of ciprofloxacin and nalidixic acid resistance or seconded low resistance. Once more, the levels of ciprofloxacin and nalidixic acid resistance on the sequence. Note the other and portugal in 2012, making the overall resistance at MS group level low at 1.3 %. Interestingly, Germany did not observe any resistance to the substances tested in the *S*. Enteritidis isolates from *Gallus gallus*.

Resistance levels in Salmonella Enteritidis isolates from broilers

Six MSs reported quantitative data on isolates of *S*. Enteritidis from broiler flocks in 2012 (Table SA20). Among the six reporting MSs, the overall resistance to ampicillin was low, at 3.2 %, and no resistance was reported by the Czech Republic and Slovakia. The overall levels of resistance to sulfonamides and tetracyclines were 2.7 % and 3.0 %, respectively. Austria was the only country to report no resistance to ciprofloxacin and nalidixic acid among isolates of *S*. Enteritidis from broiler flocks. Portugal was the only country to observe resistance to cefotaxime, at the low level of 5.3 %.

Resistance levels in Salmonella Enteritidis isolates from laying hens

Quantitative data on isolates of S. Enteritidis from laying hens were reported by 10 MSs in 2012 (Table SA20). The levels of resistance among isolates from laying hens were generally either similar (Poland, Slovakia) or higher (Romania) than those observed in broiler flocks, when considering the MSs reporting for both populations. Austria, Germany and Slovakia did not observe resistance to any of the antimicrobials tested. The occurrence of resistance to ampicillin, sulfonamides and tetracyclines was low across the reporting MSs (4.1 %, 4.7 % and 3.6 %, respectively). Only four MSs observed resistance to sulfonamides and tetracyclines with values varying from 2.3 % to 18.2 % and from 2.3 % to 15.2 %, respectively. Moderate levels of resistance to ciprofloxacin and nalidixic acid were observed at the MSs group level (18.5 % and 17.1 %, respectively). For these antimicrobials, reported resistance levels reported ranged from 0 % to 36.3 %. No resistance was observed to cefotaxime and chloramphenicol. Hungary and Romania were the only countries to report resistance to gentamicin at low levels of 4.0 % and 1.5 %, respectively.



 Table SA20. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among

 S. Enteritidis isolates from Gallus gallus in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Gent	amicin	Nalidiz	kic acid	Sulfor	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res
All Gallus gallus				· · ·	<u>.</u>	· · · ·		· · ·	. <u> </u>	•	·	· · · ·	· · · · ·	· · · ·	·	
Austria	36	2.8	36	0	36	0	36	0	36	0	36	0	36	0	36	0
Belgium	81	23.5	81	14.8	81	6.2	81	17.3	81	2.5	81	16.0	81	16.0	81	11.1
Czech Republic	251	0	251	0	251	0	251	1.2	251	0	251	0.8	251	0.4	_	-
Germany	80	0	80	0	80	0	80	0	80	0	80	0	80	0	80	0
Hungary	26	3.8	26	0	26	0	26	0	26	3.8	26	0	26	3.8	26	0
Italy	31	3.2	31	0	31	0	31	12.9	31	0	31	12.9	31	3.2	31	6.5
Latvia	11	9.1	11	0	11	0	11	0	11	0	11	0	11	0	11	9.1
Netherlands	38	2.6	38	0	38	0	38	7.9	38	0	38	7.9	38	0	38	0
Poland	496	6.9	496	0.6	496	0.4	496	43.3	496	0	496	39.3	496	3.8	496	0.8
Portugal	27	7.4	27	3.7	27	3.7	27	88.9	27	0	27	92.6	27	3.7	27	7.4
Romania	76	7.9	76	0	76	2.6	76	27.6	76	2.6	76	27.6	76	22.4	76	18.4
Slovakia	47	0	47	0	47	0	47	2.1	47	0	47	2.1	47	0	47	0
Spain	45	4.4	45	0	45	0	45	28.9	45	0	45	28.9	45	2.2	45	2.2
Total (13 MSs)	1,245	5.5	1,245	1.3	1,245	0.8	1,245	23.9	1,245	0.4	1,245	22.2	1,245	4.3	994	3.3

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Note: Data reported under 'All Gallus gallus' include data which have been reported by production level.

Table continued overleaf.



Table SA20 (continued.) Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>S. Enteritidis</u> isolates from <u>Gallus gallus</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	oicillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Gent	amicin	Nalidi	xic acid	Sulfor	namides	Tetrac	yclines
Country	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Broiler flocks																
Austria	21	4.8	21	0	21	0	21	0	21	0	21	0	21	0	21	0
Czech Republic	236	0	236	0	236	0	236	1.3	236	0	236	0.8	236	0.4	-	-
Poland	131	6.1	131	0	131	0	131	39.7	131	0	131	35.9	131	3.8	131	0
Portugal	19	10.5	19	5.3	19	5.3	19	100	19	0	19	100	19	5.3	19	10.5
Romania	10	30.0	10	0	10	20.0	10	50.0	10	10.0	10	50.0	10	50.0	10	40.0
Slovakia	21	0	21	0	21	0	21	4.8	21	0	21	4.8	21	0	21	0
Total (6 MSs)	438	3.2	438	0.2	438	0.7	438	18.3	438	0.2	438	16.9	438	2.7	202	3.0
Laying hen flocks																
Austria	15	0	15	0	15	0	15	0	15	0	15	0	15	0	15	0
Germany	21	0	21	0	21	0	21	0	21	0	21	0	21	0	21	0
Hungary	25	4.0	25	0	25	0	25	0	25	4.0	25	0	25	4.0	25	0
Italy	28	3.6	28	0	28	0	28	14.3	28	0	28	14.3	28	0	28	3.6
Latvia	11	9.1	11	0	11	0	11	0	11	0	11	0	11	0	11	9.1
Netherlands	38	2.6	38	0	38	0	38	7.9	38	0	38	7.9	38	0	38	0
Poland	91	6.6	91	0	91	0	91	36.3	91	0	91	30.8	91	3.3	91	0
Romania	66	4.5	66	0	66	0	66	24.2	66	1.5	66	24.2	66	18.2	66	15.2
Slovakia	25	0	25	0	25	0	25	0	25	0	25	0	25	0	25	0
Spain	43	4.7	43	0	43	0	43	25.6	43	0	43	25.6	43	2.3	43	2.3
Total (10 MSs)	363	4.1	363	0	363	0	363	18.5	363	0.6	363	17.1	363	4.7	363	3.6

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Note: Data reported under 'All Gallus gallus' include data which have been reported by production level.



Resistance levels in *Salmonella* Typhimurium isolates from *Gallus gallus*

Five MSs reported quantitative MIC antimicrobial susceptibility data for *S*. Typhimurium isolates from *Gallus gallus* in 2012 (Table SA21). Only three MSs provided production level information with these data: in Austria and Hungary, two isolates of *S*. Typhimurium were from broiler flocks and eight were from laying hens. Germany reported 13 isolates from broiler flocks.

The overall level of resistance to ampicillin, sulfonamides and tetracyclines, in the reporting MS group, was higher among *S*. Typhimurium isolates from *Gallus gallus* (39.5 %, 46.0 % and 35.5 %, respectively), than in *S*. Enteritidis isolates and all *Salmonella* spp. isolates as a whole. All MSs detected resistance to ampicillin, sulfonamides and tetracyclines generally at high levels, and the occurrence of resistance ranged from 7.1 % to 86.7 %, 10.0 % to 86.7 % and 7.1 % to 80.0 %, respectively. Among individual MSs, the level of ciprofloxacin resistance varied from 0 % in Austria, Germany and Hungary, to 66.7 % in Poland. Similarly, the level of resistance to nalidixic acid among individual MSs varied from 0 % in Austria and Hungary, to 60.0 % in Poland. Austria and Belgium reported resistance to gentamicin at low levels of 10.0 % and 2.1 %, respectively. Cefotaxime resistance was observed only by Belgium (10.6 %).

Resistance levels in Salmonella Infantis isolates from Gallus gallus

Nine MSs reported quantitative MIC antimicrobial susceptibility data for *S*. Infantis isolates from *Gallus gallus* in 2012 (Table SA22). Only Belgium did not provide production level information with these data.

The resistance to sulfonamides and tetracyclines varied markedly and were generally high to extremely high among the reporting MSs. A similar situation occurred regarding the resistance to ciprofloxacin and nalidixic acid. When reported, resistance to ampicillin varied between low, moderate and high levels. Cefotaxime resistance was not detected or recorded at very low levels by most reporting MSs and was observed at a high level by Italy (27.6 %), at a moderate level by Belgium (14.3 %) and at a low level by Romania (7.6 %).

Resistance levels in Salmonella Kentucky isolates from Gallus gallus

Five MSs reported quantitative MIC antimicrobial susceptibility data for S. Kentucky isolates from *Gallus gallus* in 2012 (Table SA23). All MSs provided production level information with these data.

Generally, MSs reported resistance to ampicillin, sulfonamides and tetracyclines at high to extremely high levels. Conversely, Ireland recorded low to moderate resistance to these substances. The occurrence of resistance to ciprofloxacin and nalidixic acid was generally extremely high in the reporting MSs, with the exception of Ireland detecting 3 % resistance. Resistance to chloramphenicol was low to moderate. Ireland and Italy were the only MSs not reporting resistance to gentamicin, while the three remaining MSs reported high to extremely high levels of resistance. Cefotaxime resistance was observed only in Italy and Romania at around the 10 % level, making the overall resistance at MS group level 6.3 %.



 Table SA21. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among

 S. Typhimurium isolates from Gallus gallus in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidix	cic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	10	20.0	10	0	10	20.0	10	0	10	10.0	10	0	10	30.0	10	30.0
Belgium	47	63.8	47	10.6	47	14.9	47	25.5	47	2.1	47	23.4	47	68.1	47	53.2
Germany	42	7.1	42	0	42	7.1	42	0	42	0	42	47.6	42	19.0	42	7.1
Hungary	10	10.0	10	0	10	10.0	10	0	10	0	10	0	10	10.0	10	10.0
Poland	15	86.7	15	0	15	80.0	15	66.7	15	0	15	60.0	15	86.7	15	80.0
Total (5 MSs)	124	39.5	124	4.0	124	20.2	124	17.7	124	1.6	124	32.3	124	46.0	124	35.5

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

Table SA22. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>S. Infantis</u> isolates from <u>Gallus gallus</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	oicillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidi	kic acid	Sulfor	amides	Tetrac	yclines
Country	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	25	0	25	0	25	0	25	96.0	25	0	25	96.0	25	96.0	25	96.0
Belgium	28	21.4	28	14.3	28	0	28	17.9	28	10.7	28	17.9	28	21.4	28	7.1
Czech Republic	52	15.4	52	0	52	0	52	96.2	52	0	52	96.2	52	96.2	_	-
Hungary	172	4.7	172	0.6	172	1.2	172	94.2	172	0.6	172	95.3	172	83.7	172	79.1
Italy	29	41.4	29	27.6	29	0	29	62.1	29	3.4	29	65.5	29	55.2	29	55.2
Poland	118	3.4	118	0	118	0	118	44.1	118	0	118	44.1	118	44.1	118	39.0
Romania	303	23.4	303	7.6	303	9.2	303	98.3	303	6.3	303	97.0	303	82.8	303	82.8
Slovakia	25	0	25	0	25	0	25	88.0	25	0	25	88.0	25	88.0	25	88.0
Spain	13	0	13	0	13	0	13	7.7	13	0	13	7.7	13	7.7	13	0
Total (9 MSs)	765	14.2	765	4.7	765	3.9	765	82.6	765	3.1	765	82.5	765	74.0	713	69.7



 Table SA23. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among

 <u>S. Kentucky</u> isolates from <u>Gallus gallus</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidi	kic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res
Czech Republic	11	63.6	11	0	11	0	11	54.5	11	45.5	11	54.5	11	54.5	-	-
Ireland	33	12.1	33	0	33	3.0	33	3.0	33	0	33	3.0	33	9.1	33	3.0
Italy	30	56.7	30	10.0	30	6.7	30	93.3	30	0	30	93.3	30	16.7	30	40.0
Romania	74	97.3	74	9.5	74	5.4	74	100	74	94.6	74	95.9	74	95.9	74	94.6
Spain	10	40.0	10	0	10	10.0	10	60.0	10	50.0	10	60.0	10	50.0	10	40.0
Total (5 MSs)	158	65.8	158	6.3	158	5.1	158	72.8	158	50.6	158	70.9	158	57.0	147	59.2

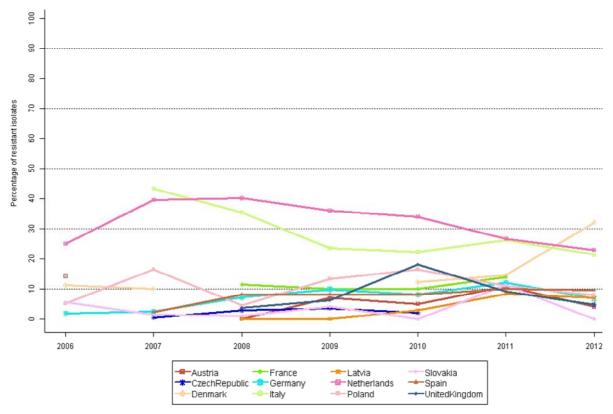


Temporal trends in resistance among *Salmonella* spp. isolates from *Gallus gallus*

Figures SA9 to SA11 indicate how the level of resistance to selected antimicrobials in *Salmonella* spp. isolates from *Gallus gallus* has changed over the period 2006–2012 in the MSs. It is important to note that because some antimicrobial resistance is associated with particular serovars or clones within serovars, fluctuations in the occurrence of resistance in *Salmonella* spp. isolates within a country may result from changes in the proportions of different *Salmonella* serovars which contribute to the total numbers of *Salmonella* spp. isolates tested.

For the majority of MSs, resistance to ampicillin decreased slightly between 2011 and 2012, although the slight increase observed in Denmark between 2010 and 2011 was continued in 2012. Across the seven years of data, levels of resistance to ampicillin remained broadly constant for most of the reporting MSs, while decreasing trends were observed in Italy and the Netherlands (Figure SA9). The level of resistance to cefotaxime in *Salmonella* spp. was generally low, very low or absent in reporting MSs between 2006 and 2012. A statistically significant decreasing trend, for five or more years, was observed in Italy and Spain (Figure SA10). Statistically significant increasing trends in resistance to ciprofloxacin and nalidixic acid were registered in three MSs for five or more years over the 2006–2012 period. Spain observed a statistically significant decreasing trends to both antimicrobials, while Italy and the Netherlands observed a significant decrease in resistance to ciprofloxacin only.

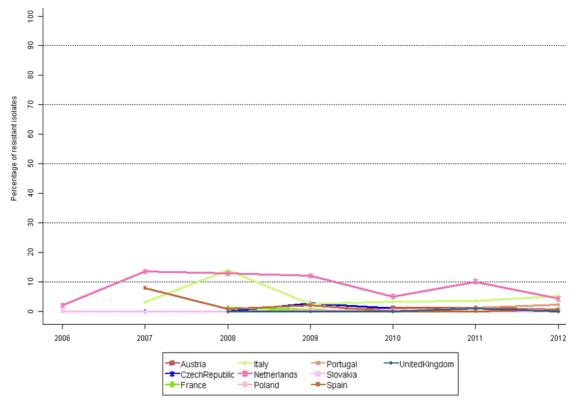
Figure SA9. Trends in <u>ampicillin</u> resistance in tested Salmonella spp. isolates from <u>Gallus gallus</u> in reporting MSs, 2006-2012, quantitative data



MS: Member State.

Note: Statistically significant increasing or decreasing trends for five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in Germany (\uparrow), Italy (\downarrow), the Netherlands (\downarrow) and Poland (\uparrow).

Figure SA10. Trends in <u>cefotaxime</u> resistance in tested Salmonella spp. isolates from <u>Gallus gallus</u> <i>in reporting MSs, 2006-2012, quantitative data

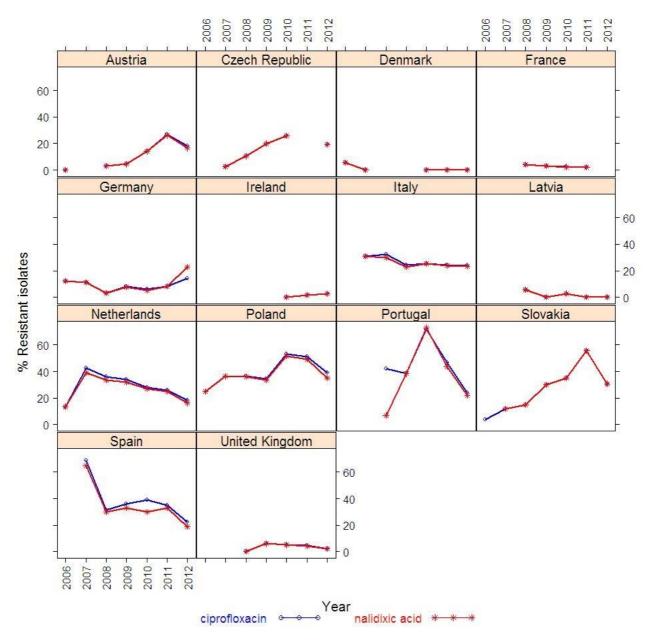


MS: Member State.

Note: A statistically significant decreasing trend for five or more years, as tested by logistic regression model ($p \le 0.05$), was observed for Italy (\downarrow) and Spain (\downarrow).



Figure SA11. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in tested <u>Salmonella spp.</u> isolates from <u>Gallus gallus</u> in reporting MSs, 2006-2012, quantitative data



MS: Member State.

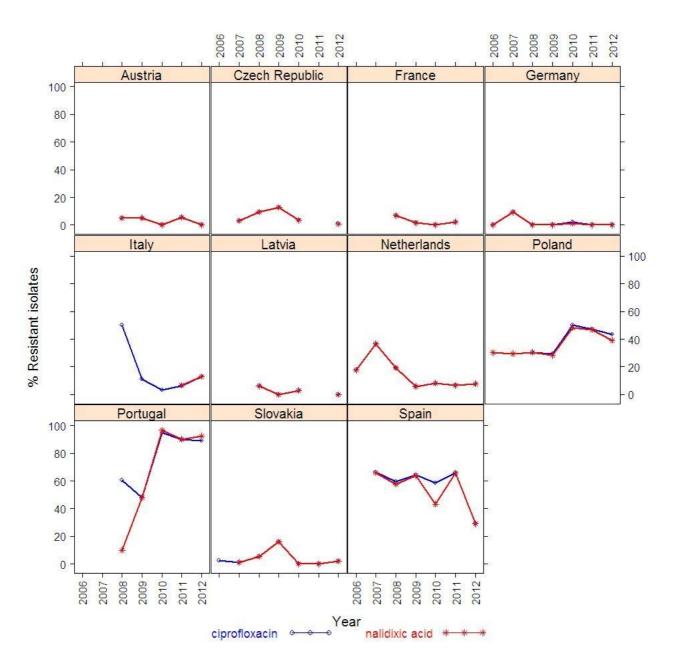
Note: A statistically significant increasing trend over five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed in Austria (↑), and Slovakia (↑) for both ciprofloxacin and nalidixic acid and in Poland (↑) for ciprofloxacin. A statistically significant decreasing trend was observed for ciprofloxacin in Italy (↓) and the Netherlands (↓), and for both ciprofloxacin and nalidixic acid in Spain (↓).



Temporal trends in resistance among *S*. Enteritidis isolates from *Gallus gallus*

Figure SA12 indicates how the level of resistance to selected antimicrobials in *S*. Enteritidis isolates from *Gallus gallus* has changed over the period 2006–2012 in the MSs. Most of the reporting MSs observed a similarity in their trends in resistance to ciprofloxacin and nalidixic acid among isolates of *S*. Enteritidis from *Gallus gallus* over the 2006-2012 period. However, statistically significant decreasing trends were observed in Germany, the Netherlands and Spain for both substances, while a significant increasing trend was observed in Poland, also for both substances.

Figure SA12. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in tested <u>S. Enteritidis</u> isolates from <u>Gallus gallus</u> in reporting MSs, 2006-2012, quantitative data



MS: Member State.

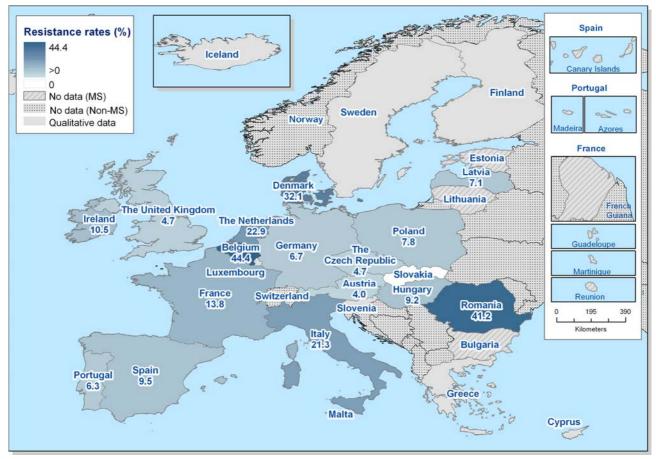
Note: A statistically significant decreasing trend for five or more years, as tested by a logistic regression model (*p* ≤ 0.05), was observed in the Czech Republic (↓), Germany (↓), the Netherlands (↓) and Spain (↓) for both ciprofloxacin and nalidixic acid. A statistically significant increasing trend was observed in Poland (↑) and Portugal (↑) for both ciprofloxacin and nalidixic acid.



Spatial distribution of resistance among Salmonella isolates from Gallus gallus

Figures SA13 to SA14 show the spatial distributions of ampicillin and nalidixic acid resistance in *Salmonella* spp. isolated from *Gallus gallus* in 2012. Figure SA13 illustrates important variability in levels of ampicillin resistance in *Salmonella* spp. across the EU and the absence of a clear spatial distribution. Figure SA14 illustrates either the continued absence or the low prevalence of resistance to nalidixic acid in *Salmonella* spp. in Northern and Western Europe, but high levels of resistance in Southern and Eastern Europe.

Figure SA13. Spatial distribution of <u>ampicillin</u> resistance among Salmonella spp. from <u>Gallus gallus</u> in countries reporting MIC data in 2012¹

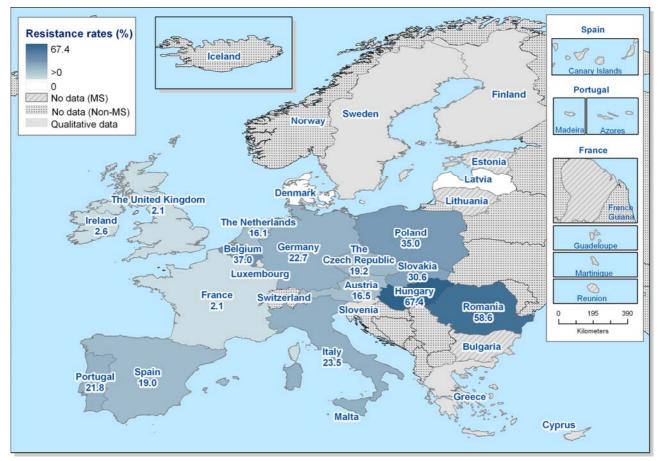


MIC: minimum inhibitory concentration; MS: Member State.

- Note:Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting either inhibition zone diameter data, MIC data for fewer than 10 isolates, or purely qualitative data (as proportion of resistant isolates).
- 1. For France, 2011 data were used.



Figure SA14. Spatial distribution of <u>nalidixic acid</u> resistance among Salmonella spp. from <u>Gallus gallus</u> in countries reporting MIC data in 2012¹



MIC: minimum inhibitory concentration; MS: Member State.

- Note:Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting either inhibition zone diameter data, MIC data for fewer than 10 isolates, or purely qualitative data (as proportion of resistant isolates).
- 1. For France, 2011 data were used.



Multi-resistance among Salmonella spp. isolates from broilers and laying hens of Gallus gallus

In 2012, 10 MSs reported isolate-based data on resistance in *Salmonella* spp. from broiler flocks, which represents 2 more reporting countries compared with 2011, while isolate-based data concerning resistance in *Salmonella* spp. from laying hen flocks were reported by 9 MSs in 2012, which accounts for 3 additional reporting countries compared with the previous year.

Complete susceptibility was typically very high (\geq 50 %) in tested isolates from laying hens and generally similar or higher than those observed in tested isolates from broilers, which varied substantially among reporting MSs from 7.4 % in Hungary to up to 81.6 % in Ireland (Tables SA24 and SA25). Similarly, multi-resistance rates in laying hens were mostly low to moderate (\leq 20 %) among reporting MSs, except in Romania, where 37.9 % of tested isolates exhibited reduced susceptibility to three or more substances, while, in broilers, multi-resistance rates were generally moderate to high, with the exception of Ireland, which reported a low level of multi-resistance, and Hungary and Romania, which recorded very high levels of multi-resistance.

Considering the multi-resistance distributions of *Salmonella* spp. isolates in broilers and laying hens (Figures SA15 and SA16), higher proportions of isolates exhibiting reduced susceptibility to an important number of different substances were generally observed in broilers compared with laying hens. While in Italy and Romania, some isolates showed reduced susceptibility to a maximum of eight and nine different substances in broilers, which were higher numbers than in certain isolates from laying hens in the same country. The opposite was true in Austria, Hungary and Spain, where the maximum numbers of substances to which a number of isolates were multi-resistant were higher in laying hens than in broilers.

Very few isolates were resistant to both ciprofloxacin and cefotaxime in *Salmonella* spp. isolates from laying hens and broilers (Tables SA24 and SA25).

Multi-resistance among S. Enteritidis isolates from broilers and laying hens of Gallus gallus

In 2012, three MSs provided isolate-based data concerning resistance in *S*. Enteritidis in broilers and six MSs provided isolate-based data concerning resistance in *S*. Enteritidis in laying hens.

Levels of complete susceptibility were commonly very high in *S*. Enteritidis isolates from broilers and laying hens and multi-resistance was either not detected or observed at low levels (Tables SA26 and SA27). An exception to this is Romania where rates of both complete susceptibility and multi-resistance were high in isolates from broilers (although assessed on a small sample of 10 isolates only) and multi-resistance was moderate in isolates from laying hens. One *S*. Enteritidis isolate from broilers exhibited multi-resistance to eight different classes.

Co-resistance to both ciprofloxacin and cefotaxime, in *S*. Enteritidis isolates, from laying hens and broilers was not observed in the reporting MSs (Tables SA26 and SA27).



Table SA24. Complete susceptibility, multi-resistance and index ofTable SA24. Complete susceptibility, multi-resistance and index ofTable SA24. Complete susceptibility, multi-resistance and index ofdiversity in Salmonella spp. from broilers in MSs reporting isolate-baseddata, 2012based

Country		eptible all		ılti- stant	Index of diversity		istant to Ind Ctx
	n	%	n	%	uiversity	n	%
Austria (N=113)	82	72.6	23	20.4	0.243	0 (0)	0 (0)
Czech Republic (N=351)	271	77.2	67	19.1	0.385	0 (0)	0 (0)
Denmark (N=24)	17	70.8	7	29.2	0	0 (0)	0 (0)
Hungary (N=175)	13	7.4	128	73.1	0.279	0 (0)	0 (0)
Ireland (N=38)	31	81.6	3	7.9	0.264	0 (0)	0 (0)
Italy (N=105)	55	52.4	43	41.0	0.67	13 (0)	12.4 (0)
Romania (N=781)	135	17.3	474	60.7	0.648	41 (2)	5.2 (0.3)
Spain (N=29)	6	20.7	10	34.5	0.394	1 (0)	3.4 (0)
United Kingdom (N=17)	5	29.4	2	11.8	0.162	0 (0)	0 (0)

Table SA25. Complete	susceptibility, multi-resistance and index o	f
diversity in Salmonella	spp. from laying hens in MSs reporting isolate)-
based data, 2012		

Country		ceptible o all		ulti- istant	Index of diversity		istant to nd Ctx
	n	%	n	%	arrenenty	n	%
Austria (N=63)	47	74.6	9	14.3	0.361	0 (0)	0 (0)
Germany (N=51)	44	86.3	1	2.0	0.221	0 (0)	0 (0)
Hungary (N=86)	59	68.6	16	18.6	0.38	0 (0)	0 (0)
Italy (N=161)	111	68.9	23	14.3	0.372	2 (0)	1.2 (0)
Romania (N=145)	72	49.7	55	37.9	0.450	0 (0)	0 (0)
Spain (N=150)	119	79.3	8	5.3	0.321	0 (0)	0 (0)
United Kingdom (N=11)	10	90.9	1	9.1	0	0 (0)	0 (0)

MS:Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella; n: number of isolates.

Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of *Salmonella* isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip:>1mg/L).



Figure SA15. Frequency distribution of <u>Salmonella spp.</u> isolates from <u>broilers</u> completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012

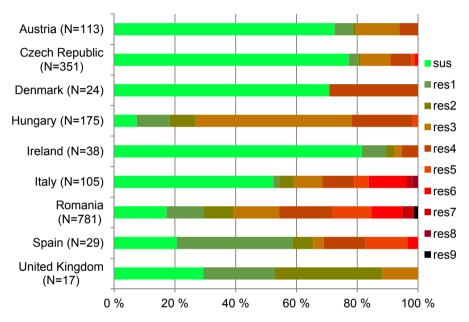
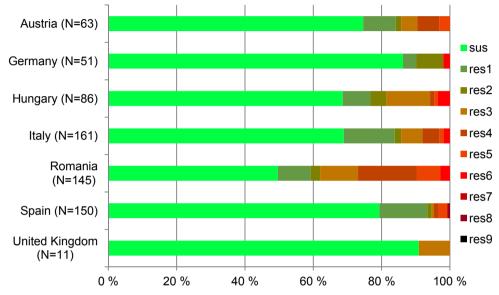


Figure SA16. Frequency distribution of <u>Salmonella spp.</u> isolates from <u>laying</u> <u>hens</u> completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*.



diversity in S. Enteriditis from broilers in MSs reporting isolate-based data, 2012

Table SA26. Complete susceptibility, multi-resistance and index of Table SA27. Complete susceptibility, multi-resistance and index of diversity in S. Enteriditis from laying hens in MSs reporting isolate-based data, 2012

Country		eptible all		ulti- stant	Index of diversity	Co-resis Cip and		Country		eptible all	Multi-r	resistant	Index of diversity		istant to Ind Ctx
	n	%	n	%	arverency	n	%		n	%	n	%	arverenty	n	%
Austria (N=21)	20	95.2	0	0	0	0 (0)	0 (0)	Austria (N=15)	15	100	0	0	NA	0 (0)	0 (0)
Czech Republic (N=236)	231	97.9	0	0	0.068	0 (0)	0 (0)	Germany (N=21)	21	100	0	0	NA	0 (0)	0 (0)
Romania (N=10)	4	40.0	5	50.0	0.315	0 (0)	0 (0)	Hungary (N=25)	23	92.0	0	0	0.086	0 (0)	0 (0)
						-		Italy (N=28)	22	78.6	0	0	0	0 (0)	0 (0)
								Romania (N=66)	49	74.2	11	16.7	0.336	0 (0)	0 (0)
								Spain (N=43)	32	74.4	1	2.3	0.14	0 (0)	0 (0)

MS:Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella; n: number of isolates.

Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set.

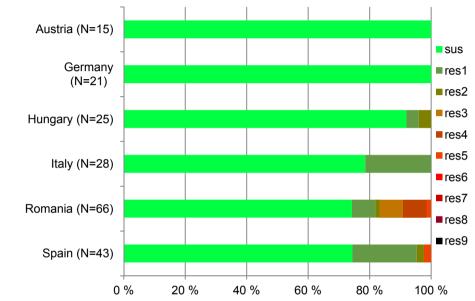
Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of Salmonella isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip: >1 mg/L).



Figure SA17. Frequency distribution of <u>S. Enteritidis</u> isolates from <u>broilers</u> completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012

Figure SA18. Frequency distribution of S. Enteritidis isolates from laying hens completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012



Austria (N=21) sus ∎res1 res2 res3 res4 Czech Republic (N=236) res5 res6

40 %

60 %

80 %

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella; sus: susceptible to all antimicrobial substances of the EFSA common set for Salmonella: res1-res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for Salmonella.

res7

res8

∎res9

100 %

Romania (N=10)

0 %

20 %



3.4.2.2. Turkeys

In 2012, nine MSs submitted quantitative antimicrobial susceptibility data for *Salmonella* spp. from turkeys, in accordance with the EU legislation. This section includes data from fattening flocks and unspecified flocks of turkeys. Seven MSs reported data on *S*. Typhimurium in turkeys and five MSs reported data on monophasic *S*. Typhimurium in turkeys; however, no country submitted sufficient data to warrant inclusion in the report.

Representative sampling and monitoring

In the reporting MSs, Austria, the Czech Republic, Germany, Hungary, Ireland, Italy, Poland, Spain and the United Kingdom, antimicrobial resistance monitoring in *Salmonella* spp. isolates from turkeys relied primarily on the national control and eradication programme of *Salmonella* based on census sampling of fattening flocks in accordance with EU regulations. Only one representative *Salmonella* isolate per positive epidemiological unit (flock), derived from environmental samples of faeces or dust, was gathered to account for clustering. In some reporting countries, representative subsets of *Salmonella* isolates were randomly selected at the laboratory for susceptibility testing, whereas, in some others, all the Salmonella isolates recovered were tested for susceptibility. Either no or incomplete information on antimicrobial resistance monitoring in turkeys was provided by Hungary, Italy, Poland and the United Kingdom.

Resistance levels in *Salmonella* spp. isolates from turkeys

Data on antimicrobial resistance among *Salmonella* spp. in turkeys were reported by nine MSs in 2012 (Table SA28). The occurrence of resistance to ampicillin, sulfonamides and tetracyclines was generally high to extremely high across the reporting MSs. An exception to this is Austria which reported low resistance to tetracyclines. For chloramphenicol and gentamicin, contrasting levels of resistance were observed. As for chloramphenicol, two-thirds of the reporting MSs did not record resistance or observed low resistance, while one-third reported moderate to high resistance, and for gentamicin, high resistance were reported by two-thirds of the reporting MSs, while the remaining MSs observed low resistance. Resistance levels to ciprofloxacin and nalidixic acid were high to extremely high among the reporting MSs, except in the United Kingdom which reported low resistance. Cefotaxime resistance was very low in the reporting group of nine MSs at 0.8 %, with only Italy, Poland and Spain reporting any cefotaxime-resistant isolates at low proportions of 4.2 %, 3.6 % and 1.2 %, respectively. In contrast to the general feature, Ireland detected resistance only to sulfonamides and tetracyclines at low levels in a small sample of 11 *Salmonella* spp. isolates.

The feature of the resistance observed in the sub-set of *Salmonella* spp. isolates obtained from fattening turkey flocks in five MSs generally paralleled that described above in *Salmonella* spp. in turkeys with high to extremely high resistance to ampicillin, ciprofloxacin, nalidixic acid, sulfonamides and tetracyclines, low to moderate resistance to gentamicin and an absence of or low resistance to cefotaxime and chloramphenicol.

Eight MSs reported resistance among *Salmonella* spp. isolates from both broilers (*Gallus gallus*) and turkeys in 2012. As previously observed in 2010 and 2011, resistance was much higher in turkeys than in broilers, in particular for ampicillin, ciprofloxacin, chloramphenicol, gentamicin, nalidixic acid, sulfonamides and tetracyclines. More reporting MSs did not detect resistance to cefotaxime in isolates from turkeys than in *Gallus gallus* and thus, overall, resistance was lower (0.8 %) in turkeys than in *Gallus gallus* (4.8 %). The difference in resistance levels between the two species needs to be interpreted with caution because, except for Spain, resistance levels in *Salmonella* spp. isolates from turkeys are estimated on lower numbers of isolates compared with *Gallus gallus*.

Resistance levels in Salmonella Kentucky isolates from turkeys

Hungary and Poland reported quantitative MIC antimicrobial susceptibility data for *S*. Kentucky isolates from turkeys in 2012 and the results are presented in Table SA29. High levels of resistance to ampicillin, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines were recorded by both MSs, while resistance was not detected in cefotaxime and chloramphenicol.



Country	Amp	oicillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Gent	amicin	Nalidiz	kic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	N	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res
Turkeys		-	-		-	-	-		-	-	-		-	-	-	
Austria	38	23.7	38	0	38	0	38	78.9	38	7.9	38	78.9	38	26.3	38	5.3
Czech Republic	27	55.6	27	0	27	0	27	81.5	27	29.6	27	81.5	27	29.6	-	-
Germany	87	55.2	87	0	87	2.3	87	32.2	87	21.8	87	28.7	87	57.5	87	48.3
Hungary	174	51.1	174	0	174	0.6	174	91.4	174	20.7	174	84.5	174	33.9	174	55.2
Ireland	14	0	14	0	14	0	14	0	14	0	14	0	14	14.3	14	14.3
Italy	48	62.5	48	4.2	48	12.5	48	37.5	48	25.0	48	37.5	48	58.3	48	97.9
Poland	55	50.9	55	3.6	55	21.8	55	52.7	55	25.5	55	45.5	55	54.5	55	58.2
Spain	169	95.9	169	1.2	169	56.8	169	89.9	169	3.0	169	21.3	169	85.2	169	91.7
United Kingdom	142	31.7	142	0	142	0.7	142	9.9	142	0.7	142	9.9	142	79.6	142	76.8
Total (9 MSs)	754	56.5	754	0.8	754	15.6	754	59.9	754	13.0	754	42.0	754	58.9	727	66.7
Fattening turkeys			-	-	-		-	-	-		-		-		-	
Austria	38	23.7	38	0	38	0	38	78.9	38	7.9	38	78.9	38	26.3	38	5.3
Czech Republic	20	40.0	20	0	20	0	20	75.0	20	10.0	20	75.0	20	10.0	_	-
Germany	12	50.0	12	0	12	0	12	0	12	8.3	12	0	12	58.3	12	66.7
Hungary	174	51.1	174	0	174	0.6	174	91.4	174	20.7	174	84.5	174	33.9	174	55.2
Spain	169	95.9	169	1.2	169	56.8	169	89.9	169	3.0	169	21.3	169	85.2	169	91.7
Total (5 MSs)	413	66.3	413	0.5	413	23.5	413	86.2	413	11.4	413	55.2	413	53.8	393	66.4

Table SA28. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella spp.</u> isolates from <u>turkeys</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Table SA29. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>S. Kentucky</u> isolates from <u>turkeys</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

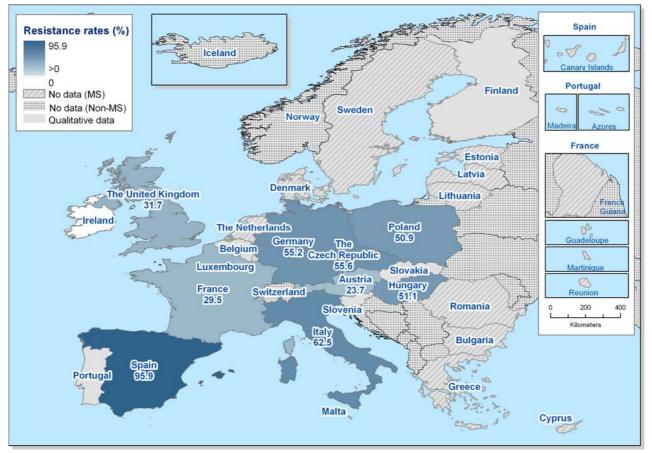
Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidia	cic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Hungary	35	100	35	0	35	0	35	100	35	97.1	35	100	35	97.1	35	97.1
Poland	11	90.9	11	0	11	0	11	100	11	100	11	100	11	100	11	100
Total (2 MSs)	46	97.8	46	0	46	0	46	100	46	97.8	46	100	46	97.8	46	97.8



Spatial distribution of resistance among Salmonella isolates from turkeys

Figures SA19 and SA20 show the spatial distributions of ampicillin and nalidixic acid resistance in *Salmonella* spp. isolated from turkeys in 2012. These illustrate great variation in levels of ampicillin and nalidixic acid resistance in *Salmonella* spp. across the EU. Higher resistance to ampicillin was recorded in Southern Europe and to a lesser extent in Central and Eastern Europe compared with Western and Northern Europe. Regarding nalidixic resistance, a number of Central European MSs reported much higher levels of resistance.

Spatial distribution of <u>ampicillin</u> resistance among Salmonella spp. from <u>turkeys</u> in countries reporting MIC data in 2012¹



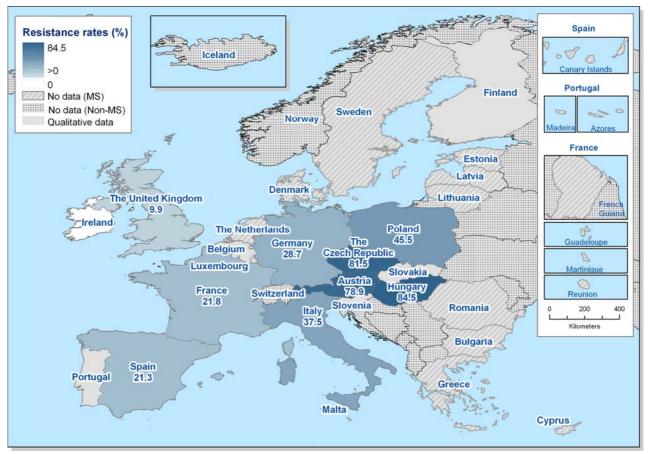
MIC: minimum inhibitory concentration; MS: Member State.

Note:Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting either inhibition zone diameter data, MIC data for fewer than 10 isolates, or purely qualitative data (as proportion of resistant isolates).

1. For France, 2011 data were used.



Figure SA19. Spatial distribution of <u>nalidixic acid</u> resistance among Salmonella spp. from <u>turkeys</u> in countries reporting MIC data in 2012¹



MIC: minimum inhibitory concentration; MS: Member State.

- Note:Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting either inhibition zone diameter data, MIC data for fewer than 10 isolates, or purely qualitative data (as proportion of resistant isolates).
- 1. For France, 2011 data were used.

Multi-resistance among Salmonella spp. isolates from turkeys

In 2012, seven MSs provided isolate-based data concerning resistance in *Salmonella* spp. from turkeys; one more than in 2011. Complete susceptibility was exhibited by less than one-quarter of the isolates tested in most of the reporting MSs with, in particular, Italy and Spain reporting low to very low levels of susceptibility. Contrastingly, Ireland reported 85.7 % of complete susceptibility, although, in this case, it was assessed on an isolate sample of small size. Multi-resistance was high to extremely high in most reporting MSs ranging from 26.3 % in Austria to 95.3 % in Spain (Table SA30), while moderate multi-resistance was reported in Ireland. The frequency distributions (Figure SA21) showed similarities among multi-resistance recorded in Austria, the Czech Republic, Germany and Hungary with some isolates showing reduced susceptibility to as many as five and six different substances, while Italy and Spain reported isolates showing reduced susceptibility to up to seven and nine antimicrobials. Ireland recorded multi-resistance to no more than three classes of antimicrobials at a maximum.

Very few isolates were resistant to both ciprofloxacin and cefotaxime (Table SA30).



Table SA30. Complete susceptibility, multi-resistance and index of diversity in Salmonella spp. from <u>turkeys</u> in MSs reporting isolate-based data, 2012

Country	Suscep	otible to all	Multi-	resistant	Index of		istant to nd Ctx
	n	%	n	%	diversity	n	%
Austria (N=38)	7	18.4	10	26.3	0.231	0 (0)	0 (0)
Czech Republic (N=27)	2	7.4	11	40.7	0.427	0 (0)	0 (0)
Germany (N=87)	20	23.0	51	58.6	0.365	0 (0)	0 (0)
Hungary (N=174)	11	6.3	96	55.2	0.426	0 (0)	0 (0)
Ireland (N=14)	12	85.7	2	14.3	0	0 (0)	0 (0)
Italy (N=48)	0	0	39	81.2	0.554	1 (0)	2.1 (0)
Spain (N=169)	3	1.8	161	95.3	0.601	2 (0)	1.2 (0)

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella; n: number of isolates.

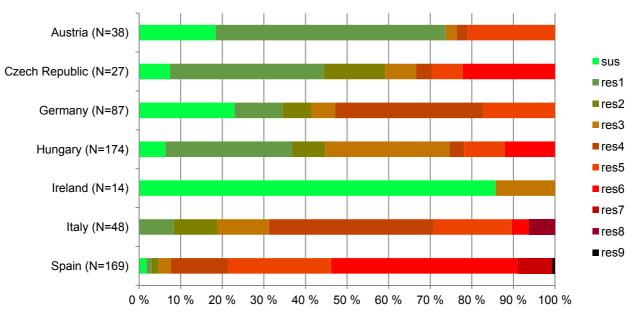
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of Salmonella isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip:>1 mg/L).

Figure SA20. Frequency distribution of Salmonella spp. in <u>turkeys</u> completely susceptible or resistant to one to nine antimicrobials, in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*.

Multi-resistance among S. Enteritidis and S. Typhimurium isolates from turkeys

Generally, the S. Enteritidis, S. Typhimurium and monophasic S. Typhimurium isolates from turkey flocks were very rare in the isolate-based dataset of the reporting MSs. Data on multi-resistance in these serovars from turkeys are therefore not presented in this report, as the inclusion criteria (more than three reporting countries providing data on more than 10 isolates per production type) were not met.



3.4.2.3. Pigs

Quantitative MIC data for *Salmonella* spp. isolated from pigs from 10 MSs in 2012 are included in the following analyses.

Representative sampling and monitoring

Isolates from Spain and some of the isolates from Estonia were collected as part of monitoring plans, whereas Germany and Italy and also Estonia tested isolates obtained through passive surveillance via diagnostic submissions. Denmark collected isolates from sub-clinical infections detected via the serological surveillance programme, from healthy pigs at slaughter. Sample types collected by MSs were generally faecal, while Estonia and Spain also tested ileocaecal lymph nodes at slaughter.

Resistance levels in Salmonella spp. isolates from pigs

Data describing the occurrence of resistance to selected antimicrobials in isolates of *Salmonella* spp. from pigs are presented in Table SA31. Isolates tested by Denmark, Germany and the Netherlands made up over 77 % of the total isolates tested in 2012, so the results from these three countries will have influenced the overall levels reported at MS group level.

Resistance to ampicillin, sulfonamides and tetracyclines was generally high to extremely high, ranging from more than 30 % to about 90 % among the reporting MSs. The only exception to this was Estonia which reported an ampicillin resistance of 18.8 %. Although lower, resistance to chloramphenicol varied between 0 % and 33.3 % among the reporting MSs, while gentamicin resistance was typically low in the reporting MSs ranging from 0 % to 10.4 %.

Three MSs detected no resistance or reported very low resistance to either ciprofloxacin or nalidixic acid in *Salmonella* spp. isolates from pigs. Among the MSs which did detect resistance, the occurrence of ciprofloxacin and nalidixic acid resistance mostly ranged from about 5 % to 25 %, with the exception of Poland which recorded a ciprofloxacin resistance of 60.0 %. Resistance to cefotaxime was generally not detected or reported at low levels in *Salmonella* spp. in pigs, with only three MSs reporting cefotaxime resistance ranging from 2.1 % to 11.2 %.

Five MSs reported quantitative data on isolates of *Salmonella* spp. from fattening pigs in 2012 (Table SA31). At the reporting MS group level, the overall resistance to ampicillin, sulfonamides and tetracyclines were high, at 35.2 %, 40.1 % and 45.6 %, respectively. Low levels of resistance to ciprofloxacin and to nalidixic acid were observed at the MS group level (4.1 % and 3.7 %, respectively), with values varying from 0 % to 23.7 %. As usually observed, the levels of resistance within each MS were generally very similar for the two compounds. In *Salmonella* spp. from fattening pigs, Denmark and Spain were the only countries to report resistance to gentamicin at levels of 2.9 % and 10.4 %, respectively. Cefotaxime resistance was observed only by Spain at a low level of 6.3 %.

Quantitative data on isolates of *Salmonella* spp. from breeding pigs were reported by one MS in 2012 (Table SA31).

Resistance levels in Salmonella Typhimurium isolates from pigs

Quantitative MIC antimicrobial susceptibility results for *S*. Typhimurium isolates from pigs were reported by five MSs in 2012 (Table SA32). More than half of isolates tested (52.6 %) were from Germany, so the results from Germany will have more bearing on the overall levels. The occurrence of resistance to ampicillin, chloramphenicol, sulfonamides and tetracyclines among *S*. Typhimurium isolates from pigs was higher than that reported in *Salmonella* spp., with the overall level of resistance in the reporting MS group being 76.7 % for ampicillin, 30.1 % for chloramphenicol, 77.5 % for sulfonamides and 71.7 % for tetracyclines. Among the individual reporting MSs, resistance to ampicillin ranged from 41.3 % to 85.5 %, resistance to chloramphenicol ranged from 14.5 % to 53.3 %, resistance to sulfonamides ranged from 46.0 % to 86.1 % and resistance to tetracyclines ranged from 36.5 % to 93.3 %. Resistance to ciprofloxacin and gentamicin was fairly similar in *S*. Typhimurium and *Salmonella* spp. (7.5 % vs. 7.6 % and 3.7 % vs 3.4 %, respectively). Resistance to cefotaxime and nalidixic acid was similar in *S*. Typhimurium and *Salmonella* spp. (2.3 % and



5.8 %, respectively). In the reporting MS group, cefotaxime resistance was detected in S. Typhimurium isolates from Belgium at a moderate level (10.7 %), and from Germany at a low level (1.1 %).

Resistance levels in monophasic Salmonella Typhimurium isolates from pigs

Table SA33 describes the resistance among isolates of monophasic *S*. Typhimurium from pigs. Six MSs reported quantitative data from pigs. As for *S*. Typhimurium, more than half of isolates tested (55.2 %) were from Germany, so the results from Germany will have more bearing on the overall levels. The levels of resistance to ampicillin, sulfonamides and tetracyclines reported in monophasic *S*. Typhimurium were much higher than the levels reported in *Salmonella* spp. from pigs (91.3 % vs. 60.2 %, 91.0 % vs. 63.3 % and 93.5 % vs 63.3 %, respectively).

Extremely high resistance to ampicillin, sulfonamides and tetracyclines was observed across all reporting MSs (91.3 %, 91.0 % and 93.5 %, respectively), and this ranged from 84.0 % to 94.3 % for ampicillin, from 84.0 % to 93.9 % for sulfonamides and from 80.0 % to 100 % for tetracyclines. Low levels of resistance to chloramphenicol (9.4 %) and gentamicin (5.1 %) were reported at the MS group level, and ranged from 2.5 % to 30.0 % and from 0 % to 21.4 %, respectively. In addition, low resistance to ciprofloxacin and nalidixic acid was reported at the MS group level (8.7 % and 5.6 %, respectively). Denmark and the Netherlands observed no resistance to both compounds, while in the remaining four MSs, resistance ranged from 3.9 % to 60.0 %. Cefotaxime resistance was observed by Belgium, Germany and Spain making the overall resistance, at MS group level, 2.7 %.

Resistance levels in Salmonella Derby isolates from pigs

Table SA34 describes the resistance among isolates of *S*. Derby from pig. Four MSs reported quantitative data from pigs. The levels of resistance to ampicillin, sulfonamides and tetracyclines reported in *S*. Derby were lower than the levels reported in *Salmonella* spp. from pigs (12.4 % vs. 60.3 %, 22.8 % vs. 63.4 % and 31.3 % vs. 63.4 %, respectively). Low levels of resistance to chloramphenicol, ciprofloxacin and gentamicin (1.2 %) were reported at the MS group level. The levels of resistance to nalidixic acid and cefotaxime were low at the MS group level.



Table SA31. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella spp.</u> isolates from <u>pigs</u> in 2012, using harmonised epidemiological cut-off values

Country	Amp	oicillin	Cefo	taxime	Chloram	phenicol	Ciprot	loxacin	Gent	amicin	Nalidi	xic acid	Sulfor	namides	Tetrac	cyclines
Country	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
All pigs																
Belgium	187	65.2	187	11.2	187	13.9	187	22.5	187	3.2	187	18.2	187	63.1	187	57.8
Denmark	374	31.3	374	0	374	4.5	374	0	374	2.9	374	0	374	36.4	374	40.6
Estonia	32	18.8	32	0	32	0	32	0	32	0	32	0	32	43.8	32	31.3
Germany	627	77.4	627	2.1	627	22.2	627	7.5	627	4.1	627	4.9	627	79.6	627	75.6
Hungary	38	57.9	38	0	38	10.5	38	23.7	38	0	38	23.7	38	68.4	38	60.5
Ireland	24	50.0	24	0	24	33.3	24	16.7	24	4.2	24	16.7	24	66.7	24	83.3
Italy	25	36.0	25	0	25	12.0	25	12.0	25	8.0	25	12.0	25	48.0	25	48.0
Netherlands	263	64.6	263	0	263	9.1	263	0.8	263	1.5	263	0.8	263	66.5	263	68.8
Poland	10	90.0	10	0	10	30.0	10	60.0	10	0	10	30.0	10	90.0	10	80.0
Spain	48	58.3	48	6.3	48	14.6	48	20.8	48	10.4	48	16.7	48	54.2	48	89.6
Total (10 MSs)	1,628	60.2	1,628	2.3	1,628	14.2	1,628	7.6	1,628	3.4	1,628	5.8	1,628	63.3	1,628	63.3
Fattening pigs	-	-	-				-		-	-	-	-	-	-	-	
Denmark	374	31.3	374	0	374	4.5	374	0	374	2.9	374	0	374	36.4	374	40.6
Estonia	14	28.6	14	0	14	0	14	0	14	0	14	0	14	35.7	14	14.3
Hungary	38	57.9	38	0	38	10.5	38	23.7	38	0	38	23.7	38	68.4	38	60.5
Netherlands	17	11.8	17	0	17	5.9	17	5.9	17	0	17	5.9	17	23.5	17	23.5
Spain	48	58.3	48	6.3	48	14.6	48	20.8	48	10.4	48	16.7	48	54.2	48	89.6
Total (5 MSs)	491	35.2	491	0.6	491	5.9	491	4.1	491	3.3	491	3.7	491	40.1	491	45.6
Breeding pigs									-		-					
Belgium	187	65.2	187	11.2	187	13.9	187	22.5	187	3.2	187	18.2	187	63.1	187	57.8



Table SA32. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>S. Typhimurium</u> isolates from <u>pigs</u> in 2012, using harmonised epidemiological cut-off values

Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidio	cic acid	Sulfor	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
All pigs																
Belgium	75	81.3	75	10.7	75	20.0	75	20.0	75	2.7	75	13.3	75	74.7	75	66.7
Denmark	63	41.3	63	0	63	17.5	63	0	63	1.6	63	0	63	46.0	63	36.5
Germany	273	82.8	273	1.1	273	37.7	273	6.6	273	5.1	273	5.5	273	86.1	273	79.5
Ireland	15	60.0	15	0	15	53.3	15	20.0	15	6.7	15	20.0	15	73.3	15	93.3
Netherlands	55	85.5	55	0	55	14.5	55	0	55	0	55	0	55	76.4	55	74.5
Total (5 MSs)	481	76.7	481	2.3	481	30.1	481	7.5	481	3.7	481	5.8	481	77.5	481	71.7
Fattening pigs					-									-		•
Denmark	63	41.3	63	0	63	17.5	63	0	63	1.6	63	0	63	46.0	63	36.5
Breeding pigs																
Belgium	75	81.3	75	10.7	75	20.0	75	20.0	75	2.7	75	13.3	75	74.7	75	66.7



Table SA33. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>monophasic S. Typhimurium</u> isolates from <u>pigs</u> in 2012, using harmonised epidemiological cut-off values

Country	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Gent	amicin	Nalidix	cic acid	Sulfor	amides	Tetrac	yclines
Country	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res	Ν	% Res
Belgium	41	90.2	41	19.5	41	14.6	41	26.8	41	7.3	41	24.4	41	87.8	41	87.8
Denmark	81	84.0	81	0	81	2.5	81	0	81	7.4	81	0	81	84.0	81	87.7
Germany	228	94.3	228	0.9	228	9.6	228	7.5	228	3.1	228	3.9	228	93.9	228	96.5
Netherlands	39	92.3	39	0	39	7.7	39	0	39	5.1	39	0	39	92.3	39	94.9
Poland	10	90.0	10	0	10	30.0	10	60.0	10	0	10	30.0	10	90.0	10	80.0
Spain	14	85.7	14	7.1	14	21.4	14	14.3	14	21.4	14	7.1	14	92.9	14	100
Total (6 MSs)	413	91.3	413	2.7	413	9.4	413	8.7	413	5.1	413	5.6	413	91.0	413	93.5

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

Table SA34. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>S. Derby</u> isolates from <u>pigs</u> in 2012, using harmonised epidemiological cut-off values

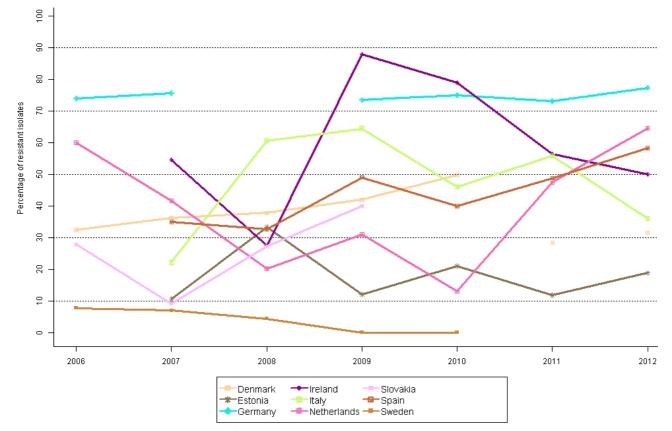
Country	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidia	kic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Belgium	14	21.4	14	7.1	14	7.1	14	7.1	14	0	14	14.3	14	50.0	14	28.6
Denmark	176	9.7	176	0	176	2.3	176	0	176	1.1	176	0	176	16.5	176	31.3
Germany	42	28.6	42	11.9	42	2.4	42	9.5	42	2.4	42	9.5	42	38.1	42	23.8
Netherlands	28	0	28	0	28	0	28	0	28	0	28	0	28	25.0	28	42.9
Total (4 MSs)	260	12.3	260	2.3	260	2.3	260	1.9	260	1.2	260	2.3	260	22.7	260	31.2



Temporal trends in resistance among Salmonella isolates from pigs

The temporal variation in the level of resistance to selected antimicrobials in *Salmonella* spp., isolated from pigs between 2006 and 2012, is presented in Figures SA22 to SA24. The figures demonstrate that, in some MSs, resistance levels have continued to fluctuate; however, in other countries, such as in Germany and Sweden, the occurrence of resistance has remained fairly stable in recent years. Over the seven reporting years, significantly decreasing trends in resistance were reported by the Netherlands for ampicillin, while Ireland, Italy and Spain reported statistically significant increasing trends in resistance to the same substance (Figure SA22). Considering resistance to (fluoro) quinolones, ciprofloxacin and nalidixic acid, Estonia and Germany reported statistically decreasing trends in resistance to both compounds over the 2006–2012 period. In contrast, Spain showed increasing trends in resistance to these two substances. Additionally, Ireland registered an increasing trend in resistance to ciprofloxacin (Figure SA24). Cefotaxime resistance among *Salmonella* spp. isolates from pigs remained either low, very low or absent in the reporting MSs between 2006 and 2012; and no significant trends were detected for MSs reporting five or more years of data (Figure SA23).

Figure SA21. Trends in <u>ampicillin</u> resistance in Salmonella spp. from <u>pigs</u> in reporting MSs, 2006-2012, quantitative data



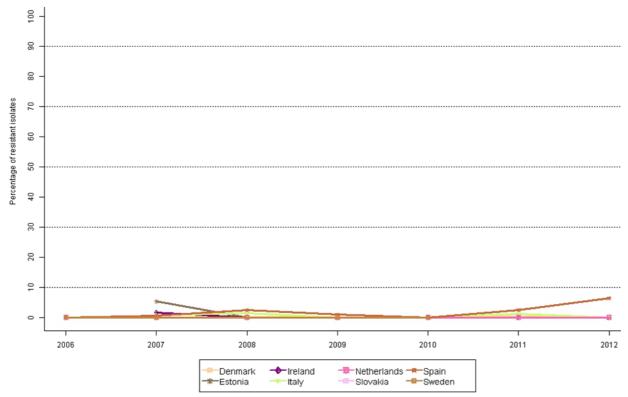
MS: Member State.

Note: Statistically significant increasing or decreasing trends for five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in Ireland (\uparrow), Italy (\uparrow), the Netherlands (\downarrow) and Spain (\uparrow).

Danish data are not comparable between years: data from 2006-2010 contained only *S*. Typhimurium isolates while all the isolates were reported in 2011 and 2012.



Figure SA22. Trends in <u>cefotaxime</u> resistance in Salmonella spp. from <u>pigs</u> in reporting MSs, 2006-2012, quantitative data

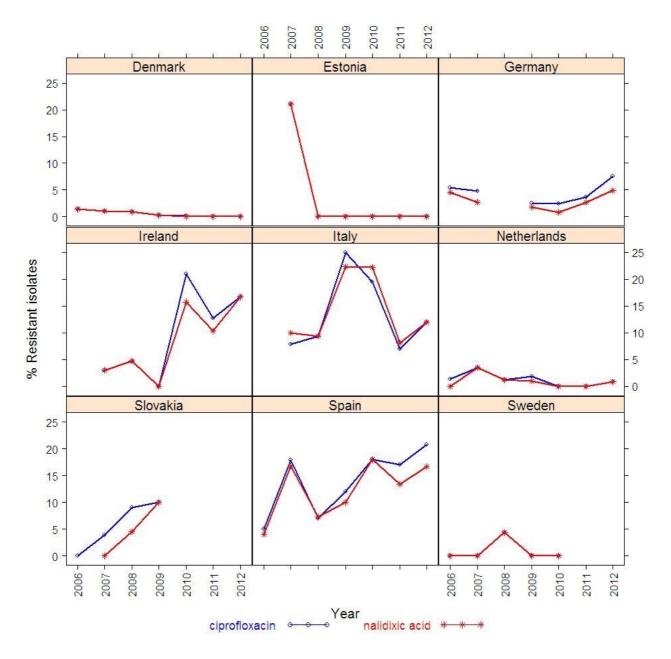


MS: Member State.

Note: No statistically significant trend for five or more years, as tested by a logistic regression model ($p \le 0.05$), was observed in any of the reporting countries.



Figure SA23. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in Salmonella spp. from <u>pigs</u> in reporting MSs, 2006-2012, quantitative data



MS: Member State.

Note: A statistically significant decreasing trend for five or more years, as tested by a logistic regression model ($p \le 0.05$), was observed in Estonia (\downarrow) and Germany (\downarrow) for both ciprofloxacin and nalidixic acid. A statistically significant increasing trend was observed in Spain (\uparrow) for both ciprofloxacin and nalidixic acid and in Ireland (\uparrow) for ciprofloxacin.

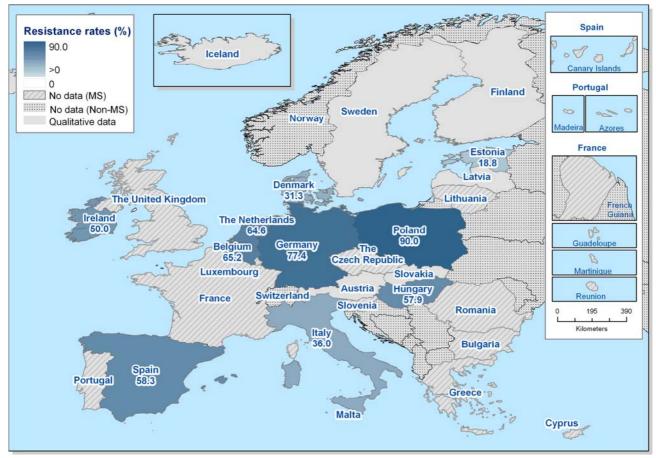
Danish data are not comparable between years: data from 2006-2010 contained only S. Typhimurium isolates while all the isolates were reported in 2011 and 2012.



Spatial distribution of resistance among Salmonella isolates from pigs

The spatial distribution of ampicillin and nalidixic acid resistance in *Salmonella* spp. from pigs in 2012 is shown in Figures SA25 and SA26. Figure SA25 emphasises the large differences in ampicillin resistance rates in different MSs, although no clear spatial distributions were observed. In most countries, nalidixic acid resistance in *Salmonella* spp. isolated from pigs was reported to be low, with no clear spatial distribution apparent (Figure SA26).



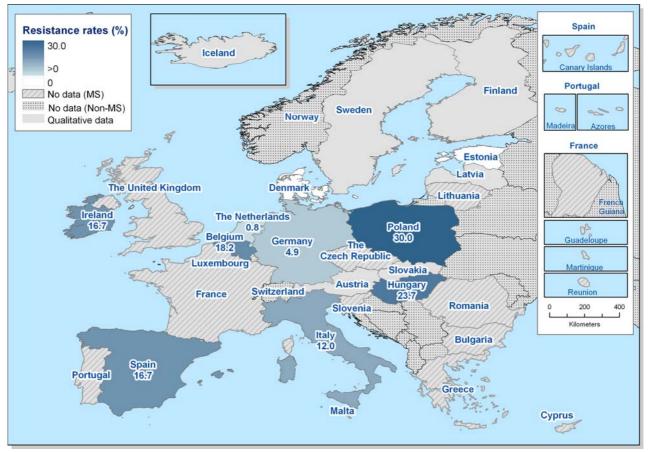


MIC: minimum inhibitory concentration; MS: Member State.

Note: Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting inhibition zone diameter data, MIC data for fewer than 10 isolates or purely qualitative data (as proportion of resistant isolates).



Figure SA25. Spatial distribution of <u>nalidixic acid</u> resistance among Salmonella spp. from <u>pigs</u> in countries reporting MIC data in 2012



MIC: minimum inhibitory concentration; MS: Member State.

Note: Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting inhibition zone diameter data, MIC data for fewer than 10 isolates or purely qualitative data (as proportion of resistant isolates).



Multi-resistance among Salmonella spp. isolates from fattening pigs

In 2012, seven MSs provided isolate-based data concerning resistance in *Salmonella* spp. from pigs. The levels of complete susceptibility varied between the reporting MSs, from 6.2 % in Spain to 64.3 % in Estonia. The multi-resistance levels were high to very high in all reporting MSs, ranging from 34.0 % in Denmark to 76.9 % in Germany (Table SA35). The frequency distributions (Figure SA27) showed discrepancies among the multi-resistance recorded in the reporting MSs with some isolates showing reduced susceptibility to up to eight different substances in Italy and Spain, while Estonia recorded multi-resistance to three classes at a maximum. Very few isolates were resistant to both antimicrobials ciprofloxacin and cefotaxime (Table SA35).

Table SA35. Complete susceptibility, multi-resistance and index of diversity in <u>Salmonella spp.</u> from <u>fattening pigs</u> in MSs reporting isolate-based data, 2012

Country	Suscept	tible to all	Multi-r	esistant	Index of		istant to Ind Ctx
, in the second s	n	%	n	%	diversity	n	%
Denmark (N=374)	179	47.9	127	34.0	0.388	0 (0)	0 (0)
Estonia (N=14)	9	64.3	5	35.7	0.151	0 (0)	0 (0)
Germany (N=627)	89	14.2	482	76.9	0.544	5 (0)	0.8 (0)
Hungary (N=38)	9	23.7	27	71.1	0.414	0 (0)	0 (0)
Ireland (N=24)	2	8.3	15	62.5	0.586	0 (0)	0 (0)
Italy (N=25)	10	40.0	12	48.0	0.586	0 (0)	0 (0)
Spain (N=48)	3	6.2	28	58.3	0.562	2 (0)	4.2 (0)

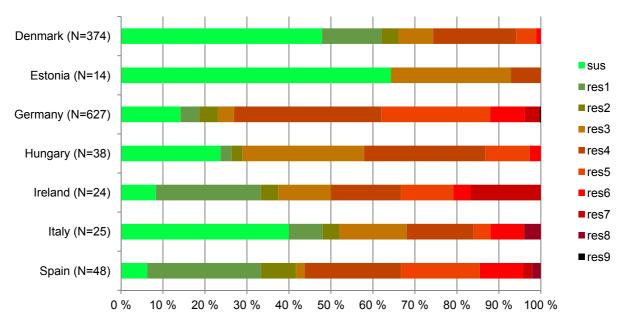
MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; n: number of isolates.

Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set. Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of *Salmonella* isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip:>1 mg/L).

Figure SA26. Frequency distribution of <u>Salmonella spp.</u> from <u>fattening pigs</u> isolates completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*.



Multi-resistance among S. Typhimurium isolates from fattening pigs

In 2012, three MSs provided isolate-based resistance data on *S*. Typhimurium from pigs concerning more than 10 isolates. Rates of full susceptibility and multi-resistance are respectively lower and higher than those assessed in *Salmonella* spp. for the three MSs reporting for both groups. Resistance to both ciprofloxacin and cefotaxime in *S*. Typhimurium isolates from pigs was not observed in the three reporting MSs (Table SA36).

Table SA36. Complete susceptibility, multi-resistance and index of diversity in <u>S. Typhimurium</u> fromfattening pigsin MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Multi-r	resistant	Index of		istant to Ind Ctx
	n	%	n	%	diversity	n	%
Denmark (N=63)	31	49.2	27	42.9	0.458	0 (0)	0 (0)
Germany (N=273)	24	8.8	221	81.0	0.519	0 (0)	0 (0)
Ireland (N=15)	0	0	10	66.7	0.593	0 (0)	0 (0)

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; n: number of isolates.

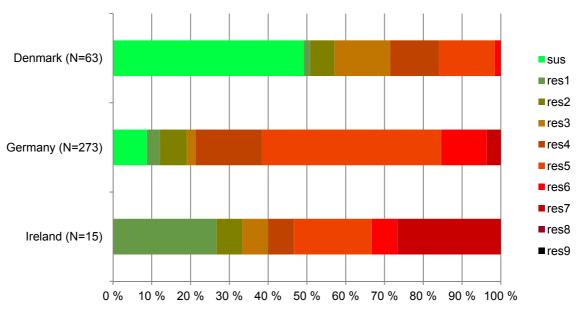
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of Salmonella isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx: >0.5 mg/L and Cip: >0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints (Ctx: >2 mg/L and Cip: >1 mg/L).

Figure SA27. Frequency distribution of <u>S. Typhimurium</u> from <u>fattening pigs</u> isolates completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*.



3.4.2.4. Cattle (bovine animals)

In this report, calves, dairy cattle, beef cows and heifers are included under the term 'cattle'. Quantitative MIC data for *Salmonella* spp. isolated from cattle in seven MSs in 2012 are included in the following analysis of antimicrobial resistance levels.

Representative sampling and monitoring

Isolates tested by Belgium, Finland, Germany, Ireland and Sweden were obtained through national monitoring programmes and generally consisted of faecal samples. Finland also tested lymph nodes at slaughter. Italy obtained isolates through passive surveillance.

Resistance levels in Salmonella spp. isolates from cattle

The levels of resistance to selected antimicrobials in isolates of *Salmonella* spp. from cattle reported by MSs in 2012 are presented in Table SA37. Isolates tested by Belgium, Germany and the Netherlands made up over 68 % of the total isolates tested in 2012, so the results from these countries will have influenced the overall levels reported at MS group level. High levels of resistance to ampicillin, sulfonamides and tetracyclines were commonly reported in *Salmonella* spp. from cattle in 2012; considering all reporting MSs, the levels of resistance were 34.5 %, 42.4 % and 36.0 %, respectively. Ampicillin resistance ranged from 13.2 % to 61.1 % across reporting MSs, the range for sulfonamides was 17.6 % to 66.7 %, while the range for tetracyclines was 5.3 % to 58.3 %. As in the previous year, only Germany and Italy reported resistance to gentamicin, making the overall resistance 1.1 %.

At MS group level, the overall occurrence of resistance to both ciprofloxacin and nalidixic acid was 9.1 %. Finland and Sweden were the only MSs to report no resistance to ciprofloxacin or nalidixic acid in *Salmonella* spp. isolates from cattle. Cefotaxime resistance was only reported by Belgium (2.4 %).

Resistance levels in Salmonella Typhimurium isolates from cattle

Table SA38 shows the level of resistance reported on *S*. Typhimurium isolates from cattle in 2012. Across the six reporting MSs, the level of resistance to ampicillin and tetracyclines was high, at 43.4 % and 44.9 %, respectively. The resistance levels reported by individual MSs varied from 11.4 % to 84.0 % for ampicillin and from 6.3 % to 83.3 % for tetracyclines. There were very high levels of resistance to sulfonamides (50.0%), ranging from 11.4 % to 91.7 %. The overall resistance to chloramphenicol was high (21.3 %) at MS group level, which varied from 0 % to 66.7 %. As in the previous year, resistance to gentamicin in *S*. Typhimurium isolates from cattle was detected only in Germany at the low level of 2.9 %.

The occurrence of resistance to both ciprofloxacin and nalidixic acid in the reporting MS group as a whole was low (2.2 % for ciprofloxacin and 2.9 % for nalidixic acid) as Belgium was the only country to report resistance (12.0 % for ciprofloxacin and 16.0 % for nalidixic acid). Cefotaxime resistance in *S*. Typhimurium isolates from cattle in 2012, as in 2011, was not reported by any MS.



Country	Amp	icillin	Cefo	taxime	Chloram	phenicol	Ciprof	floxacin	Gent	amicin	Nalidiz	kic acid	Sulfor	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Belgium	42	59.5	42	2.4	42	28.6	42	28.6	42	0	42	31.0	42	66.7	42	31.0
Finland	19	21.1	19	0	19	0	19	0	19	0	19	0	19	21.1	19	5.3
Germany	68	13.2	68	0	68	5.9	68	8.8	68	1.5	68	8.8	68	19.1	68	26.5
Ireland	36	61.1	36	0	36	44.4	36	5.6	36	0	36	5.6	36	61.1	36	58.3
Italy	14	35.7	14	0	14	21.4	14	14.3	14	14.3	14	7.1	14	35.7	14	42.9
Netherlands	68	33.8	68	0	68	5.9	68	2.9	68	0	68	2.9	68	54.4	68	50.0
Sweden	17	17.6	17	0	17	11.8	17	0	17	0	17	0	17	17.6	17	11.8
Total (7 MSs)	264	34.5	264	0.4	264	15.5	264	9.1	264	1.1	264	9.1	264	42.4	264	36.0

Table SA37. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella spp.</u> isolates from <u>cattle</u> in 2012, using harmonised epidemiological cut-off values

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

Table SA38. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among <u>Salmonella Typhimurium</u> from <u>cattle</u> in 2012, using harmonised epidemiological cut-off values

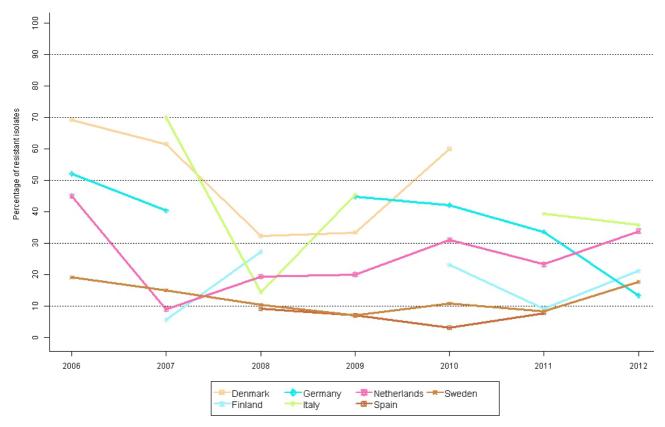
Country	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidiz	kic acid	Sulfon	amides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Belgium	25	84.0	25	0	25	20.0	25	12.0	25	0	25	16.0	25	72.0	25	48.0
Finland	16	25.0	16	0	16	0	16	0	16	0	16	0	16	25.0	16	6.3
Germany	35	11.4	35	0	35	8.6	35	0	35	2.9	35	0	35	11.4	35	28.6
Ireland	24	70.8	24	0	24	66.7	24	0	24	0	24	0	24	70.8	24	66.7
Netherlands	24	41.7	24	0	24	12.5	24	0	24	0	24	0	24	91.7	24	83.3
Sweden	12	25.0	12	0	12	16.7	12	0	12	0	12	0	12	25.0	12	16.7
Total (6 MSs)	136	43.4	136	0	136	21.3	136	2.2	136	0.7	136	2.9	136	50.0	136	44.9



Temporal trends in resistance among Salmonella isolates from cattle

It is evident from Figures SA29 and SA30 that large variations exist between MSs in the level of resistance to some antimicrobials, particularly ampicillin. The figures illustrate the trends in resistance to ampicillin ciprofloxacin and nalidixic acid among *Salmonella* isolates from cattle from 2006 to 2012. As in 2011, trends in resistance over time were mainly decreasing among *Salmonella* spp. from cattle. Germany experienced statistically significant decreasing trends in resistance to ampicillin (Figure SA29), and Germany also reported statistically significant decreasing trends in resistance to tetracyclines (data were not presented). No significant trends were observed in the reported resistance to ciprofloxacin and nalidixic acid between 2006 and 2012 (Figure SA30).

Figure SA28. Trends in <u>ampicillin</u> resistance in Salmonella spp. from <u>cattle</u> in reporting MSs, 2006-2012, quantitative data

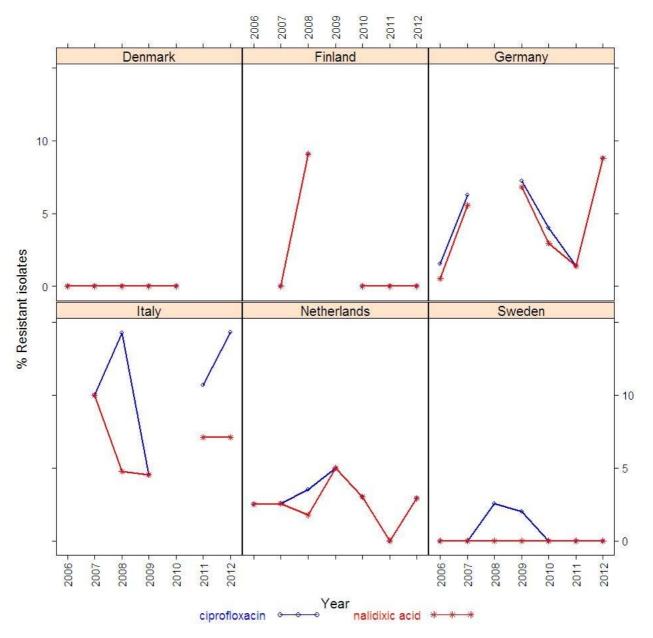


MS: Member State.

Note: A statistically significant decreasing trend for five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed in Germany (↓).



Figure SA29. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in Salmonella spp. from <u>cattle</u> in reporting MSs, 2006-2012, quantitative data



MS: Member State.

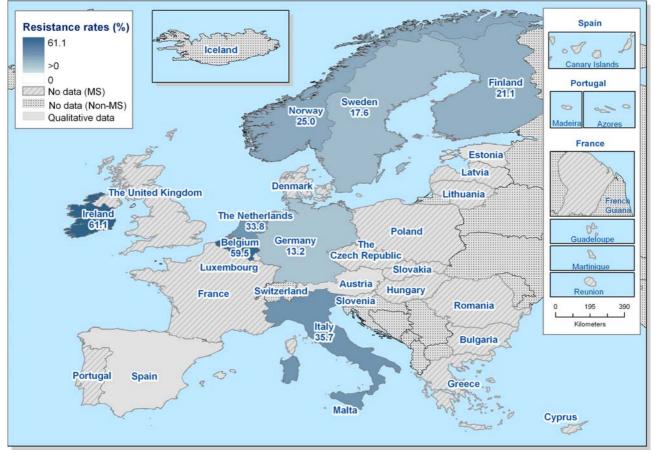
Note: For both ciprofloxacin and nalidixic acid, no statistically significant trend for five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed in any of the reporting countries.



Spatial distribution of resistance among *Salmonella* isolates from cattle

Figures SA31 and SA32 show the spatial distributions of ampicillin and nalidixic acid resistance in *Salmonella* spp. isolated from cattle in 2012. Figure SA31 illustrates the absence of a clear spatial distribution. Figure SA32 illustrates the continued absence, or low prevalence, of resistance to nalidixic acid in *Salmonella* spp. isolated from cattle in Europe.

Figure SA30. Spatial distribution of <u>ampicillin</u> resistance among Salmonella spp. from <u>cattle</u> in countries reporting MIC data in 2012¹



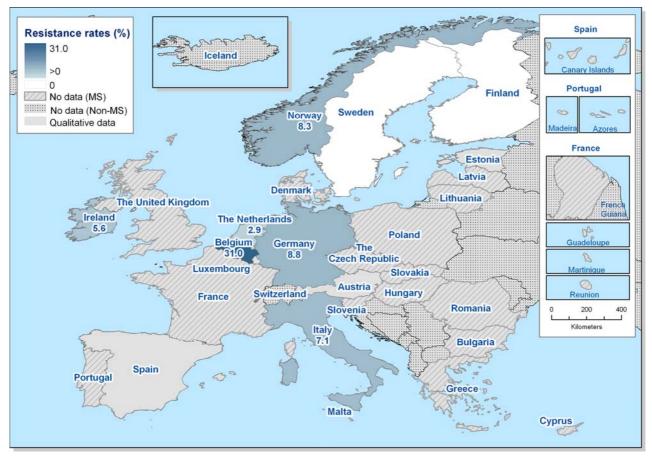
MIC: minimum inhibitory concentration; MS: Member State.

Note: Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting inhibition zone diameter data, MIC data for less than 10 isolates, or purely qualitative data (as proportion of resistant isolates).

1. For Norway, 2011 data were used.



Figure SA31. Spatial distribution of <u>nalidixic acid</u> resistance among Salmonella spp. from <u>cattle</u> in countries reporting MIC data in 2012¹



Note: Percentages shown in this map refer to countries which reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. The countries labelled as 'qualitative data' therefore include those reporting inhibition zone diameter data, MIC data for less than 10 isolates, or purely qualitative data (as proportion of resistant isolates). MIC: minimum inhibitory concentration; MS: Member State.

1. For Norway, 2011 data were used.



Multi-resistance among Salmonella spp. isolates from cattle

In 2012, six MSs reported isolate-based data concerning resistance in *Salmonella* spp. from cattle. The proportions of complete susceptible isolates were generally high to extremely high and varied between the reporting MSs, from 19.0 % in Belgium to 82.4 % in Sweden. Two reporting MSs (Finland and Sweden) reported moderate levels of multi-resistance in isolates tested from cattle, while the remaining MSs reported high levels of multi-resistance ranging between 20.6 % and 58.3 % (Table SA39). The frequency distributions (Figure SA33) showed that Belgium, Germany, Ireland and Italy detected isolates exhibiting reduced susceptibility to higher numbers of different substances (six to seven classes) than the two other MSs (five classes). Few isolates were resistant to both antimicrobials ciprofloxacin and cefotaxime (Table SA39).

Table SA39. Complete susceptibility, multi-resistance and index of diversity in <u>Salmonella spp.</u> from cattle in MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Multi-r	esistant	Index of	Co-resistant to Cip and Ctx		
	n	%	n	%	diversity	n	%	
Belgium (N=42)	8	19.0	21	50.0	0.551	1 (0)	2.4 (0)	
Finland (N=19)	15	78.9	2	10.5	0.317	0 (0)	0 (0)	
Germany (N=68)	45	66.2	14	20.6	0.409	0 (0)	0 (0)	
Ireland (N=36)	12	33.3	21	58.3	0.336	0 (0)	0 (0)	
Italy (N=14)	8	57.1	6	42.9	0.414	0 (0)	0 (0)	
Sweden (N=17)	14	82.4	3	17.6	0.203	0 (0)	0 (0)	

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; n: number of isolates.

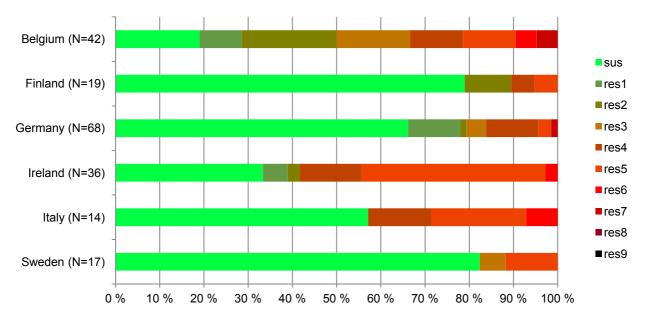
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of *Salmonella* isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip:>1 mg/L).

Figure SA32. Frequency distribution of <u>Salmonella spp.</u> from <u>cattle</u> completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*.



Multi-resistance among S. Typhimurium isolates from cattle

In 2012, five MSs reported isolate-based data concerning resistance in *S*. Typhimurium from cattle (Table SA40). These are the same as those reporting in *Salmonella* spp., except Italy which did not report any resistance data in *S*. Typhimurium in cattle. Generally, complete susceptibility and multi-resistance rates in *S*. Typhimurium isolates were similar to or lower than those in *Salmonella* spp. in the remaining MSs, with the exception of Sweden which reported higher multi-resistance in *S*. Typhimurium than in *Salmonella* spp. (Table SA39). The maximum numbers of substances to which some *S*. Typhimurium are multi-resistant are the same as those in *Salmonella* spp. (Figure SA34) No isolates were resistant to both antimicrobials ciprofloxacin and cefotaxime (Table SA40).

Table SA40. Complete susceptibility, multi-resistance and index of diversity in <u>S. Typhimurium</u> fromcattlein MSs reporting isolate-based data, 2012

Country	Suscept	tible to all	Multi-r	resistant	Index of	Co-resistant to Cip and Ctx		
	n	%	n	%	diversity	N	%	
Belgium (N=25)	1	4.0	12	48.0	0.496	0 (0)	0 (0)	
Finland (N=16)	12	75.0	2	12.5	0.317	0 (0)	0 (0)	
Germany (N=35)	24	68.6	3	8.6	0.315	0 (0)	0 (0)	
Ireland (N=24)	7	29.2	16	66.7	0.157	0 (0)	0 (0)	
Sweden (N=12)	9	75.0	3	25.0	0.203	0 (0)	0 (0)	

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; n: number of isolates.

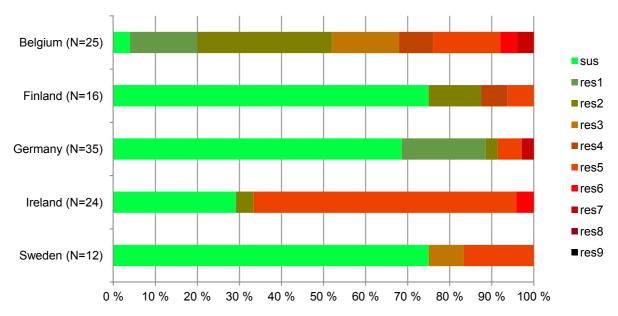
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Salmonella.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three classes from the common set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to ciprofloxacin (Cip) and cefotaxime (Ctx): the frequencies and percentages of *Salmonella* isolates not susceptible to concentrations greater than epidemiological cut-off values (Ctx:>0.5 mg/L and Cip:>0.06 mg/L). Figures in parentheses indicate the occurrence of resistance determined using the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints (Ctx:>2 mg/L and Cip:>1 mg/L).

Figure SA33. Frequency distribution of <u>S. Typhimurium</u> from <u>cattle</u> completely susceptible or resistant to one to nine antimicrobials in MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella*; sus: susceptible to all antimicrobial substances of the EFSA common set for *Salmonella*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *Salmonella*.



3.4.3. Comparison of 'clinical' and 'microbiological resistance' to ciprofloxacin

Fluoroquinolones, including ciprofloxacin, are recognised as being critically important in human medicine and often constitute the first-line treatment for invasive salmonellosis. Therefore, the high levels of ciprofloxacin resistance observed among *Salmonella* spp., from some animal species that were discussed earlier in this chapter, are of concern. Resistance levels were particularly high among *Gallus gallus* and turkeys when interpreted using the EUCAST ECOFFs.

When the data were re-analysed using the CLSI breakpoints, the resistance levels were considerably lower (Table SA41). Four countries reported very high or extremely high resistance to ciprofloxacin among *Salmonella* spp. from turkeys when using the EUCAST ECOFFS. One of the nine countries reporting more than 10 isolates detected no resistance. However, when the CLSI breakpoints were applied to analyse these data, high resistance was detected only in the Czech Republic (33.3 %), Poland (21.8 %) and Hungary (20.1 %). Among *Salmonella* spp. from *Gallus gallus*, resistance levels reached up to 69.0 % using the EUCAST ECOFFs and only three of the 17 countries reporting more than 10 isolates detected no resistance. However, using the CLSI breakpoints, resistance was only found in six countries, at low levels in Romania (8.9 %), Belgium (1.7 %), the Czech Republic (1.6 %), Hungary (1.9 %) and Spain (3.4 %) and at a very low level in Poland (0.3 %). Regarding *Salmonella* spp. from pigs, when using the EUCAST ECOFFS, one of the eight countries reporting more than 10 isolates detected no resistance. However, when the CLSI breakpoints were applied to analyse these data, low resistance was detected only in Belgium (2.7 %). Several countries reported low, moderate or high resistance among *Salmonella* spp. from cattle when EUCAST ECOFFS were used. None of these countries were found to have any resistant isolates when using the CLSI breakpoints.

The geographical distribution of the occurrence of resistance to ciprofloxacin in turkeys and the fact that this parallels the occurrence of *S*. Kentucky in farm animal species indicates how the clonal spread of one serovar can influence the overall picture.



Table SA41. Resistance (%) to ciprofloxacin among <u>Salmonella spp.</u> from <u>Gallus gallus</u>, <u>turkeys</u>, <u>pigs</u> and <u>cattle</u> in 2012, using harmonised epidemiological cut-off values or CLSI breakpoints

Country	Gallus gallus ¹		Turkeys ²			Pigs ³			Cattle ⁴			
Country	N	EUCAST % Res	CLSI % Res	N	EUCAST % Res	CLSI % Res	N	EUCAST % Res	CLSI % Res	N	EUCAST % Res	CLSI % Res
Austria	176	18.2	0	38	78.9	0	I	-	_	I	_	_
Belgium	664	39.0	1.7	Ι	-	_	187	22.5	2.7	42	28.6	0
Czech Republic	386	19.4	1.6	27	81.5	33.3	I	_	_	I	_	-
Denmark	28	0	0	-	-	-	374	0	0	_	-	-
Estonia	-	-	-	-	-	-	32	0	0	_	-	-
Finland	_	-	-	Ι	-	_	I	_	_	19	0	0
Germany	238	14.3	0	87	32.2	1.1	627	7.5	0	68	8.8	0
Hungary	261	69.0	1.9	174	91.4	20.1	38	23.7	0	_	-	-
Ireland	38	2.6	0	14	0	0	24	16.7	0	36	5.6	0
Italy	328	23.8	0	48	37.5	0	25	12.0	0	14	14.3	0
Latvia	14	0	0	-	-	-	-	_	-	_	-	-
Netherlands	192	18.2	0	-	-	-	263	0.8	0	68	2.9	0
Poland	738	39.2	0.3	55	52.7	21.8	10	60.0	0	_	_	_
Portugal	174	23.6	0	-	-	-	-	_	-	_	-	-
Romania	964	68.4	8.9	-	-	-	-	_	-	_	-	-
Slovakia	85	30.6	0	-	_	_		_	_		_	_
Spain	179	22.3	3.4	169	89.9	1.2	48	20.8	0	_	_	_
Sweden	-	_	_	_	_	_	_	_	_	17	0	0
United Kingdom	236	2.1	0	142	9.9	0		_	_	-	_	_

CLSI: Clinical and Laboratory Standards Institute; EUCAST: European Committee on Antimicrobial Susceptibility Testing; -: no data reported.

1. Gallus gallus: in Finland and Sweden, five and eight isolates were respectively sensitive to ciprofloxacin (minimum inhibitory concentration (MIC) below the epidemiological cut-off value (ECOFF)).

2. Turkeys: in Slovakia, two isolates (N=4) displayed reduced susceptibility to ciprofloxacin (MIC above both the EUCAST and CLSI thresholds), while, in Belgium, Finland and Portugal, two, one and nine isolates, respectively, were sensitive to ciprofloxacin (MIC below the EUCAST ECOFF).

3. Pigs: in Finland, Latvia and Sweden, five, three and four isolates, respectively were sensitive to ciprofloxacin (MIC below the EUCAST ECOFF).

4. Cattle: in Estonia, one isolate (N=7) displayed reduced susceptibility to ciprofloxacin (MIC above both the EUCAST and CLSI thresholds), while, in Latvia and Spain, two and nine isolates, respectively, were sensitive to ciprofloxacin (MIC below the EUCAST ECOFF).



3.4.4. Further analysis of multi-drug resistance in certain Salmonella serovars

In the paragraphs above, the 'summary indicators' of multi-drug resistance (MDR)¹⁴ have been tabulated showing notably the proportion of isolates which are fully susceptible, multi-drug resistant and co-resistant to ciprofloxacin and cefotaxime. Graphs also show the proportions of isolates resistant to different numbers of antimicrobials; corresponding data are also set out in Appendix 3. The information relating to *Salmonella* spp. from a MS often covers a variety of different serovars, each of which may have a different propensity to exhibit antimicrobial resistance and because the serovars which are prevalent in different MSs may vary, this will account for some of the pronounced variation in the recorded MDR parameters for *Salmonella* spp. which is evident in different MSs. *S.* Enteritidis in general exhibited much lower MDR than *S.* Typhimurium; however, there were marked differences between MSs in the occurrence of MDR for each of these serovars. The analysis of such patterns is most useful when applied at the serovar level, therefore MDR is considered briefly for *Salmonella* spp. and then several serovars of current importance are examined individually.

3.4.4.1. Multi-drug resistance patterns

Salmonella spp.

The patterns of resistance exhibited by all reported *Salmonella* serovars comprise aggregated data from a variety of different serovars and are presented in Appendix 4 (Appendix Tables MDRP1 to MDRP7). A range of different resistance genes, occurring in different combinations and associated with different genetic elements occurs in *Salmonella* and consequently the range of patterns obtained is large. Detailed analysis of the specific patterns of resistance detected is the most useful when it is performed at the serovar level. However, the overall data from all *Salmonella* spp. have also been examined to determine which serovars demonstrate pentavalent resistance to ampicillin, chloramphenicol, sulfonamides, streptomycin and tetracyclines. This pattern of resistance (which may occur with additional resistance) has been demonstrated by a number of *Salmonella* serovars which have spread epidemically in animals, such as *S*. Typhimurium definitive phage types 104 and 204c, as well as *S*. Newport. See Section 3.4.4.4 Analysis of pentavalent resistance.

Salmonella Enteritidis

Information on MDR was available for *S*. Enteritidis isolates from broilers and laying hens. MDR was uncommon in *S*. Enteritidis isolates, occurring in only 1.9 % of 270 isolates from broilers and 5.8 % of 206 isolates from laying hens (Appendix Tables MDRP8 and MDRP9). All MDR isolates from laying hens and the majority (4 out of 5) of MDR isolates from broilers were resistant to ciprofloxacin. Ampicillin, sulfonamide and tetracycline resistance was commonly observed as components of the MDR patterns. *S*. Enteritidis phage type 1 (PT1) from broilers commonly shows resistance to nalidixic acid and ciprofloxacin; however, other phage types (for example PT4, 7 and 21) may also demonstrate resistance to this antimicrobial, although usually less frequently (EFSA, 2007). Phage type 8 commonly showed susceptibility to the antimicrobials tested in EFSA's baseline survey of broilers in the EU in 2005-2006. Otherwise, *S*. Enteritidis showed a greatly reduced propensity to display MDR than *S*. Typhimurium did.

Salmonella Typhimurium

S. Typhimurium showed a wide diversity of MDR patterns, especially evident in fattening pigs and cattle, species from which the largest numbers of isolates were available. It is noteworthy that the commonest pattern of MDR was resistance to ampicillin, chloramphenicol, streptomycin, sulfonamides and tetracyclines in all types of animals except broilers, where it was the second most commonly observed pattern (Appendix Tables MDRP10 to MDRP13). This pattern has commonly been associated with *S*. Typhimurium definitive phage type 104 in recent years, although it can also be seen as part of the pattern in other phage types, such as DT193 and U288 (these latter types also frequently display trimethoprim resistance). Resistance to cefotaxime was not detected as part of the MDR profiles in cattle and ciprofloxacin resistance was usually uncommon in *S*. Typhimurium MDR isolates, although it was most frequent in pigs, where gentamicin resistance was also more frequently detected.

¹⁴ Multi-drug resistance is defined as resistance to any three antimicrobials in the panel recommended by EFSA – and considering resistance to nalidixic acid and ciprofloxacin as a single resistance.



Monophasic Salmonella Typhimurium

The patterns of multi-drug resistance for monophasic S. Typhimurium isolates from broilers, layers, turkeys, fattening pigs and cattle as well as meat from turkeys, pigs and cattle are shown in Appendix Tables MDRP14 to MDRP21. The most frequent pattern of resistance observed was resistance to ampicillin, streptomycin, sulfonamides and tetracyclines in all categories, except meat from turkeys, where this pattern with additional resistance to ciprofloxacin was predominant. Germany was the only MS providing data for meat from turkeys and contributed most of the isolates from turkeys; although resistance to ampicillin, streptomycin, sulfonamides and tetracyclines predominated in isolates from turkeys from this MS, this pattern with additional ciprofloxacin resistance was most commonly observed in isolates from turkey meat. More than 86 % of isolates exhibited MDR from all categories, except turkeys, where more than 70 % were MDR. The numbers of isolates received from categories other than pigs and meat from pigs were less than 20 and only two to four different resistance patterns were detected; these frequently involved acquisition of resistance to one or more of chloramphenicol, gentamicin, ciprofloxacin and trimethoprim, with or without loss of resistance to sulfonamide, ampicillin and tetracyclines. Streptomycin resistance was invariably present in the MDR isolates in all categories except from pigs and meat from pigs, where it was only absent from 5 out of 377 isolates. Among the reporting countries of isolate-based data, Germany contributed the majority of isolates for all categories, except broilers and laying hens, and the diversity of MDR patterns observed for this MS may be a reflection of the greater number of isolates examined.

Salmonella Kentucky

The patterns of MDR for *S*. Kentucky isolates from broilers, laying hens and meat from broilers are shown in Appendix Tables MDRP22 to MDRP24. More than 80 % of isolates from broilers had the core resistance pattern to ampicillin, ciprofloxacin, gentamicin, sulfonamides and tetracyclines, often with resistance to streptomycin. The core pattern with additional streptomycin resistance was also the most commonly observed pattern in meat from broilers. Although in laying hens the commonest pattern was resistance to ampicillin, ciprofloxacin and tetracyclines, the pattern of resistance to ampicillin, ciprofloxacin, gentamicin, sulfonamides and tetracyclines (in some cases with additional resistance) was also observed.

Salmonella Derby

There were very few *S*. Derby isolates reported from broilers, cattle and meat from cattle (ten or less) (Appendix Tables MDRP25 to MDRP30). *S*. Derby isolates from turkeys were 99.2 % MDR, whereas the situation for isolates from pigs and meat from pigs was markedly different with 17.4 % and 25.3 % respectively, exhibiting MDR. Resistance to streptomycin, sulfonamides and tetracyclines was the most frequently observed MDR pattern in isolates from both pigs and pig meat. Ciprofloxacin resistance was uncommon in *S*. Derby isolates from pigs occurring in only 2.4 % of the isolates for which isolate-based data was available. This contrasts with the situation in turkeys, where the most common resistance pattern was resistance to ampicillin, chloramphenicol, ciprofloxacin, sulfonamides, tetracyclines and trimethoprim, occurring in 49.6 % of isolates. Ciprofloxacin resistance was present in more than 90 % of *S*. Derby isolates from turkeys).

Salmonella Infantis

Information on MDR was available for 529 S. Infantis isolates from broilers and 162 from broiler meat (Appendix Tables MDRP33 and MDRP31). Lower numbers of isolates were available from laying hens (36), turkeys (27), fattening pigs (18) and meat from pigs (23). The number of different MDR patterns obtained was approximately in proportion to the number of available isolates from each animal and food category. A very diverse range of 43 different MDR patterns was reported for 529 S. Infantis from broilers. Resistance to ciprofloxacin was invariably present as a component of the MDR patterns in broilers and turkeys and occurred in almost all isolates from broiler meat and laying hens; it was less common in MDR isolates from pigs and pig meat. Resistance to a common core antimicrobial pattern of ciprofloxacin, sulfonamides and tetracyclines occurred in 97.5 % (158 out of 162) of S. Infantis isolates from broiler meat and 82.4 % (436 out of 529) of isolates from broilers, 77.8 % (28 out of 36) from laying hens, 92.6 % (25 out of 27) from turkeys, 39.0 % (7 out of 18) of isolates from pigs and 13.0 % (3 out of 23) of isolates from pig meat. Resistance to ciprofloxacin, streptomycin, sulfonamides and tetracyclines (a pattern associated with S. Infantis definitive phage type 213) occurred in 29.9 % (158 out of 529) of S. Infantis isolates from broilers alone or with additional resistances, while 19.4 % (7 out of 36) of isolates from laying hens, 18.5 % (5 out of 27) of isolates



from turkeys and 27.2 % (44 out of 162) of isolates from broiler meat showed this core pattern of resistance. Ciprofloxacin, streptomycin, sulfonamide and tetracycline resistance was uncommon in pigs and meat from pigs occurring in 5.6 % (1 out of 18) and 8.7 % (2 out of 23) of *S*. Infantis isolates, respectively.

3.4.4.2. Co-resistance to cefotaxime and ciprofloxacin

Table SA42 describes those *Salmonella* serovars which were co-resistant to both cefotaxime and ciprofloxacin using EUCAST clinical breakpoints. In this table, data derived from the testing of fewer than 10 isolates and from fewer than four reporting countries have been included. The table also describes the resistance patterns for such isolates (applying ECOFFs for the other antimicrobials). Co-resistance to cefotaxime and ciprofloxacin has relevance for public health, since these are frequently the two main antimicrobials used for first-line treatment of human patients with salmonellosis.

Table SA42. Co-resistance to cefotaxime and ciprofloxacin (applying EUCAST clinical breakpoints)

Source	Total number isolates for which relevant data available	Number of isolates resistant to cefotaxime and ciprofloxacin ¹ (%)	Serovar (number of isolates) and resistance pattern(s) of isolates of this serovar co- resistant to cefotaxime and ciprofloxacin ²		
Meat from			Other serovars (5) AmpCtxCipNalSuTet		
broilers	410	7 (1.7 %)	S. Infantis (1) AmpCtxChlCipNalSuTetTmp		
(Gallus gallus)			S. Rissen (1) AmpCtxChlCipNalSuTetTmp		
			S. Agona (1) AmpCtxChlCipGenNalStrSuTetTmp		
			S. Hadar (6) AmpCtxChlCipNalStrSuTmp and AmpCtxCipNalSuTetTmp		
Broilers	1,641	28 (1.7 %)	S. Infantis (14) AmpCtxCipNalSuTetTmp AmpCtxCipNalSuTet AmpCtxCipNal AmpCtxCipGenNalSuTetTmp AmpCtxCipGenNalStrSuTet and AmpCtxChlCipGenNalStrSuTetTmp		
			S. Kentucky (1) AmpCtxCipNal		
			S. Livingstone (2) AmpCtxCipGenNalStrSu and AmpCtxChlCipNalSuTetTmp		
			S. Mbandaka (1) AmpCtxCipNalSuTetTmp		
			S. Senftenberg (2) AmpCtxChlCipGenNalSu		
			S. Thompson (1) AmpCtxCipNalSuTetTmp		
Laying hens	686	2 (0.3 %)	S. Kentucky (2) AmpCtxCipNal		
Turkeys	567	1 (0.2 %)	S. Derby (1) AmpCtxChlCipNalStrSuTetTmp		
Fattening pigs and unspecified			S. Derby (4) AmpCtxCipNalStrSuTmp		
	1,155	5 (0.4 %)	S. Typhimurium, monophasic (1) AmpCtxCipNalStrSuTetTmp		
Cattle (bovine animals)	212	1 (0.5 %)	S. Livingstone (1) AmpCtxCipNalStrSuTetTmp		

1. European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints were applied for cefotaxime (R>2 mg/L) and ciprofloxacin (R>1 mg/L).

2. EUCAST epidemiological cut-off values were applied for antimicrobials other than cefotaxime and ciprofloxacin.



Considering the Salmonella serovars showing co-resistance to cefotaxime and ciprofloxacin, multi-drug resistant S. Infantis with co-resistance to cefotaxime and ciprofloxacin was infrequent among Salmonella isolates on broiler meat, comprising 0.2 % (1 out of 410) of the total available; however, it was the most common serovar which exhibited co-resistance at the clinical breakpoint for cefotaxime and ciprofloxacin in broilers, where it accounted for 0.8 % (14 out of 1,641) of the total isolates from broilers. Other serovars exhibiting co-resistance generally occurred at a low frequency and were detected as one or two MDR isolates, apart from S. Hadar in broilers (6 out of 1,641 isolates) and S. Derby in pigs (4 out of 1,155 isolates). The same co-resistant serovar (S. Infantis) was therefore detected in broilers and broiler meat. Coresistance to cefotaxime and ciprofloxacin was most frequently observed in Salmonella serovars from broilers, affecting 1.7 % of isolates and involving eight different serovars. Co-resistance affected less than 1 % of Salmonella isolates from other categories of livestock. Isolate-based data were available for fewer isolates from cattle (212) than for the other species and a single MDR isolate of S. Livingstone was detected from cattle exhibiting co-resistance to cefotaxime and ciprofloxacin. Interestingly, the resistance pattern observed (AmpCtxCipNalStrSuTetTmp), but without trimethoprim resistance was also observed in an isolate of S. Livingstone from broilers, suggesting possible transmission of this MDR serovar between these livestock sectors.

3.4.4.3. Analysis of high-level ciprofloxacin resistance

High-level resistance to ciprofloxacin in *Salmonella* of animal and food origin, as well as the serovars displaying such resistance, are shown in Appendix 5. A variety of serovars displayed high-level ciprofloxacin resistance and were frequently resistant to other antimicrobials.

Considering those serovars occurring most frequently and in a number of different MSs and types, *S*. Kentucky was represented in isolates from Romania, the Czech Republic, Spain and Hungary from broilers from Romania, Spain and Hungary from laying hens and from the Czech Republic, Spain and Hungary from turkeys. It was also detected in single isolates from meat from turkeys in the Czech Republic and from meat from broilers in Ireland. *S*. Infantis was the second most frequently encountered in a number of different MSs. A single isolate of *S*. Typhimurium showing high-level ciprofloxacin resistance was reported from meat from pigs.

A particular clone of S. Kentucky sequence type 198 with high-level ciprofloxacin resistance has been detected in Africa and the middle East (Le Hello et al., 2011) and has been subsequently detected in poultry in some European countries (Wasyl and Hoszowski, 2012). S. Kentucky showing such high-level resistance to ciprofloxacin was detected in several MSs in 2012 and from a number of different sources. The occurrence of this serovar/phenotype is only evident in MSs reporting isolate-based data. The MSs detecting such isolates in broilers include Romania (8.2 %; 64 out of 781), the Czech Republic (1.4 %; 5 out of 351), Spain (13.8 %; 4 out of 29) and Hungary (0.6 %; 1 out of 175). In laying hens, it was detected in Romania (2.7 %; 4 out of 146), Spain (1.3 %; 2 out of 150) and Hungary (2.3 %; 2 out of 86) and in turkeys in the Czech Republic (33.3 %; 9 out of 27), Spain (1.2 %; 2 out of 169) and Hungary (19.0 %; 33 out of 174). The Czech Republic detected S. Kentucky with high-level ciprofloxacin resistance in meat from turkeys (1 out of 10; 10.0 %), while Ireland detected it in meat from broilers (1.4 %; 1 out of 70). S. Kentucky with high-level ciprofloxacin resistance has therefore been reported from a larger number of MSs in 2012 than was the case in 2011 and is now being detected in all three types of poultry, which are monitored (broilers, laying hens and turkeys), as well as in broiler and turkey meat. Its occurrence may be under-estimated because not all MSs report isolate-based data and some MSs have reported the occurrence of unspecified serovars which possess this phenotype. Genotypic typing data were not available; therefore, the isolates with this phenotype cannot be definitively assigned to the ST198 clone of S. Kentucky.

Salmonella serovars displaying high-level ciprofloxacin resistance were reported in poultry only and were not detected in fattening pigs or cattle in the reporting MSs. However, single isolates of *S*. Typhimurium and *S*. Infantis displaying such resistance were recovered from pig meat in two MSs and these MSs did not examine isolates from fattening pigs. The diversity of serovars displaying high-level ciprofloxacin resistance in poultry differed between MSs and was greatest in Romania which detected it in the serovars Agona, Infantis, Kentucky, Liverpool, Livingstone, Rissen and Tennessee in broilers and in the serovars Albany, Corvallis, Hadar, Kentucky and Thompson from laying hens. Hungary also detected high-level resistance to ciprofloxacin in serovars other than Kentucky; namely in *S*. Stanley from turkeys, *S*. Saintpaul from laying hens and *S*. Infantis from broilers. High-level resistance to fluoroquinolones in *Salmonella* generally involves multiple mutations of the DNA gyrase genes which are chromosomally located and therefore not transferable between bacteria in bacterial conjugation. The results suggest that clonal spread of *S*. Kentucky with high-level ciprofloxacin resistance may be occurring in EU MSs; however, genetic confirmation would be required

to substantiate this observation. Moreover, a range of different serovars showing high-level resistance to ciprofloxacin is emerging in some MSs and this may be the result of the selective pressure of usage of fluoroquinolones.

S. Infantis with high-level ciprofloxacin resistance was detected in several MSs, raising the possibility that this serovar is also spreading clonally in poultry and possibly other species in Europe. In total, 14 isolates were recovered from Romania (broilers and broiler meat), Hungary (broilers) and the Czech Republic (pig meat). In addition to high-level ciprofloxacin resistance, all isolates were also resistant to sulfonamides and tetracyclines, with some possessing a range of additional resistances. Genetic examination would be required to investigate whether this represents clonal expansion of a high-level ciprofloxacin-resistant strain or multiple independent evolutions of several different strains. S. Infantis resistant to nalidixic acid, streptomycin, sulfonamides and tetracyclines belonging to definitive phage type 213 has been reported from several European countries (Nógrády et al., 2012). This pattern of resistance has also been noted in S. Infantis isolates from broilers in Iran (Rahmani et al., 2013), while resistance to nalidixic acid, tetracyclines and nitrofurantoin has been reported in Israel (Gal-Mor et al., 2010).

3.4.4.4. Analysis of pentavalent resistance

The occurrence of such pentavalent resistance in different serovars is shown in Table SA43. Some such serovars were detected sporadically with only one reported instance (e.g. *S.* Wisbech, *S.* Kapemba) while others were detected at higher frequencies (e.g. *S.* Infantis) and sometimes in several different types of livestock (*S.* Infantis). In view of the observation that pentavalent resistance has been shown by many of the serovars which have become highly prevalent over the last few decades; describing these serovars in this way may provide an early indication of newly emerging epidemic strains.

Origin	Serovars detected with pentavalent ¹ resistance
Meat from broilers	S. Java (2), S. Infantis (1), S. Typhimurium (3)
Meat from pigs	S. enterica subsp. enterica (2), S. Typhimurium (33), S. 1,4,[5],12:i:- (3), S. Kapemba (1), S. 4,12:i:- (1), S. Infantis (1), S. Typhimurium, monophasic (1), other serovars (1)
Meat from turkeys	Other serovars (1)
Broilers	S. Agona (1), S. Infantis (10), S. Liverpool (5), S. Tennessee (2), S. Senftenberg (2), S. Kentucky (4), S. Typhimurium (3), S. Saintpaul (1)
Laying hens	S. Typhimurium (4), S. Kentucky (1), S. Infantis (1)
Turkeys	S. 1,4,[5],12:i:- (1), S. Agona (1), S. Derby (13), S. Hadar (1), S. Typhimurium (1), S. Wisbech (1), S. London (3), S. Anatum (1)
Fattening pigs	S. Typhimurium (112), S. 1,4,[5],12:i:- (20), Other serovars (7), S. <i>enterica</i> subsp. <i>enterica</i> (2), S. 4,12:i:- (1), S. Typhimurium, monophasic (2), S. Choleraesuis (1), not typeable (1)
Cattle	S. Typhimurium (28), S. Rissen (1)

Table SA43. Salmonella serovars detected with pentavalent resistance amongst those for which isolate-based data is available

1. Resistance to ampicillin, chloramphenicol, streptomycin, sulfonamides and tetracyclines.

S. Infantis features as a serovar displaying pentavalent resistance, occurring at a relatively higher frequency in some animal categories (and in more than one different type of animal), as well as showing high-level ciprofloxacin resistance. It has also been detected in more than one MS. The detection of high-level ciprofloxacin resistance, associated with sulfonamide and tetracycline or pentavalent resistance in S. Infantis warrants further investigation to determine the relationship between those isolates currently occurring in animals and human as well as the relation to isolates detected in previous years.

Pentavalent resistance in *Salmonella* serovars can be related to the presence of the genetic structure known as *Salmonella* genomic island 1 (SGI1), which has been detected in a number of different serovars including Agona, Java, Albany and Newport in addition to *S*. Typhimurium (Velge et al., 2005). Recent genetic analysis (Beutlich et al., 2011) has shown the occurrence of several variations of the basic pattern of pentavalent resistance associated with SGI1 in European *Salmonella* isolates, related to the deletion and acquisition of different resistance genes. Thus, the 'classical' resistance pattern associated with SGI1 is conferred by the resistance genes *bla*_{PSE-1}, *floR*, *sul1*, *tet*(G) and *aadA2*, whereas the variant SGI-1A occurring in *S*. Derby, has these genes together with *dfrA10*, conferring trimethoprim resistance. SGI-1C



(also detected in *S*. Derby) has a deletion such that only *aadA2* and *sul1* remain, conferring resistance to streptomycin and sulfonamides. SGI-1M was a variant detected in *S*. Typhimurium with *bla*_{PSE-1}, *floR*, *sul1*, *tet*(G) and *aadB*, the latter gene conferring gentamicin resistance. SGI1-K1 is a variant of SGI-1 described in *S*. Kentucky ST198 (Le Hello et al., 2011) showing high-level ciprofloxacin resistance, which also carries gentamicin resistance (conferred by *aacA5*), as well as genes conferring resistance to ampicillin, sulfonamides, streptomycin and tetracyclines. This MDR pattern (with additional resistance to ciprofloxacin) was that most frequently observed in *S*. Kentucky isolates from broilers and was also observed in isolates from laying hens and meat from broilers. Another pattern detected in *S*. Kentucky with high-level ciprofloxacin resistance was also evident; namely ampicillin, sulfonamide and tetracycline resistance, which has been associated with the *Salmonella* genomic island variant SGI1-K2.

MDR was uncommon in **S. Enteritidis** isolates, occurring in only 1.9 % of 270 isolates from broilers and 5.8 % of 206 isolates from laying hens, a marked contrast to the position in *S*. Typhimurium. All MDR isolates from laying hens and the majority (4 out of 5) MDR isolates from broilers were resistant to ciprofloxacin; resistance to ampicillin, sulfonamides and tetracyclines was also generally observed in MDR isolates. There were no *S*. Enteritidis isolates recovered from animals which were resistant to the antimicrobials ampicillin, chloramphenicol, sulfonamides, streptomycin, tetracyclines and trimethoprim. This resistance phenotype has been associated with a combined virulence–resistance plasmid in *S*. Enteritidis (a hybrid plasmid carrying both virulence and resistance determinants) and allows increased intra-cellular proliferation resulting in more severe human infections; an African origin was suspected in a European study of several isolates from humans (Rodriguez et al., 2012). This resistance phenotype might occur through other genetic configurations and cannot be used to confirm the presence of the combined virulence–resistance plasmid. However, it serves to illustrate how the data may be used to assess the possible incursion of such strains into the EU and to flag isolates for possible further investigation.

S. Typhimurium displayed a diverse range of resistance patterns, including patterns which can be associated with particular definitive phage types, such as ampicillin, chloramphenicol, streptomycin, sulfonamide and tetracycline resistance (associated with DT104) and this core pattern plus additional resistance to trimethoprim (which can be associated with phage types such as DT193 and U288). Resistance to ampicillin, chloramphenicol, gentamicin, sulfonamides and tetracyclines (the 'SGI-1M pattern', see above), albeit with additional resistances, was observed in 3.2 % (11 out of 341) of S. Typhimurium isolates from pigs. Cefotaxime resistance was not observed as part of MDR patterns in S. Typhimurium and resistance to ciprofloxacin was relatively uncommon. The wide range of resistance patterns detected may reflect the considerable diversity of strain types exhibited by this serovar (which may be a very old serovar) and consequently differences in the acquisition or ability to acquire resistance genes (Lan et al., 2009). Some of the definitive phage types of *S*. Typhimurium have been considered to have complex origins (Lan et al., 2009) and therefore might also differ in their propensity to develop or acquire resistance.

The majority of monophasic S. Typhimurium demonstrated the typical pattern of 'tetravalent' resistance to ampicillin, streptomycin, sulfonamides and tetracyclines, with this pattern plus resistance to ciprofloxacin prominent in isolates from turkey meat. The core pattern with additional resistance to trimethoprim was the second most commonly observed pattern in isolates from fattening pigs. The relative frequency of occurrence of the different patterns is likely to be biased by the relative contributions from individual MSs, because different MSs have reported variations in the patterns of resistance in this serovar. Monophasic S. Typhimurium with chromosomally encoded resistance to ampicillin, streptomycin, sulfonamides and tetracyclines have become particularly common in numerous EU countries since 2000, while Spanish U302based monophasic strains have also been found to be multi-resistant, most frequently to ampicillin, chloramphenicol, gentamicin, streptomycin, sulfonamides, tetracyclines and trimethoprim (EFSA BIOHAZ Panel, 2010b). Strains showing resistance to ampicillin, streptomycin, sulfonamides and tetracyclines carrying the resistance genes blaTEM, strA-strB, sul2 and tet(B) (generally belonging to phage definitive types 120 or 193) have been considered the 'European' clone and carry Salmonella genomic island 2. Two other lineages have been described, the 'Spanish' clone referred to above and the 'US' clone, which is generally susceptible (Bugarel et al., 2013). A recent study of isolates from diverse sources in France ascribed 71 % of isolates to the Spanish clone and 2 % to the US clone (Bugarel et al., 2013). The pattern of resistance to ampicillin, chloramphenicol, gentamicin, streptomycin, sulfonamides, tetracyclines and trimethoprim was only described in 1.8 % (5 out of 279) of isolates from fattening pigs, whilst the pattern associated with the European clone was much more frequently detected. Resistance to cefotaxime as part of MDR was observed in 1.1 % of isolates from fattening pigs (3 out of 279) and one of these monophasic S. Typhimurium isolates was resistant to all antimicrobials in the test panel.



Ten isolates were detected which were resistant to all nine antimicrobials in the test panel, including the single monophasic *S*. Typhimurium isolate from fattening pigs (0.1 %; 1 out of 983) and a single isolate of *S*. Agona from turkeys (0.2 %; 1 out of 567). In broilers, 0.5 % (8 out of 1 659) of *Salmonella* isolates were resistant to all of the antimicrobials in the test panel, comprising the serovars Infantis (two), Kentucky (two), Agona, Liverpool, Senftenberg and Tennessee. The two *S*. Kentucky isolates, one of the Infantis isolates and the Agona isolate demonstrated high-level ciprofloxacin resistance with MICs >4 mg/L.

There were only very few S. Derby isolates from broilers, cattle and meat from cattle (10 or less). MDR to streptomycin, sulfonamides and tetracyclines which was observed in both of only two S. Derby isolates from cattle was the commonest pattern observed in S. Derby isolates from fattening pigs. Different MSs were involved, but, where a MDR pattern and serovar are common in one species and the same pattern but at a lower frequency is observed in another species, spill over from a major reservoir host to a minor one is a possible explanation. The findings at EU level relating to pigs are therefore broadly consistent with a French study of isolates from pigs, pig meat and humans which found that 69.2 % were simultaneously resistant to streptomycin, sulfonamides and tetracyclines (Kerouanton et al., 2013). In Spain, 85.7 % of porcine S. Derby isolates were resistant to tetracyclines, 85.7 % to streptomycin, 71.4 % to sulfonamides and 50 % to ampicillin, whilst 42.8 % showed trimethoprim/sulfonamide resistance and 7.1 % were resistant to nalidixic acid (Valdezate et al., 2005). These prevalences of resistance are consistent with the MDR patterns observed. Isolates of S. Derby from pigs were markedly different from those in turkeys in relation to their tendency to display MDR (99.2 % in turkeys; 17.4 % in pigs) and the extent to which ciprofloxacin is a component of MDR patterns (2.3 % in MDR isolates from pigs; more than 90 % in MDR isolates from turkeys). SGI1-A has been described in S. Derby and comprised resistance to ampicillin, chloramphenicol, streptomycin, sulfonamides and tetracyclines ('pentavalent' resistance) with additional resistance to trimethoprim. This pattern was observed in isolates from turkeys and - with additional resistance to ciprofloxacin - was the most commonly observed MDR pattern in that animal species.

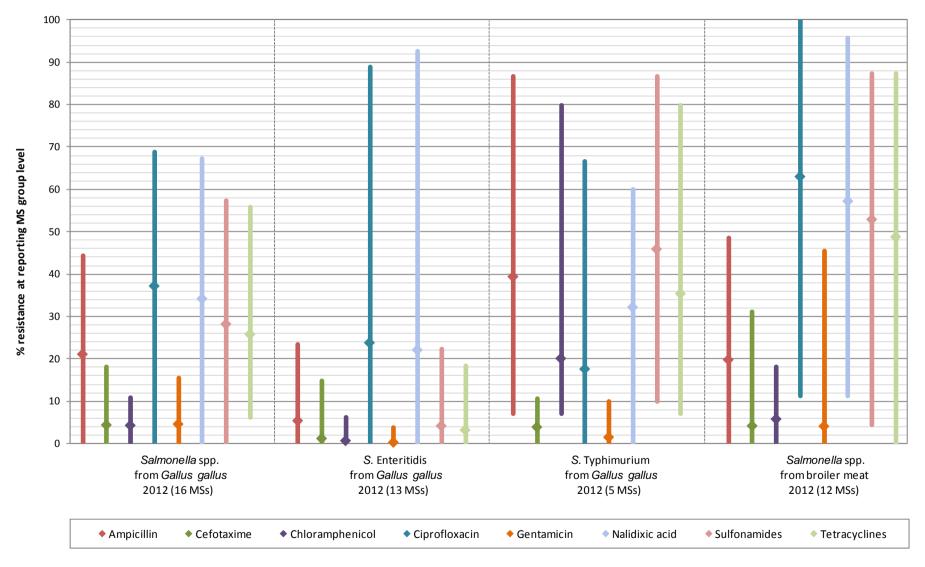
3.5. Overview of the findings of antimicrobial resistance in *Salmonella* at Member State reporting group level, 2012

Figures SA35 and SA36 illustrate the resistance levels for the groups of MSs reporting quantitative MIC data in 2012. These data were not all derived from the same group of MSs, which needs to be considered when interpreting these figures. Resistance levels to ampicillin, chloramphenicol, sulfonamides and tetracyclines in *S*. Typhimurium from *Gallus gallus* were higher than in *S*. Enteritidis from *Gallus gallus*. However, resistance to ciprofloxacin and nalidixic acid was higher in *S*. Enteritidis than in *S*. Typhimurium. In terms of all *Salmonella* spp., resistance levels in isolates from broiler meat were higher than those in isolates from *Gallus gallus*. This represents a return to the pattern observed in 2009 with an increase in resistance in isolates from broiler meat compared with the levels reported in 2010.

In a very similar pattern to that observed in 2010, resistance levels to tetracyclines, sulfonamides and ampicillin were higher in *Salmonella* isolated from turkeys, pigs and cattle than in isolates from *Gallus gallus*, whereas, for ciprofloxacin and nalidixic acid, the highest resistance was observed in turkeys and in *Gallus gallus*. The levels of resistance to sulfonamides and tetracyclines in isolates from turkeys decreased in 2012, whereas the levels of resistance to these antimicrobials in isolates from pigs increased compared with 2010.



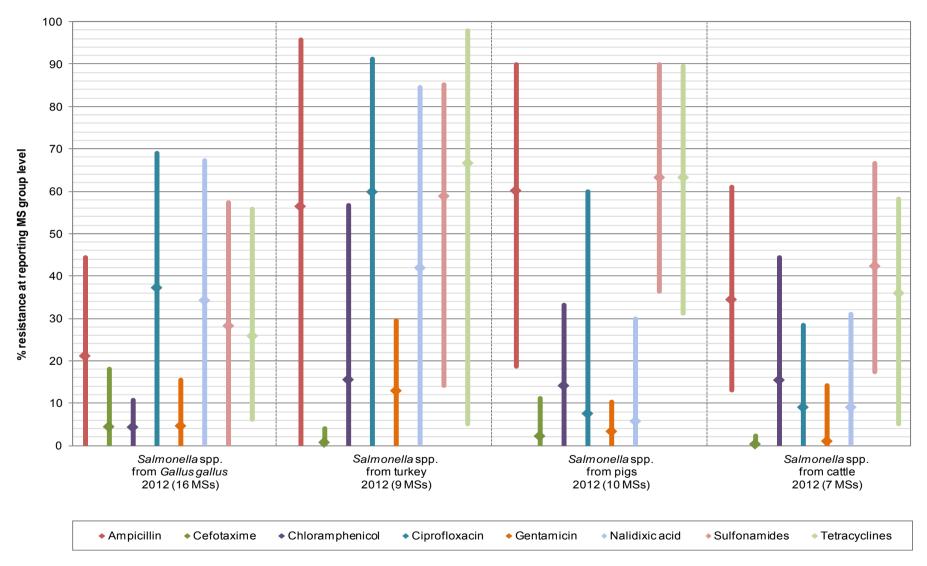
Figure SA34. Resistance to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines in <u>Salmonella spp., S. Enteritidis and S. Typhimurium</u> from <u>Gallus gallus</u> and <u>Salmonella spp.</u> from <u>meat from broilers</u> at reporting MS group level in 2012



MS: Member State



Figure SA35. Resistance to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines in <u>Salmonella spp.</u> from <u>Gallus gallus</u>, <u>turkey</u>, <u>pigs</u> and <u>cattle</u> at reporting MS group level in 2012





3.6. Discussion

Salmonellosis continues to be the second most commonly reported zoonotic disease in humans in the EU, exceeded only by campylobacteriosis, although there has been a significant decline in human salmonellosis cases over the period 2007 to 2012. This decrease is assumed to be mainly the result of a reduction in *Salmonella* prevalence in flocks of laying hens, broilers and turkeys, probably as a result of the national control and monitoring programmes implemented by the MSs in the corresponding production sectors (EFSA and ECDC, 2014).

In 2012, information on antimicrobial resistance in *Salmonella* isolates from human cases was reported by 19 MSs and two non-MS. The number of isolates submitted by these countries corresponded to one-quarter of the salmonellosis cases reported within the EU in 2012 and is considered a representative sample. MSs not reporting antimicrobial resistance data are, however, still encouraged to do so, in order to achieve the best possible assessment of the levels of antimicrobial resistance in human *Salmonella* isolates in the EU. A novelty in this year's report was that isolates from cases notified as having been acquired during travelling outside of the reporting country were excluded from all analyses except the analysis of resistance in difference geographical regions. This was done to better assess the impact on *Salmonella* isolates from food consumed within each reporting country on the antimicrobial resistance levels found in human isolates in that country. Please note, however, that imported food, which can constitute a large proportion of the food available in some countries, is not covered by this report.

Resistance in human *Salmonella* isolates was high for ampicillin, sulfonamides, streptomycin and tetracyclines and moderate for nalidixic acid. These first four compounds are antimicrobials that are or have been commonly used for treatment of animals and formerly were commonly used in humans. For these four antimicrobials, the important resistance observed was largely the result of the high to extremely high resistance levels observed among *S*. Typhimurium and particularly monophasic *S*. Typhimurium isolates. This corresponding resistance pattern (ASSuT) is that which is most commonly observed among the emerging monophasic *S*. Typhimurium definitive phage type 193 out of 120 strains (EFSA BIOHAZ Panel, 2010b). In contrast, resistance to the clinically important antimicrobials, **ciprofloxacin** and **cefotaxime**, was relatively low among the isolates tested. However, levels of resistance to ciprofloxacin were generally higher in countries using ECOFFs or other sensitive breakpoints, as both CLSI and EUCAST clinical breakpoints for ciprofloxacin resistance are significantly higher than the ECOFF. Resistance to quinolones (ciprofloxacin and nalidixic acid) was also generally higher in *S*. Enteritidis isolates than in *S*. Typhimurium isolates of human origin.

There was a rather poor correlation between ciprofloxacin and nalidixic acid resistance levels observed in human isolates in 2012. This could possibly be the result of the problem of detecting low-level ciprofloxacin resistance in *Salmonella* when using disc diffusion. In the ECDC external quality assurance scheme for *Salmonella*, it was also concluded that the use of different, sometimes non-standardised, interpretive criteria resulted in deviating results, rather than a lack of accuracy or performance in the laboratories (ECDC, 2012). In the EU protocol for harmonised monitoring of antimicrobial resistance in human *Salmonella* and *Campylobacter* isolates, ECDC is therefore promoting the use of EUCAST methods and breakpoints in all laboratories submitting AST results to TESSy, and advise laboratories using disc diffusion for ciprofloxacin to also test for nalidixic acid (ECDC, 2014).

About 30 % of human *Salmonella* spp. isolates from the 13 MSs which tested all antimicrobials in the test panel exhibited **multi-drug resistance**, meaning that they were clinically non-susceptible to at least three different antimicrobial classes. Two MSs recorded MDR levels above 50 %. However, about half of all isolates tested were susceptible to the complete range of antimicrobials reported for humans. Co-resistance to the critically important antimicrobials, ciprofloxacin and cefotaxime, was low and observed in a total of 27 isolates (out of 13,496 isolates tested) reported by five of the 12 reporting MSs.

The MDR levels observed in human isolates were sometimes higher than those observed in animals and meat. This can be explained by the fact that both clinically resistant and clinically intermediate results were combined to estimate MDR in human isolates and the clinical breakpoints for only 4 out of 10 antimicrobials (cefotaxime and ciprofloxacin regarding both CLSI and EUCAST, gentamicin and trimethoprim regarding only CLSI) had higher MIC values (i.e. being less sensitive) than the ECOFFs and for two antimicrobials (chloramphenicol and tetracyclines) the relationship was the reverse, i.e. the clinical breakpoint was more sensitive than the ECOFF. This resulted in a lower than expected difference in MDR estimates between human and animal/food isolates.

A striking observation was that many human isolates were resistant to a large number of antimicrobials, some even to all 10. This could reflect the impact of use of antimicrobials in humans, in addition to that in food-producing animals, or exposure to sources of *Salmonella* other than those associated with food-producing animals. A proportion of the *Salmonella* infections occurring in humans in a given MS is likely to be associated with food-producing animals in that MS, while other infections will be associated with a variety of other sources, including imported foods and spices, foreign travel, other types of animals (such as pet reptiles) or the environment. Some human infections can also occur through spread between affected human patients. As mentioned above, travel-associated cases (where the information was provided) were excluded from the analysis in order to reduce this bias.

In order to assess the importance of **travel-associated infections**, antimicrobial resistance was also analysed based on the most likely country of infection and aggregated by geographical region. Overall, human *Salmonella* spp. isolates acquired within the EU/EEA countries exhibited greater resistance to ampicillin, streptomycin, sulfonamides and tetracyclines than isolates from any other region, while the highest levels of resistance to ciprofloxacin were observed in isolates acquired from Asia, Africa and European countries outside of the EU/EEA.

In *Salmonella* isolates from animals and meat, information on antimicrobial resistance was reported by 19 MSs and two non-MS in accordance with EFSA's recommendations (EFSA, 2007) in 2012. The (quantitative) MIC results obtained using the methods recommended by EFSA provided the most harmonised and comparable set of data for reporting MSs, and these datasets have therefore been analysed in detail. For the second time, this EU Summary Report has examined the levels of resistance in isolates within different production types of animal species. Differences in animal husbandry and physiological differences between animals involved in different production types (e.g. fattening veal calves and dairy cattle) make evaluation of the antimicrobial resistance results at the animal species level difficult, where the production types of the species in question are not comparable. Sub-division of resistance data allows for more accurate analysis; however, this is possible only where sufficient information on production type has been submitted. In 2012, the large number of MSs providing data on isolates from *Gallus gallus* by production type allowed for more accurate analysis. However, more information is required at the production level for other animal species, particularly cattle, to improve these sections of the report in future years.

As mentioned above, some antimicrobials such as **ampicillin**, **sulfonamides** and **tetracyclines** have been widely used for many years in veterinary medicine to treat bacterial diseases. The levels of resistance to these antimicrobials are generally moderate to high among isolates from food-producing animals and meat products thereof. For ampicillin, sulfonamides and tetracyclines, as well as chloramphenicol, resistance levels were highest in isolates from pigs, followed closely by isolates from turkeys and then cattle. Isolates from *Gallus gallus* displayed the least resistance to these antimicrobials within the reported data, but were still at moderate to high levels. Considering the production level data for *Salmonella* spp. and *S*. Entertitidis from *Gallus gallus*, higher levels of resistance were generally observed among isolates from broiler flocks than in isolates from laying hen flocks. This was particularly evident for tetracyclines and sulfonamides. This may reflect the relative infrequency with which laying hens are treated with antimicrobials compared with broilers, as well as the limited numbers of antimicrobial compounds which are authorised for the treatment of laying hens in many EU MSs.

The highest occurrence of resistance to **ciprofloxacin** was noted in *Salmonella* from turkeys, fowl (*Gallus gallus*) and broiler meat. The ciprofloxacin resistance level for the *Gallus gallus* species can be further subdivided into production types and reveals a difference between *Salmonella* isolates from laying hens (19.4 % resistance) and broilers (46.3 % resistance). An equal number of MSs had significant increasing and decreasing national trends for ciprofloxacin and/or nalidixic acid resistance in *Salmonella* spp. isolates from *Gallus gallus* over the 2006 to 2012 period. These observations relating to *Salmonella* spp. may reflect the occurrence of *S*. Entertitidis phage type 1 in *Gallus gallus* within these MSs, since this phage type commonly displays resistance to nalidixic acid and ciprofloxacin. Similarly, in turkeys, the dissemination of certain serovars, such as *S*. Newport and *S*. Infantis, in some MSs, which, again, are commonly resistant to nalidixic acid and ciprofloxacin, may affect the overall levels of resistance among all *Salmonella* spp. In addition, the reporting of resistance results for an expanded number of individual serovars in this report enables some of the resistances which are associated with particular serovars to be clearly seen.

Resistance to third-generation cephalosporins, such as **cefotaxime**, was detected in *Salmonella* isolates from turkeys, fowl (*Gallus gallus*), pigs, cattle and the meat derived from broilers and pigs, but at low or very low levels when all reporting MSs were considered. However, there was some variability in third-generation cephalosporin resistance observed between the different animal or meat origins in the reporting MSs.



Belgium, Poland and Portugal detected cefotaxime resistance in *S*. Enteritidis from *Gallus gallus* in 2012, whereas, in 2011, Austria and Hungary reported resistant isolates and, in 2010, cefotaxime resistance in *S*. Enteritidis from *Gallus gallus* was reported only by the Czech Republic. As *S*. Enteritidis is one of the main serovars affecting humans, the emergence of resistance to third-generation cephalosporins is extremely undesirable.

Antimicrobial resistance in certain *Salmonella* serovars and phage types may be related not only to the selective pressure exerted by the use of antimicrobials, but also to the clonal diffusion of these *Salmonella* serovars and phage types, and may also be influenced by factors such as on-farm hygienic management and animal movements and trade. It was evident in both humans and animals that isolates of *S*. Typhimurium displayed higher levels of resistance than isolates of *S*. Entertitidis to ampicillin, chloramphenicol, streptomycin, sulfonamides and tetracyclines. This is usually observed among the data reported by MSs and is not surprising, since certain phage types of *S*. Typhimurium have an associated pattern of pentavalent resistance to these antimicrobials. The same pattern was observed in 2011, whereas in 2010 the majority of significant trends were increasing. Ultimately, it would be most useful to correlate trends and the occurrence of antimicrobial resistance with the usage of antimicrobial compounds in each animal production type that is monitored.

The **multi-drug resistance levels** (proportions of isolates showing reduced susceptibility to more than three antimicrobial classes according to ECOFFs) in *Salmonella* spp. isolates were generally high in the animal populations investigated, with notable variations between reporting countries. A striking exception to this is the multi-resistance levels recorded in isolates from laying hens, which are generally low to moderate, in particular compared with those observed in isolates from broilers. Generally, the proportions of *Salmonella* spp. isolates susceptible to all or resistant (or non-susceptible) to any one of up to nine antimicrobials differed substantially among the reporting countries, and the relative contribution of different serovars, which may exhibit particular MDR patterns, should be borne in mind when comparing the situation between the reporting countries. The occurrence of co-resistance to cefotaxime and ciprofloxacin (determined using ECOFFs) differed between MSs and was not detected in isolates from the majority of MSs reporting isolate-based data. In the MSs where it was detected, co-resistance to these antimicrobials in *Salmonella* spp. occurred at very low to moderate levels in isolates from broiler meat, broilers, layers, turkeys, pigs and cattle. Applying clinical breakpoints, co-resistance to cefotaxime and ciprofloxacin was detected in *Salmonella* isolates from meat from broilers and broilers. Co-resistance using ECOFFs was detected for a single *Salmonella* isolate from meat from pigs.

An additional goal of the monitoring programme is to highlight strains of Salmonella which may be emerging in several MSs and which have particular patterns of resistance or show specific resistances of importance. Detailed analyses of multi-drug resistance patterns in Salmonella serovars, including analysis of highlevel resistance to ciprofloxacin and pentavalent resistance and investigation of co-resistance to both cefotaxime and ciprofloxacin, was possible for MSs reporting isolate-based data and is included for the first time in this report. This is particularly important for the antimicrobial ciprofloxacin, where isolates can develop increasing resistance in a step-wise and incremental manner. Pentavalent resistance (i.e. resistance to ampicillin, chloramphenicol, streptomycin, sulfonamides and tetracyclines) is a characteristic shown by several serovars which have become highly prevalent over the last few decades. High-level ciprofloxacin resistance was observed in a limited number of S. Kentucky isolates from broilers, laying hens and turkeys, but not in isolates from pigs or cattle, although it was detected in isolates from pig meat. S. Infantis showing high-level ciprofloxacin resistance, together with resistance to a core antimicrobial pairing of sulfonamides and tetracyclines (but with other resistances which were not invariably present in all high-level ciprofloxacin Infantis isolates), was detected in meat from broilers, pig meat and broilers. The occurrence of S. Infantis with this resistance pattern in several MSs and in different types of animal or meat probably indicates that either a clone of S. Infantis showing such resistance has spread within Europe or several clones have gained such high-level ciprofloxacin resistance independently. In addition to showing high-level ciprofloxacin resistance, S. Infantis also featured as a serovar displaying 'pentavalent' resistance, that is resistance to ampicillin, chloramphenicol, streptomycin, sulfonamides and tetracyclines. A small number of serovars displayed pentavalent resistance which is potentially significant because certain Salmonella serovars which have shown epidemic spread have shown such pentavalent resistance in the past. Therefore, describing those serovars which exhibit pentavalent resistance, their frequency of occurrence and whether they are distributed across one or several different types of animals or MSs is likely to provide useful information on new and emerging MDR serovars.



4. ANTIMICROBIAL RESISTANCE IN CAMPYLOBACTER

4.1. Introduction

ZOONOTIC CAMPYLOBACTER

The Campylobacter species most commonly associated with human infection are C. jejuni followed by C. coli and C. lari, but other species are also known to cause infections in humans. The infective dose of these bacteria is generally low.

The incubation period in humans ranges from two to five days. Patients may experience mild to severe symptoms, commonly including watery, sometimes bloody, diarrhoea, abdominal pain, fever, headache and nausea. Infections are usually self-limiting and last only a few days; treatment with antimicrobials is therefore usually not required. Extra-intestinal infections, invasive infections or post-infection complications such as reactive arthritis and neurological disorders can occur, but these are infrequent. C. jejuni is a recognised antecedent cause of Guillain–Barré syndrome, a form of paralysis that can sometimes result in dysfunction of the respiratory and neurological systems and can even be fatal.

Thermotolerant Campylobacter species are widespread in nature. The primary reservoirs are the alimentary tract of birds and mammals including food-producing animals (poultry, cattle, pigs and sheep). Campylobacter species have been isolated from pet animals, including cats and dogs, from wild birds, from water and from various environmental samples. Clinical disease resulting from infection with thermotolerant Campylobacter species is rare in animals.

Campylobacter can readily contaminate various food-stuffs including meat, particularly poultry meat, raw milk and dairy products, and less frequently fish and fish products, mussels and fresh vegetables. Considering sporadic human cases, consumption of poultry meat, drinking water from untreated water sources and contact with pets and other animals have been identified as significant and major sources of infection. Drinking water contaminated with Campylobacter has caused large outbreaks.

Campylobacteriosis continues to be the most commonly reported zoonosis in humans in the EU. In 2012, the number of confirmed cases of *Campylobacter* reported in the EU however decreased by 4.3 % compared with 2011. The EU notification rate of confirmed cases of human campylobacteriosis shows a statistically significant increasing trend in the last five years, 2008 to 2012 (EFSA and ECDC, 2014). In 2012, about one quarter of the fresh broiler meat samples were reported positive, even though there were large differences between the MSs (EFSA and ECDC, 2014). In reporting countries, the prevalence of campylobacteriosis in broiler flocks remained mainly at levels similar to previous years (EFSA and ECDC, 2014).



4.2. Overview of reported data in humans, animals and food

Fourteen MSs and Iceland provided data for 2012 on *Campylobacter* isolates from human cases. These countries reported qualitative data, i.e. interpreted antimicrobial susceptibility testing (AST) results for tested isolates (S, I or R), mainly derived from diffusion methods.

In 2012, 15 MSs and 1 non-MS (Switzerland) reported quantitative dilution data on antimicrobial resistance in *Campylobacter* isolates from animals and food. AST was carried out only for *C. jejuni* and *C. coli*, all other *Campylobacter* species were excluded from the monitoring programme of antimicrobial resistance in *Campylobacter*. Twelve MSs reported data where no method was specified.

Table CA1 presents an overview of the countries reporting antimicrobial resistance data on *Campylobacter* spp. from humans and various animal and food categories in 2012.

Table CA1. Overview of countries reporting antimicrobial resistance data using MIC and disc diffusion zones on Campylobacter coli and Campylobacter jejuni from humans and various animal and food categories in 2012

Bacterial species	Method	Origin	Total number of MSs reporting	Countries
	Diffusion	Human	8	MSs: EE, FR, IT, LT, LU, NL, RO, SI
		Human	9	MSs: AT, ES, HU, LU, MT, NL, SI, SK, UK
			7	MSs: AT, CZ, ES, FR, HU, NL, RO
		Gallus gallus (fowl)	7	Non-MS: CH
		Turkeys	2	MSs: DE, NL
		Digo	F	MSs: DK,ES, FR, HU, NL
0 "		Pigs	5	Non-MS: CH
C. coli	Dilution	Cattle (heuring animale)	2	MSs: DE, ES, NL
		Cattle (bovine animals)	3	Non-MS: CH
		Meat from broilers (Gallus gallus)	7	MSs: AT, BE, CZ, EE, HU, NL, PL, RO
		Meat from turkeys	6	MSs: DE, EE, HU, NL, PL, RO
		Meat from pigs	2	MSs: BE, PL
		Meat from bovine animals	2	MSs: DE, PL
	Diffusion	Human	8	MSs: EE, FR, IT, LT, LU, NL, RO, SI
	Diffusion	numan	0	Non-MS: IS
		Human	9	MSs: AT, ES, HU, LU, MT, NL, SI, SK, UK
		Tuman	1	Non-MS: IS
		Gallus gallus (fowl)	11	MSs: AT, CZ, DK, ES, FI, FR, HU, IT, NL, RO, SE
		5 ()		Non-MS: CH
		Turkeys	2	MSs: DE, NL
C. jejuni		Digo	4	MS: HU
	Dilution	Pigs	1	Non-MS: CH
	Diration	Cattle (having animale)	5	MSs: DE, DK ,ES, FI, NL
		Cattle (bovine animals)	5	Non-MS: CH
		Meat from broilers (Gallus gallus)	8	MSs: AT, BE, CZ, DK, EE, HU ,NL, PL, RO
		Meat from turkeys	6	MSs: DE, EE, HU, NL, PL, RO
		Meat from pigs	1	MS: PL
		Meat from bovine animals	1	MS: PL

MIC: minimum inhibitory concentration.

Note: For abbreviations of Member States (MS) and other reporting countries, see Appendix 7.



4.3. Antimicrobial resistance in *Campylobacter* isolates from humans

METHODS AND INTERPRETIVE CRITERIA USED FOR ANTIMICROBIAL SUSCEPTIBILITY TESTING OF CAMPYLOBACTER ISOLATES FROM HUMANS

The method of testing for antimicrobial susceptibility varies between countries. Disc diffusion was the most common method in 2012, but often a combination of disc diffusion and dilution was used, depending on the reason for the testing. In several countries, the reference laboratories type only a fraction of the isolates. The remaining isolates are typed by hospitals or local laboratories and the methods used by these are not reported. The guidelines used for the methodology and interpretation of antimicrobial susceptibility testing for Campylobacter differed between countries and also within countries for different antimicrobials, but were more harmonised in 2012 than in 2011 (for detailed information, see Chapter 8 Materials and methods, Table MM3). The guidelines used by several national public health reference laboratories were from the French Society for Microbiology (CA-SFM), Clinical and Laboratory Standards Institute (CLSI) and European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Of the five antimicrobials tested in both human and animal/food isolates, resistance according to the EUCAST clinical breakpoints and ECOFFS were at the same MIC value or only differing by one concentration step for ciprofloxacin, erythromycin and tetracycline, while no EUCAST clinical breakpoints were available for gentamicin and nalidixic acid. The CA-SFM breakpoints differed from the ECOFFS by two concentration steps for tetracycline, the combination of C. coli/erythromycin and the combination of C. jejuni/gentamicin. In all other cases the breakpoints for CA-SFM and the ECOFFs were at the same MIC value or only differing by one concentration step. The level of resistance determined by CLSI breakpoints and ECOFFS were at the same MIC value or only differing by one concentration of C. jejuni/erythromycin where there was a two-step difference. CLSI clinical breakpoints were not available for gentamicin or nalidixic acid (Figure CA1).Owing to the variety of breakpoints used under each set of guidelines, results should be interpreted with caution in the case of antimicrobials where there are major differences in the interpretive criteria.

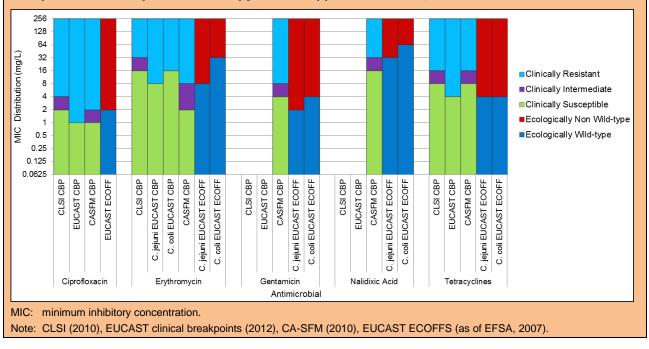


Figure CA1. Comparison of clinical breakpoints and epidemiological cut-off values used to interpret MIC data reported for Campylobacter spp. from humans, animals or food

4.3.1. Antimicrobial resistance in *Campylobacter* spp. in humans

Fourteen MSs and Iceland submitted data for 2012 on antimicrobial susceptibility of *Campylobacter* spp. isolates from human clinical cases to ECDC. This accounted for 38,835 *Campylobacter* spp. isolates, representing 17.9 % of the total number of confirmed cases of campylobacteriosis reported in the EU/EEA in 2012 (N=217,261).Thirteen MSs and Iceland reported susceptibility results for more than 20 isolates, which was the limit set for presenting the level of resistance by country. Romania reported susceptibility results for fewer than 20 isolates and was included only in the analysis totals.

In order to better assess the impact from food consumed within each reporting country on the antimicrobial resistance levels found in human *Campylobacter* isolates, the analysis focused on domestically-acquired cases. Several countries however did not provide any information on travel (or non-travel) of their cases. Cases with unknown travel status were therefore also included in the analysis. A separate analysis was made on travel-associated cases by geographical regions.

4.3.1.1. Resistance levels in *Campylobacter* spp. isolates from human cases

A large variation was observed among the reporting countries with regard to the number of antimicrobials tested, ranging from 8 countries testing for amoxicillin to all 15 countries testing for ciprofloxacin and erythromycin (Table CA2). This most likely reflects the variation in the clinical importance of the antimicrobials. The highest average level of resistance in all *Campylobacter* spp. isolates from human cases was observed for nalidixic acid (48.8 %; N=21,491) and ciprofloxacin (47.4 %; N=36,172) followed by ampicillin (36.4 %; N=7,768) and tetracyclines (32.4 %; N=6,824) (Table CA2).

The observed variability in resistance levels may reflect the differences in the *Campylobacter* population that people are exposed to in different countries. Variability may also arise due to differences in testing or sampling methods applied by individual countries as well as the use of different interpretive criteria. Sampling bias could also arise by only including isolates from hospitalised cases. This may explain some of the extreme observations.

4.3.1.2. Comparison of resistance levels in *Campylobacter* spp. isolates acquired within EU/EEA and in other geographical regions

To compare resistance levels in isolates acquired across the world, isolates from travel-associated cases were classified into seven different geographical regions¹⁵ EU/EEA, non-EU/EEA, Africa, Asia, North and Central America, South America and Oceania, based on the probable country of infection. Only a limited amount of isolates were tested and/or reported from travel-associated cases outside of Europe (Table CA3). Data on resistance to four antimicrobials (ciprofloxacin, erythromycin, nalidixic acid and tetracyclines) were reported for \geq 10 isolates from infections acquired in three geographical regions (EU/EEA, Africa and Asia). The frequencies of resistance to all four antimicrobials were noticeably higher in isolates that had been acquired in Asia and Africa compared with those acquired within the EU/EEA, with about two-fold higher levels of ciprofloxacin resistance to erythromycin (Table CA3).

Since the resistance levels differ substantially between *C. jejuni* and *C. coli*, the two most commonly reported *Campylobacter* species, further results are presented separately for these two species.

¹⁵ Regional classification from the United Nations Statistical Division. Available online: <u>http://unstats.un.org/unsd/methods/m49/m49regin.htm</u>



Country	Amox	cicillin	Ampicillin		Ciprofloxacin		Erythromycin		Genta	amicin	Nalidi>	kic acid	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res
Austria	387	0.3	387	29.2	387	62.3	387	1.8	387	0.3	387	60.7	387	31.8
Estonia	90	6.7	87	41.4	231	64.5	212	0.9	92	1.1	95	66.3	201	17.4
France	4,727	0.2	4,728	31.3	4,643	52.9	4,727	3.7	4,727	0.2	4,728	54.8	_	-
Hungary	-	-	Ι	_	71	77.5	71	1.4	-	_	-	_	-	-
Italy	-	_	184	65.8	254	62.6	251	6.4	172	2.3	146	70.5	170	52.4
Lithuania	-	-	Ι	_	195	85.1	227	0.4	-	_	-	_	-	-
Luxembourg	561	0.2	561	47.2	561	62.0	561	2.7	561	0	561	62.6	561	47.2
Malta	_	_	Ι	_	214	76.2	214	26.6	-	_	-	_	-	_
Netherlands	_	_	Ι	_	3,483	58.5	2,976	2.6	_	_	_	_	1,519	32.8
Romania	-	-	-	-	19	NA	19	NA	19	NA	19	NA	19	NA
Slovakia	3	NA	125	28.0	1,109	31.8	1,302	1.5	27	11.1	-	_	1,235	11.4
Slovenia	788	5.5	981	35.2	981	68.0	981	0.8	981	0.2	787	61.6	981	21.9
Spain	228	1.8	228	49.1	228	83.8	228	7.9	228	3.1	228	96.1	228	72.8
United Kingdom	214	14.5	487	66.1	23,796	42.6	23,868	3.0	1,048	0.4	14,540	44.2	1,523	43.8
Total (14 MSs)	6,998	1.4	7,768	36.4	36,172	47.4	36,024	3.1	8,242	0.4	21,491	48.8	6,824	32.4
Iceland	-	_	-	_	29	34.5	29	0	-	_	-	_	-	_

Table CA2. Antimicrobial resistance in Campylobacter spp. from humans per country in 2012, using clinical breakpoints¹

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM4.



Table CA3. Antimicrobial resistance in Campylobacter spp. reported to be acquired within the EU and in other geographical regions in 2012, using clinical breakpoints

Country	Amo	xicillin	Ampicillin		Ciprofloxacin		Erythromycin		Gentamicin		Nalidixic acid		Tetracyclines	
Country	Ν	% Res	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Europe (EU/EEA countries)	7,000	1.4	7,848	36.8	36,386	40.5	36,080	3.1	8,256	0.4	21,543	48.9	6,847	32.4
Europe (non-EU/EEA countries)	1	NA	3	NA	7	NA	7	NA	1	NA	4	NA	2	NA
Africa	0	NA	6	NA	35	74.3	32	9.4	7	NA	23	78.3	7	NA
Asia	4	NA	7	NA	63	81.0	62	8.1	9	NA	35	80.0	14	57.1
Northern and Central America	No obs	ervations	No obs	servations	8	NA	8	NA	1	NA	7	NA	2	NA
South America	No obs	ervations	No obs	ervations	1	NA	1	NA	No obse	rvations	1	NA	No Obse	rvations
Oceania	No obs	ervations	No obs	ervations	No obse	rvations	No obse	rvations	No obse	rvations	No obse	rvations	No Obse	rvations

N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated.



4.3.2. Antimicrobial resistance in *Campylobacter jejuni* in humans

As in previous years, *C. jejuni* was the most common *Campylobacter* species identified in 2012, with 81,621 cases reported in the EU/EEA. In 2012, 13 MSs and Iceland reported data on antimicrobial resistance in *C. jejuni* for \geq 20 isolates (ranging from 6 MSs for amoxicillin to all 14 countries for ciprofloxacin and erythromycin (Table CA4)).

4.3.2.1. Resistance levels in *Campylobacter jejuni* isolates from human cases

The highest frequencies of resistance in *C. jejuni* isolates were observed for ciprofloxacin (54.1 %; N=11,551) and nalidixic acid (53.3 %; N=6,765) (Table CA4).

Macrolides, e.g. erythromycin, are the first choice drugs for the treatment of campylobacteriosis in humans (ECDC et al., 2009). The level of resistance for erythromycin reported in humans was low, on average 1.4 % (N=11,080). In the EU, the highest proportions of resistant isolates were reported by Malta with 10.9 % (N=138) and Italy with 5.4 % (N=202) (Table CA4).

Ciprofloxacin is the second-choice drug for treatment of campylobacteriosis in humans (ECDC et al., 2009) although resistance evolves rapidly. Resistance to ciprofloxacin reported by the countries was very low to extremely high (0.3 % to 91.9 %). The highest levels of resistance, 91.9 % (N=99) and 84.1 % (N=182), were reported by Lithuania and Spain, respectively (Table CA4). Nalidixic acid is normally used as an indicator of ciprofloxacin resistance. Resistance to nalidixic acid and ciprofloxacin was comparable and the levels of resistance to nalidixic acid ranged from 40.8 % to 96.2 % (Table CA4).

4.3.2.2. Trends in resistance levels in *Campylobacter jejuni* isolates from human cases

Country-specific trends in resistance to ciprofloxacin during the period 2008 to 2012 are presented in Figure CA2. Only countries reporting data for at least three consecutive years and 10 isolates per year were included. Trends of increasing resistance could be observed in most reporting countries over the period, with the exception of Italy and the United Kingdom. Increases were most noticeable in Estonia, Iceland, Lithuania (from 2009 and onwards) and Malta.

Country-specific trends for erythromycin over the years 2008 to 2012 are presented in Figure CA3. There were few common trends between countries over the years. The exception was a peak in resistance observed in 2010 in Iceland (3.1 %; N=32), Luxembourg (3.6 %; N=528) and Slovenia (2.4 %; N=911). In the years before and after 2010, resistance levels in all three countries were markedly lower. In some other countries, a peak was instead observed in 2011: Estonia (1.2 %; N=165), France (1.7 %; N=4,171), Italy (6.9 %; N=174), Lithuania (0.3 %, N=296) and the Netherlands (2.8 %, N=2,501). In the United Kingdom, a decreasing trend could be observed over the whole period.

4.3.2.3. Multi-drug resistance among *Campylobacter jejuni* isolates from human cases

Five MSs, Austria, Estonia, Luxembourg, Slovenia and Spain, tested at least 10 isolates for the full range of antimicrobials included in the human data collection for *C. jejuni*, and these isolates were included in the multi-drug resistance analysis. Overall, 18.0 % (N=1,799) of the human *C. jejuni* isolates were susceptible to all six antimicrobials, with particularly low levels of susceptibility reported from Spain (1.6 %; N=182) (Table CA5). Multi-drug resistance was, on average, high in the five MSs (24.8 %; N=1,799; country average 24.8 %). There was large variation in the level of multi-resistance between countries ranging from 14.3 % (N=84) in Estonia to 42.3 % (N=182) in Spain (Table CA5). The proportions of *C. jejuni* isolates susceptible to all or resistant (non-susceptible) to any up to six antimicrobials by MS are presented in Figure CA4. Isolates resistant to up to five antimicrobials were reported from two MSs (Slovenia and Spain). No MS reported any isolates resistant to all six antimicrobials. Few isolates exhibited co-resistance to both ciprofloxacin and erythromycin in the five MSs (1.4 %; N=1,799) (Table CA5).



Country	Amox	cicillin	Amp	icillin	Ciprof	loxacin	Erythr	omycin	Genta	amicin	Nalidi>	kic acid	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	345	0	345	28.4	345	61.2	345	1.4	345	0	345	59.4	345	31.3
Estonia	89	5.6	86	40.7	219	63.9	200	0.5	91	1.1	93	66.7	189	16.4
France	3,852	0	3,853	30.9	3,843	50.7	3,852	0.6	3,852	0	3,853	50.8	_	_
Hungary	-	_	I	-	34	79.4	34	0	-	_	-	-	Ι	_
Italy	-	-	139	68.3	201	64.7	202	5.4	141	2.8	125	70.4	142	54.2
Lithuania	-	-	-	-	99	91.9	114	0.0	-	_	-	_	-	-
Luxembourg	493	0	493	48.5	493	60.4	493	0.8	493	0.0	493	60.6	493	43.6
Malta	-	-	-	-	138	71.7	138	10.9	-	_	-	_	-	_
Netherlands	-	-	-	-	3,076	58.0	2,624	2.0	-	_	-	_	1,249	32.5
Romania	-	-	-	-	3	NA	3	NA	3	NA	3	NA	3	NA
Slovakia	3	NA	115	27.8	1,044	30.7	1,162	1.5	13	NA	-	-	1,151	11.6
Slovenia	695	5.8	869	37.4	869	70.7	869	0.7	869	0.2	695	60.3	869	22.6
Spain	182	0.5	182	52.2	182	84.1	182	3.3	182	0.5	182	96.2	182	72.0
United Kingdom	10	NA	2	NA	1,005	43.1	862	1.5	7	NA	976	40.8	56	41.1
Total (14 MSs)	5,669	0.8	6,084	34.7	11,551	54.1	11,080	1.4	5,996	0.2	6,765	53.3	4,679	28.3
Iceland	-	_	-	_	27	33.3	27	0	-	_	-	_	-	_

Table CA4. Antimicrobial resistance in Campylobacter jejuni from humans per country in 2012, using clinical breakpoints¹

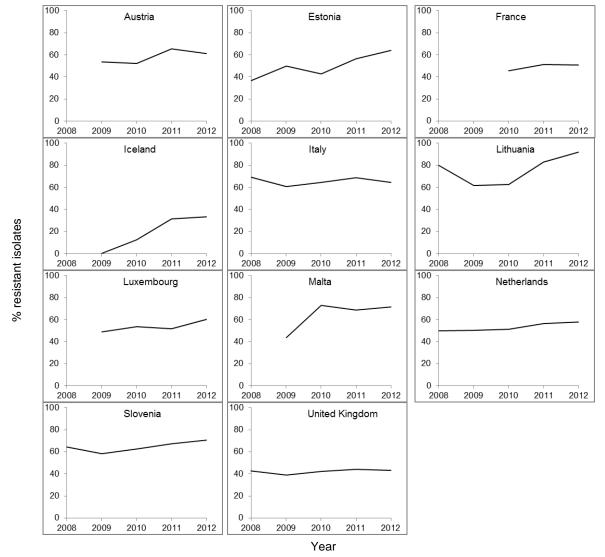
MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM4.



Figure CA2. Resistance to ciprofloxacin in C. jejuni in humans in reporting MSs and one non-MS in the EU, 2008-2012, using clinical breakpoints¹

Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²



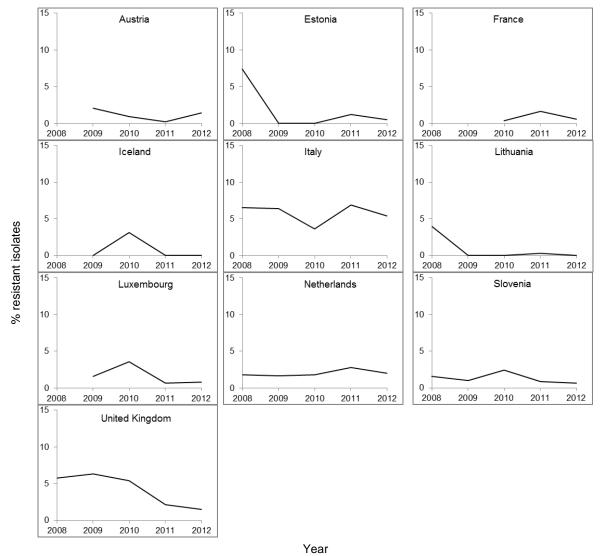
MS: Member State.

- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM4.
- Guidelines for clinical breakpoints: Austria (ECOFFs from EUCAST), Estonia (Clinical breakpoints (CBP) from CA-SFM), France (CBP from CA-SFM), Iceland (CBP from CLSI), Italy (CBP from CLSI), Lithuania (CBP from BSAC), Luxembourg (CBP from CA-SFM), Malta (CBP from HPA/CLSI), the Netherlands (unspecified), Slovenia (CBP from CLSI for dilution, CBP from CA-SFM for disc diffusion), the United Kingdom (ECOFFs from EUCAST). See also Table MM2.



Figure CA3. Resistance to erythromycin in C. jejuni in humans in reporting MSs and one non-MS in the EU, 2008-2012, using clinical breakpoints¹

Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²



MS: Member State.

- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM4.
- Guidelines for clinical breakpoints: Austria (ECOFFs from EUCAST), Estonia (CBP from CA-SFM), France (CBP from CA-SFM), Iceland (CBP from CLSI), Italy (CBP from CLSI), Lithuania (CBP from BSAC), Luxembourg (CBP from CA-SFM), the Netherlands (unspecified), Slovenia (CBP from CLSI for dilution, CBP from CA-SFM for disc diffusion), the United Kingdom (ECOFFs from EUCAST). See also Table MM3.

Table CA5. Complete susceptibility, multi-resistance and co-resistance (non-susceptibility) to ciprofloxacin and erythromycin, as determined by clinical breakpoints, in C. jejuni from humans by MS, 2012¹

Country	Susceptible to all (%)	Multi-resistant (%)	Co-resistant to Cip and Ery (%)
Austria (N=345)	28.4	14.8	1.4
Estonia (N=84)	8.3	14.3	0
Luxembourg (N=493)	21.3	28.6	2.2
Slovenia (N=695)	16.0	23.9	0.7
Spain (N=182)	1.6	42.3	2.7
Total (5 MSs) (N=1,799)	18.0	24.8	1.4

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*, Cip: ciprofloxacin; Ery: erythromycin; MS: Member State.

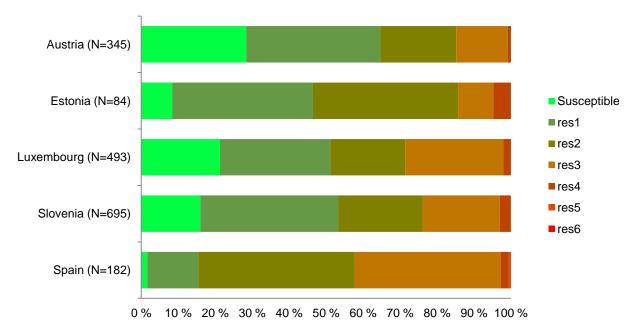
Susceptible to all: proportion of isolates susceptible to all antimicrobial substances of the ECDC common set for Campylobacter.

Multi-resistant: proportion of isolates resistant to at least three different antimicrobial substances, belonging to any three antimicrobial families from the ECDC common antimicrobial set for Campylobacter.

Co-resistant to Cip and Ery: proportion of isolates not susceptible to both Cip and Ery.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM4.

Figure CA4. Frequency distribution of completely susceptible isolates and resistant isolates to from one to six antimicrobials, as determined by clinical breakpoints, in C. jejuni from humans by MS, 2012¹



N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*, MS: Member State. Susceptible: total number of isolates susceptible to all antimicrobial substances of the common set for *C. jejuni.* res1-res6: total number of isolates resistant to between one and six antimicrobial substances of the common set for *C. jejuni.*

isolates from travel-associated, domestic and unknown cases by country please see table MM4.

Isolates from cases reported as related to travel outside the country were excluded from the graph. For the proportion of tested



4.3.3. Antimicrobial resistance in *Campylobacter coli* in humans

C. coli was the second most common *Campylobacter* species identified in 2012, with 6,231 cases reported in the EU/EEA. In 2012, nine MSs reported data on antimicrobial resistance in *C. jejuni* for \geq 20 isolates (ranging from five MSs for amoxicillin to all nine countries for ciprofloxacin and erythromycin (Table CA6)).

4.3.3.1. Resistance levels in *Campylobacter coli* isolates from human cases

The highest percentage of resistance among *C. coli* isolates was observed for nalidixic acid (69.9 %; N=1,002) and ciprofloxacin (69.0 %; N=1,322), followed by tetracyclines (49.7 %; N=328) and ampicillin (36.0 %; N=926) (Table CA6). The percentage of resistance to ciprofloxacin was highly correlated with resistance to nalidixic acid in four of the six countries which tested both antimicrobials and in two countries the percentage of resistance was higher to nalidixic acid than to ciprofloxacin. The percentage of human *C. coli* isolates resistant to erythromycin was 15.1 % (N=1,264), which was considerably higher than for *C. jejuni* (1.4 %). The highest levels of resistance to erythromycin were reported from Spain (27.3 %; N=44), but the number of isolates tested was low (Table CA6).

4.3.3.2. Trends in resistance levels in *Campylobacter coli* isolates from human cases

Country-specific trends in resistance to ciprofloxacin during the period 2008 to 2012 are presented in Figure CA5. Only countries reporting data for at least three consecutive years and 10 isolates per year were included. Increasing trends over the period were observed in Austria, Luxembourg, the Netherlands and Slovenia (from 2009). A drop in resistance was observed in several countries in 2011: France (58.1 %, N=735), Italy (46.2 %; N=13), Lithuania (74.4 %; N=39), Malta (53.8 %; N=39), Slovenia (52.4 %, N=42) and Spain (78.4 %, N=51) (Figure CA5).

There were few similarities in resistance trends for erythromycin between countries over the years 2008 to 2012 (Figure CA6) except that a peak in resistance could be observed in several countries in 2011: Austria (8.3 %; N=36), Lithuania (2.2 %; N=45), Luxembourg (23.3 %; N=60), the Netherlands (15.5 %; N=110) and Slovenia (7.1 %; N=42). A trend of increasing resistance to erythromycin was observed in the Netherlands over the five-year period and a decreasing trend was observed in Italy, although in the case of Italy only a small number of isolates were tested (N=10-30). The peak in resistance observed in Spain in 2010 could be because only 10 isolates were tested that year.

4.3.3.3. Multi-drug resistance among Campylobacter coli isolates from human cases

Four MSs, Austria, Luxembourg, Slovenia and Spain, tested at least 10 isolates for the full range of antimicrobials included in the human data collection for *C. coli* and these isolates were included in the multidrug resistance analysis. Overall, only 6.4 % (N=188) of the human *C. coli* isolates were susceptible to all six antimicrobials, with particularly low levels of susceptibility reported in Slovenia (0 %; N=36) and Spain (0 %; N=44) (Table CA7). On average, the level of multi-drug resistance was high (35.1 %; N=188; country average 33.3 %) (Table CA7). The proportions of *C. coli* isolates susceptible to all or resistant (nonsusceptible) to any one up to six antimicrobials by MS are presented in Figure CA7. All four countries reported isolates resistant to up to four antimicrobials, three MSs reported isolates resistant to up to five antimicrobials and one isolate was reported from one MS resistant to all six. The overall level of coresistance to both ciprofloxacin and erythromycin was moderate across these four countries (16.0 %; N=188) (Table CA7).



Country	Amox	cicillin	Ampicillin		Ciprof	Ciprofloxacin		omycin	Genta	amicin	Nalidiz	kic acid	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	42	2.4	42	35.7	42	71.4	42	4.8	42	2.4	42	71.4	42	35.7
Estonia	1	NA	1	NA	1	NA	1	NA	1	NA	1	NA	1	NA
France	706	0	706	35.4	705	70.5	706	17.7	706	1.3	706	70.5	_	_
Hungary	_	_	_	_	17	NA	17	NA	-	_	-	-	_	_
Italy	_	-	11	NA	10	NA	11	NA	10	NA	8	NA	8	NA
Lithuania	_	_	_	_	25	84.0	31	0	-	_	-	-	_	_
Luxembourg	66	1.5	66	39.4	66	74.2	66	16.7	66	0	66	75.8	66	75.8
Malta	_	-	_	-	37	73.0	37	13.5	_	_	-	-	_	_
Netherlands	-	-	-	-	213	65.3	163	11.7	-	_	-	-	102	36.3
Romania	-	-	-	-	2	NA	2	NA	2	NA	2	NA	2	NA
Slovakia	_	_	_	_	5	NA	4	NA	-	_	-	-	5	NA
Slovenia	36	8.3	55	34.5	55	74.5	55	1.8	55	0	36	86.1	55	30.9
Spain	44	6.8	44	38.6	44	86.4	44	27.3	44	13.6	44	95.5	44	79.5
United Kingdom	2	NA	1	NA	100	42.0	85	15.3	3	NA	97	42.3	3	NA
Total (14 MSs)	897	1.0	926	36.0	1,322	69.0	1,264	15.1	929	1.8	1,002	69.9	328	49.7

Table CA6. Antimicrobial resistance in Campylobacter coli from humans per country in 2012, using clinical breakpoints¹

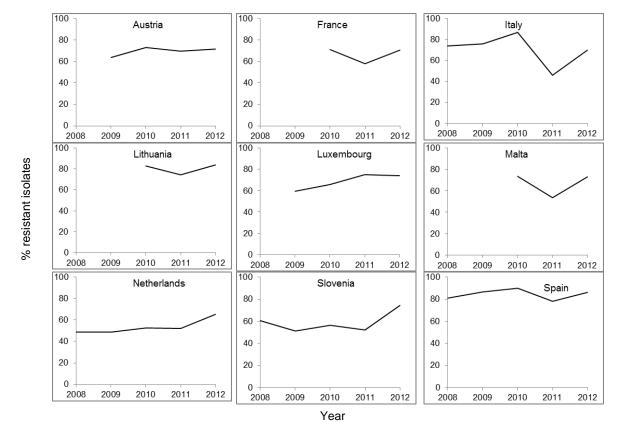
MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported; NA: not applicable - if fewer than 20 isolates were tested resistance was not calculated.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM4.



Figure CA5. Resistance to ciprofloxacin in C. coli in humans in reporting MSs in the EU, 2008-2012, using clinical breakpoints¹

Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²

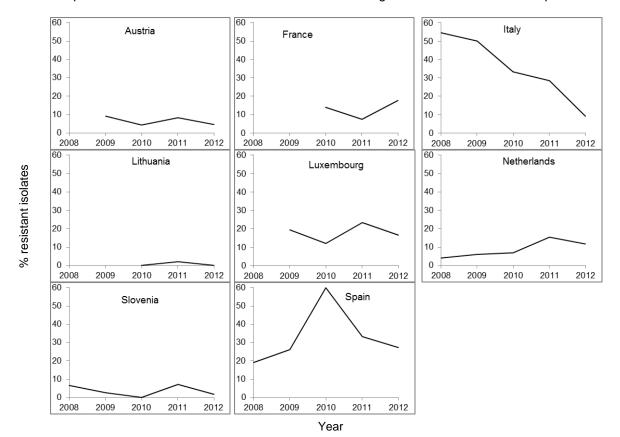


MS: Member State.

- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM4.
- Guidelines for clinical breakpoints: Austria (ECOFFs from EUCAST), France (CBP from CA-SFM), Italy (CBP from CLSI), Lithuania (CBP from BSAC), Luxembourg (CCBP from A-SFM), Malta (CBP from HPA/CLSI), the Netherlands (unspecified), Slovenia (CBP from CLSI for dilution, CBP from CA-SFM for disc diffusion), Spain (CBP from CLSI). See also Table MM3.



*Figure CA6. Resistance to erythromycin in C. coli in humans in reporting MSs in the EU, 2008-2012, using clinical breakpoints*¹



Direct comparisons between countries should be avoided owing to the use of different interpretive criteria²

MS: Member State.

- 1. Isolates from cases reported as related to travel outside the country were excluded from the graphs. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM4.
- Guidelines for clinical breakpoints: Austria (ECOFFs from EUCAST), France (CBP from CA-SFM), Italy (CBP from CLSI), Lithuania (CBP from BSAC), Luxembourg (CBP from CA-SFM), the Netherlands (unspecified), Slovenia (CBP from CLSI for dilution, CBP from CA-SFM for disc diffusion), Spain (CBP from CLSI). See also Table MM3.

Table CA7. Complete susceptibility, multi-resistance and co-resistance (non-susceptibility) to ciprofloxacin and erythromycin, as determined by clinical breakpoints, in C. coli from humans by MS, 2012¹

Country	Susceptible to all (%)	Multi-resistant (%)	Co-resistant to Cip and Ery (%)
Austria (N=42)	9.5	16.7	2.4
Luxembourg (N=66)	12.1	45.5	24.2
Slovenia (N=36)	0	19.4	5.6
Spain (N=44)	0	50.0	25.0
Total (4 MSs) (N=188)	6.4	35.1	16.0

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*, Cip: ciprofloxacin; Ery: erythromycin; MS: Member State.

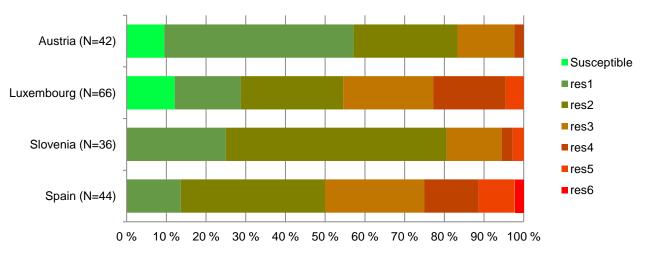
Susceptible to all: proportion of isolates susceptible to all antimicrobial substances of the ECDC common set for Campylobacter.

Multi-resistant: proportion of isolates resistant to at least three different antimicrobial substances, belonging to any three antimicrobial families from the ECDC common antimicrobial set for *Campylobacter*.

Co-resistant to Cip and Ery: proportion of isolates not susceptible to both Cip and Ery.

1. Isolates from cases reported as related to travel outside the country were excluded from this table. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see table MM4.

Figure CA7. Frequency distribution of C. coli isolates completely susceptible or resistant to one to six antimicrobials, as determined by clinical breakpoints, from humans by MS, 2012¹



N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances for *C. coli*; MS: Member States.

Susceptible: total number of isolates susceptible to all antimicrobial substances of the common set for C. coli.

res1-res6: total number of isolates resistant to between one and six antimicrobial substances of the common set for C. coli.

1. Isolates from cases reported as related to travel outside the country were excluded from this graph. For the proportion of tested isolates from travel-associated, domestic and unknown cases by country please see Table MM4.



4.4. Antimicrobial resistance in *Campylobacter* isolates from animals and food

The countries reporting on Campylobacter resistance from various animal and food sampling origins, in 2012 are presented in Table CA1. Antimicrobials selected by the different MSs, and non-MSs, for susceptibility testing of *C. jejuni* and *C. coli*, are shown in Chapter 8 Materials and methods, Table MM8.

THE REVISION OF EUCAST ECOFFS FOR CAMPYLOBACTER

There have been some recent minor revisions to the epidemiological cut-off value (ECOFF) provided by the European Committee on Antimicrobial Susceptibility Testing (EUCAST). Thus, the EUCAST ciprofloxacin ECOFF for C. coli is currently susceptible (i.e. wild-type) $\leq 0.5 \text{ mg/L}$, a decline of one log value from the previous ECOFF value of resistant (i.e. non-wild-type) >1 mg/L described on the EUCAST website and recorded in Table MM11. Similarly, the ECOFF values for C. coli and erythromycin, C. coli and nalidixic acid and C. jejuni and both ciprofloxacin and tetracyclines have declined by one dilution step. Conversely, the ECOFF has increased by one log dilution for C. jejuni versus gentamicin and streptomycin. Although deviation from wild-type susceptibility is a fixed microbiological characteristic, as greater numbers of bacterial isolates are tested, the wild-type distribution may become better defined and minor changes in the ECOFF might therefore be expected. The breakpoints used in this report to discriminate between 'microbiologically resistant' and wild-type bacteria are identical to those used in previous reports for Campylobacter and so there should be no effect of methodological changes when comparisons are made between years. When EFSA's recommendations are revised to include the latest EUCAST ECOFFs and new legislation (Decision 2013/652) incorporating those recommendations enters into force in 2014, then the historical data are likely to be re-interpreted, using the new EUCAST ECOFFs. Reference to the minimum inhibitory concentration distribution tables for C. coli and C. jejuni, which are published in the Level 3 tables, shows that the effect of these changes is in most cases likely to be small.

In this chapter, resistance to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines is described in detail. The occurrence of resistance is tabulated, a portrait of temporal evolution and spatial distribution of resistance is drawn and the multi-resistance is analysed. These analyses were performed, and the corresponding results presented, depending on whether a minimum of four or more countries reported quantitative data for a given *Campylobacter* species and the origin of sample (animal population and food category), and whether data were related to at least 10 isolates per country, origin of sample and year. *C. jejuni* and *C. coli* are both addressed, as monitoring data on the prevalence of *Campylobacter* in broilers and broiler meat in some reporting countries can reveal that *C. coli* prevalence is either not negligible or even of the same magnitude as that of *C. jejuni* (EFSA and ECDC, 2014).

- Temporal trend graphs were generated, showing percentage resistance to different antimicrobials among *Campylobacter* isolates, per sample origin, over the period 2006 to 2012, by year of sampling. Temporal trend graphs were included only for countries which had reported on four or more years in the 2006 to 2012 period.
- The spatial distributions of ciprofloxacin and erythromycin resistance rates in *C. jejuni* from *Gallus gallus* and *C. coli* from pigs are presented. For countries where resistance level figures for 2012 were not available, 2011 figures were used instead. For cattle, the number of reporting countries was lower than in the case of the other animal species monitored and, therefore, no spatial distribution maps were generated.
- Multi-resistance was analysed in the isolate-based dataset of *Campylobacter* isolates tested for the full harmonised set of five antimicrobials (ciprofloxacin, erythromycin, gentamicin, streptomycin and tetracyclines) belonging to different classes. 'Multi-resistance' was defined as non-susceptibility to at least three different antimicrobial classes. The proportions of isolates susceptible to all and resistant (non-susceptible) to any one of up to five antimicrobials are presented. Co-resistance to ciprofloxacin and erythromycin was also estimated as these two antimicrobials are of particular interest in human medicine in the treatment of campylobacteriosis.

Further information on reported MIC distributions and numbers of *C. jejuni* and *C. coli* isolates resistant to amoxicillin/clavulanic acid, ampicillin chloramphenicol, ciprofloxacin, clarithromycin, erythromycin,



gentamicin, imipenem, nalidixic acid, neomycin, streptomycin, sulfonamides, tetracyclines and tulathromycin can be found in the Level 3 tables published on the EFSA website.

4.4.1. Antimicrobial resistance in Campylobacter isolates from food

In 2012, more than four MSs provided quantitative antimicrobial resistance data for *C. jejuni* and/or *C. coli* isolates only from broiler meat, so the analysis of other food categories is not presented in this report.

4.4.1.1. Meat from broilers and spent hens (Gallus gallus)

Representative sampling and monitoring

In the reporting MSs, data on antimicrobial resistance in *Campylobacter* isolates from meat from *Gallus gallus* derived from active monitoring programmes were based mainly on the random collection of broiler meat samples obtained either at the slaughterhouse, at the processing plant or at retail outlets. In Austria, Denmark, Estonia and Romania, representative random sampling of meat from broilers, whether neck skin, fresh meat, minced meat or meat preparation, was carried out entirely or primarily at wholesale or retail outlets. In Poland, sampling of broiler meat and broiler carcases was performed at retail outlets and at the slaughterhouse, respectively, while, in Hungary, meat samples were gathered at processing plants. In Belgium, *Campylobacter* isolates were derived from carcase swabs of spent hens and chilled broiler carcases collected at the slaughterhouse.

Resistance levels among C. jejuni and C. coli isolates from meat from broilers

In 2012, eight and six MSs provided quantitative antimicrobial resistance data for C. jejuni and C. coli isolates, respectively, from broiler meat (Table CA8 and Table CA9). Although resistance is typically higher among C. coli than C. jejuni, common features in the levels of resistance to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines can be observed in the two Campylobacter species. For the commonly used antimicrobials, resistance to tetracyclines and nalidixic acid generally ranged from high to extremely high levels, whereas resistance to gentamicin varied less among reporting MSs and was either undetected or recorded at low levels. For clinically important antimicrobials, resistance to ciprofloxacin was high to extremely high in reporting MSs and closely paralleled the results obtained for nalidixic acid (as expected), whereas resistance to erythromycin was much lower considering all reporting MSs. However, the recorded levels of resistance to erythromycin were contrasting considering C. jejuni and C. coli, with higher levels generally observed in C. coli. In contrast to the other reporting MSs, Romania recorded a moderate resistance level to erythromycin in C. jejuni at 14.1 %. Belgium, the Netherlands and Romania reported moderate to high resistance to erythromycin in C. coli, whereas Austria, Hungary and Poland either did not detect resistance or reported low resistance. In Belgium, where resistance was monitored in a parallel fashion in carcases of spent hens and broilers. Campylobacter isolates from spent hens' meat exhibited either similar or lower resistance levels to those observed in isolates from broiler meat.

Table CA8. Resistance (%) to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines among <u>Campylobacter jejuni</u> from <u>meat from broilers</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Ciprof	loxacin	Erythromycin		Gentamicin		Nalidixic acid		Tetracyclines	
Country	Ν	% Res	N	% Res	Ν	% Res	Ν	% Res	N	% Res
Austria	60	63.3	60	0	60	0	60	56.7	60	33.3
Belgium ¹	75	57.3	75	1.3	75	0	75	57.3	75	64.0
Belgium ²	106	28.3	106	1.9	106	0	106	27.4	106	37.7
Denmark	66	28.8	66	0	66	0	66	28.8	66	15.2
Estonia	33	51.5	33	3.0	33	3.0	33	51.5	33	24.2
Hungary	22	81.8	22	0	22	4.5	-	_	22	54.5
Netherlands	241	55.6	241	0.8	241	0	241	55.6	241	50.6
Poland	157	88.5	157	0	157	0	157	86.0	157	59.2
Romania	64	81.3	64	14.1	64	6.3	64	82.8	64	59.4
Total (8 MSs)	824	59.5	824	1.8	824	0.7	802	57.9	824	47.5

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

1. Meat from broilers.

2. Meat from spent hens.



 Table CA9. Resistance (%)
 to
 ciprofloxacin,
 erythromycin,
 gentamicin,
 nalidixic
 acid
 and

 tetracyclines
 among
 Campylobacter
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 from
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Country	Ciprof	loxacin	Erythromycin		Gentamicin		Nalidixic acid		Tetracyclines	
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	30	73.3	30	0	30	0	30	73.3	30	33.3
Belgium ¹	30	80.0	30	26.7	30	0	30	70.0	30	86.7
Belgium ²	43	65.1	43	16.3	43	4.7	43	65.1	43	62.8
Hungary	47	87.2	47	2.1	47	0	_	-	47	34.0
Netherlands	126	83.3	126	23.8	126	0.8	126	83.3	126	62.7
Poland	116	89.7	116	5.2	116	0	116	89.7	116	64.7
Romania	81	82.7	81	32.1	81	6.2	81	80.2	81	46.9
Total (6 MSs)	473	82.7	473	16.5	473	1.7	426	81.0	473	57.3

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

1. Meat from broilers.

2. Meat from spent hens.

Multi-resistance among C. jejuni and C. coli isolates from meat from broilers and spent hens

As only three MSs and one non-MS reported isolate-based resistance data on 10 or more isolates of *C. jejuni* and *C. coli*, respectively, from broiler meat, the corresponding multi-resistance analysis is not presented in this report.

4.4.2. Antimicrobial resistance in *Campylobacter* isolates from animals

4.4.2.1. Domestic fowl (Gallus gallus): broilers

Representative sampling and monitoring

In this section, data on antimicrobial resistance in *Campylobacter* isolates from fowl (*Gallus gallus*) are completely derived from broilers. The vast majority of samples was collected from healthy broilers at the slaughterhouse. Entire caeca or caecal content samples were collected in Austria, the Czech Republic, Spain and Sweden, cloacal swabs in Denmark and Switzerland, and faeces before slaughter in Finland. In contrast, in Romania, sampling of faeces was carried out on the farm in the framework of the national control programme for *Salmonella*. For the majority of the MSs specifying details of the sampling strategy, sampling was randomised throughout the year, with the exception of Finland, where sampling was more intense over the high-risk period of the summer months. In accordance with EFSA's recommendations (EFSA, 2007), only one representative sample of caecal content per flock/batch, derived from either a unique carcase or a number of carcases, was gathered to account for clustering. Typically, given the relatively high prevalence of *Campylobacter* in broilers, representative subsets of *C. jejuni* and *C. coli* isolates recovered from caecal samples, each representing one flock, were randomly selected at the laboratory for susceptibility testing. No information on the sampling strategy was provided by two MSs (Hungary and the Netherlands).

Resistance levels among C. jejuni and C. coli isolates from broilers

For 2012, quantitative data on *C. jejuni* isolates from broilers were provided by 10 MSs and one non-MS (Table CA10), while quantitative data on *C. coli* isolates were submitted by six MSs and one non-MS (Table CA11). Generally, in both *C. jejuni* and *C. coli*, resistance to gentamicin and erythromycin was either undetected or recorded at low to moderate levels, while resistance to tetracyclines and quinolones (ciprofloxacin and nalidixic acid) was high to extremely high among reporting MSs. A striking exception to this was the low to moderate resistance in *C. jejuni* isolates reported by the Nordic countries (Denmark, Finland and Sweden); corresponding resistance data on *C. coli* were not detected by these MSs. Romania also reported low levels of resistance to quinolones in *C. jejuni*. Typically, resistance in *C. coli* was either similar or greater than that observed in *C. jejuni* in those MSs reporting results for both *Campylobacter* species. Considering *C. jejuni* and *C. coli*, levels of resistance to ciprofloxacin and nalidixic acid were rather



similar within each species as expected. Resistance to erythromycin and gentamicin in *C. jejuni* showed differences between reporting MSs, with either no resistance detected or resistance detected at low levels.

Generally, resistance in *C. coli* and *C. jejuni* from broiler meat and broilers was reported at rather similar levels in the MSs reporting data on both animal and meat origins. An exception to this was resistance to quinolones (nalidixic acid and ciprofloxacin) in Romania where resistance levels in broilers (3.6 %) proved to be much lower than those recorded in isolates from broiler meat, which were approximately 80 %.

Table CA10. Resistance (%) to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines among <u>Campylobacter jejuni</u> from <u>Gallus gallus (broilers</u>) in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Ciprof	loxacin	Erythromycin		Gentamicin		Nalidixic acid		Tetracyclines	
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	108	75.9	108	0	108	0	108	64.8	108	29.6
Czech Republic	39	84.6	39	0	39	2.6	39	76.9	39	35.9
Denmark	41	14.6	41	0	41	0	41	14.6	41	14.6
Finland	83	2.4	83	0	83	0	83	2.4	83	2.4
France	49	49.0	49	4.1	49	0	49	46.9	49	49.0
Hungary	46	87.0	46	0	46	0	46	84.8	46	43.5
Netherlands	102	61.8	102	0	102	0	102	61.8	102	59.8
Romania	83	3.6	83	0	83	2.4	83	3.6	83	51.8
Spain	32	96.9	32	3.1	32	6.3	32	93.8	32	90.6
Sweden	100	17.0	100	0	100	0	100	17.0	100	2.0
Total (10 MSs)	683	44.1	683	0.4	683	0.7	683	41.4	683	34.1
Switzerland	171	33.3	171	0	171	0	171	33.3	171	22.2

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

Table CA11. Resistance (%) to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines among <u>Campylobacter coli</u> from <u>Gallus gallus (broilers</u>) in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Ciprof	loxacin	Erythromycin		Gentamicin		Nalidixic acid		Tetracyclines	
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria	33	66.7	33	0	33	0	33	66.7	33	45.5
Czech Republic	17	94.1	17	0	17	5.9	17	76.5	17	41.2
France	78	65.4	78	15.4	78	0	78	62.8	78	92.3
Hungary	63	79.4	63	1.6	63	0	63	81.0	63	52.4
Netherlands	23	82.6	23	21.7	23	4.3	23	82.6	23	69.6
Spain	54	96.3	54	22.2	54	16.7	54	90.7	54	98.1
Total (6 MSs)	268	78.4	268	11.2	268	4.1	268	75.7	268	73.1
Switzerland	14	50.0	14	14.3	14	0	14	50.0	14	50.0

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

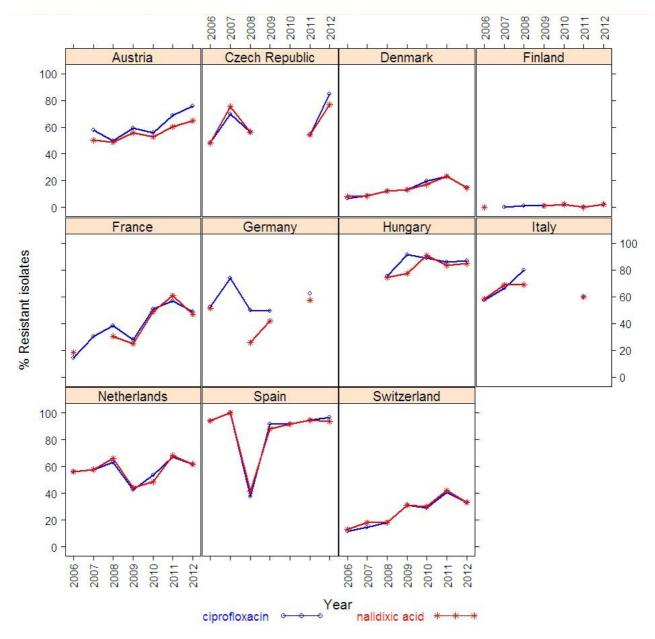
Temporal trends in resistance among *C. jejuni* isolates from broilers

Figures CA8 and CA9 present the observed temporal trends in antimicrobial resistance in *C. jejuni* isolates from *Gallus gallus* over the period 2006 to 2012. As in previous years, resistance to ciprofloxacin and nalidixic acid varied greatly among reporting MSs in 2012. When considering resistance to both ciprofloxacin and nalidixic acid, statistically significant increasing trends were observed in Austria, Denmark, France, Spain and Switzerland for five or more years (Figure CA8). For erythromycin, levels of resistance remained absent or low over the period 2006 to 2012 and a statistically significant decreasing trend in erythromycin resistance was detected in the Czech Republic and Hungary over the reporting period (Figure CA9). With regards to gentamicin, resistance remained generally at levels lower than 10 % with slight fluctuations for all reporting countries over the period 2006 to 2012 (data not shown). For tetracyclines, important variations in



resistance levels were observed among reporting countries and statistically significant increasing trends were seen in the Czech Republic, Denmark, the Netherlands and Spain over the period 2006 to 2012 (data not shown).

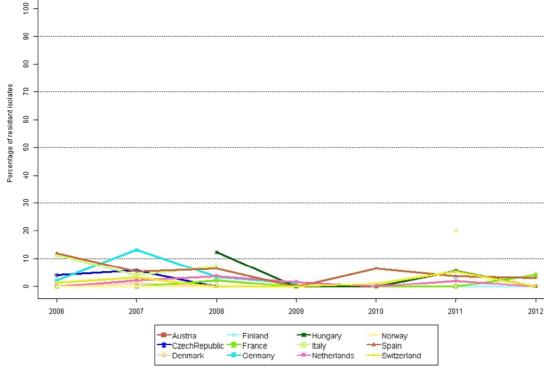
Figure CA8. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in <u>Campylobacter jejuni</u> from <u>Gallus gallus</u> in reporting MSs and non-MSs, 2006–2012, quantitative data



MS: Member State.

Note: A statistically significant increasing trend over five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed for both ciprofloxacin and nalidixic acid in Austria (↑), Denmark (↑), France (↑), Spain (↑) and Switzerland (↑), for ciprofloxacin in the Czech Republic (↑) and for nalidixic acid in Hungary (↑).

Figure CA9. Trends in <u>erythromycin</u> resistance in <u>Campylobacter jejuni</u> from <u>Gallus gallus</u> in reporting MSs and non-MSs, 2006–2012, quantitative data



MS: Member State.

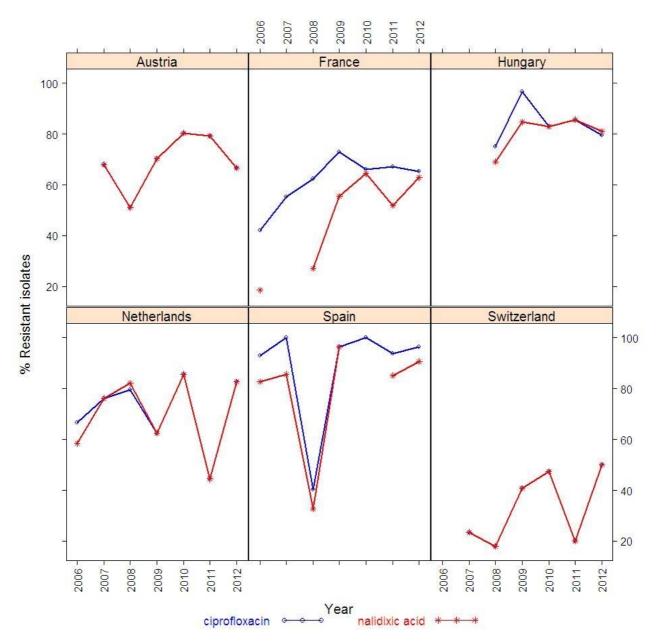
Note: A statistically significant decreasing trend over five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed in the Czech Republic (↓) and Hungary (↓).

Temporal trends in resistance among C. coli isolates from broilers

Figures CA10 and CA11 present observed trends in antimicrobial resistance in *C. coli* from *Gallus gallus*. In 2012, as was the case in previous years, a high degree of variation was observed in levels of resistance to ciprofloxacin and nalidixic acid among reporting MSs. For ciprofloxacin and nalidixic acid, statistically significant increasing trends, for the last five or more years, were observed in Austria, France, Spain and Switzerland (Figure CA10). For erythromycin (Figure CA11), resistance was generally lower over the reporting period than for the other antimicrobials presented. A similar situation was observed for gentamicin over the period with resistance levels reported lower than 10 % with the exception of Spain (data not shown). Resistance to erythromycin and gentamicin increased significantly over the seven years presented in Spain. Over the same period, resistance levels to tetracyclines varied from 30 % to 100 % for most of the reporting countries. France, Spain and Switzerland also exhibited statistically increasing trends in resistance to tetracyclines (data not shown).



Figure CA10. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in <u>Campylobacter coli</u> from <u>Gallus gallus</u> in reporting MSs and one non-MS, 2006–2012, quantitative data

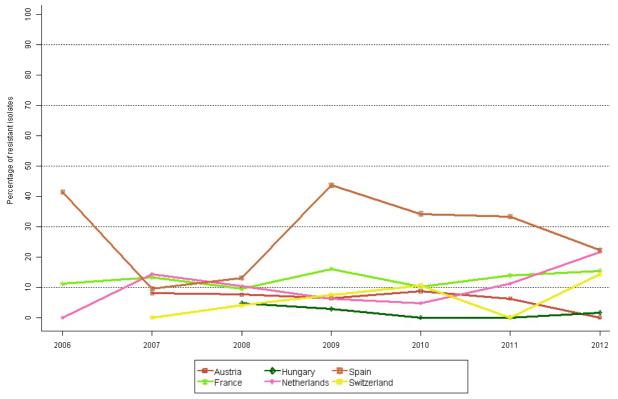


MS: Member State.

Note: A statistically significant increasing trend over five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed for both ciprofloxacin and nalidixic acid in Austria (↑), France (↑), Spain (↑) and Switzerland (↑) and for nalidixic acid in Hungary (↑).



Figure CA11. Trends in <u>erythromycin</u> resistance in <u>Campylobacter coli</u> from <u>Gallus gallus</u> in reporting MSs and one non-MS, 2006–2012, quantitative data



MS: Member State.

Note: A statistically significant increasing trend over five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed in Spain (↑).

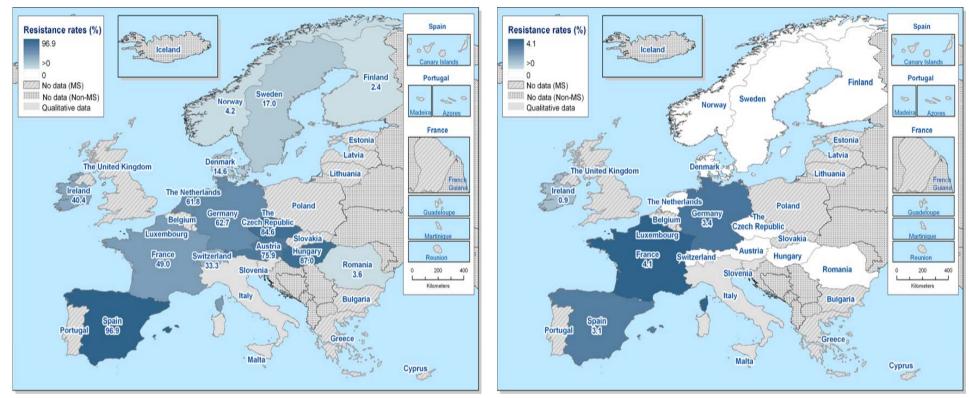
Spatial distribution of resistance among C. jejuni isolates from broilers

Figures CA12 and CA13 show the spatial distribution of ciprofloxacin and erythromycin resistance in *C. jejuni* from *Gallus gallus*. For both antimicrobials, overall resistance was lower among the reporting Nordic countries than in the rest of the European reporting countries.



*Figure CA12. Spatial distribution of <u>ciprofloxacin</u> resistance among <u>Campylobacter jejuni</u> from <u>broilers</u> of Gallus gallus in countries reporting <i>MIC data in 2012*¹

*Figure CA13. Spatial distribution of <u>erythromycin</u> resistance among <u>Campylobacter jejuni</u> from <u>broilers</u> of Gallus gallus in countries reporting <i>MIC data in 2012*¹



Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead. MIC: minimum inhibitory concentration.

1. For Germany, Ireland and Norway, 2011 data were used.



Multi-resistance among C. jejuni and C. coli isolates from broilers

In 2012, five MSs and one non-MS reported isolate-based data on resistance in *C. jejuni* isolates from broilers, while three MSs and one non-MS provided isolate-based data regarding resistance in *C. coli* isolates from broilers.

Analysis of the multi-resistance showed that there was a large variation in the levels of complete susceptibility among the reporting countries. Complete susceptibility to the common set of antimicrobials for *Campylobacter* (five antimicrobials) was generally found in more than 20 % of the *C. jejuni* isolates tested in the reporting MSs, and reached up to 78.0 % in Denmark and 80.0 % in Sweden, while in Hungary and Spain the proportion of fully susceptible isolates was much lower. In *C. coli* isolates, complete susceptibility was generally either comparable or lower than that observed in *C. jejuni*.

Multiple resistance (reduced susceptibility to three or more antimicrobial classes) was not recorded or was detected at levels lower than 5 % in *C. jejuni* isolates in most reporting countries, while in Spain 9.4 % of isolates exhibited multi-resistance (Table CA12). In *C. coli* the occurrence of multi-resistance was either comparable or greater than that reported in *C. jejuni* isolates (Table CA13). The frequency distributions (Figures CA14 and CA15) showed variation between different reporting countries. Most of the reporting countries detected resistance to a maximum of three antimicrobial classes in *C. jejuni* (Figure CA14), whereas multi-resistant *C. coli* isolates generally displayed reduced susceptibility to three to five different classes of antimicrobials (Figure CA15).

The important co-resistance for public health, i.e. resistance to both ciprofloxacin and erythromycin, was generally undetected in *C. jejuni* isolates, while a number of *C. coli* isolates from Hungary, Spain and Switzerland exhibited such co-resistance. The term co-resistance has been defined as two or more resistance genes which are genetically linked, i.e. located adjacent or close to each other on a mobile genetic element (Chapman, 2003). For brevity, the term is used slightly more loosely in this report and indicates two or more phenotypic resistances to different classes of antimicrobials, exhibited by the same bacterial isolate.



Table CA12. Complete susceptibility, multi-resistance and index of Table CA13. Complete susceptibility, multi-resistance and index of diversity in Campylobacter jejuni from broilers in MSs and non-MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Multi-re	esistant	Index of diversity	Co-resistant to Cip and Ery		
	n	%	n	%		n	%	
Austria (N=108)	22	20.4	4	3.7	0.283	0	0	
Denmark (N=41)	32	78.0	0	0	0.202	0	0	
Hungary (N=46)	4	8.7	0	0	0.21	0	0	
Spain (N=32)	1	3.1	3	9.4	0.313	1	3.1	
Sweden (N=100)	80	80.0	0	0	0	0	0	
Switzerland (N=171)	98	57.3	1	0.6	0.23	0	0	

diversity in Campylobacter coli from broilers in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscept	ible to all	Multi-r	esistant	Index of diversity	Co-resistant to Cip and Ery	
	n	%	n	%		n	%
Austria (N=33)	6	18.2	1	3.0	0.284	0	0
Hungary (N=63)	9	14.3	3	4.8	0.307	1	1.6
Spain (N=54)	1	1.9	30	55.6	0.793	12	22.2
Switzerland (N=14)	4	28.6	3	21.4	0.549	2	14.3

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Campylobacter; n: number of isolates.

Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Campylobacter.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three antimicrobial families from the common antimicrobial set.

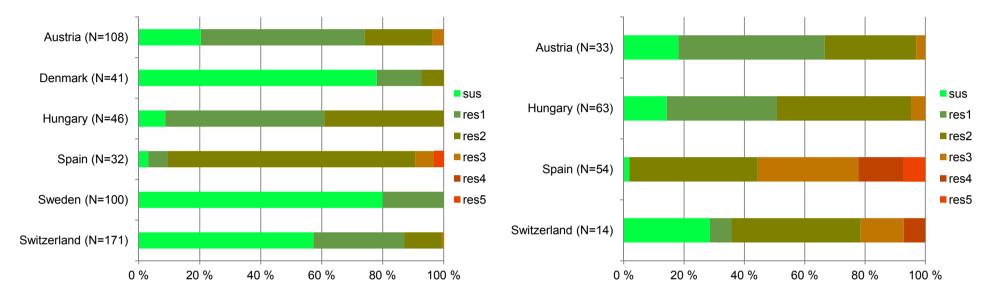
Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to Cip and Ctx: the frequencies and percentages of isolates not susceptible to ciprofloxacin concentrations >1 mg/L and erythromycin concentrations >16 mg/L for C. coli and to ciprofloxacin concentrations >1 mg/L and erythromycin concentrations >16 mg/L for C. jejuni.



completely susceptible and resistant to one to five antimicrobials in broilers in MSs and non-MSs reporting isolate-based data, 2012

Figure CA14. Frequency distribution of Campylobacter jejuni isolates Figure CA15. Frequency distribution of Campylobacter coli isolates completely susceptible and resistant to one to five antimicrobials, in broilers in MSs and one non-MS reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Campylobacter; sus: susceptible to all antimicrobial substances of the EFSA common set for Campylobacter, res1-res5: resistance to one antimicrobial substance/resistance to five antimicrobial substances of the common set for Campylobacter.



Multi-resistance patterns in *C. jejuni* and *C. coli* isolates from broilers

Considering *C. jejuni,* isolate-based data were available from five contributing MSs which reported details of 498 isolates. The isolates reported by Denmark (N=41), Hungary (N=46) and Sweden (N=100) are not addressed in this table as they were not multi-resistant. Analysis of the patterns of resistance to erythromycin, ciprofloxacin, tetracyclines, streptomycin and gentamicin was possible for 164 *C. coli* isolates from four contributing MSs which provided isolate-based data. Tables CA14 and CA15 summarise the different resistance patterns obtained and show the frequency with which these patterns were observed in each MS and for the MS group as a whole. Considering the isolates showing multi-resistance (i.e. resistance to any three or more antimicrobials of those listed above), the proportion of resistant isolates is expressed both as a proportion of the total multi-resistant isolates and as a proportion of the total number of *C. jejuni* isolates contributed by the reporting MS group.

Among the 311 *C. jejuni* isolates from broilers from the reporting group of MSs submitting isolate-based data, 2.6 % (n=8) exhibited multi-resistance (Table CA14). The commonest pattern of multi-resistance was resistance to ciprofloxacin, tetracyclines and streptomycin, occurring in six of eight resistant isolates reported by submitting MSs. A single isolate was reported which showed resistance to all of the antimicrobials in the multi-resistance panel, although this only accounted for 0.3 % of the total *C. jejuni* isolates for which isolate-based data was available. The situation differed in *C. coli* where 2.4 % of the total number of *C. coli* isolates, for which isolate-based data was available, showed multi-resistance to all of the antimicrobials in the test panel.

Table CA14. Multi-resistance patterns of interest in Campylobacter jejuni from broilers in	MSs one
non-MS reporting isolate-based data, 2012	

Multi-resistance pattern					Group reporti countr (N=31	ng ies	Austria (N=108)	Spain (N=32)	Switzerland (N=171)	
Ery	Cip	Tet	Str	Gen	n	%	group %	n	n	n
	R	R	R		6	75.0	1.9	4	1	1
	R	R		R	1	12.5	0.3	0	1	0
R	R	R	R	R	1	12.5	0.3	0	1	0
Total				8	100	2.6	4	3	1	

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*, Ery: erythromycin; Cip: ciprofloxacin; Tet: tetracyclines; Str: streptomycin; Gen: gentamicin; n: number of multi-resistant isolates; R: minimum inhibitory concentration above the European Committee on Antimicrobial Susceptibility Testing epidemiological cut-off values.

The proportion of isolates of *C. jejuni* from broilers showing multi-resistance, as well as the diversity of multi-resistance within that multiple resistant population, was lower than that observed in *C. coli* from broilers. The proportion of the total number of *C. coli* isolates from broilers showing resistance to all antimicrobials in the panel (2.4 %) was identical to the figure found for pigs for *C. coli* isolates which showed an identical pattern of resistance (2.4 %).

Among the 164 *C. coli* isolates from broilers tested in the reporting group of MSs submitting isolate-based data, 22.6 % (n=37) showed multi-resistance to three or more antimicrobials (Table CA15). The commonest resistance pattern detected was resistance to ciprofloxacin, tetracyclines and streptomycin occurring in 10.4 % of the multi-resistant isolates. This pattern of resistance together with additional erythromycin resistance occurred in 3.7 % of isolates, while a further 2.4 % of isolates demonstrated additional resistance to gentamicin. These three resistance patterns accounted for more than 70 % of the multi-resistant isolates which were detected. Gentamicin resistance has a component of multiple drug resistance patterns and was only observed in Spain. This resistance was not detected in the other reporting MSs. Interestingly, Spain was the only MS providing isolate-based data to report gentamicin resistance in *C. jejuni*, although other MSs (the Czech Republic and Romania), which did not report isolate-based data, also detected gentamicin resistance in *C. jejuni* (Table CA10). The range of different resistance patterns observed was greater in

C. coli than for *C. jejuni*; Spain contributed isolates with a greater range of different resistance patterns than other MSs, although this may have merely reflected the small isolate sample size from other MSs.

Table CA15. Multi-resistance patterns of interest in <u>Campylobacter coli</u> from <u>broilers</u> in MSs and one non-MS reporting isolate-based data, 2012

Multi-resistance pattern					Group reporti countri (N=16	ng ies	Austria (N=33)	Hungary (N=63)	Spain (N=54)	Switzerland (N=14)	
Ery	Cip	Tet	Str	Gen	n	%	group %	n	n	n	n
	R	R	R		17	46.0	10.4	1	2	13	1
R	R	R	R		6	16.2	3.7	0	0	5	1
R	R	R			4	10.8	2.4	0	1	3	0
R	R	R	R	R	4	10.8	2.4	0	0	4	0
	R	R	R	R	3	8.1	1.8	0	0	3	0
	R	R		R	2	5.4	1.2	0	0	2	0
R	R		R		1	2.7	0.6	0	0	0	1
	Total				37	100	22.6	1	3	30	3

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter* and multi-resistant; Ery: erythromycin; Cip: ciprofloxacin; Tet: tetracyclines; Str: streptomycin; Gen: gentamicin; n: number of multi-resistant isolates; R: minimum inhibitory concentration above the European Committee on Antimicrobial Susceptibility Testing epidemiological cut-off values

Comparison of resistance in broilers and meat from broilers

Considering individual MSs, the levels of ciprofloxacin resistance were generally lower in *C. coli* and *C. jejuni* isolates from meat from broilers than in isolates from broilers. Generally, resistance levels to all antimicrobials were higher in *C. coli* than in *C. jejuni* for the same host species. Similarly, the levels of multi-resistance (reduced susceptibility to at least three different antimicrobial classes) in *C. coli* isolates from broilers were much higher than those detected in *C. jejuni* isolates of the same origin. However, it should be borne in mind that, despite the high levels of resistance/multi-resistance observed, *C. coli* is much less prevalent in poultry than *C. jejuni*.



4.4.2.2. Pigs

Representative sampling and monitoring

In the reporting MSs, antimicrobial resistance monitoring in *Campylobacter* isolates from pigs was based primarily on active monitoring plans involving random sampling of healthy pig carcases at the slaughterhouse. The sampling plan was typically stratified per slaughterhouse, by allocating the number of samples collected per slaughterhouse in proportion with the annual throughput of that slaughterhouse. An approximately equal distribution of the collected samples over the year enabled the different seasons to be covered. Only one representative faecal sample per epidemiological unit (batch/farm), either derived from a unique carcase or pooled from a number of carcases, was gathered to account for clustering, in accordance with EFSA's recommendations (EFSA, 2007). In the reporting MSs, antimicrobial resistance monitoring in *Campylobacter* spp. in pigs focused on *C. coli*, as this is the more prevalent *Campylobacter* species in pigs. Because of the very low *C. jejuni* prevalence in pigs, the number of samples required to be collected to achieve a sufficient number of *C. coli* isolates would have been too large to be cost effective. In some reporting countries, representative subsets of *C. coli* isolates recovered from faecal samples were randomly selected at the laboratory for susceptibility testing, whereas, in others, all *C. coli* isolates were tested for susceptibility.

Resistance levels among *C. coli* isolates from pigs

In 2012, quantitative data were provided by five MSs and one non-MS (Switzerland) on *C. coli* isolates from pigs (Table CA16). *C. coli* isolates tested were mainly derived from fattening pigs (the production type was not specified for the Netherlands). As seen in 2011, the range of resistance to the antimicrobials studied varied greatly between the reporting countries in 2012. However, in general, the levels of resistance to tetracyclines observed were high to extremely high, while those to nalidixic acid, ciprofloxacin and erythromycin were moderate to high. Exceptions to this general pattern of resistance to these substances were observed for isolates from Denmark, which reported the lowest occurrence of resistance (at low to moderate levels), and Spain, which recorded the highest resistance, at levels classed as extremely high. Resistance to fluoroquinolones (ciprofloxacin) and quinolones (nalidixic acid) proved to be very similar in all reporting countries. In contrast, gentamicin resistance was either undetected or reported at low level. Only Spain recorded a moderate resistance to gentamicin.

Table CA16. Resistance (%) to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines among <u>Campylobacter coli</u> from <u>pigs</u>¹ in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Ciprof	loxacin	Erythr	omycin	Gent	amicin	Nalidiz	kic acid	Tetracyclines		
Country	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	
Denmark	103	11.7	103	6.8	103	0	103	11.7	103	14.6	
France	96	40.6	96	36.5	96	0	96	39.6	96	91.7	
Hungary	53	50.9	53	13.2	53	3.8	53	50.9	53	88.7	
Netherlands	232	12.5	232	11.2	232	0.4	232	12.5	232	88.4	
Spain	73	97.3	73	79.5	73	17.8	73	95.9	73	100	
Total (5 MSs)	557	32.0	557	23.9	557	2.9	557	31.6	557	76.8	
Switzerland	144	41.0	144	9.0	144	0.7	144	38.9	144	31.9	

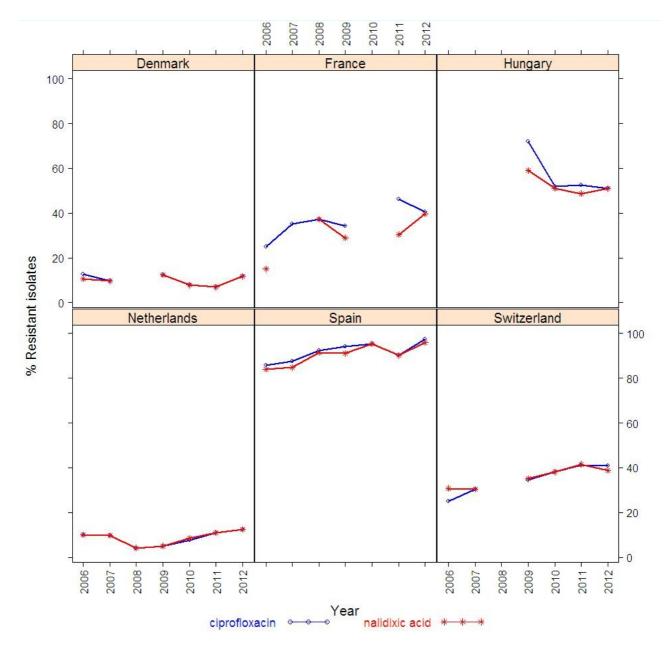
MIC: minimum inhibitory concentration; MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates.
1. The *C. coli* isolates tested were derived from fattening pigs. For the Netherlands, the production level was not specified.



Temporal trends in resistance among *C. coli* isolates from pigs

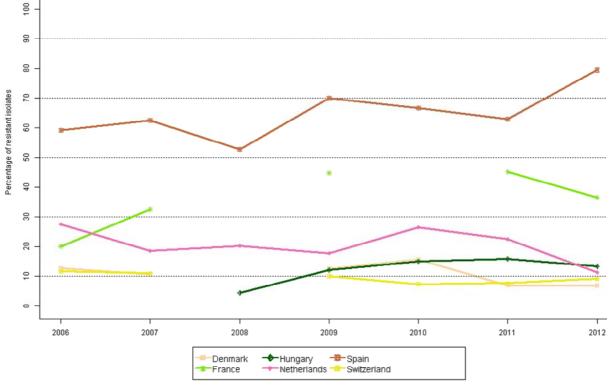
Figures CA16 and CA17 show the trends in antimicrobial resistance observed in *C. coli* from pigs over the period 2006–2012. For most of the antimicrobials considered, levels of resistance have remained relatively stable between 2006 and 2012. For ciprofloxacin and nalidixic acid, a statistically significant increasing trend was seen for Spain, while France and Switzerland reported significantly increasing levels of resistance to ciprofloxacin over the reporting period. Levels of erythromycin resistance increased significantly in France, the Netherlands and Spain and, for gentamicin, resistance increased significantly in Spain (data not shown). When considering tetracyclines (data not shown), marked differences in resistance were observed among reporting countries: Switzerland and Denmark reported resistance of around 30 % and 10 %, respectively, while France, Hungary, the Netherlands and Spain recorded levels greater than 80 %, over the 2006 to 2012 period. Resistance to tetracyclines demonstrated a significantly increasing trend in Denmark during the period 2006 to 2012.

Figure CA16. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in <u>Campylobacter coli</u> from <u>pigs</u> in reporting MSs and one non-MS, 2006–2012, quantitative data



Note: A statistically significant increasing trend over five or more years, as tested by a logistic regression model ($p \le 0.05$), was observed in Spain (\uparrow) for both ciprofloxacin and nalidixic acid and in France (\uparrow) and Switzerland (\uparrow) for ciprofloxacin; MS: Member State.





MS: Member State.

Note: A statistically significant increasing trend over five or more years, as tested by a logistic regression model (*p* ≤0.05), was observed in France (↑), the Netherlands (↑) and Spain (↑);

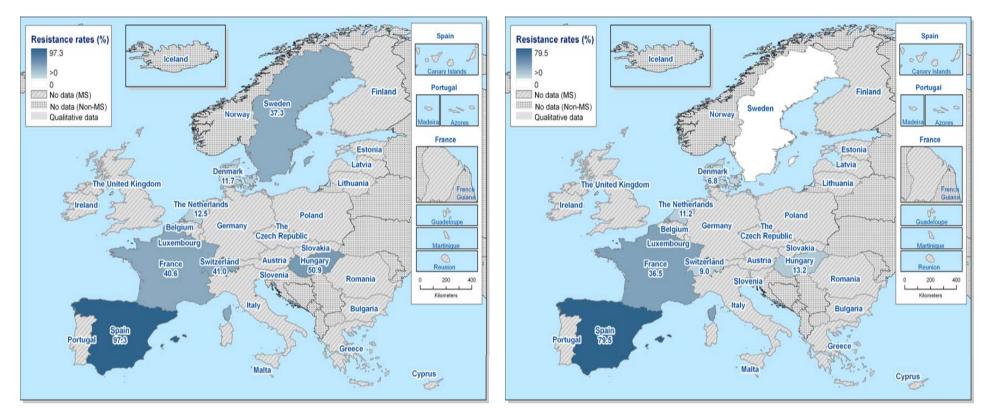
Spatial distribution of resistance among C. coli isolates from pigs

Figures CA18 and CA19 show the spatial distributions of ciprofloxacin and erythromycin resistance in *C. coli* from pigs. For both erythromycin and ciprofloxacin, the highest levels of resistance were reported by Southern countries, while Northern countries reported lower levels.



Figure CA18. Spatial distribution of <u>ciprofloxacin</u> resistance among <u>Campylobacter coli</u> from <u>pigs</u> in countries reporting MIC data in 2012¹

Figure CA19. Spatial distribution of <u>erythromycin</u> resistance among <u>Campylobacter coli</u> from <u>pigs</u> in countries reporting MIC data in 2012¹



MIC: minimum inhibitory concentration; MS: Member State.

Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data have been used instead.

1. For Sweden, 2011 data were used.



Multi-resistance among C. coli isolates from pigs

In 2012, three MSs and one non-MS reported isolate-based data on resistance in *C. coli* isolates from pigs. Analysis of the multi-resistance showed that there was a large variation in the levels of complete susceptibility and multi-resistance among the reporting countries. Isolates exhibiting complete susceptibility accounted for 33.0 % in Denmark and 14.6 % in Switzerland, while in Hungary and Spain none of the isolates tested were completely susceptible (Table CA17). Conversely, multi-resistance was low in Denmark (3.9 %), moderate in Switzerland (19.4 %), high in Hungary (49.1 %) and extremely high in Spain (97.3 %). The frequency distributions (Figure CA20) showed an important diversity between the reporting countries. Hungary, Spain and Switzerland reported isolates displaying reduced susceptibility to up to four or five different classes of antimicrobials. In addition, a high proportion of isolates showing co-resistance to ciprofloxacin and erythromycin was observed in isolates from Spain.

Table CA17. Complete susceptibility, multi-resistance and index of diversity in Campylobacter coli from <u>fattening pigs</u> in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscept	ible to all	Multi-r	esistant	Index of		istant to nd Ery
	n	%	n	%	diversity	n	%
Denmark (N=103)	34	33.0	33.0 4 3.9		0.28	3	2.9
Hungary (N=53)	0	0	26	49.1	0.662	5	9.4
Spain (N=73)	0	0	71	97.3	0.74	56	76.7
Switzerland (N=144)	21	14.6	28	19.4	0.471	6	4.2

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*, n: number of isolates; Cip: ciprofloxacin; Ery: erythromycin.

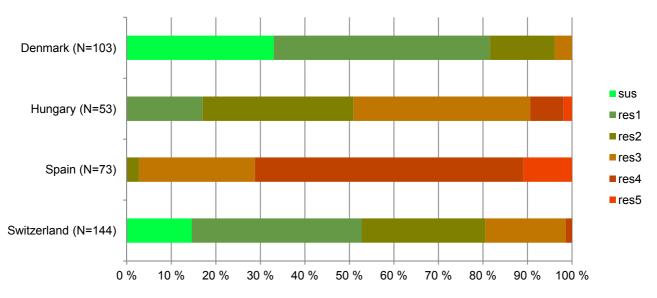
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Campylobacter.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three antimicrobial families from the common antimicrobial set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to Cip and Ery: the frequencies and percentages of *C. coli* isolates not susceptible to ciprofloxacin concentrations >1 mg/L and erythromycin concentrations >16 mg/L.

Figure CA20. Frequency distribution of <u>Campylobacter coli</u> isolates completely susceptible and resistant to one to five antimicrobials, in <u>fattening pigs</u> in MSs and one non-MS reporting isolate-based data, 2012



N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*; MS: Member State; sus: susceptible to all antimicrobial substances of the EFSA common set for *Campylobacter*; res1–res5: resistance to one antimicrobial substance/resistance to five antimicrobial substances of the common set for *Campylobacter*.



Multi-resistance pattern in C. coli isolates from fattening pigs

Isolate-based data was available for 373 C. coli isolates from fattening pigs, contributed by three reporting MSs and one non-MS, from which 34.6 % exhibited different patterns of multi-resistance (Table CA18). The commonest multi-resistance pattern observed in fattening pigs was resistance to ciprofloxacin, tetracyclines and streptomycin, occurring in 13.1 % of the total number of isolates for which isolate-based data was available. The next most common pattern of multi-resistance comprised resistance to the preceding antimicrobials, together with resistance to erythromycin. Taken together, these patterns of resistance accounted for more than 70 % of the total multiple drug resistance C. coli isolates from pigs. The range of multi-resistance patterns observed in C. coli from pigs was greater than that observed in broilers and unlike in broilers, gentamicin resistance, as part of the multi-resistance pattern, was observed in three of four reporting MSs. Although Denmark contributed a large number of isolates (N=103), the diversity of resistance patterns observed in Denmark, and the numbers of multi-resistant isolates as a proportion of the total isolates reported, was lower than for the remaining three MSs which reported data. Most (greater than 85%) multiply resistant C. coli isolates from pigs were resistant to tetracyclines and streptomycin as a component of the multi-resistance pattern. Most isolates which were resistant to gentamicin were also resistant to streptomycin. Resistance to ciprofloxacin and tetracyclines was observed in more than 90 % of multiresistance C. coli isolates from pigs.

Table CA18. Multi-resistance patterns of interest in Campylobacter coli from fattening pigs in MSs and one non-MS reporting isolate-based data, 2012

	Multi-re	esistanc	e patteri	n		Group report count (N=3	ing ries	Denmark (N=103)	Hungary (N=53)	Spain (N=73)	Switzerland (N=144)
Ery	Cip	Tet	Str	Gen	n	%	group %	n	n	n	n
	R	R	R		49	38.0	13.1	2	18	11	18
R	R	R	R		43	33.3	11.5	0	3	39	1
R	R	R			10	7.8	2.7	1	1	7	1
R	R	R	R	R	9	7.0	2.4	0	1	8	0
R		R	R		7	5.4	1.9	0	2	1	4
	R	R	R	R	4	3.1	1.1	0	1	2	1
R	R		R		4	3.1	1.1	1	0	0	3
R	R	R		R	2	1.6	0.5	0	0	2	0
R	R R R				1	0.8	0.3	0	0	1	0
	Total				129	100	34.6	4	26	71	28

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter* and multi-resistant; Ery: erythromycin; Cip: ciprofloxacin; Tet: tetracyclines; Str: streptomycin; Gen: gentamicin; n: number of multi-resistant isolates; R: minimum inhibitory concentration above the European Committee on Antimicrobial Susceptibility Testing epidemiological cut-off values.



4.4.2.3. Cattle (bovine animals)

Representative sampling and monitoring

In 2012, data on antimicrobial resistance among *C. jejuni* isolates from cattle include samples collected both at the slaughterhouse (Denmark, Finland, Germany, Spain and Switzerland) and at the farm level (the Netherlands). These countries tested different production types and ages of cattle, including veal calves, young meat production animals, adult cattle and dairy cows; Denmark and Finland did not specify the type of cattle which were tested. Slaughterhouse sampling programmes were randomised over the year and stratified by the number of slaughtered animals by abattoirs across the MSs. The sampling was evenly distributed throughout the year or a significant part of the year to account for a possible seasonal effect. Only one caecal or faecal sample per bovine animal carcase was collected. In the reporting MSs, antimicrobial resistance monitoring in *Campylobacter* spp. in cattle focused on *C. jejuni*, as this is the more prevalent *Campylobacter* species in cattle. In some reporting countries, representative subsets of *Campylobacter* isolates recovered from animal samples were randomly selected at the laboratory for susceptibility testing, while, in some others, all isolates were tested for susceptibility.

Resistance levels among C. jejuni isolates from cattle

For 2012, five MSs and one non-MS provided quantitative data on *C. jejuni* isolates from cattle (Table CA19). *C. jejuni* isolates tested were derived from veal calves (Germany, the Netherlands), young cattle (Spain, Switzerland), dairy cows (the Netherlands) and production type unspecified (Denmark and Finland). As seen in 2011, the range of resistance to the antimicrobials studied varied greatly between the reporting countries in 2012. The levels of resistance to ciprofloxacin, nalidixic acid and tetracyclines were generally high, while resistance to erythromycin and gentamicin was either not detected or recorded at low to very low levels.

Country	Cipro	loxacin	Erythr	omycin	Genta	amicin	Nalidi	xic acid	Tetrac	cyclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
All cattle										
Denmark	89	14.6	89	1.1	89	0	89	15.7	89	0
Finland	72	13.9	72	0	72	0	72	13.9	72	2.8
Germany	73	49.3	73	2.7	73	0	73	43.8	73	76.7
Netherlands	178	31.5	178	0	178	0.6	178	31.5	178	61.8
Spain	68	63.2	68	0	68	0	68	64.7	68	60.3
Total (5 MSs)	480	32.9	480	0.6	480	0.2	480	32.5	480	43.5
Switzerland	38	36.8	38	2.6	38	0	38	39.5	38	44.7
Veal calves								-		-
Germany	73	49.3	73	2.7	73	0	73	43.8	73	76.7
Netherlands	137	38.7	137	0	137	0.7	137	38.7	137	78.1
Total (2 MSs)	210	42.4	210	1.0	210	0.5	210	40.5	210	77.6
Young cattle (1	-2 years)								
Spain	68	63.2	68	0	68	0	68	64.7	68	60.3
Switzerland	38	36.8	38	2.6	38	0	38	39.5	38	44.7
Dairy cows										
Netherlands	41	7.3	41	0	41	0	41	7.3	41	7.3
Unspecified ty	be of cat	le						-	÷	-
Denmark	89	14.6	89	1.1	89	0	89	15.7	89	0
Finland	72	13.9	72	0	72	0	72	13.9	72	2.8
Total (2 MSs)	161	14.3	161	0.6	161	0	161	14.9	161	1.2

Table CA19. Resistance (%) to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines among Campylobacter jejuni from cattle¹ in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

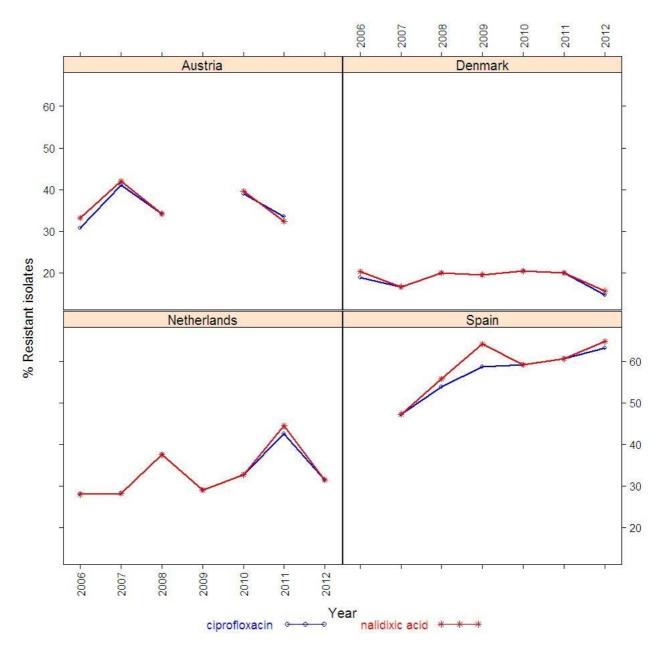
1. Data presented in this table were derived from a variety of production types. These include veal calves (Germany, the Netherlands), young cattle (Spain, Switzerland), dairy cows (the Netherlands) and production type unspecified (Denmark and Finland).



Temporal trends in resistance among C. jejuni isolates from cattle

Figures CA21 and CA22 show the temporal trends in resistance for *C. jejuni* from cattle. As seen in *C. coli* in pigs, levels of resistance for *C. jejuni* in cattle have remained relatively stable over the 2006–2012 reporting period for individual MSs. In general, resistance to ciprofloxacin, nalidixic acid and tetracyclines was relatively higher than levels of resistance to erythromycin and gentamicin for the reporting MSs. When considering trends in ciprofloxacin, nalidixic acid (Figure CA21) and gentamicin resistance, no significant changes were observed over the reporting period. For erythromycin, a significantly decreasing trend was observed in the Netherlands when tested by a logistic regression model (Figure CA22), and for tetracyclines a significantly decreasing trend was observed in Austria and a significantly increasing trend was observed in the Netherlands (data not shown).

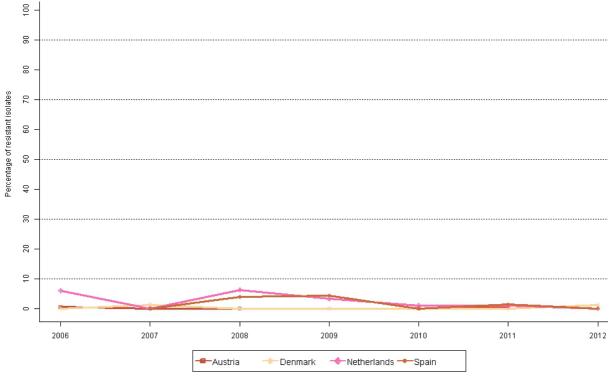
Figure CA21. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in <u>Campylobacter jejuni</u> from <u>cattle</u> in reporting MSs, 2006–2012, quantitative data



MS: Member State.

Note: No statistically significant trends over five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in any of the reporting countries;





MS: Member State.

Note: A statistically significant decreasing trend over five or more years, as tested by a logistic regression model ($p \le 0.05$), was observed in the Netherlands (\downarrow);

Multi-resistance among C. jejuni isolates from cattle

In 2012, three MSs and one non-MS reported isolate-based data on resistance in *C. jejuni* isolates from differing cattle populations (Table CA20). Difference in the cattle populations monitored may most likely partly explain the variability observed in summary indicators of multi-resistance among reporting countries. Germany, which monitored veal calves specifically, reported the highest level of multi-resistance (12.3 %) and the lowest level of complete susceptibility (16.4 %) among the reported data, while Spain and Switzerland, which monitored young cattle (1–2 years), recorded high levels of complete susceptibility and similar low levels of multi-resistance at around 5 %. Conversely, Denmark, which reported resistance data on an unspecified cattle population, recorded an extremely high level of complete susceptibility and did not detect any multi-resistant *C. jejuni* isolates. While the resistant isolates monitored were only resistant to one substance in Denmark, other reporting countries (Germany, Spain and Switzerland) detected isolates displaying reduced susceptibility to up to three or four different classes of antimicrobials (Figure CA23). In addition, only a few isolates exhibited co-resistance to both ciprofloxacin and erythromycin.



Table CA20. Complete susceptibility, multi-resistance and index of diversity in Campylobacter jejuni from cattle in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscepti	ble to all	Multi-r	esistant	Index of		istant to Ind Ery
	n	%	n	%	diversity	n	%
Denmark (N=89)	75	84.3	0	0	0	0	0
Germany (N=73)	12	16.4	9	12.3	0.434	2	2.7
Spain (N=68)	17	25.0	4	5.9	0.323	0	0
Switzerland (N=38)	18	47.4	2	5.3	0.428	1	2.6

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*, n: number of isolates; Cip: ciprofloxacin; Ery: erythromycin.

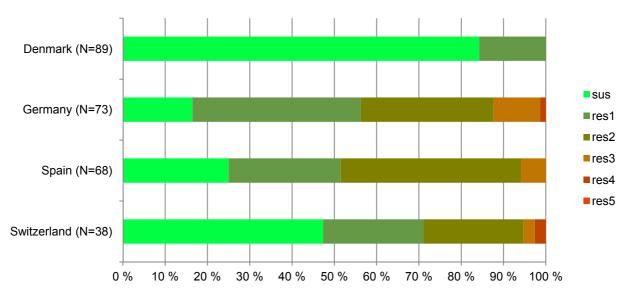
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for Campylobacter.

Multi-resistant: resistant to at least three different antimicrobial substances, belonging to any three antimicrobial families from the common antimicrobial set.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to Cip and Ery: the frequencies and percentages of *C. jejuni* isolates not susceptible to ciprofloxacin concentrations >1 mg/L and erythromycin concentrations >4 mg/L.

Figure CA23. Frequency distribution of <u>Campylobacter jejuni</u> isolates completely susceptible and resistant to one to five antimicrobials, in <u>cattle</u> in MSs and one non-MS reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter*, sus: susceptible to all antimicrobial substances of the EFSA common set for *Campylobacter*, res1–res5: resistance to one antimicrobial substance/resistance to five antimicrobial substances of the common set for *Campylobacter*.



Multi-resistance pattern in C. jejuni isolates from cattle

Isolate-based data were available for 268 *C. jejuni* isolates submitted by four reporting countries. Denmark did not detect multi-resistance in *C. jejuni* from cattle. In the remaining reporting countries the most common pattern of multi-resistance was resistance to ciprofloxacin, tetracyclines and streptomycin, occurring in 80 % of multi-resistance isolates (Table CA21). Gentamicin resistance was not detected in multi-resistance isolates of *C. jejuni* from cattle.

Table CA21. Multi-resistance patterns of interest in <u>Campylobacter jejuni</u> from <u>cattle</u> in MSs and one non-MSs reporting isolate-based data, 2012

	Multi-r	esistanc	e pattern			repo coun	up of rting ntries 179)	Spain (N=68)	Switzerland (N=38)	
Ery	Cip	Tet	Str	Gen	n	%	group %	n	n	n
	R	R	R		12	80.0	6.7	7	4	1
R	R	R	R		2	13.3	1.1	1	0	1
R	R R R				1	6.7	0.6	1	0	0
	Total				15	100	8.4	9	4	2

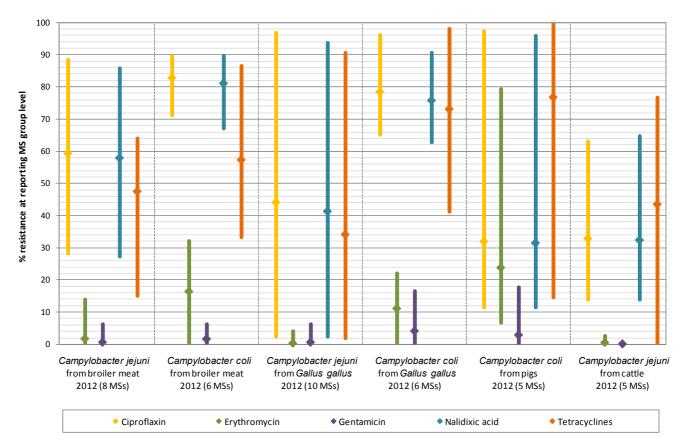
MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Campylobacter* and multi-resistant; Ery: erythromycin; Cip: ciprofloxacin; Tet: tetracyclines; Str: streptomycin; Gen: gentamicin; n: number of multi-resistant isolates; R: minimum inhibitory concentration above the European Committee on Antimicrobial Susceptibility Testing epidemiological cut-off values.



4.5. Overview of the findings on antimicrobial resistance in *Campylobacter* at reporting Member State group level, 2012

Figure CA24 shows the resistance levels in the reporting MS group based on the quantitative data submitted in 2012 for the various animal species and meat derived from those animal species. These data may derive from different MS groups, which should be considered when interpreting the figure. As was the case in previous years, *C. coli* isolates tended to be more resistant than *C. jejuni* isolates. Direct comparisons of the levels of resistance in *Campylobacter* from *Gallus gallus* and in broiler meat may not be entirely appropriate because different MSs have reported different proportions of isolates tested from meat and live fowl. The levels of resistance sometimes differ between different MSs and the relative contribution of individual MSs can affect the summary group level figures.

Figure CA24. Resistance to ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines in <u>Campylobacter jejuni</u> and <u>Campylobacter coli</u> from fowl, pigs and cattle at reporting MS group level in 2012



MS: Member State.



4.6. Discussion

Campylobacter causes a large number of human cases of gastro-enteritis and has been the most frequently reported cause of human food-borne zoonoses in the EU since 2004 (EFSA and ECDC, 2014). The majority of campylobacteriosis cases are self-limiting, however chronic symptoms (Guillain–Barré syndrome, acute transverse myelitis, myocarditis, and reactive arthritis) may also develop as a result of intestinal infection. Bacteraemia is very rare, except for infections with *C. fetus*. Resistance to antimicrobials in *Campylobacter* is of concern because of the large number of cases of human infection and the fact that some cases require treatment. Treatment of enteric infections in humans may involve administration of fluoroquinolones, such as ciprofloxacin, or macrolides, such as erythromycin (ECDC et al., 2009). The rapid increase in resistance in *C. jejuni* to fluoroquinolones concurrent with discoveries of associations between postinfectious irritable bowel syndrome and a longer duration of untreated *Campylobacter* infection (Kirkpatrick and Tribble, 2011) is therefore worrying.

In 2012, information on antimicrobial resistance in *Campylobacter* isolates from human cases of campylobacteriosis was collated from 14 MSs and one non-MS (Iceland). The data submitted by these countries represented isolates from 18 % of the human campylobacteriosis cases reported within the EU/EEA in 2012. A novelty in this year's report was that isolates from cases notified as having been acquired during travelling outside of the reporting country, were excluded from all analysis except the analysis on resistance in difference geographical regions. This was done to better assess the impact on *Campylobacter* isolates from food consumed within each reporting country on the antimicrobial resistance levels found in human isolates in that country. Please note however that imported food, which can constitute a large proportion of the food available in some countries, is not covered by this report.

There was a large variation in the guidelines used for interpreting the susceptibility tests for human *Campylobacter* isolates among countries. In two MSs, two different guidelines were also applied in order to cover all antimicrobials under monitoring. Although the clinical breakpoints used for the dilution test for *Campylobacter* were less variable than those for *Salmonella*, the breakpoints for disc diffusion differed significantly depending on the guidelines used, particularly for ciprofloxacin. Disc diffusion was still the most common method of testing for antimicrobial susceptibility in human isolates, sometimes in combination with gradient strip. Only four countries used micro-broth dilution. The disc diffusion method and clinical breakpoints established by EUCAST in 2012 are therefore much welcomed and is recommended by the ECDC in the EU protocol for harmonised monitoring of antimicrobial resistance in human *Salmonella* and *Campylobacter* isolates (ECDC, 2014).

There was also a large variation with regard to the number of antimicrobials tested among the reporting countries, which reflects the clinical importance of the antimicrobials. **Erythromycin** and **ciprofloxacin** were the antimicrobials for which the greatest numbers of human *Campylobacter* spp. isolates were tested. The levels of resistance in human *C. jejuni* isolates to erythromycin was overall low, but moderately high in *C. coli*, although the number of tested isolates for this bacterial species was small. Very high resistance levels to ciprofloxacin were reported in human *Campylobacter* isolates, with increasing trends observed in several MSs.

In order to assess the importance of **travel-associated infections**, antimicrobial resistance was also analysed based on the most likely country of infection reported. Human isolates acquired in Asia and Africa had the highest frequency of resistance to the antimicrobials tested, with two and three times higher resistance levels to ciprofloxacin and erythromycin than in isolates acquired within the EU/EEA. This could affect which antimicrobials could be applied for successful treatment of severe travel-associated cases.

Human antimicrobial susceptibility data were available for the full range of antimicrobials from five MSs for *C. jejuni* and from four MSs for *C. coli*. Overall, only one in six (18.0 %) human *C. jejuni* isolates and one in twenty (6.4 %) human *C. coli* isolates were fully susceptible to all antimicrobials. On average, one quarter of *C. jejuni* (24.8 %) and one third of *C. coli* (35.1 %) isolates exhibited **multi-drug resistance**, meaning that they were clinically non-susceptible to at least three different antimicrobial groups. The clinical breakpoints used to interpret the human data were, in some cases, more sensitive than the ECOFFs when intermediate and resistant results were combined. The human data also covered two penicillins which were not included in the animal/food testing. All these factors could explain the generally higher proportion of multi-resistance observed in humans compared with animals, particularly for *C. jejuni*, which was the most common species in humans. Co-resistance to the critically important antimicrobials ciprofloxacin and erythromycin was, on



average, low for *C. jejuni* (1.4 %) but detected in one out of six (16.0 %) *C. coli* isolates, although in the case of *C. coli* fewer isolates were tested.

The data relating to the susceptibility of *Campylobacter* of food and animal origin reported by MSs were, in general well harmonised with almost all MSs adopting the EFSA guidelines and recommendations. Overall, levels of antimicrobial resistance in *Campylobacter* isolates from animals and food were similar to those in 2011. Considering all reporting MSs, ciprofloxacin resistance in *C. jejuni* from *Gallus gallus* and cattle was 44.1 % and 32.9 % respectively, while in *C. coli* from *Gallus gallus* and pigs it was 78.4 % and 32.0 %.

Among *Campylobacter* isolates from *Gallus gallus* and broiler meat, very high to extremely high levels of resistance, to one or more antimicrobials, were reported by a number of MSs, with the exception of some Nordic countries, as well as Central and Eastern European countries, particularly when using ECOFFs as interpretive criteria of reduced susceptibility or 'microbiological resistance'. For example, extremely high resistance rates to **ciprofloxacin** were notably detected. Over the period 2009–2011, the highest levels of resistance to quinolones and fluoroquinolones were in general detected in *Campylobacter* isolates from *Gallus gallus*. In 2012, however, the highest levels of resistance to these antimicrobials were detected in *Campylobacter* isolates from both broiler meat and broilers, levels of resistance were generally similar, although *C. coli* isolates from broiler meat tended to exhibit slightly higher resistance compared with *C. coli* isolates from broilers. Interestingly, *C. jejuni* and *C. coli* isolates from meat from spent hens tested in Belgium presented lower levels of ciprofloxacin resistance than those observed in isolates from meat from broilers.

This high level of ciprofloxacin resistance in Campylobacter from broiler meat is of particular concern, since the EFSA Panel on Biological Hazards (BIOHAZ), in its recent scientific opinion on the quantification of the risk of campylobacteriosis posed to humans by broiler meat, estimated that the handling, preparation and consumption of broiler meat may account for 20 % to 30 % of human campylobacteriosis cases, while 50 % to 80 % of cases may be attributed to the chicken (broiler) reservoir as a whole (EFSA BIOHAZ Panel, 2010a). In 2012, ciprofloxacin resistance in C. coli isolates from humans was 69.0 % for all contributing MSs (range: 42.0-86.4 %) and 32.0 % in pigs (range: 11.7-97.3 %). However, the picture is clearly complex in relation to the sources of human infections because these may be related to consumption of pig or poultry meat (as well as other sources). International trade also means that consumers may be exposed to meat produced in a number of different countries. Similar considerations apply when comparing resistance levels in humans and animals for other resistances. However, resistance to gentamicin, in C. coli from humans, meat from broilers, broilers and pigs does show similarities at the MS level. While gentamicin resistance was reported in C. coli from Gallus gallus (16.7 %) and pigs (17.8 %) from Spain but was not reported in broilers (Austria, France), meat from broilers (Austria) or pigs (France), C. coli from human infections showed 13.6 %, 2.4 % and 1.3 % gentamicin resistance in Spain, Austria and France, respectively. However, Campylobacter strains from the broiler reservoir may also reach humans via routes other than food (e.g. the environment or by direct contact).

In 2012, ciprofloxacin resistance in *C. jejuni* isolates from humans was 54.1 % for all contributing MSs (range: 30.7 %–91.9 %) and 44.1 % in broilers (range: 2.4 %–96.9 %). These figures and the possible reasons why the levels of resistance in humans and animals may or may not show correlations have been discussed in detail in previous reports in a dedicated farm to fork chapter which has not been included in the analysis for 2012. However, similar factors will apply to the 2012 data and readers are referred to earlier EU Summary Reports for detailed discussion of the ciprofloxacin and erythromycin results in campylobacter isolates from man and animals.

Over the period 2006 to 2012, statistically significant increasing trends in ciprofloxacin and nalidixic acid resistance in *C. jejuni* from broilers were observed over five or more years in five reporting countries; this was also observed in *C. coli* from broilers in three MSs. Considering *C. coli* from pigs a statistically significant increasing trend for ciprofloxacin/nalidixic acid resistance, was observed for three reporting countries. Comparison of these results and trends with the relevant antimicrobial usage figures might reveal interesting insights into the development of resistance.

Regarding resistance to **erythromycin**, a representative of the macrolides (commonly used in the treatment of human campylobacteriosis) in all reporting MSs, erythromycin resistance in *C. jejuni* from *Gallus gallus* and cattle was 0.4 % and 0.6 %, respectively, while, in *C. coli* from *Gallus gallus* and pigs, resistance equalled 11.2 % and 23.9 %, respectively. This situation in which low to moderately high levels of resistance were registered is similar to that observed over the 2009 to 2011 period. In countries which reported results



for *C. coli* from both pigs and *Gallus gallus* and *C. jejuni* from *Gallus gallus*, resistance to erythromycin has usually been highest in *C. coli* isolates from pigs and lower in the isolates from the other sources, for each country, over the period 2009 to 2011. Similar results have also been observed in other studies in which macrolide-resistant isolates of *C. coli* from food animals have mainly been of porcine origin (Gibreel and Taylor, 2006).

Levels of erythromycin resistance increased in *C. coli* from broilers in some reporting countries between 2011 and 2012. Several MSs, are implementing policies to reduce the amount of antimicrobials used in livestock production and it is important to gauge the effect this may have on the occurrence of resistance.

Two MSs provided data on **multiple drug resistance** for *C. coli* and *C. jejuni* from both humans and animals. Although some lack of harmonisation¹⁶ may preclude detailed direct comparison of the multi-resistance figures in isolates from animals and humans, some trends are evident. The MS with the higher proportion of multi-resistance in broilers (in both *Campylobacter* spp.) also reported a high proportion of multi-resistance characteristics (for example gentamicin and erythromycin resistance, co-resistance to ciprofloxacin and erythromycin). In the MS that provided the most comprehensive data among the reporting MSs, it is interesting that the figures for co-resistance to erythromycin and ciprofloxacin for *C. coli* isolates from humans and animals, for ciprofloxacin and erythromycin are, however, slightly different (Figure CA1) and although this is a small difference of one dilution, ideally the criteria in this type of analysis should be fully harmonised.

Campylobacter generally develops resistance to the different antimicrobials in the common test panel by different mechanisms. Resistance to ciprofloxacin and erythromycin in Campylobacter is usually the result of mutation with or without the additional action of efflux pumps (Piddock et al., 2003; Ge et al., 2005; Luangtongkum et al., 2009). Additionally, the efflux pump CmeABC has been shown to confer a degree of resistance to erythromycin, ciprofloxacin and tetracyclines (Ge et al., 2005). Some isolates of both C. coli and C. jejuni, from animals and humans, showed resistance to erythromycin, ciprofloxacin and tetracyclines, raising the possibility that CmeABC may have been responsible, or contributed to the observed pattern of resistance. In multiple resistant isolates of C. coli and C. jejuni which were gentamicin resistant, streptomycin resistance was also observed, occurring for example, in seven of nine gentamicin resistant and multiply resistant C. coli isolates and in one of the two C. jejuni isolates from broilers which showed gentamicin resistance. Recently a cluster of aminoglycoside modifying enzymes has been reported in C. coli from broiler chickens in China (Qin et al., 2012). A novel genomic island carrying multiple aminoglycoside resistance genes on the C. coli chromosome was identified by the researchers who showed that the genomic island can be transferred experimentally to C. jejuni and was associated with particular clones of C. jejuni in broilers in China where it was first detected. The aminoglycoside modifying enzymes found to be present on this genomic island, include those conferring resistance to gentamicin and streptomycin, as well as to certain other aminoglycosides (neomycin, kanamycin and tobramycin). The occurrence of isolates of C. coli and C. jejuni, resistant to both gentamicin and streptomycin, suggests that resistance genes to each of these aminoglycosides have been acquired by these multiple resistant isolates. The genomic island described by Qin et al. (2012) contained a truncated tetracycline resistance gene, illustrating the potential of this set of aminoglycoside resistance genes to capture other resistance genes. Streptomycin and tetracycline resistance were commonly associated with each other in multiple drug-resistant strains of both C. coli and C. jejuni. Conjugative plasmids have been described in C. jejuni, which can carry clusters of aminoglycoside resistance genes (Nirdnoy et al., 2005); however, it appears that both streptomycin and gentamicin resistance can occur independently of each other in at least some C. coli and C. jejuni isolates.

¹⁶ The antimicrobial substances included in the analysis of multiple drug resistance in isolates from humans and animals and the interpretive thresholds of resistance, either clinical breakpoints or ECOFFs, have not yet been harmonised between both sectors.



5. ANTIMICROBIAL RESISTANCE IN INDICATOR ESCHERICHIA COLI

5.1. Introduction

Commensal *E. coli* is commonly chosen as an indicator of antimicrobial resistance in Gram-negative bacterium, as it is commonly present in animal faeces, is relevant to human medicine and can often acquire conjugative plasmids, which are resistance determinants transferred between enteric bacteria. Commensal *E. coli*, present in the intestine of farm animals, have a reservoir of resistance genes that can spread horizontally to zoonotic and other bacteria present in the food chain. Commensal indicator organisms, rather than pathogenic types of *E. coli*, such as enterotoxigenic *E. coli* (ETEC) or verotoxigenic *E. coli* (VTEC), are the target of the monitoring of indicator *E. coli*. The monitoring of antimicrobial resistance in indicator *E. coli*, isolated from either randomly selected healthy animals or derived carcases and meat thereof, and chosen to be representative of the general population, provides valuable data on the resistance occurring in that population.

Determining the occurrence of resistance to antimicrobials in indicator *E. coli* provides data useful for investigating the relationship with the selective pressure exerted by the use of antimicrobials on the intestinal population of bacteria in food-producing animals. Indicator *E. coli* are also useful as representatives of the *Enterobacteriaceae* to monitor the emergence and changes in the proportion of bacteria possessing ESBLs. The EFSA monitoring guidelines (EFSA, 2008) recommend that monitoring may be carried out at the farm or slaughterhouse levels and that at least 90 % of the animal population in a MS should be included in the sampling frame. Samples should be collected randomly from selected holdings or flocks, or randomly selected within the slaughterhouse. Samples collected (and subsequently tested) in accordance with the EFSA recommendations should therefore be comparable between MSs.

5.2. Antimicrobial resistance in indicator *Escherichia coli* isolates from animals and food

In total, 11 MSs and 2 non-MSs (Norway and Switzerland) reported quantitative MIC data on antimicrobial resistance in commensal (indicator) *E. coli* isolates from animals in 2012. In addition, five of these countries provided MIC data on isolates collected from food. Table EC1 shows the countries that reported data concerning indicator *E. coli* in 2012. Antimicrobials selected by the different MSs and non-MSs for MIC susceptibility testing of indicator *E. coli* are shown in Chapter 8 Materials and methods, Table MM9.

Antimicrobial susceptibility data were interpreted using ECOFFs to determine organisms exhibiting reduced susceptibility, i.e. showing 'microbiological resistance' (as opposed to 'clinical resistance'). For reasons of continuity and because other amendments have also been made to the ECOFFs for some other antimicrobial - organism combinations by EUCAST, the ECOFFs used in this report have been those adopted in EFSA's 2008 recommendations (EFSA, 2008). Of particular note is that 'microbiological resistance' to ciprofloxacin was addressed using ECOFF Cip >0.03 mg/L in this report (see Section 5.5 Discussion, for further details).

The proportions of resistant isolates to the antimicrobial agents ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines are described in detail later in this chapter. The tables of occurrence of resistance were generated, and multi-drug resistance analysis was performed if more than four countries reported quantitative data per sampling origin. In addition, only data where 10 or more isolates were available per country, per sampling origin, per year, are included in the report. In the graphs illustrating trends in the evolution of antimicrobial resistance over time, results for MIC data interpreted using ECOFFs are shown. Only a few MSs have reported data for the seven consecutive years from 2006 to 2012, as the monitoring of resistance in indicator *E. coli* is performed on a voluntary basis.

Where the minimum criteria for detailed analysis were met, multi-resistance was analysed in the isolatebased dataset on the indicator *E. coli* isolates tested for the full harmonised set of nine antimicrobials belonging to different classes. Multi-resistance is defined as non-susceptibility to at least three different antimicrobial classes. The proportions of isolates susceptible to all antimicrobial substances tested and resistant (non-susceptible) to any one up to nine substances are presented. Co-resistance to cefotaxime and ciprofloxacin was estimated as these two antimicrobials are of particular interest in human medicine. Coresistance was addressed using both ECOFFs (Ctx >0.25 mg/L and Cip >0.03 mg/L) and clinical breakpoints (Ctx >2 mg/L and Cip >1 mg/L).



Table EC1. Overview of countries reporting MIC and disc inhibition zones on indicator Escherichia coli from animals and food in 2012

Method	Origin	Total number of MSs reporting	Countries
	Cattle (bovine animals)	1	MS: PT
Diffusion	Meat from broilers (Gallus gallus)	1	MS : SI
Dillusion	Meat from pigs	1	MS: SI
	Meat from bovine animals	1	MS: SI
		8	MSs: AT, BE, DK, FR, HU, NL, PL, SE
	Gallus gallus (fowl)	0	Non-MSs: CH, NO
	Turkeys	3	MSs: DE, NL, PL
	Rigo	7	MSs: AT, BE, DK, FR, HU, NL, PL
	Pigs	1	Non-MS: CH
Dilution	Cattle (boying animals)	7	MSs: AT, BE, DE, DK, FI, NL, PL
Dilution	Cattle (bovine animals)	1	Non-MS: CH
	Maat from broilers (Callus callus)	5	MSs: DK, ES, HU, NL, SE
	Meat from broilers (Gallus gallus)	5	Non-MS: NO
	Meat from turkeys	3	MSs: DE, HU, NL
	Meat from pigs	4	MSs: DK, ES, HU, NL
	Meat from bovine animals	5	MSs: DE, DK, ES, HU, NL

MIC: minimum inhibitory concentration.

Note: For abbreviations of Member States (MS) and other reporting countries see Appendix 7.

For further information on reported MIC distributions and numbers of resistant isolates for ampicillin, apramycin, cefazolin, cefepime, cefotaxime, cefoxitin, cefpodoxime, ceftazidime, ceftiofur, ceftriaxone, cephalothin, chloramphenicol, ciprofloxacin, colistin, florfenicol, gentamicin, imipenem, kanamycin, meropenem, nalidixic acid, neomycin, spectinomycin, streptomycin, sulfonamides, trimethoprim and tetracyclines for *E. coli* in 2012, please refer to the Level 3 Tables published on the EFSA website.

5.2.1. Antimicrobial resistance in indicator *Escherichia coli* isolates from food

In 2012, six MSs and one non-MS (Norway) reported quantitative MIC data for *E. coli* isolates from meat from bovine animals, broilers (*Gallus gallus*) and pigs.

5.2.1.1. Representative sampling and monitoring

The antimicrobial resistance data in indicator *E. coli* isolates from the three kinds of meat reported by Denmark, Germany, the Netherlands, Spain, Sweden and Norway mostly derived from active and representative monitoring programmes. Only one MS did not report details on either the sampling stages or on the sampling design of meat samples. In Denmark, *E. coli* isolates originated from meat sampled at wholesale and retail outlets, collected randomly in all regions of the country and spread evenly throughout the year, in the framework of three centrally coordinated sampling plans corresponding to each kind of meat. In Sweden, *E. coli* isolates originated from broiler meat collected randomly in all regions of the country and sampled between 23 October 2012 and 24 January 2013.

5.2.1.2. Meat from broilers (Gallus gallus)

The occurrence of resistance to selected antimicrobials in indicator *E. coli* isolates from broiler meat, in four reporting MSs and one non-MS, in 2012, is presented in Table EC2.

Considering data from the four reporting MSs, resistance levels to ampicillin, sulfonamides and tetracyclines were high at 33.7 %, 28.6 % and 22.2 %, respectively. The resistance to these antimicrobials was highly variable across the reporting MSs, ranging from 18.5 % to 57.7 % for ampicillin, from 16.3 % to 45.7 % for sulfonamides and from 11.7 % to 34.3 % for tetracyclines. Conversely, resistance to chloramphenicol and gentamicin at the reporting MS group level was low at 4.0 % and 3.2 %, respectively; resistance levels



ranged from 0 % to 12.5 % for chloramphenicol and from 0 % to 6.9 % for gentamicin. Resistance to ciprofloxacin and nalidixic acid, among reporting MSs, was 29.1 % and 24.1 %, respectively. The overall level of resistance to cefotaxime across the reporting MSs was low in 2012, at 3.0 %.

5.2.1.3. Meat from pigs

Among the four reporting MSs, *E*. coli isolated from pig meat displayed high levels of resistance to ampicillin sulfonamides and tetracyclines (20.4 %, 23.2 % and 23.5 %, respectively). Chloramphenicol resistance was low at 2.6 %. Overall, gentamicin resistance was 1.5 % in the reporting group of MSs; it was not detected in two MSs. The proportion of *E. coli* isolates resistant to ciprofloxacin and nalidixic acid among the reporting MSs was low at 5.6 % and 5.1 %. The occurrence of resistance to cefotaxime among all reporting MSs was low at 1.1 %.

5.2.1.4. Meat from bovine animals

Among the four reporting MSs, *E. coli* isolated from meat bovine animals displayed moderate levels of resistance to ampicillin and sulfonamides (18.7 % for both antimicrobials) and high level of resistance to tetracyclines (20.8 %). Chloramphenicol resistance was low, at 6.6 %, for the reporting group of MSs. Overall, gentamicin resistance was 2.1 % in the reporting group of MSs; it was not detected in Denmark and Hungary. The proportion of *E. coli* isolates resistant to ciprofloxacin and nalidixic acid among the reporting MSs were 6.9 % and 5.9 %, respectively. The occurrence of resistance to cefotaxime among all reporting MSs was low at 1.7 % with one MS reporting no resistance.

5.2.1.5. Multi-resistance among Escherichia coli isolates from meat

As too few MSs reported multi-resistance isolate-based data on more than 10 isolates of indicator *E. coli* in food, tables and graphs on multi-resistance are not presented in this report.

5.2.1.6. Comparison of resistance among *Escherichia coli* isolates from meat and animals

Four MSs reported on antimicrobial resistance in meat, but those which did generally reported comparable resistance levels in meat as in the corresponding source animal species. Indeed, Denmark reported resistance in isolates from broiler meat broadly comparable to that recorded in isolates from broilers. Resistance recorded by Denmark in isolates from pig meat is roughly similar to that reported from pigs. Similarly, in cattle, resistance in isolates from meat from cattle recorded in Denmark and Germany was roughly comparable to that reported for bovine animals in the same MSs (Tables EC4 and EC15), for which Germany reported on isolates from young meat production animals and Denmark reported on isolates from unspecified cattle type.



Table EC2. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among indicator Escherichia coli from <u>meat from broilers</u> (Gallus gallus) in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country			Chloramphenicol C		Ciprofl	Ciprofloxacin		Gentamicin		Nalidixic acid		omycin	Sulfon	amides	Tetrac	yclines		
Country	Ν	% Res	N	% Res	Ν	% Res	N	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Denmark	197	21.8	197	1.0	197	0.5	197	3.6	197	0	197	3.6	197	8.1	197	16.8	197	11.7
Hungary	64	26.6	64	0	64	12.5	64	75.0	64	3.1	64	75.0	64	14.1	64	35.9	64	32.8
Netherlands	175	57.7	175	8.0	175	6.9	175	41.1	175	6.9	175	38.9	175	41.1	175	45.7	175	34.3
Sweden	92	18.5	92	0	92	0	-	-	92	3.3	92	4.3	92	6.5	92	16.3	92	14.1
Total (4 MSs)	528	33.7	528	3.0	528	4.0	436	29.1	528	3.2	528	24.1	528	19.5	528	28.6	528	22.2
Norway	197	6.1	197	0.5	197	0	_	-	197	0	197	2.0	197	3.0	197	8.1	-	-

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Table EC3. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among indicator Escherichia coli from <u>meat from pigs</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	Ampicillin Cefotaxime		Chloramphenicol Cipr		Ciprof	Ciprofloxacin Gentamicin		Nalidixic acid		Streptomycin		Sulfor	namides	Tetrac	cyclines		
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Denmark	73	32.9	73	1.4	73	1.4	73	1.4	73	0	73	0	73	35.6	73	30.1	73	27.4
Hungary	14	28.6	14	0	14	7.1	14	28.6	14	0	14	28.6	14	21.4	14	35.7	14	35.7
Netherlands	98	8.2	98	1.0	98	2.0	98	3.1	98	2.0	98	3.1	98	15.3	98	16.3	98	18.4
Spain	11	36.4	_	-	11	9.1	11	27.3	11	9.1	11	27.3	11	45.5	-	-	11	27.3
Total (4 MSs)	196	20.4	185	1.1	196	2.6	196	5.6	196	1.5	196	5.1	196	25.0	185	23.2	196	23.5

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.



Table EC4. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among indicator Escherichia coli from <u>meat from bovine animals</u> in MSs reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	npicillin Cefotaxime		Chloramphenicol Ciproflo		iprofloxacin Gentamicin I		Nalidixic acid		Streptomycin		Sulfon	amides	Tetra	cyclines			
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res
Denmark	46	4.3	46	0	46	0	46	2.2	46	0	46	2.2	46	4.3	46	4.3	46	6.5
Germany	71	39.4	71	4.2	71	16.9	71	14.1	71	4.2	71	12.7	71	35.2	71	38.0	71	43.7
Hungary	31	9.7	31	3.2	31	0	31	3.2	31	0	31	3.2	31	6.5	31	6.5	31	12.9
Netherlands	141	14.9	141	0.7	141	5.0	141	5.7	141	2.1	141	4.3	141	16.3	141	16.3	141	15.6
Total (4 MSs)	289	18.7	289	1.7	289	6.6	289	6.9	289	2.1	289	5.9	289	18.0	289	18.7	289	20.8

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.



5.2.2. Antimicrobial resistance in indicator *Escherichia coli* isolates from animals

5.2.2.1. Domestic fowl (Gallus gallus)

Representative sampling and monitoring

In this section, data on antimicrobial resistance in indicator *E. coli* isolates from fowl (*Gallus gallus*) are presented separately for broilers and laying hens. The majority of MSs collected isolates as part of their national monitoring programmes of antimicrobial resistance. In all reporting countries except Norway, monitoring programmes were based on random sampling of healthy broilers at the slaughterhouse. Indicator *E. coli* isolates were isolated from caecal contents in Austria and France, from intestinal content in Sweden and from cloacal swabs in Denmark and Switzerland, by sampling healthy broilers at slaughter. In Norway, indicator *E. coli* were isolated from faeces sampled from *Gallus gallus* on the farm. Hungary and Poland did not report information on the sample type, sampling context and sampling stage.

Resistance levels among Escherichia coli isolates from Gallus gallus

In 2012, seven MSs and one non-MS provided quantitative data concerning antimicrobial resistance in *E. coli* from broilers, among which two MSs (Poland and Sweden) also provided comparable data concerning *E. coli* from laying hens (Table EC5). In addition, Norway reported *E. coli* resistance data in parent breeders for broiler production, while Belgium reported data at *Gallus gallus* species level without distinguishing between animal populations of origin.

Generally, the occurrence of resistance in *E. coli* isolates from broilers varied markedly between reporting countries. Resistance to ampicillin, streptomycin, sulfonamides and tetracyclines was generally high to very or extremely high in most reporting countries, with the exceptions of Sweden, reporting low to moderate resistance to these substances, and Denmark, recording low to moderate resistance to streptomycin and tetracyclines. Resistance to chloramphenicol was generally low to moderate with only Poland reporting high resistance and Denmark and Sweden no or very low resistance. In all reporting countries, gentamicin resistance was reported at very low to low levels.

Resistance to ciprofloxacin and nalidixic acid was generally high to very or extremely high among the reporting countries, with the exception of Denmark recording low resistance to these substances. A side-by-side comparison of resistance to ciprofloxacin and nalidixic acid in each reporting country shows that similar levels of resistance to both antimicrobials were typically recorded. Resistance to cefotaxime was generally low in most reporting countries, although two MSs reported moderate levels of resistance.

Resistance features in *E. coli* isolates from laying hens, tested in Poland and Sweden, were similar to those observed in isolates from broilers in the same MSs, although resistance levels in laying hens were in some cases less than half those reported in broilers. However, in Sweden, where resistance is typically low, resistance levels to cefotaxime, chloramphenicol, gentamicin, sulfonamides and tetracyclines were similar in broilers and laying hens. Although lower levels of resistance were typically reported in Sweden compared with Poland, Poland recorded high resistance to ciprofloxacin in laying hens.

In addition, Norway reported resistance data on *E. coli* isolates from parent breeders for broiler production at low to moderate levels.

Belgium was the only MS to report resistance data at the species level of *Gallus gallus*, potentially including a mixture of data on broilers and laying hens: high to extremely high resistance levels to all the antimicrobials tested except to gentamicin were recorded.



Table EC5. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among indicator Escherichia coli from <u>Gallus gallus</u> in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
All Gallus gallus										
Austria	130	26.9	130	3.1	130	8.5	130	65.4	130	0.8
Belgium	325	79.7	325	28.0	325	44.6	325	79.7	325	5.8
Denmark	115	20.0	115	1.7	115	0	115	7.8	115	0
France	201	58.2	201	10.4	201	6.5	201	35.8	201	1.5
Hungary	105	52.4	105	7.6	104	9.6	104	74.0	104	1.9
Netherlands	292	69.9	292	5.8	292	16.4	292	51.4	292	8.6
Poland	328	61.0	328	10.7	328	12.5	328	63.7	328	5.5
Sweden	255	11.8	255	0.4	255	0.4	—	-	255	0.8
Total (8 MSs)	1,751	52.7	1,751	10.2	1,750	15.4	1,495	57.6	1,750	4.0
Norway	113	15.0	113	0.9	113	0	—	-	113	0
Switzerland	185	32.4	185	2.2	185	1.1	185	46.5	185	0.5
Broilers										
Austria	130	26.9	130	3.1	130	8.5	130	65.4	130	0.8
Denmark	115	20.0	115	1.7	115	0	115	7.8	115	0
France	201	58.2	201	10.4	201	6.5	201	35.8	201	1.5
Hungary	105	52.4	105	7.6	104	9.6	104	74.0	104	1.9
Netherlands	292	69.9	292	5.8	292	16.4	292	51.4	292	8.6
Poland	171	86.0	171	13.5	171	21.1	171	82.5	171	8.2
Sweden	194	14.4	194	0	194	0.5	_	_	194	0.5
Total (7 MSs)	1,208	50.4	1,208	6.2	1,207	9.9	1,013	52.7	1,207	3.8
Switzerland	185	32.4	185	2.2	185	1.1	185	46.5	185	0.5
Laying hens										
Poland	157	33.8	157	7.6	157	3.2	157	43.3	157	2.5
Sweden	61	3.3	61	1.6	61	0	_	_	61	1.6
Total (2 MSs)	218	25.2	218	6.0	218	2.3	157	43.3	218	2.3
Parent breeders	for broile	er product	ion							
Norway	113	15.0	113	0.9	113	0	-	-	113	0

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.



Table EC5 (continued). Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among indicator Escherichia coli from <u>Gallus gallus</u> in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Nalidix	ic acid	Strepto	mycin	Sulfona	mides	Tetracy	clines
Country	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res
All Gallus gallus		•				•	•	•
Austria	130	65.4	130	42.3	130	44.6	130	30.8
Belgium	325	77.8	325	81.5	325	80.0	325	68.3
Denmark	115	7.8	115	11.3	115	20.9	115	7.8
France	201	29.4	201	40.3	201	57.7	201	76.1
Hungary	104	71.2	104	26.0	104	32.7	104	36.5
Netherlands	292	50.0	292	58.2	292	62.7	292	50.7
Poland	328	54.0	328	36.3	328	46.0	328	44.5
Sweden	255	10.6	255	7.8	255	9.4	255	11.4
Total (8 MSs)	1,750	47.4	1,750	42.9	1,750	48.6	1,750	44.9
Norway	113	0.9	113	1.8	113	8.8	113	5.3
Switzerland	185	45.9	185	14.6	185	24.9	185	33.5
Broilers								
Austria	130	65.4	130	42.3	130	44.6	130	30.8
Denmark	115	7.8	115	11.3	115	20.9	115	7.8
France	201	29.4	201	40.3	201	57.7	201	76.1
Hungary	104	71.2	104	26.0	104	32.7	104	36.5
Netherlands	292	50.0	292	58.2	292	62.7	292	50.7
Poland	171	73.1	171	58.5	171	66.7	171	66.7
Sweden	194	12.4	194	8.8	194	9.8	194	10.8
Total (7 MSs)	1,207	43.2	1,207	38.4	1,207	45.4	1,207	43.3
Switzerland	185	45.9	185	14.6	185	24.9	185	33.5
Laying hens						-		
Poland	157	33.1	157	12.1	157	23.6	157	20.4
Sweden	61	4.9	61	4.9	61	8.2	61	13.1
Total (2 MSs)	218	25.2	218	10.1	218	19.3	218	18.3
Parent breeders for	r broiler pro	duction						
Norway	113	0.9	113	1.8	113	8.8	113	5.3

MS: Member State; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.



Temporal trends in resistance among indicator *Escherichia coli* isolates from broilers of *Gallus gallus*

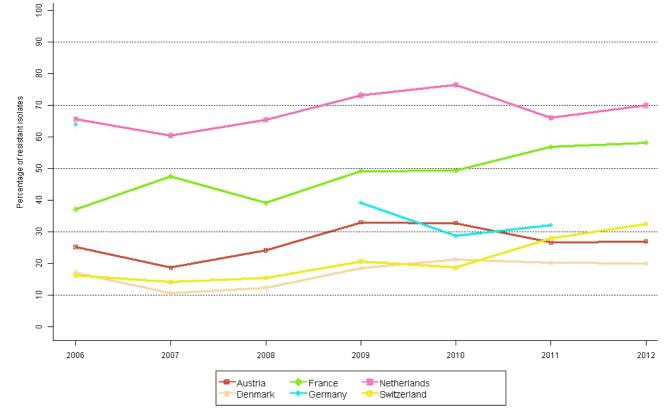
Figures EC1 to EC5 display temporal trends in resistance to selected antimicrobials in indicator *E. coli* from broilers of *Gallus gallus* over the seven-year study period of 2006 to 2012. It is of note that the 2010 and 2011 resistance levels in Germany presented in these figures combine data on broilers and laying hens, while in the other reporting countries resistance data derive from broilers only.

The figures illustrate the wide variation in resistance between MSs for many of the antimicrobials. Spain and the Netherlands tended to report the highest levels of resistance to most antimicrobials over the period, although Austria, Spain and the Netherlands reported the highest resistance to quinolones between 2010 and 2012 and France, Spain and the Netherlands reported the highest resistance to tetracyclines from 2007 to 2012. Conversely, Denmark often recorded the lowest resistance levels reported.

The resistance to ciprofloxacin reported over the study period was high to very high for all reporting countries, with the exception of Denmark for the whole period, of Germany for the years 2010 and 2011 (combine broiler and laying hen data), and of Norway for the period 2011 to 2012, which in all three cases was below 20 %. Figure EC3 clearly demonstrates the close similarity in resistance levels to ciprofloxacin and nalidixic acid in most MSs. There was less variation between countries in the resistance to cefotaxime and chloramphenicol, which, in most countries, was at a moderate or low level. However, although resistance levels in 2012 tended to be generally similar to those observed in 2011, there were a few exceptions; for example, in Poland (data not shown) and Switzerland, resistance to ampicillin and cefotaxime in broiler flocks increased from 2011 to 2012. Such inter-annual evolutions need to be confirmed by longer term trends.

Resistance levels for many of the antimicrobials were broadly stable or had shown only gradual increases or decreases. Nevertheless, there was evidence of statistically significant trends in the occurrence of resistance to some of the antimicrobials over five or more years. Austria reported significant increases in resistance to ciprofloxacin, nalidixic acid and streptomycin. France reported significant increases in resistance to ampicillin, cefotaxime, ciprofloxacin and streptomycin. Denmark also reported statistically significant increasing trends in resistance to ampicillin and cefotaxime, although resistance levels are lower than those recorded in France. Switzerland also reported an increase in resistance to ampicillin, ciprofloxacin and nalidixic acid. Contrastingly, the Netherlands reported significant declines in resistance to cefotaxime and ciprofloxacin, in particular over the last three years. There were no significant trends in resistance to tetracyclines in any of the reporting countries.

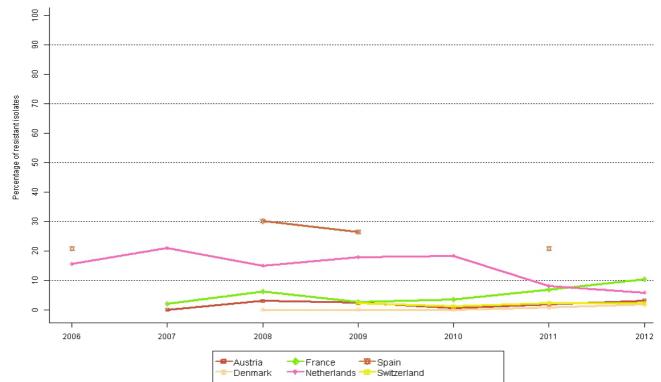




Note: Statistically significant increasing trends over five or more years, as tested by a logistic regression model (*p* ≤0.05), were observed in Denmark (↑), France (↑) and Switzerland (↑). MS: Member State.

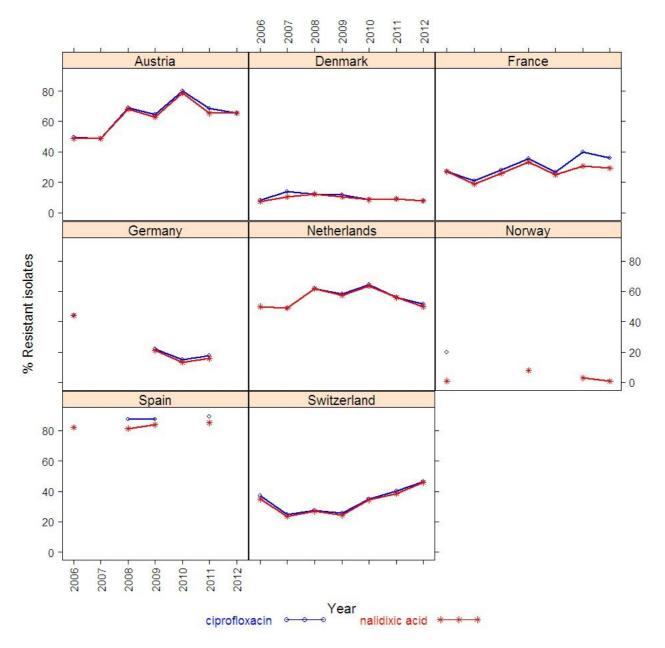
1. The data from Germany in 2010 and 2011 originated from broilers and laying hens.

Figure EC2. Trends in <u>cefotaxime</u> resistance in indicator Escherichia coli from <u>broilers</u> of Gallus gallus in reporting MSs and one non-MS, 2006–2012, quantitative data



Note: Statistically significant increasing or decreasing trends over five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in Denmark (\uparrow), France (\uparrow) and the Netherlands (\downarrow). MS: Member State.

Figure EC3. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in indicator Escherichia coli from <u>broilers</u> of Gallus gallus¹ in reporting MSs and two non-MSs, 2006–2012, quantitative data

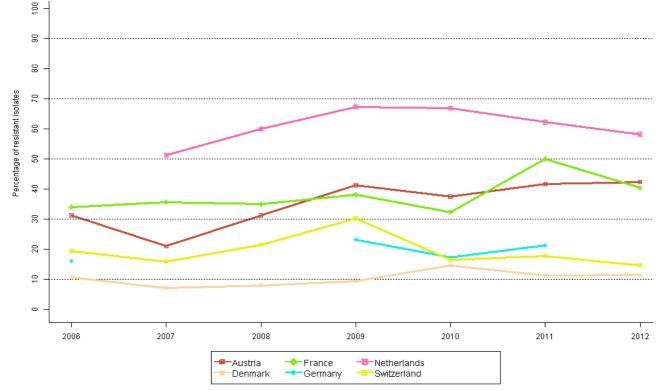


MS: Member State.

Note: Statistically significant increasing or decreasing trends over five or more years, as tested by a logistic regression model (*p* ≤0.05), were observed in Austria (↑) and Switzerland (↑) for both ciprofloxacin and nalidixic acid and in France (↑) and the Netherlands (↓) for ciprofloxacin.

1. The data from Germany in 2010 and 2011 originated from broilers and laying hens.

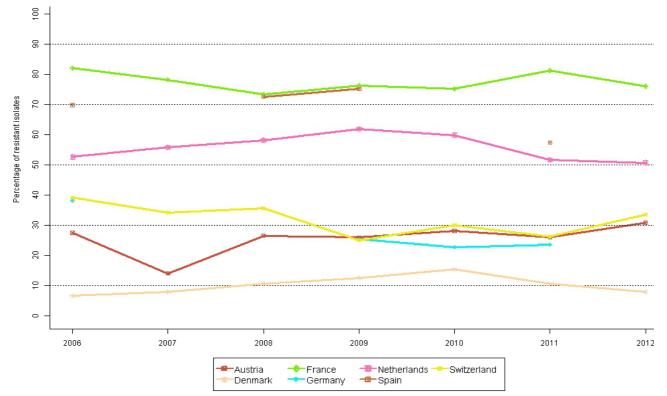




Note: A statistically significant increasing trend over six years, as tested by a logistic regression model (*p* ≤0.05), was observed in Austria (↑) and France (↑). MS: Member State.

1. The data from Germany in 2010 and 2011 originated from broilers and laying hens.

Figure EC5. Trends in <u>tetracyclines</u> resistance in indicator Escherichia coli from <u>broilers</u> of Gallus gallus¹ in reporting MSs and one non-MS, 2006–2012, quantitative data



Note: No statistically significant trends over five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in any of the reporting countries. MS: Member State.

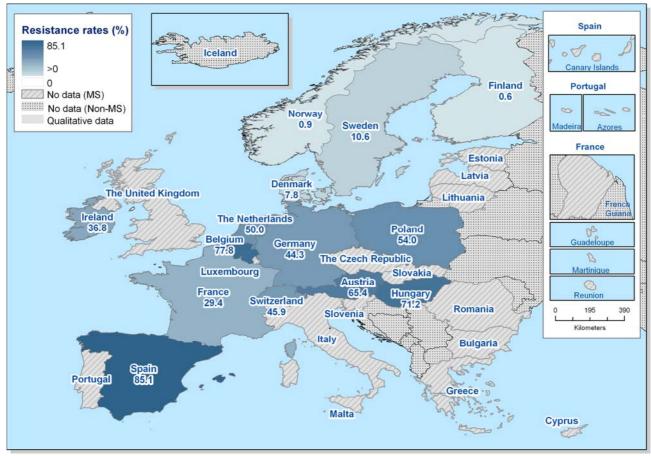
1. The data from Germany in 2010 and 2011 originated from broilers and laying hens.



Spatial distribution of resistance among indicator *Escherichia coli* from broilers of *Gallus gallus*

The spatial distributions of nalidixic acid and tetracycline resistance in *E. coli* from *Gallus gallus* are shown in Figures EC6 and EC7. The Nordic countries reported the lowest levels of resistance to both antimicrobials. The highest resistance to tetracyclines tended to be reported by the most western countries, while the spatial pattern for nalidixic acid was less clear.

Figure EC6. Spatial distribution of <u>nalidixic acid</u> resistance among indicator Escherichia coli from <u>broilers</u>¹ of Gallus gallus in countries reporting MIC data in 2012²

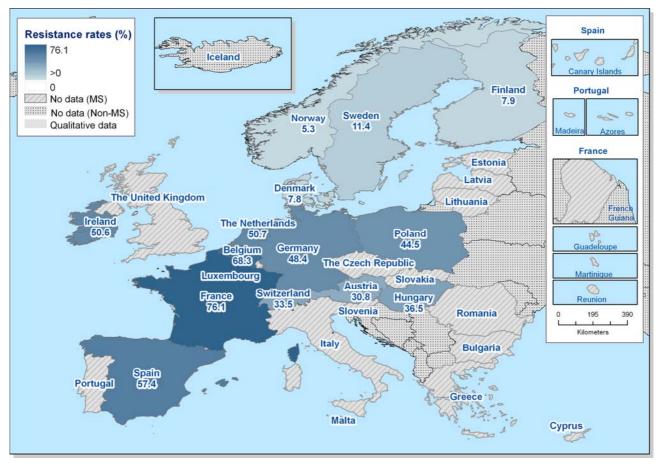


MS: Member State; MIC: minimum inhibitory concentration.

- Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data were used instead.
- 1. The data from Norway originated from parent breeders for broiler production and data from Belgium originated from unspecified production type.
- 2. For Finland, Germany, Ireland and Spain, 2011 data were used.



Figure EC7. Spatial distribution of <u>tetracycline</u> resistance among indicator Escherichia coli from <u>broilers</u>¹ of Gallus gallus in countries reporting MIC data in 2012²



MS: Member State; MIC: minimum inhibitory concentration.

Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data were used instead.

- 1. The data from Norway originated from parent breeders for broiler production and data from Belgium originated from unspecified production type.
- 2. For Finland, Germany, Ireland and Spain 2011 data were used.



Multi-resistance among indicator Escherichia coli isolates from broilers

In 2012, three MSs and one non-MS provided isolate-based data regarding resistance in indicator *E. coli* in broilers. Among the reporting countries, variations were observed in the percentages of completely susceptible isolates, which varied from 13.5 % in Hungary to 56.5 % in Denmark. Although all reporting countries recorded multi-resistant isolates, their proportions differed substantially between them, reaching up to 47.1 % in Hungary (Table EC6). The frequency distributions (Figure EC8) showed that isolates resistant to as many as five antimicrobials were reported from all reporting countries, and three MSs reported a few isolates resistant to seven substances. Co-resistance to cefotaxime and ciprofloxacin was undetected in the Nordic MS or detected at only low or very low levels in central European MSs (Table EC6).

Table EC6. Complete susceptibility, multi-resistance and index of diversity in Escherichia coli from <u>broilers</u> in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscepti	ible to all	Multi-re	esistant	Index of	Co-resistant to Cip and Ctx		
	n	%	n	%	diversity	n	%	
Austria (N=130)	22	16.9	56	43.1	0.493	1(0)	0.8(0)	
Denmark (N=115)	65	56.5	15	13.0	0.250	0(0)	0(0)	
Hungary (N=104)	14	13.5	49	47.1	0.537	7(1)	6.7(1.0)	
Switzerland (N=185)	49	26.5	46	24.9	0.383	4(0)	2.2(0)	

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *E. coli*; n: number of isolates.

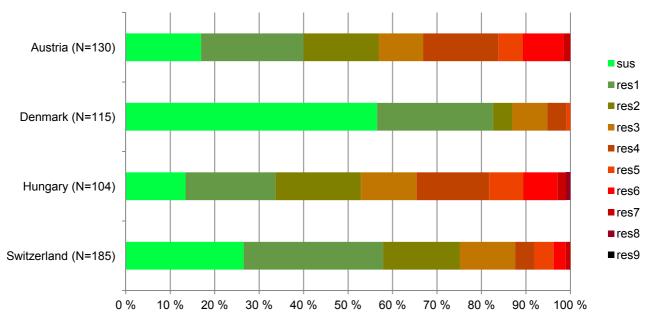
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for E. coli.

Multi-resistant: resistant to at least 3 different antimicrobial substances, belonging to any three antimicrobial families from the common antimicrobial set for *E. coli*.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to cefotaxime (Ctx) and ciprofloxacin (Cip): the effectives and percentages of *E. coli* isolates non-susceptible to concentrations greater than epidemiological cut-off values (Ctx >0.25 mg/L and Cip >0.03 mg/L). Figures in parentheses indicate the occurrence of co-resistance to Cip and Ctx determined using clinical breakpoints (Ctx >2 mg/L and Cip >1 mg/L).

Figure EC8. Frequency distribution of Escherichia coli isolates completely susceptible and resistant to one to nine antimicrobials in broilers in MSs and non-MSs reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *E. coli*; sus: susceptible to all antimicrobial substances of the EFSA common set for *E. coli*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *E. coli*.



Multi-/co-resistance patterns among indicator Escherichia coli isolates from broilers

As expected, most isolates resistant to ciprofloxacin were also resistant to nalidixic acid when using ECOFFs as thresholds. Several *E. coli* isolates from Hungary were resistant to cefotaxime but not to ceftazidime, hence they appear in Table EC6 but not in Table EC7. Considering resistance patterns in isolates corresistant to ciprofloxacin and cefotaxime (Table EC8), a number of isolates were also resistant to sulfonamides, streptomycin and tetracyclines. Trimethoprim resistance was also observed, while resistance to nalidixic acid and ampicillin was expected in isolates co-resistant to cefotaxime and ciprofloxacin. Analysing occurrence of higher levels of resistance to ciprofloxacin in *E. coli* reveals marked differences between MSs (Table EC9).

Table EC7. <u>Multi-/co-resistance</u> patterns of interest in Escherichia coli from <u>broilers</u> in MSs and one non-MS reporting isolate-based data, 2012

Multi-/co-resistance pattern							Group of reporting countries (N=534)		Austria (N=130)	Denmark (N=115)	Hungary (N=104)	Switzerland (N=185)					
Amp	Caz	Ctx	ChI	Cip	Gen	Nal	Str	Su	Tet	Tmp	n	%	group %	n	n	n	n
				R		R					253	98.1	47.4	85	9	74	85
	R	R		R							5	1.9	0.9	1	0	0	4
	Total						258	100	48.3	86	9	74	89				

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *E. coli* and multiresistant; Amp: ampicillin; Caz: ceftazidime; Ctx: cefotaxime; ChI: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; NaI: nalidixic acid; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethroprim; n: number of multi-/co-resistant isolates; R: minimum inhibitory concentration above the European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

Table EC8. Co-resistance to <u>cefotaxime</u> and <u>ciprofloxacin</u> in Escherichia coli from broilers in MSs and one non-MS reporting isolate-based data, 2012

Country	Resistance to both Cip and Ctx, applying ECOFFs ¹	Resistance to both Cip and Ctx, applying clinical breakpoints ¹	Multi-resistance patterns of isolates resistant to both Cip and Ctx, applying ECOFFs ¹			
	n (%)	n (%)	(number of isolates)			
Austria (N=130)	1 (0.8 %)	0 (0 %)	AmpCtxChlCipNalSuTet (1)			
Denmark (N=115)	0 (0 %)	0 (0 %)	NA			
			AmpCtxCipNal (3)			
			AmpCtxCipNalSuTetTmp (1)			
Hungary (N=104)	7 (6.7 %)	1 (1.0 %)	AmpCtxCipNalStrSuTmp (1)			
		AmpCtxCipNalStrSu				
			AmpCtxChlCipNalSuTmp (1)			
Total (3 MSs) (N=349)	8 (2.3 %)	1 (0.3 %)				
			AmpCtxCipNalSuTet (2)			
Switzerland (N=185)	4 (2.2 %)	0 (0 %)	A mpCtxCipNal (3) mpCtxCipNalSuTetTmp (1) mpCtxCipNalStrSuTmp (1) mpCtxCipNalStrSuTetTmp (1) mpCtxChlCipNalSuTmp (1) mpCtxCipNalSuTet (2) mpCtxCipNal (1)			
			AmpCtxCipNalSuTetTmp (1)			

MS: Member State; N: total number isolates for which relevant data are available; n: number of co-resistant isolates; Cip: ciprofloxacin; Ctx: cefotaxime.

1. European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs) and clinical breakpoints were applied.

Table EC9. <u>Ciprofloxacin</u> resistance assessed at differing thresholds in indicator Escherichia coli from <u>broilers</u> in MSs and one non-MS reporting isolate-based data, 2012

Country	Isolates resistant at >0.03mg/L ¹ Cip	Isolates resistant at >1mg/L ² Cip	Isolates resistant at >2mg/L ³ Cip	Isolates resistant at >4mg/L ⁴ Cip	Resistance patterns of isolates resistant at >4mg/L Cip (number of isolates)
	n (%)	n (%)	n (%)	n (%)	
					ChlCipNalStrSuTet (1)
Austria (N=130)	85 (65.4 %)	8 (6.2 %)	4 (3.1 %)	4 (3.1 %)	AmpCipNalStrSuTet (1)
/			. (0.1. /0)	. (0.1. /0)	AmpCipNalStrSuTetTmp (1)
					AmpChlCipGenNalSuTmp (1)
Denmark (N=115)	9 (7.8 %)	1 (0.9 %)	1 (0.9 %)	0 (0 %)	NA
					CipNal (4)
					AmpCipNal (3)
			28 (26.9 %)		AmpCipNalTet (2)
					AmpCipNalStrSuTetTmp (2)
					CipNalTet (1)
					CipNalSuTet (1)
					CipNalSuTetTmp (1)
					AmpCipNalTmp (1)
Hungary (N=104)	77 (74 %)	35 (33.7 %)		24 (23.1 %)	AmpCipNalTetTmp (1)
					AmpCipNalSuTmp (1)
					AmpCipNalSuTet (1)
					AmpChlCipNalSuTmp (1)
					AmpChlCipNalSuTetTmp (1)
					AmpChlCipNalStrSuTet (1)
					AmpChlCipNalStrSuTetTmp (1)
					AmpChlCipGenNalStrSuTetTmp (1)
					AmpCtxChlCipNalSuTmp (1)
Total (3 MSs)(N=349)	171 (49.0 %)	44 (12.6 %)	33 (9.5 %)	28 (8.0 %)	
					AmpCipNalTet (5)
					CipNal (1)
Switzerland (N=185)	86 (46.5 %)	11 (5.9 %)	10 (5.4 %)	9 (4.9 %)	CipNalSuTetTmp (1)
		. ,	. ,		AmpCipNalSuTetTmp (1)
					AmpCipGenNalStrSuTmp (1)

MS: Member State; N: total number isolates for which relevant data are available; n: number of resistant isolates; Cip: ciprofloxacin.

1. European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2. EUCAST clinical breakpoint.

3. High breakpoint.

4. Very high breakpoint.



5.2.2.2. Pigs

Representative sampling and monitoring

In 2012, seven MSs and one non-MS (Switzerland) provided quantitative antimicrobial resistance data on indicator *E. coli* in pigs which were included in the following analysis (Table EC10). These data were not split by production type, as isolates originated from either fattening pigs or breeding animals (Belgium) or the production type was not specified. The majority of MSs collected isolates as part of their national monitoring programme of antimicrobial resistance, mostly based on random sampling of healthy slaughter pig carcases at the slaughterhouse. A two-stage stratified sampling design, with slaughterhouses as primary sampling units and carcases as secondary units, with proportional allocation of the number of samples to the annual throughput of the slaughterhouse, was typically applied in the reporting countries. The sample collection was approximately evenly distributed over the year. Only one representative faecal sample per epidemiological unit (batch), either derived from a unique carcase or pooled from a number of carcases, was gathered to account for clustering. Hungary did not report detailed information on sample type and sampling context.

Resistance levels among Escherichia coli isolates from pigs

In 2012, resistance to ampicillin in *E. coli* isolates from pigs was generally high among reporting MSs, ranging from 22.0 % to 48.5 % - except in Austria which recorded a 12.9 % resistance, while resistance to streptomycin, sulfonamides and tetracyclines was high to very high in all reporting countries, ranging from 36.8 % to 59.9 %, 22.9 % to 60.0 % and 29.2 % to 69.1 %, respectively. Conversely, resistance to chloramphenicol was low to moderate in most reporting countries, with the notable exception of Belgium reporting, as in 2011, a high resistance of 30.2 %, while gentamicin resistance was generally recorded at low to very low levels. Resistance to ciprofloxacin and nalidixic acid was low to moderate among all reporting countries, ranging between 0.7 % and 16.6 %, and resistance to cefotaxime was either not detected or reported at low levels in all reporting countries.



Table EC10. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among isolates of indicator Escherichia coli from <u>pigs</u> in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Ampicillin		Cefotaxime		Chloramphenicol		Ciprof	oxacin	Gentamicin	
Country	N	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Austria ¹	140	12.9	140	0	140	2.1	140	5.0	140	0
Belgium ²	205	48.3	205	2.9	205	30.2	205	16.6	205	1.0
Denmark ¹	152	28.9	152	0.7	152	3.3	152	0.7	152	0.7
France ¹	200	22.0	200	2.0	200	9.5	200	8.5	200	3.0
Hungary ¹	68	48.5	68	1.5	68	14.7	68	13.2	68	2.9
Netherlands ¹	284	25.0	284	0	284	11.6	284	1.1	284	2.1
Poland ³	190	29.5	190	2.6	190	8.9	190	11.6	190	2.6
Total (7 MSs)	1,239	29.5	1,239	1.4	1,239	12.0	1,239	7.5	1,239	1.8
Switzerland ¹	185	20.0	185	1.1	185	4.9	185	3.2	185	1.1

Country	Nalidix	ic acid	Strepto	mycin	Sulfona	mides	Tetracyclines		
Country	Ν	% Res	N	% Res	N	% Res	N	% Res	
Austria ¹	140	5.0	140	52.1	140	22.9	140	50.7	
Belgium ²	205	12.2	205	53.7	205	60.0	205	61.5	
Denmark ¹	152	0.7	152	42.1	152	34.9	152	35.5	
France ¹	200	3.0	200	46.0	200	44.0	200	63.5	
Hungary ¹	68	10.3	68	36.8	68	35.3	68	69.1	
Netherlands ¹	284	1.1	284	59.9	284	45.4	284	56.3	
Poland ³	190	6.3	190	46.8	190	35.8	190	48.9	
Total (7 MSs)	1,239	4.9	1,239	50.3	1,239	41.7	1,239	54.7	
Switzerland ¹	185	3.2	185	46.5	185	38.9	185	29.2	

MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates; MS: Member State.

1. Fattening pigs.

2. Breeding animals.

3. Unspecified production type.

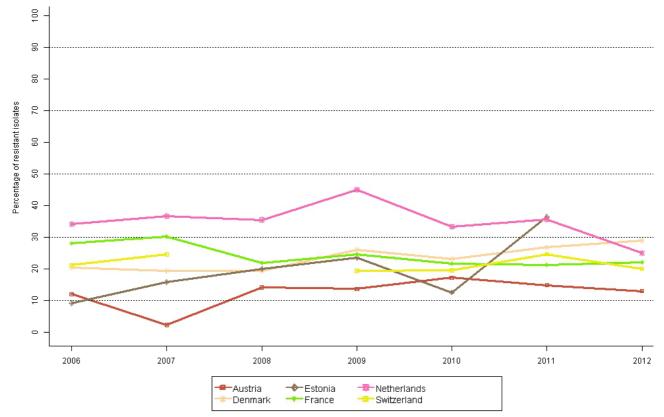


Temporal trends in resistance among indicator *Escherichia coli* isolates from pigs

Figures EC9 to EC13 display the trends in resistance to selected antimicrobials in indicator *E. coli* from pigs over the period 2006 to 2012. There was variation in the resistance levels in different MSs, particularly for tetracyclines (Figure EC13). However, the differences between MSs were often not as marked as was observed for isolates from *Gallus gallus*. In some cases, this was because the resistance levels tended to be lower than those observed in *Gallus gallus* (e.g. ampicillin; Figure EC9), whereas, for others, it was due to the resistance levels all being higher than those recorded in *Gallus gallus* (e.g. streptomycin; Figure EC12). Cefotaxime resistance has been below 5 % in all countries since 2005, and at a lower level than in *Gallus gallus* (Figure EC10). Resistance to both ciprofloxacin and nalidixic acid has also generally been at a low level since 2005 (Figure EC11).

For many of the antimicrobials, the resistance levels were relatively stable with only minor fluctuations or gradual changes. There were fewer statistically significant trends than observed among isolates from *Gallus gallus*. Denmark reported significant increases in resistance to ampicillin, Switzerland reported significant increases in resistance to ampicillin, Switzerland reported significant declines in resistance to ampicillin, ciprofloxacin and tetracyclines, and France reported significant declines in resistance to streptomycin and tetracyclines. No statistically significant trends were observed in resistance to cefotaxime.

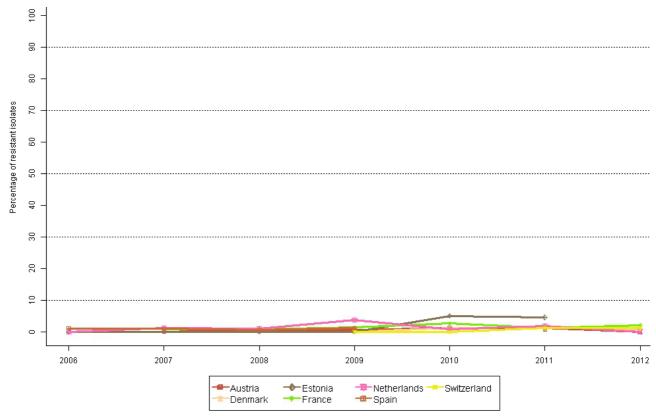
Figure EC9. Trends in <u>ampicillin</u> resistance in indicator Escherichia coli from <u>pigs</u> in reporting MSs and one non-MS, 2006–2012, quantitative data



MS: Member State.

Note: Statistically significant increasing and decreasing trends over seven years, as tested by a logistic regression model ($p \le 0.05$), were observed in Denmark (\uparrow) and the Netherlands (\downarrow).

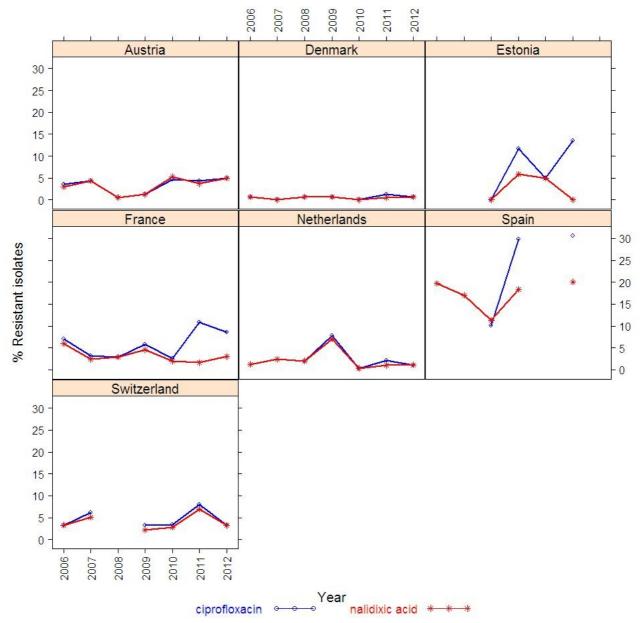
Figure EC10. Trends in <u>cefotaxime</u> resistance in indicator Escherichia coli from <u>pigs</u> in reporting MSs and non-MS, 2006–2012, quantitative data



MS: Member State.

Note: No statistically significant trends over five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in any of the reporting countries.

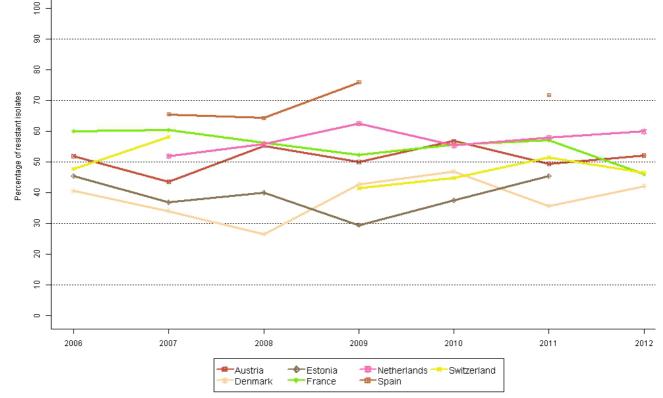
Figure EC11. Trends in ciprofloxacin and <u>nalidixic acid</u> resistance in indicator Escherichia coli from <u>pigs</u> in reporting MSs and one non-MS, 2006–2012, quantitative data



MS: Member State.

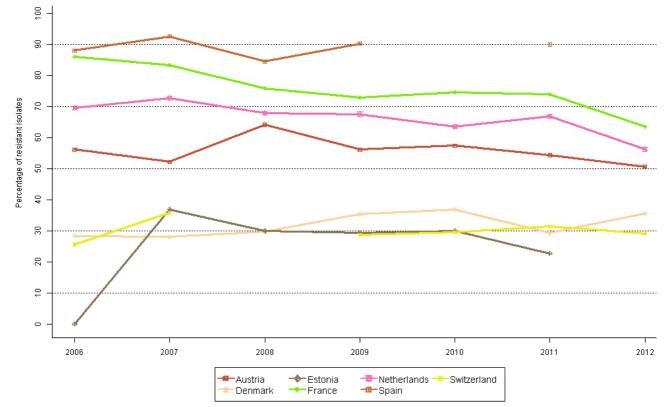
Note: Statistically significant increasing or decreasing trends over five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in Switzerland (\uparrow) for both ciprofloxacin and nalidixic acid, in France (\uparrow) and the Netherlands (\downarrow) for ciprofloxacin.





Note: A statistically significant decreasing trend over seven years, as tested by a logistic regression model (*p* ≤0.05), was observed in France (↓). MS: Member State.

Figure EC13. Trends in <u>tetracycline</u> resistance in indicator Escherichia coli from <u>pigs</u> in reporting <i>MSs and one non-MS, 2006–2012, quantitative data



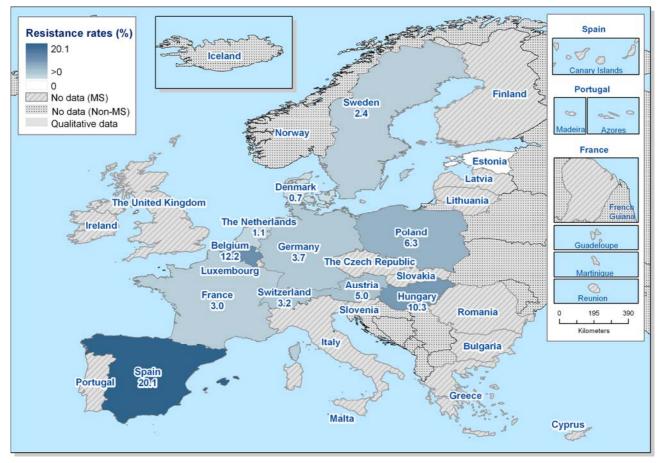
Note: Statistically significant decreasing trend over seven years, as tested by a logistic regression model (*p* ≤0.05), were observed in France (↓) and the Netherlands (↓). MS: Member State.



Spatial distribution of resistance among indicator *Escherichia coli* isolates from pigs

The spatial distribution of nalidixic acid and tetracycline resistance in indicator *E. coli* from pigs is shown in Figures EC14 and EC15, respectively. For nalidixic acid, most countries reported low levels of resistance so the spatial pattern was less clear. Figure EC15 illustrates the variability in levels of tetracyclines resistance in *E. coli* across the EU and the absence of a clear spatial distribution.

Figure EC14. Spatial distribution of <u>nalidixic acid</u> resistance among indicator Escherichia coli from <u>pigs</u> in countries reporting MIC data in 2012¹



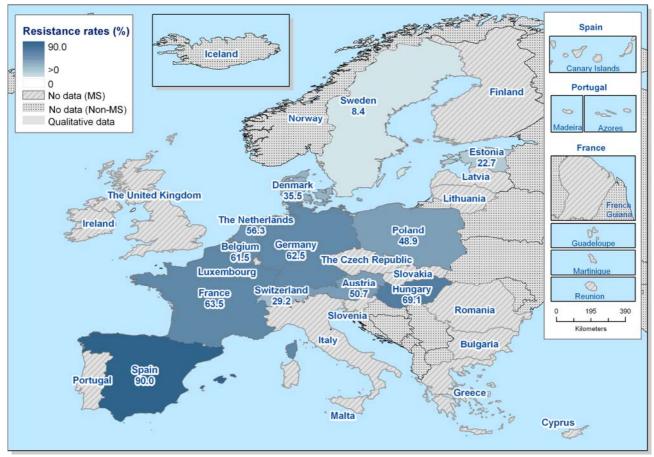
MS: Member State; MIC: minimum inhibitory concentration.

Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data were used instead.

1. For Estonia, Germany, Spain and Sweden, 2011 data were used.



Figure EC15. Spatial distribution of <u>tetracycline</u> resistance among indicator Escherichia coli from <u>pigs</u> in countries reporting MIC data in 2012¹



MS: Member State; MIC: minimum inhibitory concentration.

Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data were used instead.

1. For Estonia, Germany, Spain and Sweden, 2011 data were used.



Multi-resistance among indicator Escherichia coli isolates from fattening pigs

Three MSs and one non-MS tested the complete harmonised set of antimicrobials for *E. coli* and reported isolate-based data. Around 40 % of the isolates tested were susceptible to the nine antimicrobials of the set in three reporting countries, while the proportion was lower than 25 % in Hungary. Multi-resistance levels (i.e. reduced susceptibility to three or more antimicrobial classes) were high in all reporting countries (Table EC11), ranging between about one-quarter and half of the indicator *E. coli* isolates from pigs. The frequency distributions (Figure EC16) showed that all reporting countries detected multi-resistance to as many as six or seven antimicrobial classes. Very few isolates exhibited co-resistance to cefotaxime and ciprofloxacin using either ECOFFs or clinical breakpoints as interpretive criteria (Table EC11).

Table EC11. Complete susceptibility, multi-resistance and index of diversity in Escherichia coli from <u>fattening pigs</u> in MSs and one non-MSs reporting isolate-based data, 2012

Country	Suscept	Susceptible to all		Multi-resistant		Co-resistant to Cip and Ctx	
	n	%	n	%	diversity	n	%
Austria (N=140)	53	37.9	34	24.3	0.379	0(0)	0(0)
Denmark (N=152)	63	41.5	49	32.2	0.428	0(0)	0(0)
Hungary (N=68)	16	23.5	34	50.0	0.516	0(0)	0(0)
Switzerland (N=185)	80	43.2	64	34.6	0.420	1(0)	0.5(0)

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *E. coli*; n: number of isolates.

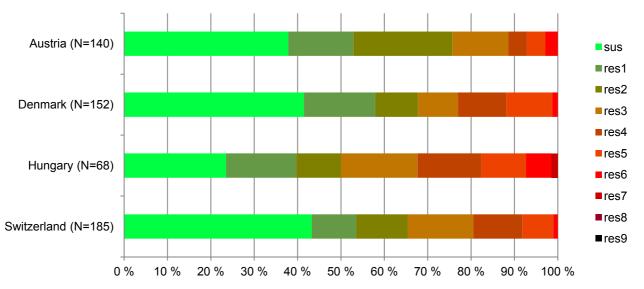
Susceptible to all: isolate susceptible to all antimicrobial substances of the EFSA common set for E. coli.

Multi-resistant: resistant to at least 3 different antimicrobial substances, belonging to any three antimicrobial families from the common antimicrobial set for *E. coli*.

Index of diversity: see definition in Section 8.4.2.1 of Chapter 8 Materials and methods.

Co-resistant to cefotaxime (Ctx) and ciprofloxacin (Cip): the effectives and percentages of *E. coli* isolates non-susceptible to concentrations greater than epidemiological cut-off values (Ctx >0.25 mg/L and Cip >0.03 mg/L). Figures in parentheses indicate the occurrence of co-resistance to Cip and Ctx determined using clinical breakpoints (Ctx >2 mg/L and Cip >1 mg/L).

Figure EC16. Frequency distribution of Escherichia coli isolates completely susceptible and resistant to one to nine antimicrobials in <u>fattening pigs</u> in MSs and one non-MS reporting isolate-based data, 2012



MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *E. coli*; sus: susceptible to all antimicrobial substances of the EFSA common set for *E. coli*; res1–res9: resistance to one antimicrobial substance/resistance to nine antimicrobial substances of the common set for *E. coli*.



Multi-/co-resistance patterns among indicator Escherichia coli isolates from fattening pigs

Indicator *E. coli* isolates resistant to cefotaxime and ciprofloxacin were only observed in Switzerland among reporting countries and streptomycin, sulphonamide and tetracycline resistance was also often present in the isolates tested (Table EC13). These additional resistances (together with trimethoprim resistance in some cases) were also noted in *E. coli* isolates showing high-level ciprofloxacin resistance (Table EC14).

Table EC12. <u>Multi-/co-resistance</u> patterns of interest in Escherichia coli from <u>fattening pigs</u> in MSs and one non-MS reporting isolate-based data, 2012

	Multi-/co-resistance pattern							Gro co	oup of rep untries (N	oorting I=545)	Austria (N=140)	Denmark (N=152)	Hungary (N=68)	Switzerland (N=185)			
Amp	Caz	Ctx	Chl	Cip	Gen	Nal	Str	Su	Tet	Tmp	n	%	group %	n	n	n	n
				R		R					20	95.2	3.7	7	1	7	5
	R	R		R							1	4.8	0.2	0	0	0	1
	Total					21	100	3.9	7	1	7	6					

MS: Member State; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *E. coli* and multiresistant; Amp: ampicillin; Caz: ceftazidime; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Nal: nalidixic acid; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethroprim; n: number of multi-/co-resistant isolates; R: minimum inhibitory concentration above European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

Table EC13. Co-resistance to cefotaxime and ciprofloxacin in Escherichia coli from fattening pigs in MSs and one non-MS reporting isolate-based data, 2012

Country	Resistance to both Cip and Ctx, applying ECOFFs ¹	Resistance to both Cip and Ctx, applying clinical breakpoints	Multi-resistance patterns of isolates resistant to both Cip and Ctx, applying ECOFFs ¹
	n (%)	n (%)	(number of isolates)
Austria (N=140)	0 (0 %)	0 (0 %)	NA
Denmark (N=152)	0 (0 %)	0 (0 %)	NA
Hungary (N=68)	0 (0 %)	0 (0 %)	NA
Total (3 MSs) (N=360)	0 (0 %)	0 (0 %)	
Switzerland (N=185)	1 (0.5 %)	0 (0 %)	AmpCtxCip (1)

MS: Member State; N: total number isolates for which relevant data are available; n: number of co-resistant isolates; Cip: ciprofloxacin; Ctx: cefotaxime; NA: not applicable.

1. European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs) and clinical breakpoints were applied.

Table EC14. <u>Ciprofloxacin</u> resistance assessed at differing thresholds in indicator Escherichia coli from <u>fattening pigs</u> in MSs and one non-MS reporting isolate-based data, 2012

Country	Isolates resistant at >0.03 mg/L ¹ Cip	Isolates resistant at >1 mg/L ² Cip	Isolates resistant at ⊳2 mg/L ³ Cip	Isolates resistant at >4 mg/L ⁴ Cip	Resistance patterns of isolates resistant at >4 mg/L Cip (number of isolates)
	n (%)	n (%)	n (%)	n (%)	
					AmpCipNalStrSuTetTmp (2)
Austria (N=140)	7 (5.0 %)	4 (2.9 %)	4 (2.9 %)	4 (2.9 %)	CipNalStrTetTmp (1)
					AmpCipNalTet (1)
Denmark (N=152)	1 (0.7 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA
(h) (h)	0 (40 0 0()				AmpCipNalTet (1)
Hungary (N=68)	9 (13.2 %)	2 (2.9 %)	2 (2.9 %)	2 (2.9 %)	AmpCipNalStrSuTetTmp (1)
Total (3 MSs) (N=360)	17 (4.7 %)	6 (1.7 %)	6 (1.7 %)	6 (1.7 %)	
Switzerland (N=185)	6 (3.2 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA

MS: Member State; N: total number isolates for which relevant data are available; n: number of resistant isolates; Cip: ciprofloxacin; NA: not applicable.

1. European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2. EUCAST clinical breakpoint.

3. High breakpoint.

4. Very high breakpoint.



5.2.2.3. Cattle (bovine animals)

Representative sampling and monitoring

In 2012, quantitative data for *E. coli* in cattle were provided by seven MSs and one non-MS (Switzerland) (Table EC15). These countries tested different production types and ages of cattle, including calves, young cattle, meat production animals, adult cattle and dairy cows; Denmark and Poland did not specify the type of cattle that were tested.

Among the reporting MSs, antimicrobial resistance monitoring in indicator *E. coli* isolates from cattle was chiefly based on monitoring plans of healthy bovine animals randomly selected within the slaughterhouses (Austria, Denmark, Finland and Switzerland). Indicator *E. coli* isolates were isolated from caecal contents in Austria, from recto-anal swabs in Switzerland and from faeces in Denmark and Finland by sampling healthy cattle at slaughter. Belgium and Poland did not report information on the sample type, sampling context and sampling stage.

The overall results for cattle presented in Table EC15 include all isolates of *E. coli* that were collected from this animal species by MSs which tested more than 10 isolates from cattle in total. Results are also presented for the specific production levels of cattle from which these *E. coli* isolates originated. Some MSs tested fewer than 10 isolates from individual production types. In such cases, the data for these production types are included in the overall results for cattle but are not presented in the production level-specific sections of this table.

Resistance levels among Escherichia coli isolates from cattle

The occurrence of resistance to gentamicin and nalidixic acid was less common, with an overall level at the reporting MS group of 3.1 %, and 8.6 %, respectively. Two countries reported no resistance to cefotaxime, with the highest resistance levels being 8.8 % and 2.6 %, recorded by Belgium and Poland among mixed herds and unspecified production type, respectively.

In indicator *E. coli* isolates from calves of less than one year of age, tested in Austria, Germany and the Netherlands, resistance to ampicillin, streptomycin, sulfonamides and tetracyclines was generally moderate to high, while resistance to chloramphenicol and gentamicin was recorded at low to moderate and low levels, respectively. The occurrence of resistance to (fluoro)quinolones and third-generation cephalosporins was less common, as ciprofloxacin and nalidixic acid resistance was reported at low to moderate levels, while resistance to cefotaxime was low to very low. Switzerland, which monitored indicator *E. coli* resistance in bovine animals for meat production, recorded similar features of resistance to the same panel of substances, although at a slightly lower level.

Austria which also submitted data concerning young cattle (aged one to two years) and adult cattle (over two years) reported lower resistance levels in these age groups than in calves of less than one year; only resistance to streptomycin, sulfonamides and tetracyclines was detected at around 5 %, while no resistance was recorded to the other substances of the common panel. The Netherlands reported much lower resistance, at around 1 % levels, among dairy cows than among veal calves.

Table EC15. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among isolates of Escherichia coli from <u>cattle</u> in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

0	Amp	icillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Gent	amicin
Country	Ν	% Res	N	% Res	Ν	% Res	N	% Res	N	% Res
All cattle	•	•	-	•	•	•	-	•	•	•
Austria	273	5.9	273	0.7	273	2.2	273	2.2	273	0.7
Belgium	364	56.0	364	8.8	364	31.0	364	33.8	364	6.0
Denmark	98	5.1	98	0	98	2.0	98	0	98	0
Finland	295	1.7	295	0	295	0	_	_	295	0.3
Germany	515	46.8	515	2.5	515	17.5	515	16.5	515	6.4
Netherlands	559	13.6	559	0.5	559	7.7	559	3.4	559	1.4
Poland	190	7.9	190	2.6	190	2.6	190	5.3	190	2.6
Total (7 MSs)	2,294	24.5	2,294	2.4	2,294	11.3	1,999	12.2	2,294	3.1
Switzerland	187	14.4	187	0.5	187	3.7	187	3.2	187	5.9
Calves (under 1 y	/ear)									
Austria	151	10.6	151	1.3	151	3.3	151	4.0	151	1.3
Germany	515	46.8	515	2.5	515	17.5	515	16.5	515	6.4
Netherlands	285	25.6	285	0.7	285	14.4	285	6.0	285	2.5
Total (3 MSs)	951	34.7	951	1.8	951	14.3	951	11.4	951	4.4
Young cattle (1–2	2 years)									
Austria	73	0	73	0	73	0	73	0	73	0
Meat production	animals									
Switzerland	187	14.4	187	0.5	187	3.7	187	3.2	187	5.9
Adult over 2 year	'S									
Austria	49	0	49	0	49	2.0	49	0	49	0
Dairy cows										
Netherlands	274	1.1	274	0.4	274	0.7	274	0.7	274	0.4
Mixed herds										
Belgium	364	56.0	364	8.8	364	31.0	364	33.8	364	6.0
Finland	295	1.7	295	0	295	0	_	-	295	0.3
Total (2 MSs)	659	31.7	659	4.9	659	17.1	364	33.8	659	3.5

MIC: minimum inhibitory concentration; MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates.

Table continued overleaf.



Table EC15 (continued). Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines among isolates of Escherichia coli from <u>cattle</u> in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

0	Nalidixi	c acid	Strepto	mycin	Sulfona	mides	Tetrac	yclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
All cattle		<u>.</u>		<u>.</u>	<u>.</u>	•		
Austria	273	1.8	273	12.5	273	11.4	273	14.7
Belgium	364	28.6	364	51.6	364	59.3	364	58.2
Denmark	98	0	98	6.1	98	6.1	98	7.1
Finland	295	0	295	5.4	295	3.4	295	2.4
Germany	515	12.0	515	43.3	515	50.1	515	53.0
Netherlands	559	2.9	559	16.3	559	17.0	559	25.2
Poland	190	5.3	190	8.9	190	17.4	190	11.6
Total (7 MSs)	2,294	8.6	2,294	25.1	2,294	28.3	2,294	30.6
Switzerland	187	3.2	187	24.6	187	26.2	187	24.6
Calves (under 1 y	ear)				-			
Austria	151	3.3	151	18.5	151	17.2	151	22.5
Germany	515	12.0	515	43.3	515	50.1	515	53.0
Netherlands	285	5.3	285	30.9	285	32.6	285	48.1
Total (3 MSs)	951	8.6	951	35.6	951	39.6	951	46.7
Young cattle (1–2	years)	·						
Austria	73	0	73	5.5	73	4.1	73	5.5
Meat production	animals	·						
Switzerland	187	3.2	187	24.6	187	26.2	187	24.6
Adult over 2 year	S							
Austria	49	0	49	4.1	49	4.1	49	4.1
Dairy cows								
Netherlands	274	0.4	274	1.1	274	0.7	274	1.5
Mixed herds								
Belgium	364	28.6	364	51.6	364	59.3	364	58.2
Finland	295	0	295	5.4	295	3.4	295	2.4
Total (2 MSs)	659	15.8	659	31.0	659	34.3	659	33.2

MIC: minimum inhibitory concentration; MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates.



Temporal trends in resistance among indicator *Escherichia coli* isolates from cattle

Figures EC17 to EC21 display the trends in resistance to selected antimicrobials in *E. coli* from cattle. It should be noted that the figures presented for each country combine the results for all cattle production types and/or ages submitted each year. As in the other livestock species, the resistance levels varied substantially between MSs for several of the antimicrobials, including ampicillin, streptomycin and tetracyclines. Austria and Denmark reported the lowest levels of resistance for many of the antimicrobials. As in pigs, cefotaxime resistance has been below 5 % in all countries since 2005 (Figure EC18).

Considering the previous years of reporting, the resistance levels reported by Denmark in 2011 and 2012 were broadly comparable. In Austria, the studied population changed in 2012 with calves being overrepresented compared with the other age of groups in the years before 2012; therefore, statistical trends were not calculated. Switzerland reported decreases in resistance to most antimicrobials between 2010 and 2012, which is most probably because the study population in 2010 was veal calves less than six months old whereas in 2011 older cattle (>12 months) were sampled. In Germany, resistance rates were much lower in 2011 than in 2010 and in 2012. However, in 2010 and 2012 veal calves were tested while in 2011 young beef animals were tested which usually differ in management and antimicrobial exposure.

Some countries, such as Denmark and the Netherlands, have shown relatively stable resistance levels or only minor fluctuations or trends since 2006 whereas other countries, such as Germany and Switzerland, have shown more substantial fluctuations in resistance levels that are, at least partially, due to the sampling of different cattle production types in different years. There have been numerous statistically significant trends in resistance levels since 2006; for example, Germany and the Netherlands showed significant declines in resistance to five of the antimicrobials. Significant decreasing trends were also observed in Denmark and Switzerland.

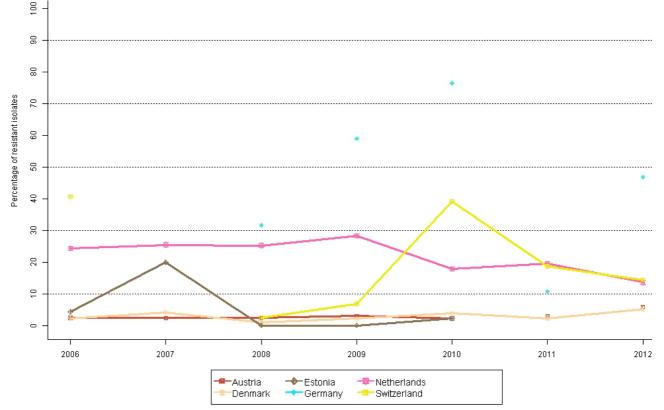
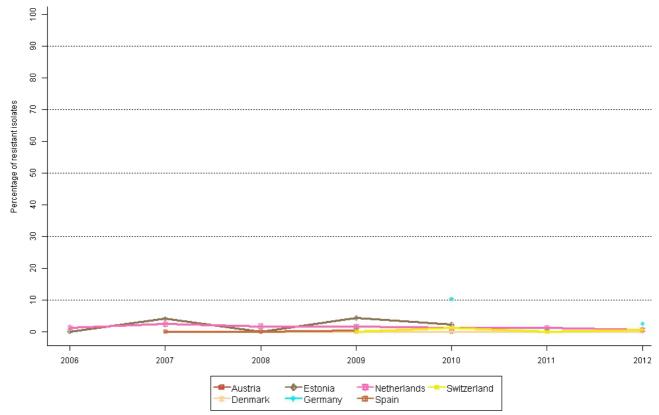


Figure EC17. Trends in <u>ampicillin</u> resistance in indicator Escherichia coli from <u>cattle</u> in reporting <i>MSs and one non-MS, 2006–2012, quantitative data

M: Member State.

Note: Statistically significant increasing or decreasing trends over five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in Germany (\downarrow), the Netherlands (\downarrow) and Switzerland (\downarrow).

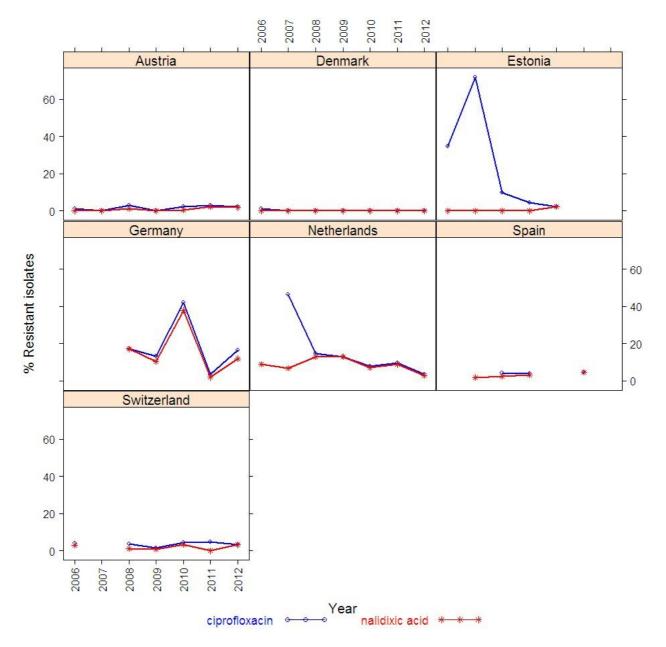
Figure EC18. Trends in <u>cefotaxime</u> resistance in indicator Escherichia coli from <u>cattle</u> in reporting MSs and one non-MS, 2006–2012, quantitative data



M: Member State.

Note: No statistically significant trends over five or more years, as tested by a logistic regression model ($p \le 0.05$), were observed in any of the reporting countries.

Figure EC19. Trends in <u>ciprofloxacin</u> and <u>nalidixic acid</u> resistance in indicator Escherichia coli from <u>cattle</u> in reporting MSs and one non-MS, 2006–2012, quantitative data

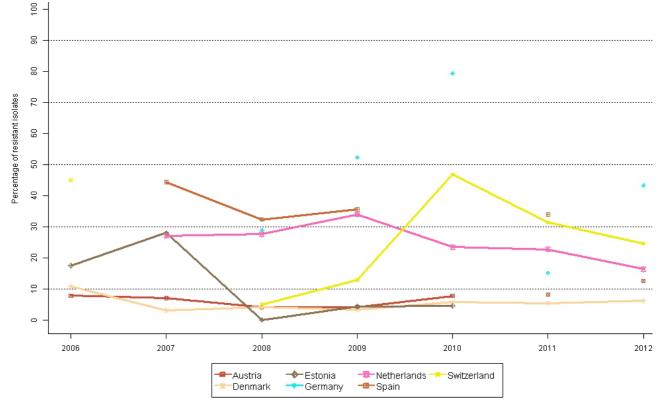


M: Member State.

Note: Statistically significant decreasing trends over five or more years, as tested by a logistic regression model (*p* ≤0.05), were observed in Germany (↓) and the Netherlands (↓) for both ciprofloxacin and nalidixic acid, in Denmark (↓) and Estonia (↓) for ciprofloxacin.

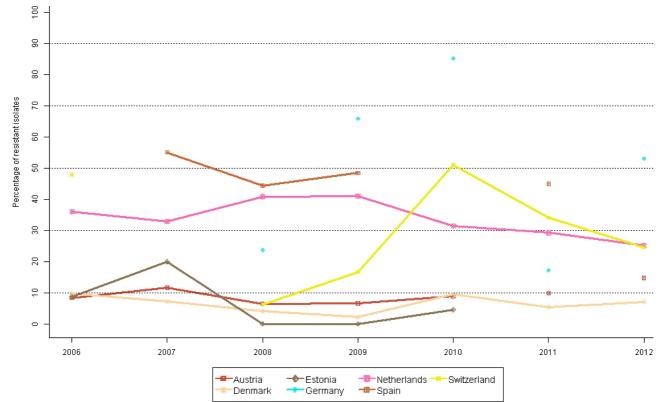






Note: Statistically significant increasing and decreasing trends over five or more years, as tested by a logistic regression model $(p \le 0.05)$, were observed in Estonia (\downarrow), Germany (\downarrow), the Netherlands (\downarrow) and Switzerland (\downarrow). MS: Member State.





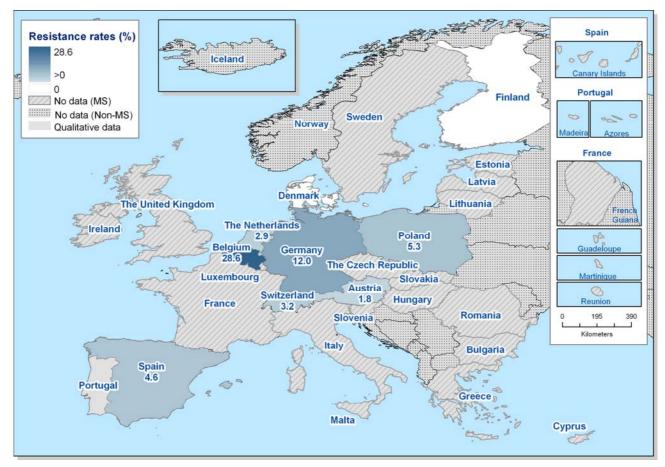
Note: Statistically significant increasing and decreasing trends over five or more years, as tested by a logistic regression model $(p \le 0.05)$, were observed in Germany (\downarrow), the Netherlands (\downarrow) and Switzerland (\downarrow). MS: Member State.



Spatial distribution of resistance among indicator *Escherichia coli* isolates from cattle

The spatial distributions of nalidixic acid and tetracycline resistance among *E. coli* from cattle are shown in Figures EC22 and EC23. Nevertheless, there was still some evidence that the lowest resistance to tetracyclines occurred in the northern countries and the highest occurred in the southern and western countries. With respect to nalidixic acid, the majority of countries reported low levels of resistance and no spatial pattern was evident.

Figure EC22. Spatial distribution of <u>nalidixic acid</u> resistance among indicator Escherichia coli from <u>cattle</u> in countries reporting MIC data in 2012¹



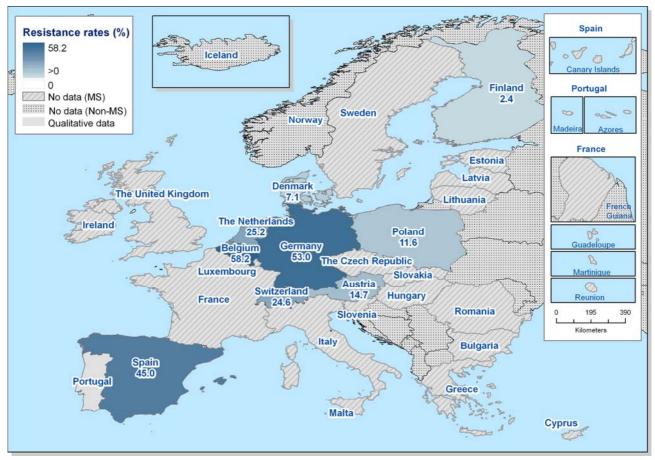
MIC: minimum inhibitory concentration; MS: Member State.

Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data were used instead. The countries labelled as 'qualitative data' therefore include those reporting MIC data for fewer than 10 isolates or purely qualitative data (as proportion of resistant isolates).

1. For Spain, 2011 data were used.



Figure EC23. Spatial distribution of <u>tetracycline</u> resistance among indicator Escherichia coli from <u>cattle</u> in countries reporting MIC data in 2012¹



MIC: minimum inhibitory concentration; MS: Member State.

- Note: Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, 2011 data were used instead. The countries labelled as 'qualitative data' therefore include those reporting MIC data for fewer than 10 isolates or purely qualitative data (as proportion of resistant isolates).
- 1. For Spain, 2011 data were used.

Multi-resistance among indicator Escherichia coli isolates from cattle

No tables and graphs on multi-resistance are presented in this report for *E. coli* in cattle because too few MSs reported multi-resistance isolate-based data on more than 10 isolates for the different production types of cattle.



5.3. Multi-drug resistance patterns in indicator *E. coli* in 2012

The MDR patterns in indicator *E. coli* from broilers and fattening pigs, in MSs reporting isolate-based data, are shown in Appendix 4, Table MDRP37, and Appendix 4, Table MDRP38, respectively.

5.3.1. Multi-drug resistance in *E. coli* isolates from broilers

A large number of different resistance patterns in indicator *E. coli* isolates from broilers were evident (57 different patterns displayed by 534 isolates), reflecting the diverse nature of the *E. coli* strains which have been tested. Although no single pattern occurred at a frequency of greater than 3 % amongst the MDR patterns obtained from broilers, a common core of resistance to ampicillin, sulfonamides and tetracyclines, generally with resistance to ciprofloxacin and frequently with resistance to streptomycin and trimethoprim, was discernible. Several resistance patterns which occurred at a higher frequency included resistance to cefotaxime; however, cefotaxime resistance also occurred as a component of infrequent resistance patterns. Ciprofloxacin resistance frequently occurred as a component of MDR in *E. coli* from broilers and was observed in 72.3 % of MDR isolates (120 out of 166).

5.3.2. Multi-drug resistance in *E. coli* isolates from fattening pigs

The overall range of different patterns observed in indicator *E. coli* isolates from pigs in MSs reporting isolate-based data was similar to that seen in broilers, with a large number of different resistance patterns evident (54 different patterns displayed by 565 isolates), again reflecting the diverse nature of the *E. coli* strains which have been tested. Particular MDR patterns were predominant in fattening pigs, with two patterns occurring at a frequency of greater than 14 % amongst the MDR patterns obtained, and each of these patterns comprising more than 5 % of the *E. coli* isolates for which isolate-based data was available. In pigs, *E. coli* with either of two MDR patterns (pattern one: streptomycin, sulfonamides, tetracyclines, ampicillin and trimethoprim), therefore, accounted for approximately 10 % of the total number of *E. coli* isolates for which isolate-based data were available. Resistance to streptomycin, sulfonamides and tetracyclines formed a common core in both of these MDR patterns and also occurred as a recurring core pattern in isolates showing additional resistances. Considering those resistance patterns occurring at a higher frequency in pigs, these did not generally include resistance to cefotaxime; however, cefotaxime resistance did occur as a component of infrequent resistance patterns. Ciprofloxacin resistance occurred infrequently as a component of MDR in pigs and was present in 16 % of porcine MDR *E. coli* isolates (32 out of 200).

5.4. Overview of findings on indicator *E. coli* resistance at reporting MS group level, 2012

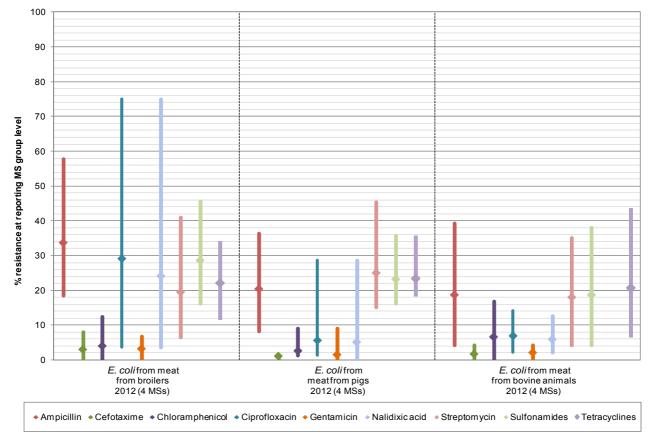
Figures EC24 and EC25 display the resistance levels among *E. coli* isolates in the reporting MS group, based on quantitative data submitted in 2012. These data were not all derived from the same group of MSs, which needs to be considered when interpreting these figures.

The levels of resistance were broadly similar for meat from broilers, pigs and bovine animals for all reporting MSs for these antimicrobials (Figure EC24). The situation was different for ciprofloxacin and nalidixic acid, where resistance was high in meat from broilers considering all reporting MSs at 29.1 % and 24.1 % respectively, but low in meat from pigs and meat from bovine animals at less than 7 %.

The resistance levels observed in *E. coli* isolates from cattle were lower than in *E. coli* isolates from either *Gallus gallus* or pigs, most notably for ampicillin, streptomycin, sulfonamides and tetracyclines (Figure EC25). The variations at the reporting MS group level between years could be attributable to different MSs contributing data and different production types of livestock being sampled. The MSs that provided data for all three livestock species in both 2011 and 2012 usually reported the lowest resistance levels among cattle.

As in previous years, isolates from pigs had the highest levels of resistance to streptomycin and tetracyclines, while isolates from *Gallus gallus* had the highest resistance to ampicillin, ciprofloxacin, nalidixic acid and sulfonamides. Resistance to chloramphenicol and gentamicin was relatively low in all types of livestock, with the highest resistance level occurring in *Gallus gallus*. This differs from the situation in 2010, when the highest resistance levels for these two antimicrobials were observed in cattle. Chloramphenicol has not been used for food production animals in the EU for several years; thus, the resistance observed must either indicate persistence of resistance genes or co-selection resulting from use of related compounds (such as florfenicol). The lowest levels of resistance were usually observed to cefotaxime; the highest level of resistance to this antimicrobial occurred in isolates from *Gallus gallus*, which was also the case in previous years.

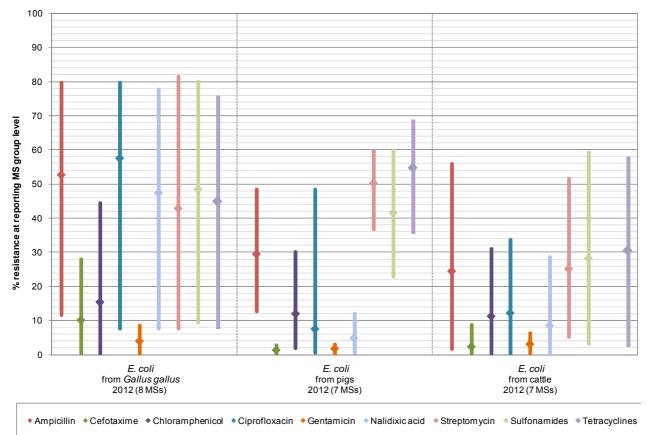
Figure EC24. Resistance in indicator Escherichia coli *from <u>meat from broilers</u>, <u>meat from pigs</u> and <u>meat from bovine animals</u> to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, <i>nalidixic acid, streptomycin, sulfonamides and tetracyclines at reporting MS group level, in 2012*



MS: Member State.



Figure EC25. Resistance in indicator Escherichia coli from <u>fowl</u>, <u>pigs</u> and <u>cattle</u> to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, streptomycin, sulfonamides and tetracyclines at reporting MS group level, in 2012



MS: Member State.



5.5. Discussion

Antimicrobial resistance in indicator commensal *E. coli* from animals and food can be used to examine the reservoir of resistance genes occurring in those bacteria that could be transferred to bacteria that are pathogenic for humans and/or animals. The major factor influencing the occurrence of resistance to antimicrobials in indicator *E. coli* is likely to be the selective pressures exerted by use of antimicrobials in the different food animal populations; variations in usage between animal species may also contribute to the observed differences in resistance levels between the animal species. Indicator *E. coli* are thus also of interest when investigating possible associations between the usage of antimicrobials in a given country and the occurrence of resistance in an animal species, because of their ubiquity in food-producing animals. Multi-resistance data, available for the first time in 2012, indicates that the co-resistance in animals.

A total of 11 MSs and 2 non-MSs provided quantitative MIC data, in 2012, on at least one of the livestock species. Reported antimicrobial resistance data in *E. coli* isolates from food-producing animals and food, derived mainly from active and representative monitoring programmes, were chiefly based on randomised sampling performed at the slaughterhouse. At the reporting MS group level, a high level of resistance was observed to several antimicrobials among food-producing animals, with some countries reporting a very or extremely high occurrence of resistance. As resistance levels tend to vary substantially between countries, the variation in resistance in *Gallus gallus*, pigs and cattle observed between the years 2009 and 2012, at the overall MS group level, may partly result from different MSs contributing to data as well as different production types of livestock being sampled.

In 2012, four MSs reported on antimicrobial resistance in each category of meat, but those which did generally reported comparable resistance levels in meat as in the corresponding source animal species. Resistance levels were generally higher among *E. coli* isolates from *Gallus gallus* and pigs than isolates from cattle. This was the second year that resistance data were reported separately for different production types of *Gallus gallus* and cattle. However, only two countries provided data on laying hens, and both of these MSs also provided data on broilers. Although there is limited information available for 2011 and 2012 on which to draw firm conclusions, resistance levels were generally higher among broilers than in laying hens. Similarly, in 2012, only two MSs reported on more than one production type or age group of cattle. The Netherlands reported much higher resistance levels among younger animals and a similar trend to higher resistance in young animals was also observed in Austria. Regional differences in the occurrence of resistance were evident for some antimicrobials for indicator *E. coli* from broilers and pigs. This may reflect differences in the structure of the respective livestock sectors, with pyramidal production systems more common in pigs and broilers.

Generally, the highest resistance levels were identified for **ampicillin**, **sulfonamides and tetracyclines**, which are commonly used therapeutically in animals. Moreover, some countries have shown statistically significant increasing or decreasing trends in resistance to these antimicrobials over five or more years since 2006. At the MS group level, resistance to gentamicin was highest in *Gallus gallus* (4.0 %) and lowest in pigs (1.8 %). Gentamicin is an interesting antimicrobial because there are differences in the degree of usage in different MSs of this and other antimicrobials to which cross-resistance may occur (for example apramycin).

Resistance was also identified to fluoroguinolones (ciprofloxacin), a class of antimicrobials recognised to be critically important in human medicine. Although resistance was generally similar in E. coli from meat from broilers, pigs and cattle, for all of the antimicrobials tested for the group of reporting MSs, resistance to nalidixic acid and ciprofloxacin was a notable exception, as much higher levels of resistance were recorded in E. coli isolates from meat from broilers than from meat from the other species. Similarly, the occurrence of resistance to nalidixic acid and ciprofloxacin was higher in E. coli from broilers than in isolates from pigs and cattle. As resistance to fluoroquinolones commonly includes a mutational component, this suggests that either E. coli from broilers are exposed to greater selective pressure from the overall use of fluoroquinolones, or the use of fluoroquinolones at a particular part of the production pyramid (which selects for mutational resistance there) engenders resistance which is subsequently disseminated to flocks lower in the pyramid by the spread and transfer of resistant bacterial clones. Although the occurrence of high-level fluoroquinolone resistance is likely to be influenced by the degree of fluoroquinolone usage, it is also likely to be influenced by the degree to which terminal hygiene and disinfection procedures allow strains, which have developed some resistance, to persist and colonise the subsequent group of animals. The occurrence of resistance to nalidixic acid was often similar then that for ciprofloxacin, suggesting that mutation in the topoisomerase enzymes (gyrA or parC) may, in those cases, have been responsible for resistance. However, in some MSs, the occurrence of resistance to ciprofloxacin was slightly higher than that obtained for nalidixic acid. In these

cases, mechanisms such as transferable fluoroquinolone resistance conferred by *qnr* genes may have been responsible for resistance; as such plasmid-mediated mechanisms can result in that phenotypic pattern of resistance. One country has shown statistically significant decreasing trends in resistance to ciprofloxacin in all species over five or more years since 2006.

REVISION OF EPIDEMIOLOGICAL CUT-OFF VALUES FOR CIPROFLOXACIN FOR E. COLI

The epidemiological cut-off value (ECOFF) for E. coli versus ciprofloxacin has been recently revised by the European Committee on Antimicrobial Susceptibility Testing (EUCAST). Wild-type isolates are now considered to have a ciprofloxacin minimum inhibitory concentration (MIC) lower than or equal to 0.06 mg/L, an increase from the previous ECOFF of 0.03 mg/L. The proportion of isolates showing microbiological resistance according to this breakpoint will alter when the new breakpoint is adopted and in fact will be reduced. For reasons of continuity and to comply with the current legislation where applicable, the ECOFFs used in this report have been those adopted in EFSA's recommendations (EFSA, 2007, 2008). For these reasons, the most recent revisions by EUCAST have not been included in this report. The report for 2013 will incorporate all of these changes in a comprehensive revision, which will also re-evaluate the historical data using the revised ECOFFs, as well as taking into account revised EU legislation in this area, which will include revised ECOFFs.

The reported MIC distributions for *E. coli* versus ciprofloxacin are available on the EFSA website in 'Level 3 Tables'. From these it can be seen that the total number of *E. coli* reported was 8,580. Of these, 32.7 % were resistant considering the ECOFF of >0.03 mg/L whereas 25.6 % were resistant considering the ECOFF of >0.06 mg/L.

Resistance to third-generation cephalosporins (**cefotaxime**), another class categorised as critically important in human medicine, was infrequently detected in 2012. There were few countries which reported significant increasing or decreasing trends in resistance to cefotaxime over five or more years since 2006. EFSA has published recommendations for surveillance of indicator *E. coli* resistant to cefotaxime, which would extend the scope of the current monitoring by including selective culture for such organisms (EFSA, 2012b). Current procedures rely on random selection of indicator *E. coli* isolates from primary culture plates; selective culture could additionally be used to detect the presence or absence of isolates resistant to cefotaxime in a sample (within the detection limit of the chosen method). Monitoring using selective media for cefotaxime resistance would thus detect cefotaxime-resistant *E. coli* present as a minor component of the total bacterial flora in the test sample and which might only occasionally be detected by random sampling from non-selective culture plates. The occurrence of third-generation cephalosporin resistance was still generally low, although a number of reporting MSs recorded high to moderate levels in *E. coli* from *Gallus gallus*, and resistance was typically higher in isolates from *Gallus gallus* than in pigs or cattle. The findings in relation to third-generation cephalosporin resistance are discussed further in Chapter 7.

Although the levels of multi-resistance¹⁷ were relatively high in indicator *E. coli* isolates from both broilers and pigs in most reporting countries, co-resistance to cefotaxime and ciprofloxacin was detected at very low or low levels in broilers, but not in pigs, in the reporting MSs in 2012. This year, for the first time in this report, the multi-drug resistance patterns shown by indicator E. coli from broilers and pigs from MSs reporting isolate-based data have been included. A common core of resistance to ampicillin, sulfonamides and tetracyclines, generally with resistance to ciprofloxacin and frequently with resistance to streptomycin and trimethoprim, was discernible in broilers where no single pattern or patterns of resistance occurred at a high frequency. In fattening pigs, two MDR patterns were predominant (pattern one: streptomycin, sulfonamides, tetracyclines and streptomycin, and pattern two: sulfonamides, tetracyclines, ampicillin and trimethoprim) and each accounted for more than 5 % of the total number of E. coli isolates from fattening pigs for which isolatebased data were available. The occurrence of these particular patterns might reflect spread of particular clones of bacteria which exhibit that pattern of resistance or dissemination of plasmids carrying those resistances and possibly being transmitted between different strains of E. coli. In broilers, ciprofloxacin resistance was particularly noted in MDR patterns and resistance to this compound can be mediated through chromosomal mutations or through transferable mechanisms of resistance. Ciprofloxacin resistance frequently occurred as a component of MDR in E. coli from broilers and was observed in 72.3 % of MDR

¹⁷ Proportions of isolates showing reduced susceptibility to at least three antimicrobial classes according to epidemiological cut-off values.



isolates (120 out of 166), whereas ciprofloxacin resistance occurred infrequently as a component of MDR in pigs and was present in 16.0 % (32 out of 200) of porcine MDR *E. coli* isolates.

Several resistance patterns, which occurred at a higher frequency in *E. coli* from broilers, included resistance to cefotaxime; however, cefotaxime resistance also occurred as a component of infrequent resistance patterns in this species. Considering the resistance patterns occurring at a higher frequency in pigs, these did not generally include resistance to cefotaxime; however, cefotaxime resistance did occur as a component of infrequent resistance patterns. Resistance to cefotaxime may be conferred by ESBL or AmpC beta-lactamase enzymes; in *E. coli* the former are generally carried on plasmids, whereas AmpC resistance may be carried on plasmids or relate to mutations of the promoter region of the endogenous beta-lactamase carried by *E. coli*. Further characterisation of cefotaxime resistance shown by these isolates, which has been recommended by EFSA (EFSA, 2012b) in particular phenotypic and genetic characterisation of the resistance patterns to be identified which are associated with the carriage of ESBL or AmpC enzymes. This has potential implications for control through the identification of antimicrobials which might encourage the spread of cefotaxime resistance through co-selection.

A recent study in Spain examined the integrons carried by *E. coli* isolates recovered from healthy broilers and pigs (Marchant et al., 2013). Integrons can be associated with particular antimicrobial resistance genes and in the Spanish study both class 1 and class 2 integrons were detected in pigs and chickens. Class 1 integrons classically carry the resistance gene *sul1*; additionally, both types of integrons in the Spanish study often carried genes associated with streptomycin and trimethoprim resistance, while resistance genes conferring chloramphenicol and gentamicin resistance were detected in the variable region of class 1 integrons only. The widespread occurrence of integrons and their associated antimicrobial resistance genes in animal *E. coli* is likely to account for some of the resistance patterns (or associations between resistances) which are evident in the MDR tables and probably explains why sulfonamide, streptomycin and trimethoprim resistance are common components of MDR patterns. The Spanish study also reported that the presence of integrons was associated with resistance to amoxicillin (equivalent to ampicillin for resistance purposes) and tetracyclines. The common core patterns of resistance to ampicillin, streptomycin, sulfonamides, tetracyclines and trimethoprim (and combinations thereof) frequently observed in the monitoring of *E. coli* isolates are probably therefore related to the presence of integrons.

There may be numerous permutations in the ways in which resistance genes may be arranged or accumulated by *E. coli*, and attempting to relate the resistance phenotype to the likely underlying mechanisms of resistance is therefore difficult. Full resistance to all of the antimicrobials in the test panel was noted only for a single *E. coli* isolate from pigs and there might be numerous ways in which such resistance might develop. It is interesting, however, that such resistance has been described before in *E. coli* isolates from animals and has been reported, for example, in *E. coli* recovered from cattle in France (Meunier et al., 2010).



6. METHICILLIN-RESISTANT STAPHYLOCOCCUS AUREUS

6.1. Introduction

METHICILLIN-RESISTANT STAPHYLOCOCCUS AUREUS (MRSA)

MRSA has been recognised as an important cause of healthcare-associated infections in humans for decades. Strains of MRSA have also emerged which are particularly associated with communityassociated infections in humans. Moreover, in recent years, MRSA has also been detected in several animal species, notably including pigs and companion animals as well as some other farm animal species. Hospital-associated MRSA and community-associated MRSA are those strains predominantly affecting humans, while they generally do not involve food-producing animals; however, livestock-associated MRSA may also be harboured by humans, especially where there is occupational contact with affected livestock. Livestock-associated MRSA may cause illness in humans, although transmissibility between humans has been shown to be very limited, even in healthcare facilities.

Antimicrobial susceptibility in European invasive Staphylococcus aureus isolates is reported by the Member States (MSs) to the European Antimicrobial Resistance Surveillance Network (EARS-Net) (ECDC, 2013). Molecular typing data are not reported and, thus, where there may be possible links to the animal reservoir, these cannot be detected easily with current monitoring procedures, at least at the European level. Recent EARS-Net data showed decreasing or stabilising percentages of MRSA in reported invasive S. aureus isolates (both healthcare and community associated) in most European countries, which might reflect the beneficial impact of improved infection control routines implemented in several countries. However, MRSA remains a human public health priority, as the percentage of MRSA remains above 25 % in 7 out of 30 countries, mainly in Eastern and Southern Europe (ECDC, 2013).

Molecular typing techniques, such as spa-typing and multi-locus sequence typing (MLST), are commonly used in S. aureus to sub-type strains and determine lineages. In spa-typing, different genetic types or strains of MRSA are designated by a number with the prefix t, while in MLST, strains are designated by a sequence type (ST) number. Using such typing results, often in conjunction with certain other virulence and antimicrobial resistance characteristics, it is possible to sub-divide strains of MRSA into groups characterised by differing epidemiology. These techniques are of particular relevance, for instance, in the investigation of outbreaks, such as in the clinical case of hospital-associated MRSA, in transmission events, for example of livestock-associated MRSA, and in the detection of emerging strains showing new and/or multiple resistance patterns.

Pigs are acknowledged as an important source of colonisation of a particular strain of MRSA (designated ST398-multi-locus sequence type 398) for pig farmers and veterinarians, and their families, through direct or indirect contact with pigs. In some countries, other livestock (for example poultry and veal calves) have also become colonised. This recently recognised strain, MRSA ST398, which appears to be primarily acquired by occupational exposure to colonised pigs (or other colonised farmed livestock), can, on occasion, cause infections in humans and can be introduced into healthcare settings. In order to increase awareness and to assess the occurrence of MRSA in pig primary production across the EU, the occurrence and diversity of MRSA and MRSA ST398 in pig holdings in MSs were assessed through an EU-wide baseline survey (EFSA, 2009b, 2010a).

The EFSA's assessment of the public health significance of MRSA in animals and food (EFSA, 2009c) and the Joint Scientific Report of ECDC, EFSA and EMEA on MRSA in livestock, companion animals and food (EFSA, 2009a) provide more background information and recommendations on MRSA. A principal recommendation is that monitoring of food-producing animals, in particular intensively reared animals, is carried out periodically in conjunction with a systematic surveillance of MRSA in humans so that trends in the diffusion and evolution of zoonotically-acquired MRSA in humans can be identified. In particular, isolate samples, representative of various animal and food origins, should be analysed for lineage determination, antimicrobial susceptibility and virulence-associated traits. These issues were reviewed in the recent EFSA Scientific Report proposing technical specifications to improve the harmonisation of the monitoring and reporting of the prevalence, genetic diversity and multi-resistance profile of MRSA in food-producing animals and food thereof (EFSA, 2012c) (see Section 6.3 Discussion below for further information).

6.2. Methicillin-resistant *Staphylococcus aureus -* reports from individual MSs

Livestock-associated MRSA isolates are the principal focus of this chapter, which summarises the prevalence of MRSA results in various food-producing animal species and food reported by MSs to EFSA in 2012. The chapter also includes prevalence data on companion animals reported by an increasing number of MSs compared with 2011. Ten MSs - Belgium (cattle), Finland (pigs), Germany (cattle, turkeys and food), Hungary (cattle, *Gallus gallus*, geese, goats, pigs, sheep and turkeys), Ireland (dogs), the Netherlands (birds, cats, cattle, dogs, goats, pigs, sheep and solipeds), Poland (food), Slovakia (cats, cattle, chinchillas, dogs, *Gallus gallus*, goats, pigs and sheep), Slovenia (food) and Spain (food) - and one non-MS, Switzerland (pigs), submitted data on MRSA prevalence in animals and food in their national zoonoses reports for 2012 (Table MRSA1). The methods for the isolation of MRSA from animals and food to date have not been harmonised at the EU level and, therefore, the methods used by individual reporting MSs may differ in sensitivity.

In addition, data on antimicrobial resistance of MRSA isolates and *Staphylococcus aureus* from foodproducing animals were reported by only two countries in 2012.

Data	Origin	Total number of MSs reporting	Countries
	Companion animals: cats and dogs	3	MSs: IE, NL, SK
	Cattle (bovine animals)	5	MSs: BE, DE, HU, NL, SK
	Sheep and goats	3	MSs: HU, NL, SK
A i		4	MSs: FI, HU, NL, SK
Animals	Pigs	4	Non-MS: CH
	Poultry: Gallus gallus (fowl), geese and turkeys	3	MSs: DE, HU, SK
	Solipeds, domestic	1	MS: NL
	Other animals: birds and chinchillas	2	MSs: NL, SK
	Cheeses, dairy products and milk	3	MSs: DE, ES, PL
	Fish, fishery products and crustaceans	1	MS: PL
Food	Fruits	1	MS: ES
	Meat	3	MSs: DE, ES, SI
	Other food	1	MS: DE

Table MRSA1. Overview of countries reporting data on MRSA in animals and food in 2012

Note: For abbreviations of Member States (MS) and other reporting countries see Appendix 7. MRSA: Methicillin-resistant Staphylococcus aureus.

6.2.1. Methicillin-resistant Staphylococcus aureus in food

In 2012, four MSs - Germany, Poland, Slovenia and Spain - reported information on the occurrence of MRSA in various categories of food, as summarised in Table MRSA2. Germany investigated a wide range of meat from turkeys and bovine animals for MRSA. Poland examined samples of raw fish and raw cows' milk for the presence of MRSA. Slovenia investigated 74 samples of meat from pigs, among which 14 samples tested positive for MRSA. Spain examined a range of food products for MRSA and the positive isolates were obtained from fruits (two isolates: 1.8 %), meat from broilers (25 isolates: 12.4 %) and meat from pigs (one isolate: 0.8 %). The corresponding *spa*-typing data were not available from those reporting MSs, positive isolates being reported of unspecified *spa*-type.

Generally, meat from several different sources proved positive for MRSA, including meat from poultry, pigs and cattle. Certain types of poultry meat tested positive for MRSA most frequently, reaching a prevalence greater than 60 % in turkey meat, exceeding that observed in the other kinds of meat tested (beef and pork).



Table MRSA2. MRSA in food, 2012

Food species/ country	Description	Sample unit	Number of units tested	Number (%) positive for MRSA
Meat from broile	rs (Gallus gallus)			
Spain	Meat preparation	Single	201	25 (12.4)
Meat from turke	ys			
Germany	Carcase, at slaughterhouse ¹ , monitoring	Slaughter Batch	353	242 (68.6)
Connaily	Fresh, at retail, monitoring	Single	749	282 (37.7)
Spain	Minced meat/meat products/meat preparation ²	Single	11	0
Meat from pigs				
Slovenia	Fresh, at cutting plant, monitoring	Single	74	14 (18.9)
Spain	Fresh/minced meat/meat products ³	Single	122	1 (0.8)
Meat from bovin	e animals			•
Cormony	Carcase, at slaughterhouse, monitoring	Single	312	96 (30.8)
Germany	At retail, monitoring	Single	421	44 (10.5)
Milk, cows'			-	•
Poland	Raw milk for manufacture-intended for manufacture of Raw or low heat-treated products, monitoring	Single	12	1 (8.3)
Milk, goats'				
Spain	Raw milk for manufacture, at processing plant, Monitoring	Single	5	0
Fish				
Poland	Raw, at processing plant	Single	100	4 (4.0)
Fruits				
Spain	Pre-cut and ready-to-eat, at retail, monitoring	Single	109	2 (1.8)

MRSA: Methicillin-resistant Staphylococcus aureus.

1. Sample type: neck skin.

2. Meat preparation (N=6, n=0); minced meat (N=1, n=0); meat products (N=4, n=0).

3. Fresh (N=60, n=1); minced meat (N=3, n=0); meat products (N=59, n=0).



6.2.2. Methicillin-resistant Staphylococcus aureus in animals

6.2.2.1. Monitoring MRSA in food-producing animals

Belgium, Germany, Finland, the Netherlands and Switzerland reported information on the prevalence of MRSA in food-producing animals and/or their immediate environment within a monitoring, surveillance or unspecified sampling context. The results are summarised in Table MRSA3.

Regarding the monitoring approaches, in Finland, a national prevalence survey was performed in the 68 pig breeding holdings with a specific pathogen-free status through nasal swabbing of 60 randomly selected animals, per holding, and boot swabbing of every unit of the holding between October 2011 and March 2013. In Switzerland, the monitoring of MRSA in pigs was based on random sampling of 397 healthy slaughter pig carcases in the slaughter plants accounting for 85 % of the total pig production in the country. The sampling plan was stratified proportionally to the annual production of the slaughterhouse and evenly distributed over the year to address any seasonal effect.

MRSA prevalence in fattening pigs in Switzerland was comparatively moderate at 18.1 % (72 isolates from 397 samples tested), while in Finland MRSA was not detected in either boot swabs or nasal swabs collected at farms in national monitoring. Of particular note was the extremely high MRSA prevalence recorded in the Netherlands in slaughter pigs (99.0 %) and cattle (79.0 %), both sampled at the slaughterhouse by nasal swabbing in 2012. Germany reported a high prevalence (45.0 %) in calves under one year of age sampled at slaughter, while, considering the same animal population sampled on the farm by means of environmental dust sampling, the prevalence of positive herds was 19.2 %. A flock prevalence of 12.8 % was also recorded in fattening flocks of turkeys in Germany. Belgium also reported a high prevalence (47.1 %) in calves under one year of age sampled at the slaughterhouse, while the prevalences in herds of calves under one year sampled on the farm and in dairy cows were of the same magnitude (10.2 % and 9.9 %, respectively).

A number of different *spa*-types were reported (Table MRSA3, shown as footnotes). The majority of isolates from pigs in Switzerland were *spa*-type t034, with lower numbers of t011; both of these *spa*-types are associated with MRSA CC398 and accounted for 97 % (70 out of 72) of the MRSA isolates from fattening pigs in Switzerland, with the remaining MRSA isolates belonging to *spa*-type t208, which is associated with ST49. Belgium provided *spa*-type data for MRSA isolates from calves less than one year old on the farm and at the slaughterhouse, as well as from adult dairy cattle on farms. *Spa*-type t011 was predominant in samples from calves as well as in adult dairy cows and is associated with ST398. The other *spa*-types reported from Belgian cattle were present in much lower numbers and of these, t1456 and t1451 were both detected in the baseline survey of breeding pigs (EFSA, 2009b) and are associated with CC398 and related to t011. *Spa*-types t1985, t3423 and t6228 are also all associated with MLST ST8, while t037 and t388 are associated with ST239 and both of these MRSA sequence types (ST8 and ST239) are considered to be hospital-associated strains of MRSA. t037 ST239 was also recovered from Belgian poultry in 2011 (Butaye and Nemeghaire, 2012).

Animal species/ country	Production type/description	Sample unit	Number of units tested	Number (%) positive for MRSA
Turkeys				
Germany	Breeding flocks, unspecified, at farm, dust, monitoring	Flock	16	0
-	Meat production flocks, at farm, dust, monitoring	Flock	235	30 (12.8)
Pigs				
Finland	Breeding animals, at farm, boot swabs, survey - national survey	Holding	68	0
Finianu	Breeding animals, at farm, nasal swabs, survey - national survey	Holding	68	0
Netherlands	At slaughterhouse, nasal swabs, monitoring	Herd ¹	104	103 (99.0)
Switzerland ²	Fattening pigs, at slaughterhouse, nasal swabs, monitoring	Animal	397	72 (18.1)
Cattle (bovine a	nimals)			
	Calves (under 1 year), at farm, nasal swabs ³	Herd	187	19 (10.2)
Belgium	Calves (under 1 year), at slaughterhouse, nasal swabs ⁴	Slaughter batch	104	49 (47.1)
	Dairy cows, at farm, nasal swabs, monitoring ⁵	Animal	141	14 (9.9)
	Calves (under 1 year), at farm, dust, monitoring	Herd	240	46 (19.2)
Germany	Calves (under 1 year), at slaughterhouse, nasal swabs, monitoring	Animal	320	144 (45.0)
Netherlands	At slaughterhouse, nasal swabs, monitoring	Herd ¹	100	79 (79.0)
Goats		-		
Netherlands	At farm, monitoring	Animal	221	0
Sheep				-
Netherlands	At farm, monitoring	Animal	467	0

Table MRSA3. MRSA in food-producing animals (excluding clinical investigations), 2012

MRSA: Methicillin-resistant Staphylococcus aureus.

1. 10 animals per herd.

2. 61 and 9 isolates were of the spa-types t034 and t011, respectively, which belonged to CC398 and 2 of the spa-type t208 (ST49).

3. Isolates belonged to the *spa*-types t011 (16), t1456 (1), t121 (1) and t1985 (1).

4. Isolates belonged to the *spa*-types t011 (40), t1456 (1), t1451 (3), t1985 (3), t3423 (1) and one was not typed.

5. Isolates belonged to the *spa*-types t011 (8), t1456 (1), t6228 (2), t037 (1), t388 (1) and one was not typed.

6.2.2.2. Clinical investigations for MRSA in food-producing animals

Clinical investigations often differ from monitoring data in food-producing animals or meat in that selective culture methods may not be used, the number of units tested may be low and the sample may involve a biased sample population. These data are not prevalence data and cannot be extrapolated at the population/group level. However, the results were nevertheless presented in this report, because it is considered important to report the range of animals/animal populations which can be affected.

In 2012, three MSs (Hungary, the Netherlands and Slovakia) reported information on results of clinical investigations for MRSA in different kinds of food-producing animals, which tested, most frequently, negative (Table MRSA4). Only the Netherlands reported six positive for MRSA (out of 11 samples tested) in a clinical investigation context in pigs.

Animal species/ country	Production type/description	Sample unit	Number of units tested	Number (%) positive for MRSA	
Gallus gallus (fo	wi)				
Hungary	At farm	Flock	24	0	
Slovakia	Broilers day-old chicks, at farm, organ/tissue	animal	3	0	
Turkeys					
Hungary	At farm	Flock	7	0	
Geese					
Hungary	At farm	Flock	13	0	
Pigs					
Hungary	At farm	Holding	11	0	
Netherlands	-	Animal	11	6 (54.5)	
Slovakia	Fattening pigs, at farm	Animal	7	0	
Cattle (bovine a	nimals)	-			
Hungary	At farm	Herd	39	0	
Netherlands	_	Animal	11	0	
Slovakia	Calves (under 1 year), at farm	Animal	3	0	
SIOVAKIA	Dairy cows, at farm, milk	Animal	130	0	
Goats					
Hungary	At farm	Herd	2	0	
Netherlands	_	Animal	14	0	
Slovakia	Animals over 1 year, at farm, milk	Animal	6	0	
Sheep					
Hungary	At farm	Herd	2	0	
Netherlands	_	Animal	1	0	
Slovakia	Milk ewes, at farm, milk	Animal	2	0	

Table MRSA4. MRSA in food-producing animals, clinical investigations, 2012

MRSA: Methicillin-resistant Staphylococcus aureus.

6.2.2.3. Clinical investigations for MRSA in companion animals

Three MSs reported data on MRSA in companion animals compared with one in 2011 (Table MRSA5). Ireland, the Netherlands and Slovakia reported MRSA data from samples taken from pets and horses. In all of these cases, the bacteria were isolated from clinical specimens sent for routine bacteriology. MRSA was confirmed in 14 horses, 10 dogs and 3 cats in the Netherlands and in 1 dog in Ireland, in 2012. The corresponding *spa*-typing data were not available.



Animal species/ country	Production type/description	Sample unit	Number of units tested	Number (%) positive for MRSA				
Cats								
Netherlands	Pet animals	Animal	32	3 (9.4)				
Slovakia	Pet animals	Animal	8	0				
Dogs								
Ireland	At farm	Animal	92	1 (1.1)				
Netherlands	Pet animals	Animal	63	10 (15.9)				
	Pet animals, faeces	Animal	5	0				
Slovakia	Pet animals, organ/tissue	Animal	2	0				
	Pet animals	Animal	133	0				
Solipeds, dome	Solipeds, domestic							
Netherlands	Horses	Animal	48	14 (29.2)				

Table MRSA5. MRSA in companion animals, clinical investigations, 2012

MRSA: Methicillin-resistant Staphylococcus aureus.

6.2.2.4. Temporal trends in the occurrence of MRSA

Although methodological differences may occur between reporting countries, where repeat studies were performed in countries the same methods were usually employed. This was, for example, the case in Switzerland where a pre-enrichment in Mueller-Hinton broth, supplemented with 6.5 % salt, followed by culture through selective broth, containing cefoxitin and aztreonam, and final plating onto an MRSA-selective agar, were typically performed.

Germany reported annual results on the occurrence of MRSA in calves, at the herd/farm level, in 2010 and 2012 and, in both years, similar moderate levels of prevalence were registered at 19.6 % (of 296 samples tested) and 19.2 % (of 240 samples tested), respectively (Table MRSA6). Germany also reported results on the occurrence of MRSA in fattening turkeys for meat production, at the flock level, in 2010 and 2012, and the prevalence reported was moderate in both years (19.6 % in 2010 and 12.8 % in 2012). No data on the genotypes of the strains of MRSA isolated were reported to EFSA.

The Netherlands also monitored, consistently, the prevalence of MRSA in goats and sheep over the period 2011 to 2012. Interestingly, out of around 200 samples tested in goats and the 450 samples tested in sheep in 2012, none of them tested positive.

Switzerland reported results on the yearly prevalence of MRSA in fattening pigs over the period 2009 to 2012. Prevalence had significantly increased in 2012 compared with the previous years, when it was low: from 2.2 % in 2009, the prevalence increased threefold in 2010 and 2011 (5.9 % and 5.6 %, respectively) and reached 18.1 % in 2012. The marked increase is primarily the result of the diffusion of clones of *spa*-types t034 and t011, both belonging to the clonal complex CC398, within the Swiss population of fattening pigs. Switzerland also noted the continuing presence of ST49 t208 at a much lower frequency in pigs in 2012. Switzerland was the first country to describe the presence of this previously undescribed clonal lineage of MRSA in pigs (Overesch et al., 2011), suggesting that selection may have occurred within the Swiss pig population. Evidence to support the fact that MRSA ST49 t208 may have emerged in pigs in Switzerland includes the observation that methicillin-susceptible *S. aureus* (MSSA), belonging to *spa*-type t208, had previously been described in Switzerland in pigs. However, occurrence data for the years 2011 and 2012 suggest that ST49 strains did not show a spread, similar to CC398 strains, in the sampled population over the same period. This *spa*-type is otherwise rarely recorded in Europe, having been described in only one human infection in the United Kingdom and in three cases of skin infection and laryngeal ulceration in wild squirrels (Overesch et al., 2011).

Country	Year	Production type/description	Sample unit	Number of units tested	Number (%) positive for MRSA
Germany	2010	Cattle (bovine animals), calves (under 1 year), at farm, monitoring	Herd	296	58 (19.6)
Germany	2012	Cattle (bovine animals), calves (under 1 year), at farm, monitoring	Herd	240	46 (19.2)
Cormony	2010	Turkeys, meat production flocks, at farm, dust, monitoring	Flock	112	22 (19.6)
Germany	2012	Turkeys, meat production flocks, at farm, dust, monitoring	Flock	235	30 (12.8)
Netherlands	2011	Goats, at farm, monitoring	Animal	214	0
Nethenanus	2012	Goats, at farm, monitoring	Animal	221	0
Netherlands	2011	Sheep, at farm, monitoring	Animal	564	0
Nethenanus	2012	Sheep, at farm, monitoring	Animal	467	0
	2009	Fattening pigs, at slaughter, nasal swabs	Animal	405	8 (2.2) ¹
Switzerland	2010	Fattening pigs, at slaughter, nasal swabs	Animal	392	23 (5.9) ²
Switzenand	2011	Fattening pigs, at slaughterhouse, nasal swabs, monitoring	Animal	392	22 (5.6) ³
	2012	Fattening pigs, at slaughterhouse, nasal swabs, monitoring	Animal	397	72 (18.1) ⁴

Table MRSA6. Temporal occurrence of MRSA in animals

MRSA: Methicillin-resistant *Staphylococcus aureus*.

1. In 2009, isolates were reported as unspecified genotypes.

2. In 2010, 17 isolates were of genotype ST398-t034-V, one was of genotype ST398-t011-V and five were of genotype ST49-t208-V.

- 3. In 2011, 19 isolates were of genotype ST398-t034-V, one was of genotype ST398-t011-V, one was of genotype ST49-t208-V and one was of genotype ST1-t2279-IVc.
- 4. In 2012, 61 isolates belonged to genotype CC398-t034, nine belonged to genotype CC398-t011 and two belonged to genotype ST49-t208.

6.2.3. Susceptibility testing of methicillin-resistant *Staphylococcus aureus* isolates

In 2012, data on the susceptibility of MRSA and *S. aureus* isolates were reported only by Belgium and Switzerland. Both countries used a broth dilution method and EUCAST ECOFFs to determine the susceptibility of isolates to cefotoxin, chloramphenicol, ciprofloxacin, clindamycin, erythromycin, fusidic acid, gentamicin, kanamycin, linezolid, mupirocin, quinupristin/dalfopristin, sulfamethoxazole, tetracyclines, tiamulin and vancomycin. All of the 78 MRSA strains isolated from cattle in Belgium and the 72 MRSA isolates from pigs in Switzerland were resistant to cefoxitin, as expected (data not shown).

6.2.3.1. MRSA isolates from cattle

Considering the susceptibility of MRSA isolates from cattle reported by Belgium, 3 isolates belonged to *spa*-type t1456 (CC398), 3 isolates belonged to *spa*-type t1451 (CC398) and 64 isolates belonged to *spa*-type t011 (CC398). For the remaining eight isolates tested in 2012, no *spa*-typing data were available.

Among MRSA isolates (N=78) from different types of cattle (dairy cows, meat production animals, young cattle (one to two years)) in Belgium tested in 2012, with the exception of resistance to vancomycin, resistance was detected for all antimicrobials tested (Table MRSA7). Resistance was reported at extremely high levels for tetracycline (96.2 % resistant), clindamycin and erythromycin (88.5 % resistant), kanamycin (82.1 % resistant), gentamicin (78.2 % resistant); at high levels for ciprofloxacin (42.3 % resistant), fusidic acid (28.2 % resistant), quinupristin/dalfopristin and sulfamethoxazole (24.4 % resistant); at moderate levels for tiamulin (17.9 % resistant), chloramphenicol (12.8 % resistant) and mupirocin (11.5 % resistant); and at low levels for linezolid (1.3 % resistant).



6.2.3.2. MRSA isolates from fattening pigs

Considering the susceptibility of MRSA isolates from fattening pigs reported by Switzerland, 2 isolates belonged to the genotype *spa*-type t208 (ST49), 9 isolates belonged to *spa*-type t011 (CC398) and 61 isolates belonged to *spa*-type t034 (CC398). Thirty-four isolates belonging to the most commonly detected genotype CC 398-t034 shared an identical resistance profile. These showed resistance to beta-lactams, tetracyclines, macrolides, lincosamides, trimethoprim, pleuromutilins, streptomycin and quinupristin/dalfopristin. Twenty-one additional isolates were resistant to all these antimicrobials except for streptomycin, whereas two isolates had additional resistance to all tested aminoglycosides.

Among MRSA isolates (N=72) from pigs in Switzerland, no resistance was detected to chloramphenicol, fusidic acid, linezolid and vancomycin (Table MRSA7). Resistance was reported at extremely high levels to tetracycline (100 % resistant), clindamycin and erythromycin (90.3 % resistant), tiamulin (86.1 % resistant) and quinupristin/dalfopristin (84.7 % resistant), and at low levels for gentamicin and kanamycin (6.9 % resistant), ciprofloxacin (4.2 % resistant), sulfamethoxazole (2.8 % resistant) and mupirocin (1.4 % resistant).

Thirty-four isolates, belonging to the most commonly detected genotype CC 398-t034, in fattening pigs in Switzerland shared an identical resistance profile exhibiting resistance to beta-lactams, tetracyclines, macrolides, lincosamides, trimethoprim, pleuromutilins, streptomycin and quinupristin/dalfopristin. Genes conferring multiple antibiotic resistance have been detected in MRSA ST398 isolates, such as vga genes conferring resistance to pleuromutilins, streptogramin A and lincosamides (Hauschild et al., 2012) and the cfr gene providing resistance to pleuromutilins, streptogramin A, lincosamides, phenicols and oxazolidinones (Kehrenberg et al., 2009). Only a single MRSA isolate from Belgian cattle was resistant to linezolid, suggesting that cfr genes, if present, were rare in this population. The genes vga(A) and vga(C) confer resistance to the lincosamide clindamycin but not to the macrolide erythromycin (Kadlec et al., 2010); however, resistance to the lincosamide clindamycin and the macrolide erythromycin occurred at the same level in both Swiss pigs and Belgian cattle, perhaps suggesting that erm genes may be responsible and that they are constitutively expressed. Indeed, erm genes were frequently detected in bovine MRSA CC398 isolates in a recent Belgian study (Vandendriessche et al., 2013). Tetracycline resistance is common in MRSA CC398 (De Neeling et al., 2007). Considering the aminoglycosides gentamicin and kanamycin, there are differences in the occurrence of resistance in Swiss pigs and Belgian cattle, which may reflect the exposure of these different animal species to antimicrobials. Resistance to ciprofloxacin and fusidic acid can arise by mutation and again differences in the occurrence of resistance are evident in cattle and pigs in the different countries. Studies in Belgium have also detected differences in the occurrence of resistance in MRSA isolates from veal calves and pigs, with the levels of resistance proving higher in veal calves (Vandendriessche et al., 2013). Vancomycin is one of the antimicrobials of last resort for treating S. aureus infections in humans and resistance to this antimicrobial is currently infrequent. None of the isolates from cattle and pigs, tested for susceptibility, was resistant to vancomycin.

Table MRSA7. Resistance (%) to chloramphenicol, ciprofloxacin, clindamycin, erythromycin, fusidic acid, gentamicin, kanamycin, linezolid, mupirocin, quinupristin/dalfopristin, sulfamethoxazole, tetracyclines, tiamulin and vancomycin among MRSA from food and animals in countries reporting MIC data in 2012, using harmonised epidemiological cut-off values

Country	Chloramphenicol		Ciprofloxacin		Clindamycin		Erythromycin		Fusidic acid	
	N	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res
Cattle (bovine animals)										
Belgium	78	12.8	78	42.3	78	88.5	78	88.5	78	28.2
Pigs										
Switzerland	72	0	72	4.2	72	90.3	72	90.3	72	0

Country	Gentamicin		Kanamycin		Linezolid		Mupirocin		Quinupristin/ Dalfopristin	
	Ν	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res
Cattle (bovine animals)										
Belgium	78	78.2	78	82.1	78	1.3	78	11.5	78	24.4
Pigs										
Switzerland	72	6.9	72	6.9	72	0	72	1.4	72	84.7

Country	Sulfamet	hoxazole	Tetrac	yclines	Tian	nulin	Vancomycin		
Country	N	% Res	Ν	% Res	N	% Res	N	% Res	
Cattle (bovine animals)									
Belgium	78	24.4	78	96.2	78	17.9	78	0	
Pigs									
Switzerland	72	2.8	72	100	72	86.1	72	0	

MRSA: Methicillin-resistant Staphylococcus aureus; MIC: minimum inhibitory concentration; N: number of isolates tested; % Res: percentage of resistant isolates.

Note: All MRSA isolates tested were also resistant to cefoxitin, as expected.



6.3. Discussion

Although food is not currently considered to be a relevant source of MRSA infection or colonisation of humans (EFSA, 2009c), the monitoring of MRSA in various food products performed in several MSs consistently indicates that MRSA can be detected, quite frequently, in different types of food. In situations where livestock are colonised by MRSA, meat or raw milk produced from these animals may also on occasion be contaminated with MRSA, as generally shown by the monitoring results. The detection of MRSA in low numbers of samples of raw fish (Poland) and fruit (Spain) may also possibly indicate contamination from colonised workers preparing such food. Nevertheless, it is of note that the laboratory techniques used to detect MRSA employ selective bacterial culture and, therefore, low levels of contamination can be detected. In each case, molecular typing would be very useful in investigating the strains of MRSA involved, which might assist in interpreting the findings and unravelling the epidemiology.

Where data are available in classes of animals tested both on the farm and at slaughter, comparison of the proportion of MRSA-positive animals generally reveals a higher prevalence when animals are tested at slaughter compared with animals tested on farms. This may reflect either cross-colonisation of animals during transport to abattoirs (or while in temporary lairage pens at the slaughterhouse) or acquisition of the organism from various sources encountered during transport and lairage (pens, human contact, vehicles and so forth). It is therefore of note that the high prevalence of MRSA in pigs (99.0 %) and in cattle (79.0 %) in the Netherlands was assessed at slaughter.

Livestock-associated (LA)-MRSA is considered a poor coloniser of humans and occurs uncommonly in persons without contact with livestock (Graveland et al., 2010). However, considering those people who do have contact with livestock, recent research has shown that livestock veterinarians can carry MRSA CC398 for prolonged periods (Verkade et al., 2013), while studies of pig farmers and their household members (Garcia-Graells et al., 2013) showed that the risk of acquisition of MRSA by household members was strongly dependent on their exposure to pigs on the farm. Similar findings have been reported in relation to fattening turkeys and persons living on fattening turkey farms in Germany, where people with frequent access to the turkey accommodation were found to be more likely to carry MRSA (Richter et al., 2012). An increased risk of MRSA carriage has also been reported in personnel working at broiler slaughterhouses, especially where personnel have direct contact with live birds (Mulders et al., 2010). The technique of wholegenome mapping has recently been used in the Netherlands to show that veterinarians can carry and transmit different LA-MRSA strains at the same time (Bosch et al., 2013). LA-MRSA is not generally considered to spread via food and the selective techniques used for isolation mean that low levels of contamination can be identified. The public health risk posed by LA-MRSA has been assessed to be of minor importance as long as the occurrence of strains with acquired toxin genes remains very uncommon (Vandendriessche et al., 2013).

Two MSs reported *spa*-type data for MRSA isolates from food-producing animals and the results from Belgium for cattle were interesting in that *spa*-types t121 (associated with MLST ST8) and t037 and t388 (associated with ST239) are considered to be hospital-associated strains of MRSA. t037 ST239 was also recovered from Belgian poultry in 2011 (Butaye and Nemeghaire, 2012). The occurrence of ST239 in different food-producing animals in different years indicates that this is a further strain of MRSA able to colonise different animal species and reinforces the value of such ongoing surveillance.

EFSA TEHNICAL SPECIFICATION ON HARMONISED MONITORING OF MRSA

In 2012, EFSA published a Scientific Report describing technical specifications for the harmonised monitoring and reporting of antimicrobial resistance in methicillin-susceptible Staphylococcus aureus (MRSA) in food-producing animals and food (EFSA, 2012c). The technical specifications should enable collection of harmonised data from Member States on both the degree on which food-producing animals (and food produced from them) are colonised with MRSA, and the strains of MRSA involved. A new definition of MRSA includes those strains harbouring the mecC gene and the laboratory methods adapted accordingly, so that those strains can be also targeted by the harmonised routine monitoring. The situation with regard to MRSA and some food-producing animal species has changed substantially over the last decade; therefore, the proposed monitoring aims to provide a means to detect, without undue delay, further developments which may occur, in particular regarding the possible emergence of MRSA strains displaying particular virulence or resistance patterns and/or their potential transfer and diffusion between human and animal populations. Ongoing evolution and development of the situation relating to MRSA in food-producing animals may be exemplified by the recent description of MRSA ST49 in pigs in Switzerland (Overesch et al.. 2011).



7. THIRD-GENERATION CEPHALOSPORIN RESISTANCE IN ESCHERICHIA COLI AND SALMONELLA

7.1. Introduction

RESISTANCE TO THIRD-GENERATION CEPHALOSPORINS: THE IMPORTANCE OF ESBLS AND AMPC ENZYMES

Extended-spectrum beta-lactamases (ESBLs) are considered to be an important emerging issue in Gram-negative bacteria of public health significance. Bacteria which possess ESBL resistance are usually resistant to third-generation cephalosporins, which are critically important antibiotic drugs for the treatment of systemic or invasive Gram-negative bacterial infections in humans. These drugs play a critical role in the treatment of certain invasive Salmonella infections, particularly in children, in whom the use of fluoroquinolones may not be favoured because of certain potential adverse effects. A low level of resistance in Salmonella may therefore still constitute an important finding. Commensal bacteria, such as indicator Escherichia coli, may contribute to the dissemination of ESBL resistance because such resistance is usually transferable.

Salmonella and E. coli may become resistant to third-generation cephalosporins by several different mechanisms. Among these different mechanisms, the most common is the acquisition of beta-lactamase enzymes on plasmids (small covalently closed circles of DNA which can be transferred between bacteria during bacterial conjugation). There are several different types of beta-lactamase which can confer resistance to third-generation cephalosporins. These are conveniently sub-divided into four classes, designated A to D: ESBL enzymes of the TEM, SHV and CTX-M families belong to class A, while class C includes the AmpC beta-lactamases.

Wild-type Salmonella isolates never possess a beta-lactamase of any class. For beta-lactamases to occur in Salmonella, acquisition must generally have occurred by conjugation, usually with other Enterobacteriaceae through transfer of plasmids. Although all four different types of beta-lactamase classes have been described in Salmonella globally, within the EU, the most important types of beta-lactamase resistance acquired by Salmonella are primarily ESBL resistance and, secondly, AmpC resistance. E. coli can acquire beta-lactamases from other bacteria in a similar fashion to Salmonella but, since it also possesses an endogenous AmpC beta-lactamase, in some circumstances this can be activated, conferring resistance to third-generation cephalosporins.

The position has been further complicated in recent years by the emergence of resistance to carbapenems in human medicine. Carbapenems are used for the treatment of highly resistant infections in humans, including, for example, the treatment of infections with Gram-negative bacteria which possess ESBL enzymes. These compounds are not used in food-producing animals anywhere within the EU. Resistance to carbapenems in Gram-negative bacteria is usually related to the acquisition of carbapenemase enzymes and a number of different types are recognised. Although carbapenem antimicrobials are not used in food-producing animals in the EU, resistance has occasionally been detected in bacteria carried by animals (Woodford et al., 2013) and dissemination from humans to animals directly or through environmental routes is suspected. In view of the great importance of the carbapenem compounds, they have been added to the panels of antimicrobials recommended for testing by Member States to improve surveillance for resistance (EFSA, 2012b).

The EFSA guidelines for monitoring resistance in indicator *E. coli* (EFSA, 2008) state that cefotaxime is a good substrate for what are currently the most common and important ESBLs in humans in Europe, the CTX-M enzymes, which can therefore be used as an indicator for ESBL resistance. ECOFFs for *Salmonella* and *E. coli* for the antimicrobial cefotaxime facilitate detection of CTX-M ESBLs, but resistance to cefotaxime may, of course, be conferred by mechanisms of resistance other than ESBLs, such as certain other types of beta-lactamases, including AmpC beta-lactamases. In this chapter, the occurrence of resistance is given, where available, for both cefotaxime and ceftazidime. As very few MSs reported data on resistance to ceftiofur, and because this compound is not considered optimal for the detection of ESBL enzymes, results for ceftiofur are not included in this chapter. Furthermore, because this report covers only phenotypic monitoring, it is not possible to determine the class or exact type of beta-lactamase enzyme which is responsible for conferring the resistance detected to third-generation cephalosporins.

The monitoring reported here and performed in accordance with EFSA's guidelines (EFSA, 2008), does not utilise selective primary isolation media containing cephalosporins so the results generally relate to organisms chosen effectively at random from primary culture media. In certain types of monitoring, selective media containing cephalosporins may be used to investigate the presence or absence of cephalosporin-resistant organisms in a particular sample (within the limit of detection) and, in that case, a different type of result would be obtained from such monitoring, which has a greater sensitivity. Ideally, the establishment of optimum phenotypic testing systems for sensitive, specific and rapid detection of ESBLs would be a very important component of antimicrobial resistance monitoring programmes. Recommendations for such monitoring recently developed by EFSA (EFSA, 2012b) notably put forward further testing of isolates which are resistant to third-generation cephalosporins, including testing to establish whether isolates have an ESBL- or AmpC-producing phenotype.

7.2. Third-generation cephalosporin resistance in *Salmonella* isolates from animals and food

7.2.1. Third-generation cephalosporin resistance in Salmonella isolates from food

In 2012, the results of testing for resistance to third-generation cephalosporins in *Salmonella* spp. isolates recovered from meat from broilers, meat from pigs and meat from bovine animals were reported by 11, 12 and 4 MSs, respectively (Table ESBL1). In most reporting MSs, resistance was either not detected or reported at low levels in the three kinds of meat. Resistance to cefotaxime was typically equal or similar to that observed to ceftazidime at the MS level. Considering all MSs, the apparent difference in resistance to each compound in meat from broilers largely reflects differences in the number of sensitive isolates contributing to the denominator.



Table ESBL1. Resistance (%) to cefotaxime and ceftazidime in Salmonella spp. isolates from meat from broilers, pigs and bovine animals tested by MSs in 2012

0	Cefc	otaxime	Ceftazidime		
Country	N	% Res	N	% Res	
Meat from broilers			<u>-</u>	•	
Belgium	109	0	109	0	
Czech Republic	47	0	47	0	
Germany	94	12.8	94	11.7	
Hungary	168	0.6	_	_	
Ireland	70	2.9	70	2.9	
Latvia	32	0	_	_	
Netherlands	74	31.1	74	29.7	
Poland	93	0	93	0	
Portugal	37	0	_	_	
Romania	189	1.1	_	_	
Slovakia	14	0	14	0	
Total (11 and 7 MSs)	927	4.3	501	7.0	
Meat from pigs		-	Ł		
Belgium	262	1.5	262	1.5	
Czech Republic	33	3.0	33	3.0	
Denmark	41	0	_	_	
Estonia	22	0	22	0	
Germany	163	0.6	163	0.6	
Hungary	16	0	_	_	
Ireland	69	0	69	0	
Italy	85	0	85	0	
Latvia	13	0	_	_	
Netherlands	52	0	52	0	
Poland	22	0	20	0	
Romania	125	1.6	_	_	
Total (12 and 8 MSs)	903	0.9	706	0.8	
Meat from bovine animals			•		
Germany	16	0	16	0	
Ireland	24	0	24	0	
Italy	13	0	13	0	
Netherlands	18	5.6	18	5.6	
Total (4 MSs)	71	1.4	71	1.4	

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.



Similar observations of resistance to cefotaxime or ceftazidime generally not detected or reported at low levels may be made in *S*. Enteritidis from meat from broilers (Table ESBL2) and in *S*. Typhimurium isolates and monophasic *S*. Typhimurium isolates from meat from pigs (Tables ESBL3 and ESBL4). Exceptions to this are resistance to cefotaxime and ceftazidime in *Salmonella* spp. isolates from broiler meat recorded at levels greater than 10 % in Germany and around 30 % in the Netherlands (Table ESBL1).

Table ESBL2. Resistance (%) to cefotaxime and ceftazidime in S. Enteritidis isolates from meat from broilers tested by MSs in 2012

Country	Ce	efotaxime Ceftazidime		eftazidime
	N	% Res	N	% Res
Belgium	31	0	31	0
Latvia	23	0	-	-
Poland	26	0	26	0
Total (3 and 2 MSs)	80	0	57	0

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Table ESBL3. Resistance (%) to cefotaxime and ceftazidime in S. Typhimurium isolates from meat from pigs tested by MSs in 2012

Country	Cefo	Cefotaxime		zidime
	N	% Res	N	% Res
Belgium	105	1.0	105	1.0
Denmark	18	0	_	-
Germany	58	0	58	0
Hungary	10	0	_	-
Ireland	22	0	22	0
Italy	18	0	18	0
Latvia	13	0	-	-
Netherlands	16	0	16	0
Poland	11	0	-	-
Romania	43	2.3	_	-
Total (10 and 5 MSs)	314	0.6	219	0.5

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Table ESBL4. Resistance (%) to cefotaxime and ceftazidime in monophasic S. Typhimurium isolates from meat from pigs tested by MSs in 2012

Country	Ce	Cefotaxime Cefta		ftazidime
	N	% Res	N	% Res
Denmark	22	0	-	-
Germany	37	0	37	0
Ireland	23	0	23	0
Total (3 and 2 MSs)	82	0	60	0

7.2.2. Third-generation cephalosporin resistance in *Salmonella* isolates from animals

7.2.2.1. Resistance levels in Gallus gallus (fowl)

Resistance to third-generation cephalosporins in *Salmonella* spp. from *Gallus gallus* is shown in Table ESBL5. A low level of resistance to cefotaxime of 4.5 %, and to ceftazidime of 4.8 %, was reported in *Salmonella* spp. isolates from all reporting MSs, reflecting either no or very low to low resistance recorded in nearly all reporting countries. Only Belgium recorded much higher levels of resistance at 18.1 % to cefotaxime and 17.3 % to ceftazidime.

Table ESBL5. Resistance (%) to cefotaxime and ceftazidime in Salmonella spp. isolates fromGallus gallus tested by MSs in 2012

Country	Cefota	axime	Ceftaz	idime
Country	N	% Res	N	% Res
Austria	176	0.6	176	0.6
Belgium	664	18.1	664	17.3
Czech Republic	386	0.3	386	0.3
Denmark	28	0	-	-
Germany	238	0.8	238	0.8
Hungary	261	0.4	-	-
Ireland	38	0	38	0
Italy	328	5.2	328	3.4
Latvia	14	0	-	-
Netherlands	192	4.2	192	4.2
Poland	739	0.7	739	0.8
Portugal	174	2.3	-	-
Romania	964	5.3	964	5.7
Slovakia	85	0	85	0
Spain	179	0.6	179	0.6
United Kingdom	236	0	216	0
Total (16 and 12 MSs)	4,702	4.5	4,205	4.8

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Considering differing populations of *Gallus gallus* separately, levels of resistance to third-generation cephalosporins in *Salmonella* ssp. isolates from broiler, laying hen and breeding flocks were reported by 13, 12 and 4 MSs, respectively, in 2012 (Table ESBL6) and were generally either not detected or recorded at low levels. The levels of resistance in broilers were generally slightly higher than those reported when all *Gallus gallus* were considered, as most reporting MSs detected resistance, while, in laying hen and breeding flocks, resistance was generally not recorded. In *Salmonella* spp. from laying hen and breeding flocks, only three (Austria, Italy and Romania) and two MSs (Italy and Romania) detected low level resistance to third-generation cephalosporins out of the 12 and 4 reporting MSs, respectively. In broilers, three MSs reported levels of resistance to third-generation cephalosporins greater than 5 %.

Table ESBL6. Resistance (%) to cefotaxime and ceftazidime in Salmonella spp. isolates from broilers, laying hens and breeding hens of Gallus gallus tested by MSs in 2012

Country	Cefotaxime		Ceftazi	dime
Country	Ν	% Res	N	% Res
Broilers				-
Austria	113	0	113	0
Czech Republic	351	0.3	351	0.3
Denmark	24	0	_	_
Hungary	175	0.6	_	_
Ireland	38	0	38	0
Italy	105	13.3	105	7.6
Netherlands	130	6.2	130	6.2
Poland	189	0.5	189	0.5
Portugal	122	3.3	_	_
Romania	784	6.4	784	6.3
Slovakia	55	0	55	0
Spain	29	3.4	29	3.4
United Kingdom	170	0	153	0
Total (13 and 10 MSs)	2,285	3.5	1,947	3.5
Laying hens				-
Austria	63	1.6	63	1.6
Germany	51	0	51	0
Hungary	86	0	_	_
Italy	161	1.2	161	1.2
Latvia	14	0	_	_
Netherlands	54	0	54	0
Poland	132	0	132	0
Portugal	32	0	_	_
Romania	145	0	145	2.8
Slovakia	29	0	29	0
Spain	150	0	150	0
United Kingdom	66	0	63	0
Total (12 and 9 MSs)	983	0.3	848	0.8
Breeding flocks				
Czech Republic	27	0	27	0
Italy	36	2.8	36	2.8
Poland	15	0	15	0
Romania	32	3.1	32	6.3
Total (4 MSs)	110	1.8	110	2.7



The resistance to cefotaxime and ceftazidime in *S*. Enteritidis isolates from *Gallus gallus*, broilers and laying hens, reported, respectively, by 13, 6 and 10 MSs (Table ESBL7), was most generally not detected in reporting MSs. In contrast, resistance to both cefotaxime and ceftazidime was reported at moderate levels (14.8 %) in Belgium. Among the reporting MSs on broilers, Portugal observed resistance to cefotaxime, at the low level of 5.3 %. Considering isolates from laying hens, Romania was the only MS to report resistance to ceftazidime at a level of 6.1 %, while resistance to cefotaxime was not detected in this country.

Table ESBL7. Resistance (%) to cefotaxime and ceftazidime in S. Enteritidis isolates from Gallus gallus tested by MSs in 2012

	Cefot	axime	Ceftaz	zidime
Country	N	% Res	N	% Res
All Gallus gallus			•	-
Austria	36	0	36	0
Belgium	81	14.8	81	14.8
Czech Republic	251	0	251	0
Germany	80	0	80	0
Hungary	26	0	-	_
Italy	31	0	31	0
Latvia	11	0	-	_
Netherlands	38	0	38	0
Poland	496	0.6	496	0.8
Portugal	27	3.7	-	_
Romania	76	0	76	5.3
Slovakia	47	0	47	0
Spain	45	0	45	0
Total (13 and 10 MSs)	1,245	1.3	1,181	1.7
Broilers				
Austria	21	0	21	0
Czech Republic	236	0	236	0
Poland	131	0	131	0
Portugal	19	5.3	-	_
Romania	10	0	10	0
Slovakia	21	0	21	0
Total (6 and 5 MSs)	438	0.2	419	0
Laying hens				
Austria	15	0	15	0
Germany	21	0	21	0
Hungary	25	0	-	_
Italy	28	0	28	0
Latvia	11	0	-	-
Netherlands	38	0	38	0
Poland	91	0	91	0
Romania	66	0	66	6.1
Slovakia	25	0	25	0
Spain	43	0	43	0
Total (10 and 8 MSs)	363	0	327	1.2

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Resistance to third-generation cephalosporins in *S.* Typhimurium isolates from *Gallus gallus* (Table ESBL8) was reported by five MSs. Belgium was the only MS to detect resistance at levels of 10.6 % to cefotaxime and 14.9 % to ceftazidime.

Table ESBL8. Resistance (%) to cefotaxime and ceftazidime in S. Typhimurium isolates from Gallus gallus tested by MSs in 2012

Country	Cefo	Cefotaxime		azidime
	N	% Res	N	% Res
Austria	10	0	10	0
Belgium	47	10.6	47	14.9
Germany	42	0	42	0
Hungary	10	0	-	_
Poland	15	0	15	0
Total (5 and 4 MSs)	124	4.0	114	6.1

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

7.2.2.2. Resistance levels in turkeys

Resistance to cefotaxime and ceftazidime in *Salmonella* spp. isolates from turkeys, mainly fattening turkeys, is shown in Table ESBL9. Nine MSs reported results for cefotaxime and eight MSs reported results for ceftazidime. Resistance to both cefotaxime and ceftazidime was reported by Italy and Poland at low proportions, whereas Spain reported 1.2 % resistance to cefotaxime only.

Table ESBL9. Resistance (%) to cefotaxime and ceftazidime in Salmonella spp. isolates from turkeys tested by MSs in 2012

Country	Cefc	otaxime	Cefta	azidime
Country	N	% Res	Ν	% Res
Turkeys				
Austria	38	0	38	0
Czech Republic	27	0	27	0
Germany	87	0	87	0
Hungary	174	0	-	-
Ireland	14	0	14	0
Italy	48	4.2	48	4.2
Poland	55	3.6	55	3.6
Spain	169	1.2	169	0
United Kingdom	142	0	141	0
Total (9 and 8 MSs)	754	0.8	579	0.7
Fattening turkeys				
Austria	38	0	38	0
Czech Republic	20	0	20	0
Germany	12	0	12	0
Hungary	174	0	-	-
Spain	169	1.2	169	0
Total (5 and 4 MSs)	413	0.5	239	0



7.2.2.3. Resistance levels in pigs

Resistance to cefotaxime and ceftazidime in *Salmonella* spp. isolates from pigs is shown in Table ESBL10. Ten MSs reported results for cefotaxime and eight MSs reported results for ceftazidime; however, most of them did not detect any resistance making the overall level of resistance at the MS group level in pigs low at 2.3 % for cefotaxime and 2.6 % for ceftazidime. Three MSs detected both cefotaxime and ceftazidime resistance in *Salmonella* spp. isolates from pigs. In fattening pigs, cefotaxime and ceftazidime resistance was observed only by Spain (6.3 % and 2.1 %, respectively) while, in breeding pigs, Belgium recorded cefotaxime and ceftazidime resistance at similar levels of 11.2 % and 9.6 %, respectively. Germany reported low levels of resistance to both cefotaxime and ceftazidime in pigs of unspecified production type.

Table ESBL10. Resistance (%) to cefotaxime and ceftazidime in Salmonella spp. isolates from pigs tested by MSs in 2012

Country	Cefotaxime		Ceftazidime	
Country	N	% Res	N	% Res
All pigs				
Belgium	187	11.2	187	9.6
Denmark	374	0	-	-
Estonia	32	0	32	0
Germany	627	2.1	627	2.1
Hungary	38	0	_	_
Ireland	24	0	24	0
Italy	25	0	25	0
Netherlands	263	0	263	0
Poland	10	0	10	0
Spain	48	6.3	48	2.1
Total (10 and 8 MSs)	1,628	2.3	1,216	2.6
Fattening pigs				
Denmark	374	0	_	_
Estonia	14	0	14	0
Hungary	38	0	_	_
Netherlands	17	0	17	0
Spain	48	6.3	48	2.1
Total (5 and 3 MSs)	491	0.6	79	1.3
Breeding pigs				
Belgium	187	11.2	187	9.6



Results relating to resistance to third-generation cephalosporins in *S*. Typhimurium from pigs, were reported by five and four MSs for cefotaxime and ceftazidime, respectively, and are shown in Table ESBL11. Among reporting MSs, Belgium and Germany were the only countries to report cefotaxime and ceftazidime resistance in *S*. Typhimurium, at the same levels, for both substances, of 10.7 % and 1.1 %, respectively. The overall levels of resistance for all reporting MSs were, therefore, at low levels of 2.3 % for cefotaxime and 2.6 % for ceftazidime.

Table ESBL11. Resistance (%) to cefotaxime and ceftazidime in S. Typhimurium isolates from pigs tested by MSs in 2012

Country	Ce	fotaxime	Cef	tazidime
Country	N	% Res	N	% Res
All pigs	-	-	-	
Belgium	75	10.7	75	10.7
Denmark	63	0	-	-
Germany	273	1.1	273	1.1
Ireland	15	0	15	0
Netherlands	55	0	55	0
Total (5 and 4 MSs)	481	2.3	418	2.6
Fattening pigs	-		-	
Denmark	63	0	-	-
Breeding pigs	-		-	
Belgium	75	10.7	75	10.7

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

The occurrence of resistance to cefotaxime and ceftazidime in monophasic *S*. Typhimurium isolates from pigs is shown in Table ESBL12. Six MSs reported results for cefotaxime and five MSs reported results for ceftazidime. Resistance to third-generation cephalosporins was detected by Belgium at moderate levels for both antimicrobials (19.5 % for cefotaxime and 14.6 % for ceftazidime) and by Germany at a very low level (0.9 % for both antimicrobials), while Spain recorded resistance only to cefotaxime at 7.1 %.

Table ESBL12. Resistance (%) to cefotaxime and ceftazidime in monophasic S. Typhimuriumisolates from pigs tested by MSs in 2012

Country	Cefotaxime		Cefta	azidime
	N	% Res	N	% Res
Belgium	41	19.5	41	14.6
Denmark	81	0	-	-
Germany	228	0.9	228	0.9
Netherlands	39	0	39	0
Poland	10	0	10	0
Spain	14	7.1	14	0
Total (6 and 5 MSs)	413	2.7	332	2.4



7.2.2.4. Resistance levels in cattle

Among the seven MSs reporting data on resistance to cefotaxime and five for ceftazidime in *Salmonella* spp. from cattle (Table ESBL13), only Belgium reported resistance of 2.4 % to both antimicrobials in 2012. None of the *S*. Typhimurium isolates tested exhibited any resistance to these compounds amongst the six and four, respectively, reporting MSs (Table ESBL14).

Table ESBL13. Resistance (%) to cefotaxime and ceftazidime in Salmonella spp. isolates from cattle tested by MSs and non-MS in 2012

Country	Cefotaxime		Ceftazidime	
	N	% Res	Ν	% Res
Belgium	42	2.4	42	2.4
Finland	19	0	-	_
Germany	68	0	68	0
Ireland	36	0	36	0
Italy	14	0	14	0
Netherlands	68	0	68	0
Sweden	17	0	_	_
Total (7 and 5 MSs)	264	0.4	228	0.4

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Table ESBL14. Resistance (%) to cefotaxime and ceftazidime in S. Typhimurium isolates from cattle tested by MSs in 2012

Country	Cefot	axime	Cefta	zidime
Country	N	% Res	N	% Res
Belgium	25	0	25	0
Finland	16	0	-	-
Germany	35	0	35	0
Ireland	24	0	24	0
Netherlands	24	0	24	0
Sweden	12	0	-	-
Total (6 and 4 MSs)	136	0	108	0



7.2.3. Salmonella serovars from animals demonstrating resistance to third-generation cephalosporins

Third-generation cephalosporin resistance was identified in a range of *Salmonella* serovars in 2012. Reporting MSs do not necessarily list all of the *Salmonella* serovars identified, and so the list of affected serovars is likely to be incomplete. In 2010, 2011 and 2012, the following third-generation cephalosporin-resistant serovars were identified from one or more sources (pigs, *Gallus gallus* and/or cattle) and from one or more MSs: *S.* Derby, *S.* Enteritidis, *S.* Infantis, *S.* Kentucky, *S.* Livingstone, *S.* London, *S.* Java and *S.* Typhimurium. In addition to these serovars, *S.* <u>1</u>,9,12:I,v:-, *S.* Choleraesuis, *S.* Lamberhurst, *S.* Montevideo and *S.* Ordonez with third-generation cephalosporin resistance were identified in 2011. Isolates from turkeys (*S.* Bovismorbificans from France, *S.* Bredeney from Hungary and *S.* Muenchen from Spain) and domestic solipeds (*S.* Typhimurium DT104 from Ireland) were also found to express resistance to third-generation cephalosporins, monophasic *S.* Typhimurium was identified in pigs from Belgium, Germany and Spain in 2012 (Table ESBL 12).

7.2.4. Reporting of specific data on ESBL in Salmonella

In 2012, EFSA published a report (EFSA, 2012b) providing detailed recommendations and discussions relating to how future surveillance for third-generation cephalosporin, ESBL, AmpC and carbapenem resistance monitoring could be enhanced. Using for the first time the new functionalities of the EFSA Data Collection Framework recently introduced, one MS (Spain) reported data on the identity of the ESBL and other enzymes detected in *Salmonella* isolates from fattening pigs. Two isolates (of serovars *S*. Derby and monophasic *S*. Typhimurium) were reported producing CTX-M-1 and CTX-M-14, respectively. CTX-M-1 is an ESBL enzyme which has been previously recognised in pigs in several MSs. One *S*. Typhimurium isolate produced the enzyme OXA-1, which confers resistance to the action of clavulanic acid, a beta-lactamase enzyme inhibitor, by breaking down the clavulanate compound.

7.3. Third-generation cephalosporin resistance in indicator E. coli from food and animals

7.3.1. Third-generation cephalosporin resistance in indicator *E. coli* isolates from food

Three MSs (Denmark, Hungary and the Netherlands) reported results for resistance to cefotaxime in *E. coli* isolates from meat from broilers, meat from pigs and meat from bovine animals in 2012, and Sweden and Germany also reported data from meat from broilers and meat from bovine animals, respectively. Germany and the Netherlands tested ceftazidime and recorded similar resistance levels to those obtained for cefotaxime. Overall, resistance to third-generation cephalosporins was either not detected or reported at low levels ranging between 0.5 % and 8.0 %. Interestingly, the 2012 results for broiler meat from the Netherlands (8.0 % for cefotaxime and 6.3 % for ceftazidime) represent a decrease in the figures obtained for 2011, when 31.9 % of isolates were resistant to both antimicrobials.

Table ESBL15. Resistance (%) to cefotaxime and ceftazidime in E. coli isolates from meat from broilers, meat from pigs and meat from bovine animals tested by MSs in 2012

0	Cefo	taxime	Cefta	azidime
Country	N	% Res	Ν	% Res
Meat from broilers		·		-
Denmark	197	1.0	_	_
Hungary	64	0	_	_
Netherlands	175	8.0	175	6.3
Sweden	92	0	_	_
Total (4 and 1 MSs)	528	3.0	175	6.3
Norway	197	0.5	_	_
Meat from pigs		-		
Denmark	73	1.4	_	_
Hungary	14	0	_	_
Netherlands	98	1.0	98	1.0
Total (3 and 1 MSs)	185	1.1	98	1.0
Meat from bovine animals	-	-		
Denmark	46	0	_	_
Germany	71	4.2	71	4.2
Hungary	31	3.2	_	_
Netherlands	141	0.7	141	0.7
Total (4 and 2 MSs)	289	1.7	212	1.9

7.3.2. Third-generation cephalosporin resistance in indicator *E. coli* isolates from animals

7.3.2.1. Resistance levels in Gallus gallus (fowl)

Table ESBL16 summarises data on resistance in indicator *E. coli* isolates from *Gallus gallus* tested by eight reporting MSs, as well as Norway and Switzerland, distinguishing, where possible, between broilers and laying hens. All reporting countries tested resistance to cefotaxime and five reporting MSs also tested isolates for ceftazidime resistance. The levels of resistance reported were generally low, although Belgium reported 28 % resistance in *Gallus gallus*. The level of cefotaxime resistance (5.8 %) in *E. coli* from broilers reported by the Netherlands in 2012 represents a decline compared with 2011 and 2010, when 8.1 % and 18 % resistance, respectively, was reported. In the case of Poland where resistance to cefotaxime and ceftazidime was reported in isolates from both broilers and laying hens, the levels of resistance, when considering isolates from broiler flocks, were approximately twice that reported from laying hens.

Table ESBL16. Resistance (%) to cefotaxime and ceftazidime in indicator E. coli isolates from Gallus gallus tested by MSs and non-MSs in 2012

Country	Cefota	xime	Ceftazidime				
Country	N	% Res	N	% Res			
All Gallus gallus		· · · ·		- <u>-</u>			
Austria	130	3.1	130	3.1			
Belgium	325	28.0	325	24.9			
Denmark	115	1.7	_	_			
France	201	10.4	201	9.5			
Hungary	105	7.6	_	_			
Netherlands	292	5.8	292	6.2			
Poland	328	10.7	328	6.1			
Sweden	255	0.4	_	_			
Total (8 and 5 MSs)	1,751	10.2	1,276	11.1			
Norway	113	0.9	_	_			
Switzerland	185	2.2	185	2.2			
Broilers	÷	· · ·					
Austria	130	3.1	130	3.1			
Denmark	115	1.7	_	_			
France	201	10.4	201	9.5			
Hungary	105	7.6	_	_			
Netherlands	292	5.8	292	6.2			
Poland	171	13.5	171	8.2			
Sweden	194	0	-	_			
Total (7 and 4 MSs)	1,208	6.2	794	6.9			
Switzerland	185	2.2	185	2.2			
Laying hens							
Poland	157	7.6	157	3.8			
Sweden	61	1.6	-	_			
Total (2 and 1 MSs)	218	6.0	157	3.8			



7.3.2.2. Resistance levels in pigs

Table ESBL17 shows resistance to cefotaxime and ceftazidime in indicator *E. coli* from pigs. Overall, the levels of resistance in reporting countries were low generally. Austria and the Netherlands detected no resistance in indicator *E. coli* from pigs.

Table ESBL17. Resistance (%) to cefotaxime and ceftazidime in indicator E. coli isolates from pigs tested by MSs and non-MSs in 2012

Country	Cefota	ixime	Cefta	zidime
Country	N	% Res	N	% Res
Austria	140	0	140	0
Belgium	205	2.9	205	3.4
Denmark	152	0.7	-	_
France	200	2.0	200	2.0
Hungary	68	1.5	-	-
Netherlands	284	0	284	0
Poland	190	2.6	190	2.6
Total (7 and 5 MSs)	1,239	1.4	1,019	1.6
Switzerland	185	1.1	185	1.1



7.3.2.3. Resistance levels in cattle

The results of examinations for third-generation cephalosporin resistance in indicator *E. coli* from cattle are shown in Table ESBL18. Seven MSs and Switzerland tested indicator *E. coli* isolates from cattle for cefotaxime and five MSs and Switzerland tested these isolates for ceftazidime resistance. The overall occurrence of resistance to cefotaxime was 2.4 % and to ceftazidime was 2.6 % in all reporting MSs and this represents an increase on the figures of 0.9 % and 0.6 %, respectively, reported in 2011. Denmark and Finland did not detect cefotaxime resistance in indicator *E. coli* from cattle in 2012, and in the remaining MSs a low or very low level (0.4-9.9 %) of resistance to both antimicrobials was detected.

Table ESBL18. Resistance (%) to cefotaxime and ceftazidime in indicator E. coli isolates tested from cattle by MSs and non-MS in 2012

	Cefota	xime	Ceftazi	dime	
Country	N	% Res	Ν	% Res	
All cattle				-	
Austria	273	0.7	273	0.4	
Belgium	364	8.8	364	9.9	
Denmark	98	0	_	_	
Finland	295	0	_	_	
Germany	515	2.5	515	1.7	
Netherlands	559	0.5	559	0	
Poland	190	2.6	190	2.1	
Total (7 and 5 MSs)	2,294	2.4	1,901	2.6	
Switzerland	187	0.5	187	0	
Calves (under 1 year)					
Austria	151	1.3	151	0.7	
Germany	515	2.5	515	1.7	
Netherlands	285	0.7	285	0	
Total (3 MSs)	951	1.8	951	1.1	
Young cattle (1–2 years)					
Austria	73	0	73	0	
Meat production animals					
Switzerland	187	0.5	187	0	
Adult over 2 years					
Austria	49	0	49	0	
Dairy cows					
Netherlands	274	0.4	274	0	
Mixed herds		· · · · · · · · · · · · · · · · · · ·			
Belgium	364	8.8	364	9.9	
Finland	295	0	_	_	
Total (2 and 1 MSs)	659	4.9	364	9.9	



7.4. Comparison of cefotaxime resistance in *Salmonella* spp. and indicator *E. coli* isolates from animals

Indicator commensal *E. coli* in healthy animals may constitute a reservoir of resistance genes which can be transferred to zoonotic organisms, such as *Salmonella*, and this process may be particularly enhanced in some circumstances, for example under selection pressure resulting from antimicrobial usage. Once *Salmonella* isolates have acquired plasmids carrying genes conferring resistance to third-generation cephalosporins (either ESBL or AmpC resistance genes), the dissemination of such resistant *Salmonella* clones will also play a major part in influencing the occurrence of third-generation cephalosporin resistance.

Considering the prevalence of resistance to cefotaxime and resistance in MSs to Salmonella spp. and *E. coli* in all species for which relevant data are available, then in all reporting MSs where resistance was detected in 2012, the prevalence of resistance is higher in *E. coli* than it is in Salmonella spp. with the sole exception of Belgium and isolates from pigs. Table ESBL19 summarises the data and illustrates some interesting observations relating to the occurrence of cefotaxime resistance in Salmonella spp. and *E. coli* in MSs.

Where resistance is detected in *Salmonella* spp. in a MS, it is also invariably present in *E. coli* in that reporting MS and usually occurs at a higher level (with only one exception). Some MSs do not report cefotaxime resistance in *Salmonella* spp. or in *E. coli* for some food-producing animals. The degree of resistance observed in *Salmonella* spp. and *E. coli* may be correlated, in those MSs, which have a high level of resistance in *Salmonella* spp. and have a high level of resistance in *Salmonella* spp. and have a high level of resistance in *Salmonella* spp. and have a high level of resistance in *Salmonella* spp. and have a high level of resistance in *Salmonella* spp. and have a high level of resistance in *Salmonella* spp. It tends to appear not always hold true and would not be expected to hold where clonal dissemination of particular strains of *Salmonella* were responsible for the observed prevalence of resistance in *Salmonella* spp. It tends to appear that, in most MSs, commensal *E. coli* is the primary reservoir of beta-lactamase resistance, which is less frequently observed in *Salmonella* spp.

The data reported by the Netherlands are interesting because occurrence of resistance is assessed on a large number of samples tested. Resistance in *E. coli* isolates was detected at low levels in meat from pigs (1.0 %) and bovine animals (0.7 %), but at higher levels in meat from broilers (8.0 %). The figure for broilers represents a considerable decline from the figure reported in 2011 (31.9 %). It is desirable that these resistance figures are as low as possible. In the last years, a number of initiatives have occurred in the Netherlands, including changes to relevant antimicrobial authorisations and strenuous efforts to reduce antimicrobial consumption in livestock, which may have influenced the observed resistance levels. The decline in cefotaxime resistance observed in broilers in the Netherlands (18 %, 8.1 % and 5.8 % in 2010, 2011 and 2012, respectively) parallels the decline observed in broiler meat. In other MSs, a low occurrence of resistance to cefotaxime was generally recorded, since many countries tested a rather small sample size or reported only single-resistant isolates. As a result, minor fluctuations in the observed percentage of resistance are probably to be expected.

All MSs detected resistance to cefotaxime in broilers except Sweden; the prevalence ranged from 1.7 % to 13.5 %. Where MSs reported data for both pigs and broilers, the levels observed in pigs were consistently lower. In cattle, some MSs reported that cefotaxime resistance in cattle was intermediate, lying between that observed in pigs and broilers (Austria, Belgium and the Netherlands), while Poland found that levels in pigs and cattle were very similar and Switzerland observed that cefotaxime resistance in pigs exceeded that observed in cattle. However, it is of note that differences in the types of cattle sampled may make direct comparisons between MSs inappropriate.



Table ESBL19. Resistance (%)to cefotaxime in Salmonella spp. and indicator E. coli isolates in MSs in 2012 testing both bacterial species in either Gallus gallus, pigs or cattle

		Gallus	gallus			Pig	JS	Cattle				
Country	Salmon	ella spp.	E. coli		Salmor	Salmonella spp.		E. coli		Salmonella spp.		coli
	N	% Res	Ν	% Res	Ν	% Res	N	% Res	N	% Res	Ν	% Res
Austria	176	0.6	130	3.1	_	_	_	_	-	_	_	_
Belgium	664	18.1	325	28.0	187	11.2	205	2.9	42	2.4	364	8.8
Denmark	28	0	115	1.7	374	0	152	0.7	_	_	_	_
Finland	-	_	_	_	_	_	_	_	19	0	295	0
Germany	_	_	_	_	_	_	_	_	68	0	515	2.5
Hungary	261	0.4	105	7.6	38	0	68	1.5	-	_	-	_
Netherlands	192	4.2	292	5.8	263	0	284	0	68	0	559	0.5
Poland	739	0.7	328	10.7	10	0	190	2.6	_	_	_	_



7.5. Discussion

In 2012, as in 2011, resistance to third-generation cephalosporins was generally detected at low levels in *Salmonella* and indicator *E. coli* isolates recovered from major food-producing animals and meat thereof. In most MSs, the prevalence of resistance to cefotaxime in both *Salmonella* spp. and *E. coli* was equal to that observed for ceftazidime. Although resistance assessed using ECOFFs tends to usually detect resistance to both compounds, this is not always the case and differences in resistance to each compound may be observed, reflecting whether the ESBL enzyme conferring resistance is primarily a cefotaximase or a ceftazidimase. ESBLs belonging to the CTX-M family (primarily, though not entirely, cefotaximases) are currently the most important types of ESBL in both animals and humans in the majority of MSs. However, EFSA has recommended that both cefotaxime and ceftazidime are included in future harmonised mandatory monitoring to ensure optimal detection of all ESBLs (EFSA, 2012b) (see box below), as surveillance procedures should anticipate possible changes in the status of different ESBL enzymes.

While resistance to cefotaxime in *Salmonella* spp. isolates recovered from meat from broilers was reported at low or moderate levels in most reporting MSs, one MS recorded around 30 % resistance to both cefotaxime and ceftazidime in 2012 and 2011. In most MSs, the observed levels of resistance in meat from broilers and *Gallus gallus* (or broilers) showed many similarities; however, in a number of MSs, differences in the levels of cefotaxime resistance in meat and the species from which the meat was produced were observed. For example, in Belgium, resistance to cefotaxime or ceftazidime was not detected in *Salmonella* spp. from meat from broilers, while resistance to the same compounds was reported at moderate levels in *Gallus gallus*. Conversely, in Germany and the Netherlands, the cefotaxime resistance was recorded at a lower level in *Salmonella* spp. isolates from *Gallus gallus* than in isolates from poultry meat. The reasons underlying the absence of a clear correlation between the prevalence of resistance observed in livestock and then in meat derived from those animals within a MS may be related to the lack of direct comparability between the target populations used for the monitoring in retail meat and in broilers. The retail meat monitored may notably include not only domestic poultry meat, but also imported meat from other countries.

Considering all MSs, resistance to cefotaxime in *Salmonella* spp. recovered from meat from pigs and cattle was 0.9 % and 1.4 % respectively, which are both lower than the figure of 4.3 % reported for meat from broilers. Two factors contribute to this observed difference: (1) the proportion of reporting MSs which did not report resistance to cefotaxime in meat from pigs and cattle was higher than that for meat from broilers, and (2) in MSs which did report resistance, the levels of resistance reported for *Salmonella* spp. from meat broilers were higher than for meat from pigs and cattle.

The results have been presented by animal production type (where available and relevant) and this is the second year in which animal production type has been included in this way. Differences in the occurrence of resistance may be related to husbandry methods, age or stage of production, the degree of antimicrobial usage or the influence the structure of the particular livestock industry may have on clonal spread of resistant organisms. The prevalence of resistance to cefotaxime in *Salmonella* spp. was higher in broilers than in laying hens (when resistance was detected) for all MSs (except Austria). Laying hens tend to be infrequently treated with antimicrobials, especially once in lay. *S.* Enteritidis from broilers and layers were susceptible to cefotaxime from most MSs, although Belgium reported moderate cefotaxime resistance in *S.* Enteritidis from *Gallus gallus* and was the only MS to report moderate resistance in *S.* Typhimurium from *Gallus gallus*. Romania reported isolates resistant to ceftazidime, but susceptible to cefotaxime in laying hens, suggesting that a ceftazidimase enzyme may have been present. *Salmonella* spp. resistant to cefotaxime was most frequently observed in broilers and the proportion of MSs observing any degree of resistance was higher than that for other animal species (turkeys, pigs and cattle).

Breeding animals may play a role in the clonal dissemination of resistance in particular serovars of *Salmonella* when animals colonised at breeding units are moved to fattening farms. A proportion of the reporting MSs observed cefotaxime resistance in *Salmonella* spp. isolates from breeding flocks of *Gallus gallus* and from breeding pigs. Belgium, Germany and Spain all reported cefotaxime resistance in *S.* Typhimurium or monophasic *S.* Typhimurium from pigs, and both of these serovars are significant causes of human salmonellosis. In cattle, the level of resistance for *Salmonella* spp. from cattle was very low and the *S.* Typhimurium isolates tested were all susceptible to cefotaxime. Only one MS (Belgium) reported third-generation cephalosporin resistance in *Salmonella* spp. isolates from cattle.



Some Salmonella serovars have particular public health significance because they either are common causes of human salmonellosis or have acquired resistance to a large number of different antimicrobial compounds (or even exhibit both of these traits). Resistance to third-generation cephalosporins was detected in a number of serovars of particular public health importance, including S. Typhimurium, S. Enteritidis, S. Infantis, S. Kentucky, S. Java and monophasic S. Typhimurium. Among those serovars showing multiple antimicrobial resistance, S. Kentucky and S. Java are particularly important because they also often show other resistances, including resistance to fluoroquinolones (the other main antimicrobial used in the first-line treatment of human salmonellosis). Previous outbreaks of ESBL-producing salmonellae affecting poultry and humans have occurred, for example involving S. Virchow (Weill et al., 2004; Bertrand et al., 2006), and it is important that the monitoring performed can identify such serovars. It may be assumed that, even though the monitoring has not been designed to detect outbreaks, it should hopefully reflect indirectly serovars involved in a large outbreak(s). Considering these Salmonella serovars of particular public health importance, no resistance to cefotaxime was detected in S. Typhimurium in meat from pigs in the period 2010 to 2011, while two MSs (Belgium and Romania) detected cefotaxime resistance in the same serovar of the same origin in 2012. Belgium, Poland and Portugal detected cefotaxime resistance in S. Enteritidis from Gallus gallus in 2012, as Austria and Hungary did in 2011; however, no cefotaxime resistance was detected in S. Enteritidis in meat from broilers in 2012.

Although thorough cooking and appropriate food hygiene procedures kills any bacteria present on food and prevents cross-contamination of foods with resistant or susceptible bacteria, it is highly desirable that the level of resistance in zoonotic organisms is very low or zero, especially in relation to important antimicrobials for human treatment. Among the strains of *E. coli* occurring in animals, some may be able to cause infections in humans (many will be largely harmless animal commensals) and some, although they are primarily commensals of animals, may be able to transiently or permanently colonise the human intestine. During transient colonisation or passage through the human intestine, *E. coli* may be able to exchange their resistance plasmids with the commensal *E. coli* flora of humans. Therefore, it is also desirable that resistance to important antimicrobials for human treatment is also very low or zero in animal strains of *E. coli*, which might otherwise form a reservoir of resistance genes.

FUTURE HARMONISED MONITORING OF ESBLS AND AMPC ENZYMES

EFSA's Scientific Report on the harmonised monitoring and reporting of antimicrobial resistance in Salmonella, Campylobacter and indicator E. coli and Enterococcus spp. bacteria transmitted through food (EFSA, 2012b) has discussed further enhancement of surveillance in this area. In particular, detailed recommendations have been made for the isolation and identification of extended-spectrum beta-lactamases (ESBL) and AmpC E. coli and methods have been described which would promote a harmonised and, therefore, comparable approach to monitoring across the EU. Further characterisation of isolates in this way will allow possible links between animals and humans to be investigated and provide a better indication of the overall significance of any resistance to third-generation cephalosporins which is detected. Based on the proposals of EFSA, the European Commission has put forward and discussed with the Member States a new piece of legislation on the harmonised monitoring of antimicrobial resistance in food-producing animals and food, including details provisions on the monitoring of ESBL- and AmpC-producing Salmonella and indicator commensal E. coli. The new legislation (Decision 2013/652) will enter into force in 2014.



8. MATERIALS AND METHODS

8.1. Antimicrobial susceptibility data from humans available in 2012

MSs report results from AST to ECDC through TESSy. The data used in this report were submitted in connection to the annual data collection for the European Union Summary Report of Trends and Sources of Zoonoses and Zoonotic Agents end of May 2013.

8.1.1. Salmonella data of human origin

Nineteen MSs, as well as Iceland and Norway, provided data for 2012. The antimicrobials reported on for *Salmonella* are ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, kanamycin, nalidixic acid, streptomycin, sulfonamides, tetracyclines and trimethoprim. Some countries reported on all of these and others on only a few. Countries reported qualitative data, i.e. interpreted AST results for tested isolates (susceptible (S), intermediate (I) or resistant (R)), with the exception of Norway which used the isolate-based reporting under piloting at ECDC to report measured IZDs. The Norwegian data were then interpreted by ECDC, using the most recent clinical breakpoints from EUCAST where applicable and otherwise breakpoints from CLSI (Table MM1).

The public health reference laboratories were asked via e-mail to provide an update about which methods and which guidelines were being used for testing and interpretation. It should be noted that the public health reference laboratories in several countries type only a fraction of the isolates. The remaining isolates are then typed by hospitals or local laboratories, and the methods used by these are often unknown. Eight MSs plus Iceland and Norway used only disc diffusion methods, six other MSs used only dilution methods and another four MSs used a combination of the two depending on the situation and the antimicrobial (Table MM1 (continued)). The method used in one MS was unknown.

The guidelines used for the interpretation differed between countries (Table MM1). Nine countries used guidelines from CLSI, two used guidelines from EUCAST, five used a combination of the two and another five countries used another guideline (often national). Compared with 2011, two MSs and Iceland had changed to EUCAST from CLSI (and more followed in 2013). Clinical breakpoints were applied in all countries but two, which used EUCAST ECOFFs. For 4 of the 11 antimicrobials addressed, the MIC value for the CLSI breakpoints and the EUCAST ECOFFs are equivalent: chloramphenicol, nalidixic acid, sulfonamides and tetracyclines. Please note that CLSI define clinical resistance breakpoints as 'more than or equal to' a certain MIC value, while EUCAST use 'more than' a certain value. This means that a CLSI breakpoint of \geq 32 mg/L is the same as a EUCAST breakpoint of >16 mg/L.; see Table MM1 and Figure SA1. For three antimicrobials (cefotaxime, ciprofloxacin and gentamicin), the MIC values or zone diameters differ markedly between the clinical breakpoints and the ECOFFs. This is particularly the case for ciprofloxacin, for which the ECOFF is three times more sensitive than the EUCAST clinical breakpoint and five times more sensitive than the CLSI clinical breakpoint (Figure SA1). Results for these three antimicrobials must therefore be interpreted with caution, and no direct comparison between countries should be made.

Results are presented for countries reporting data for more than 20 isolates for the antimicrobial in question. Trend lines within the period 2008 to 2012 are shown for those countries reporting data for at least three consecutive years and 10 isolates per year. Countries which did not detect any resistant isolates during the period are mentioned but not shown in the graphs. Results are presented separately for the top three most important serovars: *S.* Enteritidis, *S.* Typhimurium and monophasic *S.* Typhimurium.

In order to better assess the impact from food consumed within each reporting country on the antimicrobial resistance levels found in human *Salmonella* isolates, the analysis focused on domestically acquired cases. However, since several countries had not provided any information on travel (or non-travel) of their cases, cases with unknown travel status were included in the analysis. The proportion of travel-associated, domestic and unknown cases among the tested *Salmonella* isolates is presented in Table MM2. An analysis was also made on the most likely country of infection of each disease case to compare resistance levels in human *Salmonella* infections acquired within the EU/EEA with those acquired when travelling in regions outside the EU/EEA.



Table MM1. Breakpoints used by MSs for the interpretation of 2012 susceptibility data on Salmonella of human origin

0	Amp	icillin	Cefota	axime	Chlora	mphenicol	Ciprofle	oxacin	Gent	amicin	Kanan	nycin	Ostidations
Country	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	Guidelines
Austria	_	≤13	_	≤16	_	≤16	_	≤18	_	≤13	_	≤13	EUCAST 2012 for Amp, Ctx, Chl, Cip, Gen. CLSI 2012 for Kan
Denmark	>8	-	>0.5	-	>16	_	>0.06	_	>2	-	>4	_	EUCAST ECOFFS. Neomycin is reported under kanamycim
Estonia	≥32	≤13	≥1	≤27	≥32	≤12	≥0.125	<30	≥16	≤12	Ι	≤13	WHO Collaborating Centre 2010, DTU Food (breakpoints based on CLSI)
France		<16	_	<23	_	<23	>1	<22	-	<16	-	<15	CA-SFM 2011
Germany	>8	-	>8	_	NA	NA	>2	_	>4	-	>16	_	German DIN standard
Greece ¹	_	≤13	-	≤22	-	≤12	-	≤15	_	≤12	-	≤13	CLSI 2011
Hungary	>8	<14	>2	<17	>8	<17	>0.064	-	>4	<14	-	<14	EUCAST 2012 except for Kan where CLSI 2012 was applied
Ireland	>8	-	>2	-	>8	_	>1	_	>4	Ι	≥64	_	EUCAST 2012 except for Kan where CLSI 2012 was applied
Italy	≥32	≤13	≥64	≤14	≥32	≤12	≥4	≤15	≥16	≤12	≥64	≤13	CLSI M100-S17-S19
Latvia ¹	≥32	≤13	NA	NA	NA	NA	≥4	≤15	NA	NA	NA	NA	CLSI
Lithuania ¹	≥32	≤13	≥64	≤14	≥32	≤12	≥4	≤15	≥16	≤12	≥64	≤13	CLSI M100-S17-S19
Luxembourg	_	≤13	-	≤22	-	≤12	-	≤15	_	≤12	-	≤13	CLSI 2012
Malta	≥16	_	≥4	NA	≥16	NA	≥2	_	≥8	-	NA	NA	Biomerieux Vitek II system; follows EUCAST 2010
Netherlands ¹	>4	_	>0.5	_	>16	_	>0.06	_	>2	Ι	NA	NA	EUCAST ECOFFS from 2007. For Str EFSA and Su CLSI
Romania ¹	≥32	≤13	≥4	≤22	≥32	≤12	≥4	≤15	≥16	≤12	≥64	≤13	CLSI 2012
Slovakia	>8	<14	>2	<17	>8	<17	>2	<19	>4	<14	NA	NA	EUCAST
Slovenia	-	≤13	-	≤22	-	≤12	-	≤15	-	≤12	-	≤13	CLSI M100-S22, 2012
Spain	_	≤13	-	≤22	-	≤12	-	≤15	_	≤12	-	≤13	CLSI-2010 M100-S-20 Vol.30
United Kingdom	≥8	_	≥1	_	≥8	_	≥0.125	_	≥4	-	≥16	_	HPA methodology based on Frost (1994)
Iceland	-	<14	NA	NA	_	<17	-	<19	NA	NA	NA	NA	EUCAST 2012
Norway	-	<14	NA	NA	_	<17	_	<19	NA	NA	NA	NA	EUCAST 2012

MS: Member State; MIC, minimum inhibitory concentration; –: this method is not used for the antimicrobial in question; NA: not applicable since this antimicrobial is not reported to The European Surveillance System or fewer than 20 isolates tested; EUCAST: European Committee on Antimicrobial Susceptibility Testing; ECOFF: epidemiological cut-off value; WHO: World Health Organization; DTU: Technical University of Denmark; CA-SFM: French Society for Microbiology; DIN: Deutsches Institut für Normung; CLSI: Clinical and Laboratory Standards Institute; HPA: Health Protection Agency.

1. Interpretive criteria and comments shown are from the 2010 or 2011 report as clinical breakpoints for 2012 were not reported.

Table continued overleaf.



	Nalidix	ic acid	Strepto	mvcin	Sulfon	amides	Tetracy	/clines	Trimet	noprim		
Country	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	Guidelines	Method
Austria	_	≤13	_	≤11	_	≤12	_	≤11	_	≤14	EUCAST 2012 for Tmp. CLSI 2012 for Nal, Str, Su, Tet	Disc diffusion
Denmark	>16	_	>16	_	>256	_	>8	_	>2	_	EUCAST ECOFFS. EFSA ECOFF for sulfonamides	Dilution
Estonia	≥32	≤13	≥32	≤11	≥512	≤12	≥16	≤11	≥16	≤10	WHO Collaborating Centre 2010, DTU Food (breakpoints based on CLSI)	Disc diffusion
France	>16	<15	-	<15	-	<12	_	<17	_	<13	CA-SFM 2011. Combination trimethoprim/sulfamethoxazole tested	Disc diffusion and dilution (gradient strip)
Germany	>16	_	>16	_	NA	NA	NA	NA	> 16	_	German DIN standard. Combination trimethoprim/sulfamethoxazole tested	Dilution
Greece ¹	_	≤13	-	≤11	NA	NA	-	≤11	_	≤10	CLSI 2011	Disc diffusion
Hungary	_	<14	_	<12	-	<13	_	<12	>4	<11	CLSI 2012 except for Stx EUCAST 2012. Combination trimethoprim/sulfamethoxazole tested	Disc diffusion and dilution (gradient strip)
Ireland	>16	_	>32	_	≥512	_	≥16	_	>4	_	EUCAST 2012 except for Su and Tet CLSI 2012 and Str EFSA 2007	Dilution
Italy	≥32	≤13	_	≤11	≥512	≤12	≥16	≤11	≥16	≤10	CLSI M100 S17 S19	Disc diffusion
Latvia ¹	NA	NA	NA	NA	NA	NA	NA	NA	≥16	≤10	CLSI	Not reported
Lithuania ¹	≥32	≤13	-	≤11	≥512	≤12	≥16	≤11	≥16	≤10	CLSI M100-S17-S19	Disc diffusion
Luxembourg	-	≤13	-	≤11	-	≤12	-	≤11	_	≤10	CLSI 2012. Combination trimethoprim/sulfamethoxazole tested	Disc diffusion
Malta	NA	NA	NA	NA	NA	NA	NA	NA	≥160	_	Biomerieux Vitek II system; follows EUCAST 2010 except for Stx. Combination trimethoprim/sulfamethoxazole tested	Dilution
Netherlands	>16	_	>32	_	-	_	>8	_	NA	NA	EUCAST ECOFFS. For Str EFSA and Su CLSI	Dilution
Romania ¹	≥32	≤13	≥32	≤11	≥512	≤12	≥16	≤11	≥16	≤10	CLSI 2012	Disc diffusion and dilution (gradient strip)
Slovakia	>16	<12	NA	NA	>256	<13	>8	<12	>4	<13	CLSI except for Stx EUCAST. Combination trimethoprim/sulfamethoxazole tested	Disc diffusion and dilution
Slovenia	_	≤13	-	≤11	_	≤12	_	≤11	_	≤10	CLSI M100-S22, 2012	Disc diffusion
Spain	_	≤13	_	≤11	-	_	_	≤11	_	≤10	CLSI-2010 M100-S-20 Vol.30. Combination trimethoprim/sulfamethoxazole tested	Disc diffusion
United Kingdom	≥16	_	≥16	_	≥64	_	≥8	_	≥2	_	HPA methodology based on Frost (1994)	Dilution (in agar breakpoint)
Iceland	_	NA	NA	NA	NA	NA	NA	NA	_	< 13	EUCAST 2012	Disc diffusion
Norway	_	≤13	-	≤11	-	≤12	-	≤11	_	<13	CLSI 2012 except for Stx EUCAST 2012	Disc diffusion
											•	

Table MM1 (continued). Breakpoints used by MSs for the interpretation of 2012 susceptibility data on Salmonella of human origin

MS: Member State; MIC: minimum inhibitory concentration; -: this method is not used for the antimicrobial in question; NA: not applicable since this antimicrobial is not reported to The European Surveillance System or fewer than 20 isolates tested; EUCAST: European Committee on Antimicrobial Susceptibility Testing; ECOFF: epidemiological cut-off value; WHO: World Health Organization; DTU: Technical University of Denmark; CA-SFM: French Society for Microbiology; DIN: Deutsches Institut für Normung; CLSI: Clinical and Laboratory Standards Institute; HPA: Health Protection Agency. 1. Interpretive criteria and comments shown are from the 2010 or 2011 report as clinical breakpoints for 2012 were not reported.



Table MM2. Proportion of tested Salmonella spp. isolates from human cases associated with travel, domestic cases and cases with unknown travel information by country in 2012

Country	Tested isolates	Travel-associated	Domestic	Unknown	
Country	N	%	%	%	
Austria	1,888	4	96	0	
Denmark	609	36	64	0	
Estonia	238	8	92	0	
France	1,278	0	0	100	
Germany	2,039	2	98	0	
Greece	112	5	88	7	
Hungary	588	0	100	0	
Ireland	306	29	12	59	
Italy	134	1	13	86	
Latvia	60	2	98	0	
Lithuania	1,744	0	0	100	
Luxembourg	135	0	0	100	
Malta	88	0	100	0	
Netherlands	1,160	11	89	0	
Romania	137	0	0	100	
Slovakia	1,010	1	99	0	
Slovenia	392	0	0	100	
Spain	1,880	0	76	23	
United Kingdom	8,644	28	21	51	
Total (19 MSs)	22,442	13	47	39	
Iceland	33	33	45	21	
Norway	1,322	73	17	10	

MS: Member State; N: number of the tested isolates.

Multi-drug resistance of human *Salmonella* spp. to 10 antimicrobials were also analysed. The 10 antimicrobials included were ampicillin, cefotaxime, chloramphenicol, ciprofloxacin/nalidixic acid, gentamicin, kanamycin, streptomycin, sulfonamides, tetracyclines and trimethoprim. Of these, only kanamycin is not on the list of antimicrobials tested for in food and animal isolates. Resistance to nalidixic acid and ciprofloxacin was addressed together: in the event that an isolate was resistant or exhibited intermediate resistance to either of these antimicrobials, the isolate was classified as non-susceptible to the combined antimicrobial ciprofloxacin/nalidixic acid, as the two substances belong to the same antimicrobial family.

Multi-drug resistance of an isolate was defined as non-susceptibility to at least three different antimicrobial classes (Magiorakos et al., 2012). Co-resistance to ciprofloxacin and cefotaxime was also estimated as these two antimicrobials are considered the most important for treatment of severe salmonellosis (ECDC et al., 2009). Only countries which reported the results of tests on the full range of antimicrobials were included in the analysis.



8.1.2. Campylobacter data of human origin

Fourteen MSs and Iceland provided data for 2012. The antimicrobials reported on for *Campylobacter* were amoxicillin, ampicillin, ciprofloxacin, erythromycin, gentamicin, nalidixic acid and tetracyclines. Some countries reported on all of these and others on only a few (Table MM3). Countries reported qualitative data, i.e. interpreted AST results for tested isolates (S, I or R).

National public health reference laboratories were asked to provide the methods and guidelines used for testing and interpretation. AST for *Campylobacter* seems to be performed in local or regional laboratories to a higher extent than for *Salmonella* and the methods and guidelines used outside of the national public health laboratories may therefore not be known. Five countries used disc diffusion for their routine testing, while six countries used dilution or gradient strip (Table MM3). Four countries used both disc diffusion and dilution, depending on the circumstances and the antimicrobial.

The guidelines used for the interpretation at the national level differed between countries (Table MM3). Four countries used guidelines from the French Society for Microbiology (CA-SFM), three used guidelines from CLSI (M45-A), two used guidelines from EUCAST, two used a combination of CA-SFM and CLSI and four used other guidelines. Of the five antimicrobials reported on from both human and animal/food isolates, the EUCAST clinical breakpoints and ECOFFS were at the same MIC values except for ciprofloxacin and the combination *C. coli*/erythromycin, where the ECOFF was one dilution higher than the clinical breakpoint. No clinical breakpoints were available for gentamicin and nalidixic acid. The CA-SFM breakpoints were generally also in the same range or one dilution higher or lower than the ECOFF except for the combination *C. coli*/erythromycin, and tetracycline, where it was two dilution steps higher than the ECOFFs. The CLSI breakpoints were often set at up to two dilutions higher than the ECOFF (Figure CA1). Results for the antimicrobials for which there are major differences in the interpretive criteria should be interpreted with caution, and direct comparisons between countries should be avoided.

Results are presented for countries reporting data for more than 20 isolates for the antimicrobial in question. Trend lines within the period 2008 to 2012 are shown for those countries reporting data for at least three consecutive years and 10 isolates per year. Countries which did not detect any resistant isolates during the period are mentioned but not shown in the graphs. Results are presented separately for the two most important *Campylobacter* species: *C. jejuni* and *C. coli*.

In order to better assess the impact from food consumed within each reporting country on the antimicrobial resistance levels found in human *Campylobacter* isolates, the analysis focused on domestically acquired cases. However, since several countries had not provided any information on travel (or non-travel) of their cases, cases with unknown travel status were included in the analysis. The proportion of travel-associated, domestic and unknown cases among the tested *Campylobacter* isolates is presented in Table MM4. An analysis was also made on the most likely country of infection of each disease case to compare resistance levels in human *Campylobacter* infections acquired within the EU/EEA with those acquired when travelling in regions outside the EU/EEA.

Multi-drug resistance of human *C. jejuni* and *C. coli* to six antimicrobials was also analysed. The six antimicrobials were amoxicillin, ampicillin, ciprofloxacin/nalidixic acid, erythromycin, gentamicin and tetracyclines. Of these, amoxicillin and ampicillin are not on the list of antimicrobials tested for in food and animal isolates. Resistance was addressed to both nalidixic acid and ciprofloxacin together: in the event that an isolate was resistant or exhibited intermediate resistance to either of these antimicrobials, the isolate was classified as non-susceptible to the combined antimicrobial ciprofloxacin/nalidixic acid, as the two substances belong to the same antimicrobial family.

Multi-drug resistance of an isolate was defined as non-susceptibility to at least three different antimicrobial classes (Magiorakos et al., 2012). Co-resistance to ciprofloxacin and erythromycin was also estimated as these two antimicrobials are considered the most important for treatment of severe salmonellosis (ECDC et al., 2009). Only countries which reported the results of tests on the full range of antimicrobials were included in the analysis.



Country	Amox	ticillin	Amp	icillin	Ciprofl	oxacin	Erythr	omycin	Genta	micin	Nalidia	kic acid	Tetrac	yclines	O dilations	
Country	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	Guidelines	Method
Austria	>8	-	>8	-	>0.5	_	>4 ¹ >8 ²	_	>2	-	>16	-	>2	-	EUCAST 2012 Combination amoxicillin/clavulanate tested	Dilution
Estonia	>16/2	< 14	>16	<14	>1	<22	>4	<17	>4	<16	>16	<15	>8	<17	CA-SFM 2010 Combination amoxicillin/clavulanate tested	Disc diffusion
France	-	< 14	-	<14	_	<22	-	<17	_	<16	-	<15	NA	NA	CA-SFM. Combination amoxicillin/clavulanate tested	Disc diffusion
Italy	NA	NA	-	≤6	_	≤6	-	≤6	_	≤6	-	≤6	_	≤6	CLSI M45-A Vol.26 no 19 for Cip and Ery. Same criteria applied for remaining antimicrobials	Disc diffusion
Hungary	NA	NA	NA	NA	>4	-	>4 ¹ >16 ²	NA	NA	NA	NA	NA	NA	NA	Unspecified reference	Dilution (gradient strip)
Lithuania ³	NA	NA	NA	NA	_	≤17	-	≤19	NA	NA	NA	NA	NA	NA	BSAC for disc diffusion	Disc diffusion
Luxembourg	-	< 14	-	< 14	>1	_	>4	_	-	<16	-	<15	NA	<17	CA-SFM 2012	Disc diffusion and dilution (gradient strip)
Malta	NA	NA	NA	NA	≥1	-	≥4	-	NA	NA	NA	NA	NA	NA	Personal communication from HPA in 2004	Dilution (gradient strip)
Netherlands ³	NA	NA	NA	NA	≥1.0–1.5	≤ 19–20	≥1.5–2.0	<13–≤23	NA	NA	NA	NA	≥2–8	≤17–28	Survey in 12 clinical labs in the Netherlands in 2009	Disc diffusion and dilution
Romania ³	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	CA-SFM 2010 Combination amoxicillin/clavulanate tested	Disc diffusion

Table MM3. Breakpoints used by MSs for the interpretation of 2012 susceptibility data on Campylobacter of human origin

MS: Member State; MIC: minimum inhibitory concentration; -: this method is not used for the antimicrobial in question; NA: not applicable since this antimicrobial is not reported to The European Surveillance System or fewer than 20 isolates tested; EUCAST: European Committee on Antimicrobial Susceptibility Testing; CA-SFM: French Society for Microbiology; CLSI: Clinical and Laboratory Standards Institute; BSAC: British Society for Antimicrobial Chemotherapy; HPA: Health Protection Agency.

1. Breakpoint used for C. jejuni.

2. Breakpoint used for C. coli.

3. Interpretive criteria and comments shown are from the 2010 or 2011 report; clinical breakpoints for 2012 were not reported.

Table continued overleaf.



Table MM3 (continued). Breakpoints used by MSs for the interpretation of 2012 susceptibility data on Campylobacter of human origin

Country	Amox	icillin	Ampi	illin	Ciprofl	oxacin	Erythro	omycin	Genta	micin	Nalidix	ic acid	Tetracy	clines	Guidelines	Mathad
Country	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	MIC	mm	Guidelines	Method
Slovakia ³	NA	NA	NA	NA	≥4	-	≥32	-	NA	NA	NA	NA	≥16	_	CLSI	Dilution
Slovenia	_	< 14	_	<14	≥4	<22	≥32	<17	_	<16	-	<15	_	<17	CA-SFM 2010 for disc diffusion and CLSI M45-A for Cip and Ery gradient strip. Combination amoxicillin/clavulanate tested	Disc diffusion and dilution (gradient strip)
Spain	>16/2	_	NA	NA	≥4	_	≥32	_	>4	_	>16	_	≥16	_	CA-SFM 2010 for Amx, Gen, Nal. CLSI-2010 M45AE for Cip, Ery, Tet Combination amoxicillin/clavulanate tested	Dilution (gradient strip)
United Kingdom ³	-	Ι	>16	Ι	>1	Ι	>4	Ι	>1	Ι	>16	Ι	>2	_	EUCAST	Dilution (microbroth and gradient strip)
Iceland	NA	NA	NA	NA	≥4	≤6	≥32	≤6	NA	NA	NA	NA	NA	NA	CLSI M45A2 2010	Disc diffusion and dilution (gradient strip)

MS: Member State; MIC: minimum inhibitory concentration; NA: not applicable since this antimicrobial is not reported to The European Surveillance System or fewer than 20 isolates tested; -: this method is not used for the antimicrobial in question; CA-SFM: French Society for Microbiology; CLSI: Clinical and Laboratory Standards Institute; EUCAST: European Committee on Antimicrobial Susceptibility Testing.

1. Breakpoint used for C. jejuni.

2. Breakpoint used for C. coli.

3. Interpretive criteria and comments shown are from the 2010 or 2011 report; clinical breakpoints for 2012 were not reported.

Table MM4. Proportion of tested Campylobacter spp. isolates from human cases associated with travel, domestic cases and cases with unknown travel information by country in 2012

Country	Tested isolates	Travel-associated	Domestic	Unknown
Country	N	%	%	%
Austria	387	0	0	100
Estonia	246	6	94	0
France	4,728	0	0	100
Hungary	71	0	100	0
Italy	319	9	24	67
Lithuania	228	0	72	27
Luxembourg	561	0	0	100
Malta	214	0	100	0
Netherlands	3,708	6	94	0
Romania	19	0	0	100
Slovakia	1,371	1	99	0
Slovenia	982	0	2	98
Spain	228	0	75	25
United Kingdom	25,715	1	11	89
Total (14 MSs)	38,777	1	22	77
Iceland	58	50	34	16

MS: Member State; N: number of the tested isolates.

8.2. Antimicrobial susceptibility data from animals and food available in 2012

8.2.1. Data reported under Directive 2003/99/EC in 2012

MSs generated data on antimicrobial susceptibility through the testing of zoonotic and indicator bacteria isolated from various animal species/production types and food categories, sampled through a number of different national schemes. Isolates may have been collected by different monitoring approaches, either by active monitoring of animals and foods or, in some cases, by passive monitoring based on diagnostic submission of samples from clinical cases of disease in animals, or from foods sampled as part of investigatory work. In the case of passive monitoring, the isolates tested often constituted a sub-sample of the total isolates available at the National Reference Laboratory (NRL). Clinical investigation data were not accounted for in this report.

Dilution and disc diffusion testing methods were used by reporting MSs for susceptibility testing, and both quantitative and qualitative data were reported at the EU level.

- 'Quantitative data' derived from dilution methods consisted of the number of isolates having a specific MIC value (measured in mg/L) relative to the total number of isolates tested, for each antimicrobial agent and in each specific food/animal category.
- 'Quantitative data' derived from diffusion methods comprised the number of isolates having a specific IZD (measured in mm) relative to the total number of isolates tested, for each antimicrobial agent and in each food/animal category.
- 'Qualitative data' consisted of the number of isolates out of the total number of isolates that were resistant to each antimicrobial agent in each food/animal category; qualitative data can be generated either from MIC determination or from disc diffusion testing.

For the year 2012, 26 MSs and 3 non-MSs reported data on antimicrobial resistance in tested *Salmonella* and *Campylobacter*, commensal *E. coli* or MRSA isolates from food-producing animals and/or food. Data on antimicrobial resistance in tested *Salmonella* and *Campylobacter* have been reported on a mandatory basis under Directive 2003/99/EC and data on antimicrobial resistance in tested commensal *E. coli* and commensal enterococci or MRSA isolates have been reported by the MSs on a voluntary basis. An overview of the MSs and non-MSs reporting antimicrobial resistance data (which were included in this report) in 2012 is shown in Table MM5.

	Number of MSs and non-MSs	Number of the tested i	ncluded in the report
Bacteria	reporting quantitative or qualitative data	MIC dilution	Diffusion
Salmonella	24 MSs+3 non-MSs	73,840	3,821
Campylobacter	15 MSs+1 non-MS	18,191	-
Indicator Escherichia coli	13 MSs+2 non-MSs	63,298	-

2,100

Table MM5. MSs reporting data in 2012 from animals and food and description of data included in the report

MS: Member State; MIC: minimum inhibitory concentration; MRSA: methicillin-resistant *Staphylococcus aureus*.

1 MS+1 non-MS

1. In 2012, 10 MSs and one non-MS reported data on the occurrence of MRSA.

For the purpose of this report, only quantitative dilution and quantitative disc diffusion data have been considered.

MRSA¹



8.2.1.1. Resistance data in *Salmonella* and *Campylobacter* from animals and food

Quantitative (MIC) results on antimicrobial resistance in *Salmonella* isolates from animals and food were reported by 20 MSs and 2 non-MSs (Norway and Switzerland) in 2012. The information collected by these countries was in accordance with EFSA's recommendations (EFSA, 2007); these data are described in Chapter 3. The countries reported results for only low numbers of isolates (fewer than 10) have been excluded from the analysis.

In 2012, 15 MSs and 1 non-MS (Switzerland) reported data on antimicrobial resistance in *Campylobacter*. All *Campylobacter* results were reported as MIC values in accordance with EFSA's recommendations (EFSA, 2007). These data are described in Chapter 4.

8.2.1.2. Resistance data in indicator bacteria

For indicator (commensal) *E. coli*, a total of 11 MSs and 2 non-MSs (Norway and Switzerland) reported quantitative dilution (MIC) results from animals or meat derived from those animals; these data are described in Chapter 5. Some countries reported results for only low numbers of isolates (fewer than 10); these data have been excluded from the analysis. Slovenia and Portugal reported quantitative results for indicator *E. coli* isolates tested according to CLSI recommendations and using the CLSI disc diffusion method.

8.2.1.3. Resistance data to third-generation cephalosporins

In relation to third-generation cephalosporin resistance in indicator *E. coli* and *Salmonella* spp., EFSA's recommendations suggest the use of cefotaxime alone to detect important types of resistance (EFSA, 2007). Most MSs reported results for cefotaxime; some also reported results for ceftazidime; these data are described in Chapter 7. Cefotaxime is likely to detect the presence of most cefotaximases (CTX-M enzymes), which appear to be currently the most prevalent type of ESBL enzymes in bacteria isolated from food-producing animals in the EU. The use of cefotaxime will also detect the presence of AmpC enzymes in *Salmonella* or *E. coli*. Some ESBLs are ceftazidimases rather than cefotaximases (particularly enzymes in the TEM and SHV families of ESBLs). Although testing both cefotaxime and ceftazidime is therefore optimal for the detection of all ESBLs and AmpC enzymes, EFSA's guidelines have recommended testing cefotaxime to detect all CTX-M enzymes mainly for reasons of affordability.

8.2.1.4. Data on methicillin-resistant *Staphylococcus aureus* (MRSA)

Data relating to MRSA prevalence were reported by 10 MSs and one non-MS (Switzerland). Among these, Switzerland reported data on resistance in MRSA isolates from pigs and Belgium in MRSA isolates from cattle. The methods for collecting and testing samples for MRSA are not harmonised between MSs and as a result MSs may use differing procedures. Owing to the variety of methods employed by MSs, these are explained in detail within Chapter 6 to enable readers to better follow the procedures carried out by individual countries.

There is an important difference between the methods used to isolate *Salmonella*, *Campylobacter* and indicator *E. coli* and that used to isolate MRSA. For the former group of organisms, there is no selective medium used to isolate from primary samples organisms possessing a particular resistance, whereas, for MRSA, antimicrobials are used to selectively isolate only those *Staphylococcus aureus* isolates which are resistant to methicillin. Some MSs may have sampled particular production types of animals (for example laying hens in *Gallus gallus* or veal calves in cattle), and this introduces another source of possible variation which may account for observed differences between MSs.



8.3. Antimicrobials used for susceptibility testing in animals and food

The antimicrobials incorporated in this summary analysis were selected based on their relative public health importance and as representatives of different antimicrobial classes, taking into account EFSA's reports and recommendations on the harmonised monitoring and reporting of antimicrobial susceptibility data (EFSA 2007, 2008).

8.3.1. Antimicrobials for susceptibility testing of Salmonella

In 2012, both dilution and disc diffusion methods were used to test the susceptibility of *Salmonella* isolates from animals and food by MSs. Tables MM6 and MM7 show the antimicrobials selected by the different countries for susceptibility testing. Quantitative dilution results allowed MIC distributions to be reported for *Salmonella* for the following antimicrobials: ampicillin, apramycin, cefotaxime, ceftazidime, ceftiofur, chloramphenicol, ciprofloxacin, colistin, florfenicol, gentamicin, kanamycin, nalidixic acid, neomycin, spectinomycin, streptomycin, sulfonamides, trimethoprim and tetracyclines. For further information on reported MIC distributions and the number of resistant isolates, refer to the Level 3 tables published on the EFSA website.

Data on *Salmonella* which were reported as disc diffusion data are presented in Appendix 2. Although results may not be directly comparable between MSs, it is anticipated that in most cases procedures will not have changed markedly over time within a country, and therefore comparisons of the proportion of resistant isolates over time in that country may be possible.



Table MM6. Antimicrobials selected for susceptibility testing of Salmonella isolates from animals and food by MSs and non-MSs reporting quantitative data as MIC distributions, in 2012

Country	Ampicillin	Apramycin	Cefotaxime	Ceftazidime	Ceftiofur	Chloramphenicol	Ciprofloxacin	Colistin	Florfenicol	Gentamicin	Kanamycin	Nalidixic acid	Neomycin	Spectinomycin	Streptomycin	Sulfonamides	Trimethoprim	Tetracyclines
Austria	٠		٠	٠		٠	٠	•	•	٠	•	•			•	•	٠	•
Belgium	٠		٠	٠		٠	٠	•	•	٠	•	•			•	•	٠	•
Czech Republic	٠		٠	٠		٠	٠			٠		٠			٠	٠	٠	•
Denmark	٠	٠	٠		•	•	•	•	•	•		•	•	•	•	•	•	•
Estonia	•		•	٠		٠	٠	•	•	•	•	•			•	•	٠	•
Finland	٠		٠			٠	٠		٠	•	٠	٠			٠	٠	٠	•
Germany	٠		٠	٠		٠	٠		٠	•	٠	٠			٠	٠	٠	•
Hungary	٠		٠			٠	٠			٠		٠			٠	٠	٠	٠
Ireland	٠		٠	٠		٠	٠		٠	•	٠	٠			٠	٠	٠	٠
Italy	٠		٠	٠		٠	٠		٠	٠	٠	٠			٠	٠	٠	٠
Latvia	٠		٠			٠	٠		٠	•	٠	٠			٠	٠	٠	٠
Netherlands	٠		٠	٠		٠	٠	٠	٠	•	٠	٠			٠	٠	٠	•
Norway	٠		٠			٠				•		•			٠	٠	٠	•
Poland	٠		٠	٠		٠	٠		٠	•	٠	٠			٠	٠	٠	٠
Portugal	٠		•			٠	٠			•		٠			٠	٠	٠	•
Romania	٠		٠	٠		٠	٠	٠	٠	٠	٠	٠			•	٠	٠	•
Slovakia	٠		٠	•		٠	٠	٠	٠	٠	٠	٠			•	•	٠	•
Spain	٠		٠	•		٠	٠	٠	٠	٠	٠	٠			•	•	٠	•
Sweden	•		•			•	•		•	•	•	•			•	•	•	•
Switzerland	٠		٠	•		•	•	٠	٠	٠	٠	٠			•	•	•	•
United Kingdom	•		•	•		•	•			•		•			•	•	•	•

MS: Member State; MIC: minimum inhibitory concentration. Note: Sulfonamides may include a variety of substances.

Table MM7. Antimicrobials selected for susceptibility testing of Salmonella isolates from animals and food by MSs reporting quantitative data as disc inhibition zones, in 2012

Country	Amikacin	Ampicillin	Cefepime	Cefotaxime	Chloramphenicol	Ciprofloxacin	Ertapenem	Gentamicin	Imipenem	Levofloxacin	Meropenem	Nalidixic acid	Piperacillin	Streptomycin	Sulfonamides	Tetracyclines	Tobramycin	Trimethoprim
Greece		•		٠	•	•		•				٠		•	•	٠		•
Spain	•	•	•	٠	•	•	•	•	٠	•	•	٠	•	•	•	٠	•	•

MS: Member State.

Note: Sulfonamides may include a variety of substances.



8.3.2. Antimicrobials for susceptibility testing of *Campylobacter*

In 2012, all quantitative *Campylobacter* data were reported as MIC values, generated by dilution methods. Table MM8 shows the antimicrobials selected by the different countries for susceptibility testing of *Campylobacter* isolates. In this report, antimicrobial resistance was reported separately for *C. jejuni* and *C. coli*.

MIC distributions were analysed for the following antimicrobials: ciprofloxacin, chloramphenicol, erythromycin, gentamicin, nalidixic acid, streptomycin and tetracyclines. These antimicrobials were selected based on public health relevance and as representatives of different classes of antimicrobials. For further information on reported MIC distributions and the number of resistant isolates, refer to the Level 3 tables published on the EFSA website.

Table MM8. Antimicrobials selected for susceptibility testing of Campylobacter isolates from animals and food by MSs and non-MSs reporting quantitative data as MIC distributions, in 2012

Country	Amoxicillin	Ampicillin	Chloramphenicol	Ciprofloxacin	Clarithromycin	Erythromycin	Gentamicin	Imipenem	Nalidixic acid	Neomycin	Streptomycin	Sulfonamides	Tetracyclines	Tulathromycin
Austria	•	•	•	•		•	•	•	•	•	•		•	
Belgium			•	٠		٠	٠		•		٠		•	
Czech Republic				•		٠	•		•		•		•	
Denmark			•	٠		•	•		٠		٠		•	
Estonia				•		•	٠		•		•		•	
Finland				•		•	٠		•		•		•	
France				•		•	•		•		•		•	
Germany			•	•		•	•		•		•		•	
Hungary			•	•		•	•		•		•		•	
Italy			•	•		•	٠		•		•		•	
Netherlands		•	•	•	•	•	٠		•	•	•	•	•	•
Poland				•		•	•		•		•		٠	
Romania			٠	٠		٠	٠		•		٠		٠	
Spain		٠	٠	٠		٠	٠		•		٠		٠	
Sweden				•		•	٠		•		•		•	
Switzerland			•	٠		٠	٠		٠		•		•	

MS: Member State; MIC: minimum inhibitory concentration.

Note: Sulfonamides may include a variety of substances.



8.3.3. Antimicrobials for susceptibility testing of *Escherichia coli*

In 2012, both dilution and disc diffusion methods were used to test the susceptibility of *E. coli* isolates from animals and food. Tables MM9 and MM10 show the antimicrobials selected by the different countries for susceptibility testing. In this report, susceptibility data from food and animal isolates are presented.

MIC distributions were analysed for the following antimicrobials: ampicillin, apramycin, cefotaxime, ceftazidime, ceftiofur, chloramphenicol, ciprofloxacin, colistin, florfenicol, gentamicin, kanamycin, meropenem, nalidixic acid, neomycin, spectinomycin, streptomycin, sulfonamides, trimethoprim and tetracyclines. These antimicrobials were selected based on their public health relevance and as representatives of different antimicrobial classes. For further information on reported MIC distributions and the number of resistant isolates, refer to the Level 3 tables published on the EFSA website.

Table MM9. Antimicrobials selected for susceptibility testing of Escherichia coli isolates from animals and food by MSs and non-MSs reporting quantitative data as MIC distributions, in 2012

Country	Ampicillin	Apramycin	Cefotaxime	Ceftazidime	Ceftiofur	Chloramphenicol	Ciprofloxacin	Colistin	Florfenicol	Gentamicin	Kanamycin	Meropenem	Nalidixic acid	Neomycin	Spectinomycin	Streptomycin	Sulfonamides	Trimethoprim	Tetracyclines
Austria	•		•	•		٠	•			•		•	•			•	٠	•	•
Belgium	•		•	•		•	•	•	•	•	•		•			•	•	•	•
Denmark	•	•	٠		•	٠	•	•	•	•			•	•	•	•	٠	•	•
Finland	•		٠			٠	•	•	•	•	•		•			•	٠	•	•
France	•		٠	•		٠	٠		•	•			•			•	٠	•	٠
Germany	•		٠	•		٠	٠		•	•	•		•			•	٠	٠	•
Hungary	•		٠			٠	٠			•			•			•	٠	•	•
Netherlands	•		٠	•		٠	٠	•	•	•	•		•			•	٠	•	•
Norway	•		•			٠				•			•			•	٠	•	•
Poland	•		٠	•		٠	•		•	•	•		•			•	٠	•	•
Spain	•		•			•	•		•	•	•		•			•	•	•	•
Sweden	•		•			•	•	•	•	•	•		•			•	•	•	•
Switzerland	٠		٠	•		٠	•	•	•	•	•		•			•	٠	٠	•

MS: Member State; MIC: minimum inhibitory concentration.

Note: Sulfonamides may include a variety of substances.

Table MM10. Antimicrobials selected for susceptibility testing of Escherichia coli isolates from animals and food by one MS reporting quantitative data as disc inhibition zones, in 2012

Country	Amikacin	Ampicillin	Cefazolin	Cefotaxime	Cefoxitin	Cefpodoxime	Ceftazidime	Cephalothin	Chloramphenicol	Ciprofloxacin	Colistin	Enrofloxacin	Ertapenem	Florfenicol	Gentamicin	Imipenem	Kanamycin	Marbofloxacin	Meropenem	Moxifloxacin	Nalidixic acid	Neomycin	Nitrofurantoin	Norfloxacin	Streptomycin	Sulfonamides	Trimethoprim	Tetracyclines	Tobramycin
Portugal	٠	٠	٠	•	•	•	•		٠	٠	٠	٠	•		•	٠			٠	٠			٠	٠	•		٠	٠	•
Slovenia		•		•				•	•	•				•	•		•	•			•	•			•	٠	•	•	

MS: Member State.

Note: Sulfonamides may include a variety of substances.



8.3.4. Antimicrobials for susceptibility testing of MRSA

In 2012, Belgium reported data on susceptibility testing of MRSA isolates from cattle and Switzerland reported data from pigs. Details of the antimicrobials selected by Belgium and Switzerland are provided in Chapter 6. For further information on reported MIC distributions and the number of resistant isolates, refer to the Level 3 tables published on the EFSA website.

8.4. Data description and analysis

8.4.1. Description and analysis of antimicrobial resistance data

Methods to interpret, describe, and analyse antimicrobial resistance data were presented in detail in the 2004 to 2007 Community Summary Report on Antimicrobial Resistance (EFSA, 2010b).

Overview tables of the resistance data reported

Quantitative MIC data, generated by dilution methods recommended by EFSA, have been reported and analysed together; quantitative IZD data, which constitute a relatively small fraction of the total data, have not been included in the analysis of quantitative data and have been described separately in Appendix 2. The IZD data reported by MSs under Directive 2003/99/EC for the years 2004 to 2007 were interpreted as described in previous EU Summary Reports. Some MSs reported antimicrobial resistance data as both quantitative and qualitative data; in that case, only the quantitative data have been included. Data generated from the antimicrobial susceptibility testing and reported as quantitative/qualitative by MSs have been described in the overview tables of individual chapters.

MIC distributions, ECOFFs and the occurrence of resistance

For each combination of microorganism, antimicrobial and food or animal category tested, MIC distributions have been presented as frequency tables, giving the number of isolates tested having a given MIC at each test dilution (mg/L) of the antimicrobial. MIC distributions are available as Level 3 tables on the EFSA website.

Quantitative MIC data for *Salmonella* were, wherever possible, interpreted using ECOFFs as listed in Decision 2007/407/EC (corresponding to those published by EUCAST at the time of publication of the Decision) and presented in Table MM11. Subsequent amendments by EUCAST to the ECOFFs have not yet been incorporated; this will be achieved by issue of a revised Decision. An isolate was defined as 'microbiologically resistant' (i.e. displaying a decreased susceptibility) to a selected antimicrobial when its MIC value was above the ECOFF. A more sensitive MIC breakpoint or ECOFF (i.e. a lower MIC breakpoint or ECOFF) might be expected to result in more isolates being defined as clinically or microbiologically resistant, respectively; the number of isolates affected in that way will of course depend on the distribution of MIC results.

REVISION OF ECOFFS

The epidemiological cut-off value (ECOFF) for E. coli versus ciprofloxacin has been recently revised by the European Committee on Antimicrobial Susceptibility Testing (EUCAST). Wild-type isolates are now considered to have a ciprofloxacin minimum inhibitory concentration greater than 0.06 mg/L, an increase from the previous ECOFF of greater than 0.03 mg/L. The proportion of isolates showing microbiological resistance according to this breakpoint will alter when the new breakpoint is adopted and in fact will be reduced. For reasons of continuity and to comply with the current legislation where applicable, the ECOFFs used in this report have been those adopted in EFSA's recommendations (EFSA 2007, 2008) and quoted in Commission Decision 2007/407/EC. For these reasons, the most recent revisions by EUCAST have not been included in this report. The report for 2013, will incorporate all of these changes in a comprehensive revision, which will also re-evaluate the historical data using the revised ECOFFs.



The occurrence of resistance to a number of antimicrobials was determined (giving the percentage of isolates 'microbiologically resistant' out of those tested) for *Salmonella*, *Campylobacter*, indicator *E. coli* and enterococcal isolates from *Gallus gallus*, turkeys, pigs and cattle, and meat from *Gallus gallus*, pigs and cattle and are presented and analysed in tables on the occurrence of resistance in this report. These are the animal and food categories most frequently reported on by most MSs. Additionally, for the first time, data have been presented at the production-type level where possible. Data are included only if quantitative MIC data are provided by more than four MSs or disc diffusion data are provided by more than two MSs for the bacterium–animal/food category combination. An exception to this rule has nevertheless been made in the chapters on *Salmonella* serovars of public health importance (see below) and on MRSA. Data reported from fewer than 10 tested isolates per combination and per MS are not included. Data are reported in separate chapters dedicated to each microorganism and in Appendix 1 for *Salmonella* data obtained from disc diffusion. In addition, the occurrence of resistance (i.e. resistance levels) in reporting MS groups was calculated as totals (the total number of resistant isolates out of the total number of tested isolates across reporting MSs), and not the weighted means.

Resistance in Salmonella serovars of public health importance

In this report, antimicrobial resistance in tested *Salmonella* isolates were aggregated to give a value for *Salmonella* spp. for each country and food/animal category. In addition, whenever sufficient data were transmitted by MSs for a particular food/animal category, the most prevalent *Salmonella* serovars, *S*. Enteritidis and *S*. Typhimurium, were also reported separately for that food/animal category. Additional tables have been included in this year's report to describe the occurrence of antimicrobial resistance among *Salmonella* serovars of public health importance. In order to present a complete overview of the animal populations and food categories in which specific *Salmonella* serovars of public health importance have been recovered, data derived from fewer than four reporting countries have been included.

Data description

Throughout the report, the following definitions apply:

- Level or occurrence of antimicrobial resistance means the percentage of resistant isolates as a proportion of the isolates tested of that microorganism.
- **MS reporting group** means the MSs that provided data and were included in the relevant table of antimicrobial resistance for that bacterium–food or animal category–antimicrobial combination.

Terms used to describe the antimicrobial resistance levels are:

rare:	<0.1 %
very low:	0.1 % to 1 %
low:	>1 % to 10 %
moderate:	>10 % to 20 %
high:	>20 % to 50 %
very high:	>50 % to 70 %
extremely high:	>70 %

These terms are applied to all antimicrobials. However, the significance of a given level of resistance will depend on the particular antimicrobial and its importance in human and veterinary medicine.

Temporal trends in resistance

Where the minimum criteria were met for the inclusion of data in this report (i.e. more than 10 isolates tested by a MS and more than four MSs reporting results for that antimicrobial, microorganism, food or animal category), then temporal trend graphs were generated showing the resistance to different antimicrobials over the 2006 to 2012 period, by plotting the level of resistance for each year of sampling. Only countries which had reported for four or more years in the 2006 to 2012 period were included.

In order to assess the statistical significance of temporal trends, the proportions of resistance were modelled against time in a logistic regression. Results were provided only where there were five years or more of available data to use in the model, and where the likelihood ratio test suggested that the model was meaningful. This analysis was carried out in SAS9.2 using the PROC LOGISTIC function for each country

where temporal trend data were presented in the report. The PROC LOGISTIC function uses a logit transform to model proportion of prevalence against year, and provides estimates for both intercepts and slope. Models resulting in a p-value of <0.05 were considered to be significant.

For ciprofloxacin and nalidixic acid, resistance trends over time were visually explored for *Salmonella*, *Campylobacter* and indicator *E. coli* by *trellis* graphs, using the *lattice* package in the R software (http://www.r-project.org). Graphs were created for those countries for which resistance data were available for four or more years, for at least one of the two antimicrobials. MS-specific resistance levels trend graphs use a unique scale and countries are shown in alphabetical order.

Spatial analysis of resistance through maps

MS-specific antimicrobial resistance levels for selected bacterium/food or animal category combinations were plotted in maps for 2012, using ArcGIS 9.3. In the maps, resistance levels are presented with colours reflecting the continuous scale of resistance to the antimicrobial of interest among reporting MSs; thus, there might be some apparent discrepancies between the colours and resistance levels between maps. Percentages shown in this map refer to countries that reported quantitative MIC data for more than 10 isolates in 2012. When quantitative 2012 data were not available, the 2011 level of resistance was used instead and referred by a footnote to the map. The countries labelled as 'qualitative data' therefore include those reporting IZD data, MIC data for fewer than 10 isolates or purely qualitative data (as proportion of resistant isolates).

Table MM11. Epidemiological cut-off values used to interpret MIC distributions (mg/L) for bacteria from animals and food – the given values define the microbiologically resistant isolates

Antimicrobial agent	Salmonella	Escherichia coli	Campylobacter jejuni	Campylobacter coli
	mg/L	mg/L	mg/L	mg/L
Ampicillin	>4	>8		
Apramycin	>16	>16		
Avilamycin				
Cefotaxime	>0.5	>0.25		
Ceftazidime	>2	>0.5		
Ceftiofur	>2	>1		
Chloramphenicol	>16	>16	>16	>16
Ciprofloxacin	>0.06	>0.03	>1	>1
Erythromycin			>4	>16
Florfenicol	>16	>16		
Gentamicin	>2	>2	>1	>2
Linezolid				
Nalidixic acid	>16	>16	>16	>32
Neomycin	>4	>8		
Spectinomycin		>64		
Streptomycin	>32	>16	>2	>4
Sulfonamides	>256 ¹	>64		
Quinupristin/dalfopristin				
Tetracyclines	>8	>8	>2	>2
Trimethoprim	>2	>2		
Vancomycin				

MIC: minimum inhibitory concentration.

1. Cut-off values were not defined by the European Committee on Antimicrobial Susceptibility Testing; instead cut-off values defined by the European Union Reference Laboratory on antimicrobial resistance (Technical University of Denmark) were used.



8.4.2. Analysis of multi-resistance and co-resistance data

As a consequence of the availability of antimicrobial resistance data at an isolate-based level in an important number of MSs, the analysis of multi-resistance and co-resistance data becomes a feasible and important exercise in the light of the public health relevance of the emergence of multi-resistant bacteria. As a matter of fact, the analysis and reporting on multi-resistance in the 2012 EU Summary Report on antimicrobial resistance was previously recommended and endorsed by the Task Force on Zoonoses Data Collection at its meeting on antimicrobial resistance in April 2013.

The intention is to focus mainly on multi-/co-resistance patterns involving critically important antimicrobials according to the bacterial species, such as cephalosporins, fluoroquinolones and macrolides, and to summarise important information in the EU Summary Report. The occurrence of the isolates of a serotype/resistance pattern of interest is studied at the MS level and at the reporting MS group/EU level, as the overall picture for all MSs might show a more definite pattern of emergence and spread. In addition, the analysis of data may reveal the existence of new or emerging patterns of multi-resistance, particularly in *Salmonella* serotypes.

8.4.2.1. Analysis of multi-resistance patterns

Definitions

For the purpose of this analysis, a multi-resistant isolate is one defined as resistant to at least three different antimicrobial substances, belonging to any three antimicrobial families listed in the harmonised set of antimicrobials included in the EFSA recommendations (EFSA, 2007, 2008). Table MM12 lists those recommended antimicrobials.

Resistance to nalidixic acid and ciprofloxacin is addressed together: an isolate that is resistant to either of the two will be termed resistant to the combined antimicrobial ciprofloxacin/nalidixic acid, as the two substances belong to the same antimicrobial family.

By contrast, a fully susceptible isolate is one defined as non-resistant to all of the antimicrobial substances included in the set of substances recommended for *Salmonella*, *Campylobacter* and indicator *E. coli*.

The term co-resistance has been defined as two or more resistance genes which are genetically linked, i.e. located adjacent or close to each other on a mobile genetic element (Chapman, 2003). For brevity, the term is used slightly more loosely in this report and indicates two or more phenotypic resistances to different classes of antimicrobials, exhibited by the same bacterial isolate.

Table MM12. Harmonised set of antimicrobials listed in the EFSA recommendations

Zo	onotic bacteria	Indicator bacteria
Salmonella	Campylobacter coli/C. jejuni	Escherichia coli
Ampicillin (Amp)	Ciprofloxacin (Cip)	Ampicillin (Amp)
Cefotaxime (Ctx)	Erythromycin (Ery)	Cefotaxime (Ctx)
Chloramphenicol (Chl)	Gentamicin (Gen)	Chloramphenicol (Chl)
Ciprofloxacin (Cip)	Streptomycin (Str)	Ciprofloxacin (Cip)
Gentamicin (Gen)	Tetracycline (Tet)	Gentamicin (Gen)
Nalidixic acid (Nal)		Nalidixic acid (Nal)
Streptomycin (Str)		Streptomycin (Str)
Sulfonamides (Su)		Sulfonamides (Su)
Tetracycline (Tet)		Tetracycline (Tet)
Trimethoprim (Tmp)		Trimethoprim (Tmp)



Data analysis

The frequency and percentage of isolates that are considered susceptible/resistant to all of the antimicrobials tested were determined for *Salmonella* (*Salmonella* spp., *S.* Enteritidis, *S.* Typhimurium and monophasic *S.* Typhimurium), *Campylobacter* species and indicator *E. coli* for each country and each animal population/food category. Isolates for which no susceptibility data were provided for some of the antimicrobial substances were disregarded. Data analysis was presented for a particular country only when the number of tested isolates was at least 10, except for monophasic *Salmonella* Typhimurium.

Summary indicators of multi-resistance

To illustrate the relative proportions of multi-resistant isolates and the diversity of the resistance to multiple antimicrobials, graphical illustration was chosen. The percentages of isolates susceptible and resistant to one, two, three, etc., antimicrobials are shown using a composite bar graph displaying stacked bars, but only for certain combinations of bacterium–animal population or food category–MS of particular interest.

The objective is first to give an overview of the situation on multi-resistance through summary indicators:

- the proportion of fully susceptible isolates;
- the proportion of multi-resistant isolates;
- an index/indices of diversity, such as the entropy measure¹⁸, summarising the distributions of isolate frequencies and, thus, the diversity among the different categories of multi-resistance (resistance to one, two, three, etc., antimicrobials).

The 'summary indicators' of multi-resistance can be calculated and reported yearly and, therefore, used to follow evolution of the multi-resistance situation across animal populations/food categories and MSs over time.

Diversity of multi-resistance

Resistance can be limited to resistance to only one or two antimicrobial substances, or resistance can be equally spread out from resistance from the lower to the higher number of antimicrobial substances. In other words, the frequencies across the categories resistant to one, two, three substances, and so on, can follow different types of distributions: skewed to the right with higher frequencies for the lower numbers resistant; highly peaked or fully spread out; or even, at least in theory, with higher frequencies for the larger numbers resistant. The entropy measure quantifies the degree of diversity of resistance. The standardised **unweighted entropy** takes values between 0 and 1. It takes the value 0 if all resistance is of one single type (e.g. resistance to exactly two antimicrobial substances) and takes the maximal value 1 if resistance to any number of antimicrobial substances is occurring equally often. The unweighted version does not take any order into account. Therefore, particular frequencies at the lower numbers resistant lead to the same entropy value when having these particular frequencies at the higher number resistant. The **weighted entropy** takes higher numbers of antimicrobial substances.

8.4.2.2. Analysis of co-resistance

The co-resistance patterns of interest

Co-resistance to cefotaxime and ciprofloxacin was estimated in *Salmonella* and *E. coli* isolates, as these two antimicrobials are of particular interest in human medicine. Co-resistance was addressed using both ECOFFs and clinical breakpoints in isolates of these bacteria. In *C. jejuni* and *C. coli* isolates, co-resistance to ciprofloxacin and erythromycin was estimated as these two antimicrobials are of particular interest in human medicines. The interpretive ECOFFs used to address co-resistance to ciprofloxacin and erythromycin were, for *C. jejuni*, Cip>1 mg/L and Ery>4 mg/L and, for *C. coli*, Cip>1 mg/L and Ery>16 mg/L. These values may be considered as very similar to clinical breakpoints.

¹⁸ Weighted or unweighted entropy measures may be considered.



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APPENDIX 1. Frequency distributions of *salmonella* serovars in animals and food in 2012 antimicrobial resistance data

Appendix Table SER1. Frequency distribution of Salmonella serovars in meat from broilers (Gallus gallus), in 2012

Serovars	Countries and total number of MSs reporting		Meat from broilers (Gallus gallus)		
		N	% f		
S. Infantis	8 MSs: CZ, EE, DE, HU, IT, PL, RO, SK	339	45.81		
S. Enteritidis	7 MSs: BE, CZ, DE, LV, NL, PL, SK	99	13.38		
S. Kentucky	3 MSs: HU, IE, RO	67	9.05		
S. Paratyphi B	2 MSs: BE, NL	39	5.27		
S. Java	1 MS: DE	31	4.19		
S. Indiana	2 MSs: CZ, PL	23	3.11		
S. Newport	2 MSs: CZ, PL	19	2.57		
S. Heidelberg	1 MS: NL	15	2.03		
S. Virchow	1 MS: RO	13	1.76		
S. Colindale	olindale 1 MS: RO		1.62		
S. Agona	2 MSs: CZ, SK	11	1.49		
S. Typhimurium	3 MSs: IE, IT, LV	11	1.49		
S. Hadar	2 MSs: IT, RO	9	1.22		
S. Bredeney	1 MS: RO	7	0.95		
S. Ohio	1 MS: CZ	6	0.81		
S. Ruzizi	1 MS: RO	6	0.81		
S. Rissen	1 MS: RO	5	0.68		
S. 6,7:-:1,5	1 MS: CZ	4	0.54		
S. Djugu	1 MS: RO	4	0.54		
S. Grampian	1 MS: RO	4	0.54		
S. Brandenburg	1 MS: RO	3	0.41		
S. Montevideo	1 MS: CZ	3	0.41		
S. Derby	2 MSs: CZ, DE	2	0.27		
S. Gloucester	1 MS: RO	1	0.14		
S. Isangi	1 MS: EE	1	0.14		
S. Kottbus	1 MS: CZ	1	0.14		
S. Livingstone	1 MS: RO	1	0.14		
S. Mbandaka	1 MS: IT	1	0.14		
S. Minnesota	1 MS: LV	1	0.14		
S. Thompson	1 MS: IT	1	0.14		
S. enterica subsp. enterica	1 MS: DE	1	0.14		



Appendix Table SER2. Frequency distribution of Salmonella serovars in meat from pigs, in 2012

Sorovoro	Countries and total number of MSs reporting	Meat fr	om pigs
Serovars	Countries and total number of MSs reporting	Ν	% f
S. Typhimurium	12 MSs: BE, CZ, DK, EE, DE, HU, IE, IT, LV, NL, PL, RO	322	47.08
S. Derby	8 MSs: CZ, EE, DE, HU, IE, IT, NL, RO	109	15.94
S. Typhimurium, monophasic	8 MSs: CZ, DK, DE, HU, IE, IT, NL, SK	108	15.79
S. Infantis	6 MSs: CZ, EE, DE, IE, IT, RO	27	3.95
S. Bredeney	2 MSs: IT, RO	16	2.34
S. Rissen	2 MSs: IT, RO	15	2.19
S. enterica subsp. enterica	enterica subsp. enterica 4 MSs: CZ ,EE, DE, IT		2.19
S. Ruzizi	1 MS: RO	11	1.61
S. Brandenburg	2 MSs: IT, NL	8	1.17
S. Gloucester	1 MS: RO	6	0.88
S. Kortrijk	1 MS: RO	5	0.73
S. Agona	1 MS: EE	4	0.58
S. Virchow	1 MS: RO	4	0.58
S. Choleraesuis	2 MSs: CZ, IT	3	0.44
S. Livingstone	3 MSs: EE, IT, RO	3	0.44
S. Panama	1 MS: IT	3	0.44
S. Bovismorbificans	1 MS: RO	2	0.29
S. Colindale	1 MS: RO	2	0.29
S. Enteritidis	2 MSs: CZ, RO	2	0.29
S. Give	1 MS: IT	2	0.29
S. Kapemba	1 MS: IT	2	0.29
S. Thompson	1 MS: IT	2	0.29
S. enterica subsp. salamae	1 MS: IT	2	0.29
S. Bareilly	1 MS: EE	1	0.15
S. Bsilla	1 MS: RO	1	0.15
S. Dublin	1 MS: IE	1	0.15
S. Kottbus	1 MS: IT	1	0.15
S. London	1 MS: IT	1	0.15
S. Manhattan	1 MS: IT	1	0.15
S. Minnesota	1 MS: EE	1	0.15
S. Montevideo	1 MS: RO	1	0.15
S. Muenchen	1 MS: IT	1	0.15
S. Muenster	1 MS: IT	1	0.15
S. enterica subsp. diarizonae	1 MS: IT	1	0.15

N: number of isolates tested; % f: percentage frequency of isolates tested.



Appendix Table SER3. Frequency distribution of Salmonella serovars in meat from bovine animals, in 2012

Serovars	Countries and total number of MSs reporting		Meat from bovine animals		
		N	% f		
S. Typhimurium	7 MSs: CZ, EE, FI, DE, IE, IT, NL	24	39.34		
S. Derby	3 MSs: CZ, IE, IT	10	16.39		
S. Typhimurium, monophasic	3 MSs: DE, IE, NL	7	11.48		
S. Dublin	1 MS: IE	5	8.20		
S. enterica subsp. enterica	2 MSs: CZ, DE	3	4.92		
S. Infantis	2 MSs: HU, IE	2	3.28		
S. Rissen	1 MS: IT	2	3.28		
S. 9,12:lv:-	1 MS: CZ	1	1.64		
S. Hadar	1 MS: IT	1	1.64		
S. Kentucky	1 MS: IE	1	1.64		
S. London	1 MS: CZ	1	1.64		
S. Montevideo	1 MS: CZ	1	1.64		
S. Muenster	1 MS: IT	1	1.64		
S. Newport	1 MS: IE	1	1.64		
S. Saintpaul	1 MS: IE	1	1.64		



Appendix Table SER4. Frequency distribution of Salmonella serovars in Gallus gallus (fowl), in 2012

Serovars	Countries and total number of MSs reporting	Gallus gallus (fowl)		
		N	% f	
S. Enteritidis	15 MSs: AT, BE, CZ, DK, DE, HU, IT, LV, NL, PL, PT, RO, SK, ES, UK	1,253	28.96	
S. Infantis	14 MSs: AT, BE, CZ, DK, DE, HU, IT, LV, PL, RO, SK, ES, SE, UK	781	18.05	
S. Mbandaka	9 MSs: AT, BE, CZ, HU, IT, PL, RO, ES, UK	237	5.48	
S. Tennessee	5 MSs: AT, BE, CZ, HU, RO	203	4.69	
S. Kentucky	8 MSs: BE, CZ, DK, HU, IE, IT, RO, ES	168	3.88	
S. Paratyphi B	2 MSs: BE, NL	166	3.84	
S. Typhimurium	12 MSs: AT, BE, DK, FI, DE, HU, IT, PL, RO, ES, SE, UK	161	3.72	
S. Senftenberg	7 MSs: AT, BE, HU, IT, RO, ES, UK	156	3.61	
S. Minnesota	1 MS: BE	126	2.91	
S. Livingstone	8 MSs: AT, BE, HU, IT, RO, ES, SE, UK	117	2.70	
S. Agona	7 MSs: AT, BE, HU, IT, RO, ES, UK	95	2.20	
S. Montevideo	9 MSs: AT, BE, CZ, DK, HU, IT, RO, ES, UK	84	1.94	
S. Liverpool	1 MS: RO	70	1.62	
S. Thompson	6 MSs: AT, BE, HU, IT, RO, ES	48	1.11	
S. Hadar	5 MSs: AT, BE, IT, RO, ES	42	0.97	
S. Typhimurium, monophasic	9 MSs: AT, BE, DK, DE, IT, NL, PT, ES, UK	40	0.92	
S. enterica subsp. enterica	3 MSs: CZ, DE, IT	39	0.90	
S. Rissen	5 MSs: BE, IT, RO, ES, UK	37	0.86	
S. Ohio	4 MSs: AT, CZ, ES, UK	33	0.76	
S. Corvallis	3 MSs: IT, RO, ES	32	0.74	
S. Kottbus	5 MSs: AT, BE, CZ, HU, IT	32	0.74	
S. Kedougou	2 MSs: IT, UK	28	0.65	
S. Derby	4 MSs: CZ, DK, IT, RO	21	0.49	
S. Indiana	4 MSs: CZ, HU, PL, UK	19	0.44	
S. Taksony	1 MS: RO	17	0.39	
S. 6,7:-:1,5	2 MSs: CZ, ES	16	0.37	
S. Virchow	5 MSs: BE, DE, IT, PT, ES	14	0.32	
S. Bredeney	5 MSs: BE, HU, IT, RO, ES	13	0.30	
S. Bovismorbificans	3 MSs: AT, DK, HU	12	0.28	
S. Braenderup	4 MSs: AT, BE, CZ, ES	12	0.28	
S. Muenchen	2 MSs: IT, ES	11	0.25	
S. Newport	7 MSs: BE, CZ, HU, IT, RO, ES, UK	11	0.25	
S. Anatum	4 MSs: BE, IT, ES, UK	10	0.23	
S. Give	4 MSs: AT, BE, IT, UK	10	0.23	
S. Uganda	1 MS: RO	10	0.23	
S. Cerro	4 MSs: BE, HU, IT, ES	9	0.20	
S. Havana	4 MSs: BE, IT, ES, UK	9	0.21	
S. Lille	1 MS: CZ	9	0.21	
S. 6,7:z29	1 MS: BE	8	0.21	
S. Blockley	2 MSs: IT, RO	8	0.18	
S. Java	2 MSs : DE, UK	8	0.18	
S. Lexington	2 MSs : BE, IT	7	0.16	
S. Amsterdam	2 MSs : BE, RO	6	0.10	
S. Coeln	2 MSs: AT, IT	6	0.14	
S. Gallinarum biovar Pullorum	2 MSs : AT, TT 2 MSs : IT, RO	6	0.14	
S. Albany	1 MS: RO	5	0.14	
S. Dabou	1 MS: ES	5	0.12	
S. Dublin	3 MSs: AT, BE, UK	5	0.12	
S. Glostrup	1 MS: RO	5	0.12	
S. Isangi	1 MS: IT	5	0.12	
S. Isang	1 mo. 11	5	0.12	

Table continued overleaf.



Appendix Table SER4 (continued). Frequency distribution of Salmonella serovars in Gallus gallus (fowl), in 2012

Serovars	Countries and total number of MSs reporting	Gallus gallus (fowl)		
		N	% f	
S. London	3 MSs: AT, IT, ES	5	0.12	
S. Saintpaul	4 MSs: AT, BE, HU, IT	5	0.12	
S. Stanley	2 MSs: CZ, HU	5	0.12	
S. 6,7:-:-	2 MSs: BE, UK	4	0.09	
S. Abony	1 MS: HU	4	0.09	
S. Bareilly	1 MS: DK	4	0.09	
S. Haifa	1 MS: IT	4	0.09	
S. 3,19:-:-	1 MS: BE	3	0.07	
S. 9:-:-	1 MS: BE	3	0.07	
S. Cubana	2 MSs: RO, ES	3	0.07	
S. Goverdhan	1 MS: DK	3	0.07	
S. Hessarek	1 MS: BE	3	0.07	
S. Idikan	1 MS: BE	3	0.07	
S. Orion	1 MS: UK	3	0.07	
S. Worthington	2 MSs: AT, BE	3	0.07	
S. 4:i:-	1 MS: BE	2	0.05	
S. Altona	1 MS: ES	2	0.05	
S. Brandenburg	2 MSs: BE, ES	2	0.05	
S. Gallinarum biovar Gallinarum	1 MS: RO	2	0.05	
S. Goldcoast	2 MSs: ES, UK	2	0.05	
S. Llandoff	2 MSs: AT, BE	2	0.05	
S. Oranienburg	1 MS: AT	2	0.05	
S. Ouakam	1 MS : BE	2	0.05	
S. Szentes	1 MS: CZ	2	0.05	
S. Veneziana	1 MS: IT	2	0.05	
S. 1,3,19:-:-	1 MS: ES	1	0.02	
S. 13,23:i:-	1 MS : BE	1	0.02	
S. 4,12:-:-	1 MS : BE	1	0.02	
S. 4,12:-:1,2	1 MS: SK	1	0.02	
S. 4,12:d:-	1 MS : BE	1	0.02	
S. 6,7:d:-	1 MS : BE	1	0.02	
S. 6,7:z10:-	1 MS: UK	1	0.02	
S. 6,8:z10:-	1 MS : BE	1	0.02	
S. Agama	1 MS : UK	1	0.02	
S. Chester	1 MS : RO	1	0.02	
S. Djugu	1 MS : BE	1	0.02	
S. Durham	1 MS : UK	1	0.02	
S. Gallinarum	1 MS: AT	1	0.02	
S. Heidelberg	1 MS: IT	1	0.02	
S. Jerusalem	1 MS : BE	1	0.02	
S. Manhattan	1 MS: IT	1	0.02	
S. Meleagridis	1 MS: UK	1	0.02	
S. Mikawasima	1 MS: ES	1	0.02	
S. Muenster	1 MS: IT	1	0.02	
S. Orion var. 15	1 MS: UK	1	0.02	
S. Sandiego	1 MS: BE	1	0.02	
S. Schwarzengrund	1 MS: UK	1	0.02	
S. Soerenga	1 MS: ES	1	0.02	
S. Toulon	1 MS: IT	1	0.02	
S. Weltevreden	1 MS: IT	1	0.02	
S. Yoruba	1 MS: BE	1	0.02	



Appendix Table SER5. Frequency distribution of Salmonella serovars in turkeys, in 2012

Serovars	Countries and total number of MSs reporting	Turkeys		
Serovars	Countries and total number of MSs reporting	Ν	% f	
S. Derby	5 MSs: CZ, IE, IT, ES, UK	172	25.15	
S. Stanley	3 MSs: AT, CZ, HU	78	11.40	
S. Kentucky	5 MSs: CZ, HU, PL, SK, ES	59	8.63	
S. Saintpaul	6 MSs: AT, CZ, DE, HU, IT, PL	51	7.46	
S. Newport	5 MSs: CZ, HU, IT, SK, UK	47	6.87	
S. Typhimurium, monophasic	4 MSs: DE, IT, ES, UK	28	4.09	
S. Infantis	3 MSs: AT, DE, HU	27	3.95	
S. Bredeney	2 MSs: HU, IT	25	3.65	
S. Kedougou	1 MS: UK	25	3.65	
S. Hadar	3 MSs: DE, IT, ES	23	3.36	
S. Kottbus	4 MSs: CZ, HU, IT, UK	20	2.92	
S. Indiana	2 MSs: ES, UK	17	2.49	
S. London	1 MS : ES	17	2.49	
S. Typhimurium	7 MSs: AT, BE, FI, DE, IT, ES, UK	17	2.49	
S. Blockley	1 MS: IT	11	1.61	
S. Mbandaka	1 MS: UK	10	1.46	
S. Bovismorbificans	3 MSs: AT, HU, UK	8	1.17	
S. Enteritidis	5 MSs: AT, CZ, DE, HU, SK	8	1.17	
S. Agona	4 MSs: AT, HU, ES, UK	7	1.02	
S. Senftenberg	3 MSs: HU, ES, UK	5	0.73	
S. Schwarzengrund	1 MS: IT	4	0.58	
S. Orion var. 15	1 MS: UK	3	0.44	
S. 6,7:z10:-	1 MS: UK	2	0.29	
S. Bardo	1 MS: UK	2	0.29	
S. Haifa	1 MS: IT	2	0.29	
S. Montevideo	2 MSs: AT, ES	2	0.29	
S. enterica subsp. enterica	2 MSs: DE, IT	2	0.29	
S. 3,15:-:-	1 MS: UK	1	0.15	
S. 4,12:-:-	1 MS: BE	1	0.15	
S. 4,12:b:-	1 MS: ES	1	0.15	
S. Agama	1 MS: UK	1	0.15	
S. Anatum	1 MS: IT	1	0.15	
S. Dabou	1 MS: ES	1	0.15	
S. Dembe	1 MS: ES	1	0.15	
S. Java	1 MS: DE	1	0.15	
S. Ohio	1 MS: UK	1	0.15	
S. Tennessee	1 MS: HU	1	0.15	
S. Wisbech	1 MS: ES	1	0.15	
S. Worthington	1 MS: AT	1	0.15	



Appendix Table SER6. Frequency distribution of Salmonella serovars in pigs, in 2012

Serevere	Countries and total number of MSs reporting	P	Pigs		
Serovars	Countries and total number of MSs reporting	N	% f		
S. Typhimurium	11 MS: BE, DK, EE, FI, DE, HU, IE, LV, NL, ES, SE	515	35.49		
S. Typhimurium, monophasic	8 MSs: BE, DK, DE, IE, IT, NL, PL, ES	421	29.01		
S. Derby	8 MSs: BE, DK, EE, DE, HU, IT, NL, ES	280	19.30		
S. Infantis	5 MSs: BE, DK, EE, DE, HU	45	3.10		
S. Livingstone	3 MSs: BE, DK, IT	22	1.52		
S. Agona	2 MSs: BE, EE	21	1.45		
S. Rissen	3 MSs: BE, IT, ES	19	1.31		
S. enterica subsp. enterica	3 MSs: EE, DE, IT	13	0.90		
S. group O:4	1 MS: HU	12	0.83		
S. Choleraesuis	2 MSs: EE, IT	11	0.76		
S. London	3 MSs: DK, IT, ES	11	0.76		
S. Enteritidis	5 MSs: BE, DK, EE, DE, HU	10	0.69		
S. Mbandaka	2 MSs: BE, DK	10	0.69		
S. 4:i:-	1 MS: BE	8	0.55		
S. Worthington	1 MS: EE	6	0.41		
S. Anatum	1 MS: BE	5	0.34		
S. Bovismorbificans	1 MS: HU	3	0.21		
S. Brandenburg	2 MSs: BE, DK	3	0.21		
S. Bredeney	2 MSs: HU, ES	3	0.21		
S. Senftenberg	2 MSs: BE, IT	3	0.21		
S. Give	2 MSs: BE, DK	2	0.14		
S. Kapemba	1 MS: ES	2	0.14		
S. Kentucky	2 MSs: IE, ES	2	0.14		
S. Lexington	1 MS: EE	2	0.14		
S. Montevideo	1 MS: EE	2	0.14		
S. Paratyphi B	1 MS : BE	2	0.14		
S. 4,12:-:1,2	1 MS: DK	1	0.07		
S. 4,5,12:-:1,2	1 MS: DK	1	0.07		
S. 4,5:b	1 MS: ES	1	0.07		
S. 6,7:-:l,w	1 MS: DK	1	0.07		
S. Abony	1 MS: IT	1	0.07		
S. Brikama	1 MS: ES	1	0.07		
S. Coeln	1 MS: IT	1	0.07		
S. Gloucester	1 MS: BE	1	0.07		
S. Goldcoast	1 MS: DK	1	0.07		
S. Heidelberg	1 MS: DK	1	0.07		
S. Jerusalem	1 MS: BE	1	0.07		
S. Kedougou	1 MS: IT	1	0.07		
S. Minnesota	1 MS: BE	1	0.07		
S. Muenchen	1 MS: DK	1	0.07		
S. Panama	1 MS: IT	1			
S. Rideau	1 MS: 11 1 MS: BE	1	0.07		
		-	0.07		
S. Stanley	1 MS: DK	1	0.07		
S. enterica subsp. salamae	1 MS: IT	1	0.07		



Appendix Table SER7. Frequency distribution of Salmonella serovars in bovine animals, in 2012

Serovars	Countries and total number		Cattle (bovine animals)		
	of MSs reporting	N	% f		
S. Typhimurium	9 MSs: BE, EE, FI, DE, IE, IT, NL, ES, SE	166	55.33		
S. Dublin	7 MSs: BE, EE, DE, IE, LV, NL, SE	66	22.00		
S. Typhimurium, monophasic	5 MSs: DE, IE, IT, NL, ES	29	9.67		
S. Enteritidis	2 MSs: BE, EE	8	2.67		
S. Montevideo	2 MSs: BE, ES	5	1.67		
S. Java	1 MS: DE	4	1.33		
S. Rissen	1 MS: ES	3	1.00		
S. enterica subsp. enterica	2 MSs: EE, IT	3	1.00		
S. 9:-:-	1 MS: BE	2	0.67		
S. Chester	1 MS: EE	2	0.67		
S. Derby	1 MS: ES	2	0.67		
S. Livingstone	1 MS: BE	2	0.67		
S. London	1 MS: IT	2	0.67		
S. Agona	1 MS: SE	1	0.33		
S. Duesseldorf	1 MS: SE	1	0.33		
S. Hadar	1 MS: DE	1	0.33		
S. Lille	1 MS: ES	1	0.33		
S. Muenchen	1 MS: IT	1	0.33		
S. enterica subsp. diarizonae	1 MS: SE	1	0.33		



APPENDIX 2. Antimicrobial resistance in *Salmonella* - qualitative data

2.1. Introduction

In 2012, two MSs, Greece and Spain, reported on antimicrobial resistance in *Salmonella* from animals (Greece: *Gallus gallus*) and food (Greece, meat from broilers and meat from pig, and Spain, egg products) as quantitative disc diffusion data, which have been analysed as qualitative data and presented in this chapter. In food, both countries reported less than 10 isolates. These disc diffusion data have been analysed using the breakpoints for resistance specified by the reporting MS and in accordance with the method used (Appendix Tables QSA1–QSA3).

In the case of data reported exclusively as qualitative data, when information on the thresholds used to interpret the resistance was also available, it has been possible to pool the data submitted by MSs and present them in this section. It should, however, be noted that countries may not have used the same threshold values or qualitative methods and so direct comparisons between the proportions of resistant isolates in MSs reporting only qualitative data should be interpreted with caution. For this reason, tables do not show the summary figure for the reporting MS group and the spatial distributions of the levels of resistance for *Salmonella* based on qualitative data are not shown here; this is in accordance with previous reports. Furthermore, for those countries that reported quantitative data on antimicrobial resistance as presented in Chapter 3, corresponding qualitative data have been excluded from the overview tables and analyses presented in this chapter.

Resistance to the following antimicrobial agents are described in detail below: ampicillin, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines.

	Qualitative data							
Origin	Total number of MSs reporting	Countries						
	3	MSs: CY, LU, SI						
Gallus gallus (fowl)	3	Non-MS: IS						
Turkeys	3	MSs: CY, IE, SI						
Dian	2	MSs: AT, IE, LV						
Pigs	3	Non-MS: IS						
Cattle (bovine animals)	3	MSs: AT, IE, LU						
Meat from broilers (Gallus	r.	MSs: AT, ES, LT, LU, SI						
gallus)	5	Non-MS: IS						
Meat from other poultry species	5	MSs: AT, ES, LT, PL, SI						
Most from pige	Λ	MSs: AT, ES, LT, LU						
Meat from pigs	4	Non-MS: IS						
Meat from bovine animals	3	MSs: AT, LU, ES						
Meat, mixed meat	2	MSs: ES, SI						
Fishery products	1	MS: ES						
Foodstuffs (unspecified)	1	MS: PT						

Appendix Table QSA1. Overview of MSs reporting qualitative data on Salmonella spp. from animals and food in 2012

Note: For abbreviations of Member States (MS) and other reporting countries, see Appendix 7.



Appendix Table QSA2. Overview of MSs reporting qualitative data on Salmonella Typhimurium from animals and food in 2012

	Qualitative data						
Origin	Total number of MSs reporting	Countries					
Gallus gallus (fowl)	1	MS: CY					
Pigs	2	MSs: AT, IE					
Cattle (bovine animals)	3	MSs: AT, IE, LU					
Meat from pigs	1	MS: AT					
Meat, mixed meat	1	MS : SI					
Foodstuffs (unspecified)	1	MS: PT					

Note: For abbreviations of Member States (MS) and other reporting countries, see Appendix 7.

Appendix Table QSA3. Overview of MSs reporting qualitative data on Salmonella Enteritidis from animals and food in 2012

	(Qualitative data
Origin	Total number of MSs reporting	Countries
Gallus gallus (fowl)	3	MSs: CY, LU, SI
Meat from broilers (Gallus gallus)	3	MSs: AT, LT, LU
Meat from other poultry species	2	MSs: AT, LT
Meat, mixed meat	1	MS: SI
All foodstuffs	1	MS : PT

Note: For abbreviations of Member States (MS) and other reporting countries, see Appendix 7.



2.2. Antimicrobial resistance in *Salmonella* isolates from food (qualitative data)

Meat from broilers (Gallus gallus), meat from pigs, meat from bovine animals

Resistance levels among Salmonella spp. isolates

Austria, Slovenia and Spain reported qualitative data on resistance among *Salmonella* spp. from meat from broilers in 2012. Austria and Spain, and the Netherlands and Spain, reported qualitative data on resistance among *Salmonella* spp. from meat from pigs and from meat from bovine animals, respectively. The results are presented in the table Appendix Tables QSA4.

2.3. Antimicrobial resistance in *Salmonella* isolates from animals (qualitative data)

2.3.1. Fowl (Gallus gallus)

2.3.1.1. Resistance levels among Salmonella

Four MSs and one non-MS reported qualitative data for isolates of *Salmonella* from *Gallus gallus*. The results are presented in the table Appendix Tables QSA5.

2.3.2. Pigs

Resistance levels among Salmonella

Austria and Iceland were the only countries to report qualitative data for isolates of *Salmonella* spp. from pigs. The results are presented in the table Appendix Tables QSA6.

2.3.3. Cattle (bovine animals)

Resistance levels among Salmonella

Austria was the only MS reported qualitative data for isolates of *Salmonella* spp. from cattle in 2012.The results are presented in the table Appendix Tables QSA6.



Appendix Table QSA4. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among Salmonella spp. isolates from meat from broilers, from pigs and from bovine animals in MSs reporting qualitative data in 2012

Country	Am	oicillin	Cefot	axime	Chloram	phenicol	Ciprof	loxacin	Genta	amicin	Nalidi	xic acid	Sulfor	namides	Tetra	cyclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Meat from broi	lers															
Austria	61	1.6	61	0	61	3.3	61	1.6	61	0	61	73.8	61	67.2	61	68.9
Slovenia	15	6.7	_	_	15	0	15	100	15	0	15	100	15	100	15	100
Spain	40	10.0	_	_	22	0	40	2.5	40	2.5	40	17.5	_	_	22	18.2
Meat from pigs	;															
Austria	19	57.9	19	0	19	15.8	19	0	19	0	19	10.5	19	57.9	19	68.4
Spain	67	43.3	18	5.6	51	25.5	50	0	67	0	67	10.4	37	70.3	51	86.3
Meat from bov	ine anima	als														
Netherlands	18	33.3	18	5.6	18	0	18	11.1	18	0	18	11.1	18	61.1	18	44.4
Spain	44	11.4	29	0	40	2.5	43	0	44	2.3	44	13.6	34	5.9	39	5.1

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.

Appendix Table QSA5. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among Salmonella spp. isolates from Gallus gallus in MSs reporting qualitative data in 2012

Country	Am	Ampicillin Cefotaxi		taxime	me Chloramphenicol		Ciprofloxacin		Gen	tamicin	Nalid	ixic acid	Sulfo	namides	Tetra	cyclines
Country	N	% Res	N	% Res	N	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Cyprus	27	37.0	-	_	27	3.7	27	40.7	27	29.6	27	37.0	27	33.3	27	40.7
Greece	72	0	72	0	72	0	72	0	71	0	72	13.9	72	100	72	8.3
Luxembourg	15	0	_	_	13	0	15	13.3	15	0	13	0	15	0	15	0
Slovenia	63	12.7	-	_	63	1.6	63	77.8	63	0	63	77.8	63	77.8	63	77.8
Iceland	10	0	-	_	10	0	10	0	_	-	-	-	10	0	-	-

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.



Appendix Table QSA6. Resistance (%) to ampicillin, cefotaxime, chloramphenicol, ciprofloxacin, gentamicin, nalidixic acid, sulfonamides and tetracyclines among Salmonella spp. isolates from pigs and cattle in countries reporting qualitative data in 2012

Country	Am	Ampicillin (Cefotaxime		Chloramphenicol		floxacin	Gent	amicin	Nalidi	xic acid	Sulfo	namides	Tetra	cyclines
Country	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res	Ν	% Res
Pigs	•															
Austria	40	12.5	40	0	40	0	40	0	40	0	40	2.5	40	20.0	40	22.5
Iceland	11	45.5	_	_	11	0	11	0	_	_	-	_	-	_	_	_
Cattle	•				•											
Austria	38	0	38	0	38	0	38	0	38	5.3	38	0	38	0	38	0

MS: Member State; N: number of isolates tested; % Res: percentage of resistant isolates; -: no data reported.



2.4. Discussion

Very few countries reported qualitative data for *Salmonella* in 2012. Furthermore, it is difficult to compare accurately the data collected using disc diffusion techniques with those deriving from dilution methods and collected quantitatively as MIC data. Therefore, as in previous years, a detailed analysis and interpretation of the results has not been undertaken.

Greece used CLSI disc diffusion methods to test the *Salmonella* isolates recovered from *Gallus gallus*, and interpreted the results using CLSI breakpoints. The results will not be directly comparable to the results obtained by MSs performing broth microdilution MIC determinations and applying EUCAST ECOFFs to interpret those results and have therefore been presented separately.



APPENDIX 3. Frequency distribution of complete susceptibility and multiple resistance

Appendix Table MDR1. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella spp. from meat from broilers (Gallus gallus) in MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Resistant to 1 AMB		Resistant to 2 AMB		Resistant	to 3 AMB	Resistant	to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Czech Republic (N=47)	30	63.8	1	2.1	0	0	9	19.2	6	12.8
Germany (N=94)	23	24.5	7	7.5	3	3.2	22	23.4	13	13.8
Ireland (N=70)	51	72.9	3	4.3	0	0	8	11.4	1	1.4
Romania (N=188)	13	6.9	4	2.1	8	4.3	45	23.9	53	28.2

Country	Resistant	to 5 AMB	Resistant	to 6 AMB	Resistant	to 7 AMB	Resistant	to 8 AMB	Resistant	to 9 AMB
Country	n	%	n	%	n	%	n	%	n	%
Czech Republic (N=47)	1	2.1	0	0	0	0	0	0	0	0
Germany (N=94)	9	9.6	15	16.0	1	1.1	1	1.1	0	0
Ireland (N=70)	6	8.6	1	1.4	0	0	0	0	0	0
Romania (N=188)	42	22.3	19	10.1	3	1.6	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR2. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella spp. from meat from pigs in MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Resistant to 1 AMB		Resistant to 2 AMB		Resistant	to 3 AMB	Resistant	to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Czech Republic (N=33)	12	36.4	2	6.1	5	15.2	2	6.1	6	18.2
Denmark (N=41)	12	29.3	3	7.3	2	4.9	9	22.0	14	34.2
Estonia (N=22)	19	86.4	0	0	0	0	2	9.1	1	4.6
Germany (N=163)	62	38.0	15	9.2	7	4.3	9	5.5	44	27.0
Ireland (N=69)	15	21.7	6	8.7	7	10.1	5	7.3	21	30.4
Italy (N=85)	32	37.6	9	10.6	1	1.2	9	10.6	21	24.7
Romania (N=125)	30	24.0	24	19.2	1	0.8	33	26.4	18	14.4

Country	Resistant	to 5 AMB	Resistant to 6 AMB		Resistant to 7 AMB		Resistant	to 8 AMB	Resistant	to 9 AMB
Country	n	%	n	%	n	%	n	%	n	%
Czech Republic (N=33)	5	15.2	1	3.0	0	0	0	0	0	0
Denmark (N=41)	0	0	1	2.4	0	0	0	0	0	0
Estonia (N=22)	0	0	0	0	0	0	0	0	0	0
Germany (N=163)	17	10.4	9	5.5	0	0	0	0	0	0
Ireland (N=69)	6	8.7	7	10.1	2	2.9	0	0	0	0
Italy (N=85)	7	8.2	5	5.9	1	1.2	0	0	0	0
Romania (N=125)	14	11.2	5	4.0	0	0	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR3. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella spp. from broilers in MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Resistant to 1 AMB		Resistant to 2 AMB		Resistant	to 3 AMB	Resistant	to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=113)	82	72.6	7	6.2	1	0.9	16	14.2	7	6.2
Czech Republic (N=351)	271	77.2	11	3.1	2	0.6	35	10.0	24	6.8
Denmark (N=24)	17	70.8	0	0	0	0	0	0	7	29.2
Hungary (N=175)	13	7.4	19	10.9	15	8.6	90	51.4	35	20.0
Ireland (N=38)	31	81.6	3	7.9	1	2.6	1	2.6	2	5.3
Italy (N=105)	55	52.4	2	1.9	5	4.8	10	9.5	11	10.5
Romania (N=781)	135	17.3	99	12.8	76	9.7	117	15.0	137	17.5
Spain (N=29)	6	20.7	11	37.9	2	6.9	1	3.5	4	13.8
United Kingdom (N=17)	5	29.4	4	23.5	6	35.3	2	11.8	0	0

Country	Resistant	to 5 AMB	Resistant to 6 AMB		Resistant to 7 AMB		Resistant	to 8 AMB	Resistant	to 9 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=113)	0	0	0	0	0	0	0	0	0	0
Czech Republic (N=351)	4	1.1	4	1.1	0	0	0	0	0	0
Denmark (N=24)	0	0	0	0	0	0	0	0	0	0
Hungary (N=175)	3	1.7	0	0	0	0	0	0	0	0
Ireland (N=38)	0	0	0	0	0	0	0	0	0	0
Italy (N=105)	5	4.8	13	12.4	2	1.9	2	1.9	0	0
Romania (N=781)	100	12.8	79	10.1	26	3.3	7	0.9	8	1.0
Spain (N=29)	4	13.8	1	3.5	0	0	0	0	0	0
United Kingdom (N=17)	0	0	0	0	0	0	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR4. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella spp. from laying hens in MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Resistant to 1 AMB		Resistant to 2 AMB		Resistant	to 3 AMB	Resistant	to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=63)	47	74.6	6	9.5	1	1.6	3	4.8	4	6.4
Germany (N=51)	44	86.3	2	3.9	4	7.8	0	0	0	0
Hungary (N=86)	59	68.6	7	8.1	4	4.7	11	12.8	1	1.2
Italy (N=161)	111	68.9	24	14.9	3	1.9	10	6.2	8	5.0
Romania (N=145)	72	49.7	14	9.7	4	2.8	16	11.0	25	17.2
Spain (N=150)	119	79.3	21	14.0	2	1.3	1	0.7	2	1.3
United Kingdom (N=11)	10	90.9	0	0	0	0	1	9.1	0	0

Country	Resistant	to 5 AMB	Resistant to 6 AMB		Resistant to 7 AMB		Resistant	to 8 AMB	Resistant	to 9 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=63)	2	3.2	0	0	0	0	0	0	0	0
Germany (N=51)	0	0	1	2.0	0	0	0	0	0	0
Hungary (N=86)	1	1.2	3	3.5	0	0	0	0	0	0
Italy (N=161)	2	1.2	3	1.9	0	0	0	0	0	0
Romania (N=145)	10	6.9	4	2.8	0	0	0	0	0	0
Spain (N=150)	4	2.7	0	0	0	0	1	0.7	0	0
United Kingdom (N=11)	0	0	0	0	0	0	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR5. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella Enteritidis from broilers in MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Resistant	to 1 AMB	Resistant to 2 AMB		Resistant	to 3 AMB	Resistant	to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=21)	20	95.2	1	4.8	0	0	0	0	0	0
Czech Republic (N=236)	231	97.9	4	1.7	1	0.4	0	0	0	0
Romania (N=10)	4	40.0	0	0	1	10.0	4	40.0	0	0

Country	Resistant	to 5 AMB	Resistant to 6 AMB		Resistant	to 7 AMB	Resistant	to 8 AMB	Resistant	to 9 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=21)	0	0	0	0	0	0	0	0	0	0
Czech Republic (N=236)	0	0	0	0	0	0	0	0	0	0
Romania (N=10)	0	0	0	0	0	0	1	10.0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR6. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella Enteritidis from laying hens in MSs reporting isolate-based data, 2012

Country	Suscept	ible to all	Resistant	to 1 AMB	Resistant	to 2 AMB	Resistant	to 3 AMB	Resistant	to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=15)	15	100	0	0	0	0	0	0	0	0
Germany (N=21)	21	100	0	0	0	0	0	0	0	0
Hungary (N=25)	23	92.0	1	4	1	4.0	0	0	0	0
Italy (N=28)	22	78.6	6	21.4	0	0	0	0	0	0
Romania (N=66)	49	74.2	5	7.6	1	1.5	5	7.6	5	7.6
Spain (N=43)	32	74.4	9	20.9	1	2.3	0	0	0	0

Country	Resistant	to 5 AMB	Resistant	to 6 AMB	Resistant	to 7 AMB	Resistant	to 8 AMB	Resistant	to 9 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=15)	0	0	0	0	0	0	0	0	0	0
Germany (N=21)	0	0	0	0	0	0	0	0	0	0
Hungary (N=25)	0	0	0	0	0	0	0	0	0	0
Italy (N=28)	0	0	0	0	0	0	0	0	0	0
Romania (N=66)	1	1.5	0	0	0	0	0	0	0	0
Spain (N=43)	1	2.3	0	0	0	0	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR7. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella spp. from turkeys in MSs reporting isolate-based data, 2012

Country	Suscep	tible to all	Resistar	nt to 1 AMB	Resistan	t to 2 AMB	Resistant	to 3 AMB	Resistan	t to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=38)	7	18.4	21	55.3	0	0	1	2.6	1	2.6
Czech Republic (N=27)	2	7.4	10	37.0	4	14.8	2	7.4	1	3.7
Germany (N=87)	20	23.0	10	11.5	6	6.9	5	5.8	31	35.6
Hungary (N=174)	11	6.3	53	30.5	14	8.1	52	29.9	6	3.5
Ireland (N=14)	12	85.7	0	0	0	0	2	14.3	0	0
Italy (N=48)	0	0	4	8.3	5	10.4	6	12.5	19	39.6
Spain (N=169)	3	1.8	2	1.2	3	1.8	5	3.0	23	13.6

Country	Resistan	t to 5 AMB	Resistant	to 6 AMB	Resistant	to 7 AMB	Resistan	t to 8 AMB	Resistant	to 9 AMB
Country	n	%	n	%	n	%	n	%	n	%
Austria (N=38)	8	21.1	0	0	0	0	0	0	0	0
Czech Republic (N=27)	2	7.4	6	22.2	0	0	0	0	0	0
Germany (N=87)	15	17.2	0	0	0	0	0	0	0	0
Hungary (N=174)	17	9.8	21	12.1	0	0	0	0	0	0
Ireland (N=14)	0	0	0	0	0	0	0	0	0	0
Italy (N=48)	9	18.8	2	4.2	0	0	3	6.3	0	0
Spain (N=169)	42	24.9	76	45.0	14	8.3	0	0	1	0.6

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR8. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella spp. from fattening pigs in MSs reporting isolate-based data, 2012

Country	Suscep	tible to all	Resistan	t to 1 AMB	Resistan	t to 2 AMB	Resistar	nt to 3 AMB	Resistan	t to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Denmark (N=374)	179	47.9	53	14.2	15	4.0	31	8.3	74	19.8
Estonia (N=14)	9	64.3	0	0	0	0	4	28.6	1	7.1
Germany (N=627)	89	14.2	28	4.5	28	4.5	24	3.8	220	35.1
Hungary (N=38)	9	23.7	1	2.6	1	2.6	11	29.0	11	29.0
Ireland (N=24)	2	8.3	6	25.0	1	4.2	3	12.5	4	16.7
Italy (N=25)	10	40.0	2	8.0	1	4.0	4	16.0	4	16.0
Spain (N=48)	3	6.2	13	27.1	4	8.3	1	2.1	11	22.9

Country	Resistan	t to 5 AMB	Resistant to 6 AMB		Resistant to 7 AMB		Resistan	t to 8 AMB	Resistant to 9 AMB	
Country	n	%	n	%	n	%	n	%	n	%
Denmark (N=374)	18	4.8	4	1.1	0	0	0	0	0	0
Estonia (N=14)	0	0	0	0	0	0	0	0	0	0
Germany (N=627)	162	25.8	52	8.3	23	3.7	0	0	1	0.2
Hungary (N=38)	4	10.5	1	2.6	0	0	0	0	0	0
Ireland (N=24)	3	12.5	1	4.2	4	16.7	0	0	0	0
Italy (N=25)	1	4.0	2	8.0	0	0	1	4.0	0	0
Spain (N=48)	9	18.8	5	10.4	1	2.1	1	2.1	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR9. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella Typhimurium from fattening pigs in MSs reporting isolate-based data, 2012

Country	Suscep	tible to all	Resistan	t to 1 AMB	Resistan	t to 2 AMB	Resistan	t to 3 AMB	Resistan	t to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Denmark (N=63)	31	49.2	1	1.6	4	6.4	9	14.3	8	12.7
Germany (N=273)	24	8.8	9	3.3	19	7.0	6	2.2	47	17.2
Ireland (N=15)	0	0	4	26.7	1	6.7	1	6.7	1	6.7

Country	Resistan	t to 5 AMB	Resistan	t to 6 AMB	Resistan	t to 7 AMB	Resistan	t to 8 AMB	Resistant to 9 AMB	
Country	n	%	n	%	n	%	n	%	n	%
Denmark (N=63)	9	14.3	1	1.6	0	0	0	0	0	0
Germany (N=273)	126	46.2	32	11.7	10	3.7	0	0	0	0
Ireland (N=15)	3	20.0	1	6.7	4	26.7	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR10. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella spp. from cattle in MSs and non-MS reporting isolate-based data, 2012

Country	Suscep	tible to all	Resistan	Resistant to 1 AMB		t to 2 AMB	Resistant to 3 AMB		Resistant to 4 AMB	
Country	n	%	n	%	n	%	n	%	n	%
Belgium (N=42)	8	19.0	4	9.5	9	21.4	7	16.7	5	11.9
Finland (N=19)	15	78.9	0	0	2	10.5	0	0	1	5.3
Germany (N=68)	45	66.2	8	11.8	1	1.5	3	4.4	8	11.8
Ireland (N=36)	12	33.3	2	5.6	1	2.8	0	0	5	13.9
Italy (N=14)	8	57.1	0	0	0	0	0	0	2	14.3
Sweden (N=17)	14	82.4	0	0	0	0	1	5.9	0	0

Country	Resistan	t to 5 AMB	Resistan	nt to 6 AMB	Resistan	t to 7 AMB	Resistan	t to 8 AMB	Resistant to 9 AMB	
Country	n	%	n	%	n	%	n	%	n	%
Belgium (N=42)	5	11.9	2	4.8	2	4.8	0	0	0	0
Finland (N=19)	1	5.3	0	0	0	0	0	0	0	0
Germany (N=68)	2	2.9	0	0	1	1.5	0	0	0	0
Ireland (N=36)	15	41.7	1	2.8	0	0	0	0	0	0
Italy (N=14)	3	21.4	1	7.1	0	0	0	0	0	0
Sweden (N=17)	2	11.8	0	0	0	0	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR11. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in Salmonella Typhimurium from cattle in MSs reporting isolate-based data, 2012

Country	Suscept	tible to all	Resistan	t to 1 AMB	Resistan	t to 2 AMB	Resistan	t to 3 AMB	Resistan	t to 4 AMB
Country	n	%	n	%	n	%	n	%	n	%
Belgium (N=25)	1	4.0	4	16.0	8	32.0	4	16.0	2	8.0
Finland (N=16)	12	75.0	0	0	2	12.5	0	0	1	6.3
Germany (N=35)	24	68.6	7	20.0	1	2.9	0	0	0	0
Ireland (N=24)	7	29.2	0	0	1	4.2	0	0	0	0
Sweden (N=12)	9	75.0	0	0	0	0	1	8.3	0	0

Country	Resistan	t to 5 AMB	Resistan	Resistant to 6 AMB		t to 7 AMB	Resistan	t to 8 AMB	Resistant to 9 AMB		
Country	n	%	n	%	n	%	n	%	n	%	
Belgium (N=25)	4	16.0	1	4.0	1	4.0	0	0	0	0	
Finland (N=16)	1	6.3	0	0	0	0	0	0	0	0	
Germany (N=35)	2	5.7	0	0	1	2.9	0	0	0	0	
Ireland (N=24)	15	62.5	1	4.2	0	0	0	0	0	0	
Sweden (N=12)	2	16.7	0	0	0	0	0	0	0	0	

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR13. Frequency distribution of completely susceptible isolates and resistant isolates to from one to five antimicrobials, in Campylobacter jejuni from broilers in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscep	otible to all	Resistant to 1 AMB		Resistant to 2 AMB		Resistant to 3 AMB		Resista	nt to 4 AMB	Resistant to 5 AMB	
Country	n	%	n	%	n	%	n	%	n	%	n	%
Austria (N=108)	22	20.4	58	53.7	24	22.2	4	3.7	0	0	0	0
Denmark (N=41)	32	78.0	6	14.6	3	7.3	0	0	0	0	0	0
Hungary (N=46)	4	8.7	24	52.2	18	39.1	0	0	0	0	0	0
Spain (N=32)	1	3.1	2	6.3	26	81.3	2	6.3	0	0	1	3.1
Sweden (N=100)	80	80.0	20	20.0	0	0	0	0	0	0	0	0
Switzerland (N=171)	98	57.3	51	29.8	21	12.3	1	0.6	0	0	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.

AMB: antimicrobial substance(s).

Appendix Table MDR14. Frequency distribution of completely susceptible isolates and resistant isolates to from one to five antimicrobials, in Campylobacter coli from broilers in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscep	Susceptible to all		Resistant to 1 AMB		Resistant to 2 AMB		Resistant to 3 AMB		nt to 4 AMB	Resistant to 5 AMB	
Country	n	%	n	%	n	%	n	%	n	%	n	%
Austria (N=33)	6	18.2	16	48.5	10	30.3	1	3.0	0	0	0	0
Hungary (N=63)	9	14.3	23	36.5	28	44.4	3	4.8	0	0	0	0
Spain (N=54)	1	1.9	0	0	23	42.6	18	33.3	8	14.8	4	7.4
Switzerland (N=14)	4	28.6	1	7.1	6	42.9	2	14.3	1	7.1	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR15. Frequency distribution of completely susceptible isolates and resistant isolates to from one to five antimicrobials, in Campylobacter coli from fattening pigs in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscep	tible to all	Resistant to 1 AMB		Resistant to 2 AMB		Resistant to 3 AMB		Resistar	nt to 4 AMB	Resistant to 5 AMB	
Country	n	%	n	%	n	%	n	%	n	%	n	%
Denmark (N=103)	34	33.0	50	48.5	15	14.6	4	3.9	0	0	0	0
Hungary (N=53)	0	0	9	17.0	18	34.0	21	39.6	4	7.6	1	1.9
Spain (N=73)	0	0	0	0	2	2.7	19	26.0	44	60.3	8	11.0
Switzerland (N=144)	21	14.6	55	38.2	40	27.8	26	18.1	2	1.4	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.

AMB: antimicrobial substance(s).

Appendix Table MDR16. Frequency distribution of completely susceptible isolates and resistant isolates to from one to five antimicrobials, in Campylobacter jejuni from cattle in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscep	tible to all	Resistant to 1 AMB		Resistant to 2 AMB		Resistant to 3 AMB		Resistar	nt to 4 AMB	Resistant to 5 AMB	
Country	n	%	n	%	n	%	n	%	n	%	n	%
Denmark (N=89)	75	84.3	14	15.7	0	0	0	0	0	0	0	0
Germany (N=73)	12	16.4	29	39.7	23	31.5	8	11.0	1	1.4	0	0
Spain (N=68)	17	25.0	18	26.5	29	42.7	4	5.9	0	0	0	0
Switzerland (N=38)	18	47.4	9	23.7	9	23.7	1	2.6	1	2.6	0	0

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR17. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in E. coli from broilers in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscept	tible to all	Resistant to 1 AMB		Resistan	t to 2 AMB	Resistan	t to 3 AMB	Resistant to 4 AMB		
Country	n	%	n	%	n	%	n	%	n	%	
Austria (N=130)	22	16.9	30	23.1	22	16.9	13	10.0	22	16.9	
Denmark (N=115)	65	56.5	30	26.1	5	4.4	9	7.8	5	4.4	
Hungary (N=104)	14	13.5	21	20.2	20	19.2	13	12.5	17	16.4	
Switzerland (N=185)	49	26.5	58	31.4	32	17.3	23	12.4	8	4.3	

Country	Resistan	t to 5 AMB	Resistan	t to 6 AMB	Resistan	t to 7 AMB	Resistan	t to 8 AMB	Resistant to 9 AMB		
Country	n	%	n	%	n	%	n	%	n	%	
Austria (N=130)	7	5.4	12	9.2	2	1.5	0	0	0	0	
Denmark (N=115)	1	0.9	0	0	0	0	0	0	0	0	
Hungary (N=104)	8	7.7	8	7.7	2	1.9	1	1.0	0	0	
Switzerland (N=185)	8	4.3	5	2.7	2	1.1	0	0	0	0	

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



Appendix Table MDR18. Frequency distribution of completely susceptible isolates and resistant isolates to from one to nine antimicrobials, in E. coli from fattening pigs in MSs and one non-MS reporting isolate-based data, 2012

Country	Suscept	tible to all	Resistan	t to 1 AMB	Resistan	t to 2 AMB	Resistan	t to 3 AMB	Resistant to 4 AMB		
Country	n	%	n	%	n	%	n	%	n	%	
Austria (N=140)	53	37.9	21	15.0	32	22.9	18	12.9	6	4.3	
Denmark (N=152)	63	41.5	25	16.5	15	9.9	14	9.2	17	11.2	
Hungary (N=68)	16	23.5	11	16.2	7	10.3	12	17.7	10	14.7	
Switzerland (N=185)	80	43.2	19	10.3	22	11.9	28	15.1	21	11.4	

Country	Resistan	t to 5 AMB	Resistan	t to 6 AMB	Resistan	t to 7 AMB	Resistan	t to 8 AMB	Resistant to 9 AMB		
Country	n	%	n	%	n	%	n	%	n	%	
Austria (N=140)	6	4.3	4	2.9	0	0	0	0	0	0	
Denmark (N=152)	16	10.5	2	1.3	0	0	0	0	0	0	
Hungary (N=68)	7	10.3	4	5.9	1	1.5	0	0	0	0	
Switzerland (N=185)	13	7.0	2	1.1	0	0	0	0	0	0	

N: total number of isolates tested for susceptibility against the whole common set of antimicrobial substances.

n: number of resistant isolates per category of susceptibility or multiple resistance.

%: percentage of resistant isolates per category of susceptibility or multiple resistance.



APPENDIX 4. Multi-resistance patterns

Appendix Table MDRP1. Multi-resistance patterns of interest in Salmonella spp. from meat from broilers in MSs reporting isolate-based data, 2012

		Mult	i-resi	stand	ce pa	ttern				MS group (N=399))	Czech Republic (N=47)	Germany (N=94)	lreland (N=70)	Romania (N=188)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
			R			R	R		65	25.3	16.3	8	14	0	43
			R			R	R	R	43	16.7	10.8	0	2	0	41
			R		R	R	R	R	26	10.1	6.5	0	0	0	26
			R		R	R	R		18	7.0	4.5	6	2	0	10
R		R	R			R	R	R	16	6.2	4.0	0	14	0	2
R	R		R			R	R		10	3.9	2.5	0	8	2	0
R			R	R	R	R	R		8	3.1	2.0	0	0	1	7
			R		R			R	7	2.7	1.8	0	5	2	0
R			R		R	R	R		7	2.7	1.8	1	0	0	6
R			R			R		R	6	2.3	1.5	0	6	0	0
R						R		R	5	1.9	1.3	0	1	2	2
R		R				R			5	1.9	1.3	0	1	4	0
		R	R		R	R	R		5	1.9	1.3	0	0	0	5
R			R		R	R	R	R	5	1.9	1.3	0	0	0	5
		R	R		R	R	R	R	4	1.6	1.0	0	0	0	4
		R	R			R	R	R	3	1.2	0.8	0	0	0	3
R	R					R	R		3	1.2	0.8	0	3	0	0
R		R	R		R	R	R	R	2	0.8	0.5	0	1	0	1
			R	R		R	R	R	2	0.8	0.5	0	0	0	2
R			R			R	R	R	2	0.8	0.5	0	0	0	2
R			R	R		R		R	2	0.8	0.5	0	0	2	0
R	R	R	R			R	R	R	2	0.8	0.5	0	0	0	2
			R	R	R	R	R		1	0.4	0.3	0	1	0	0
R		R			R	R	R		1	0.4	0.3	0	0	1	0
R					R	R	R		1	0.4	0.3	0	0	0	1

Table continued overleaf.



Appendix Table MDRP1 (continued). Multi-resistance patterns of interest in Salmonella spp. from meat from broilers in MSs reporting isolate-based data, 2012

		Mult	i-resi	stand	ce pa	ttern				MS grou (N=399)	ip	Czech Republic (N=47)	Germany (N=94)	lreland (N=70)	Romania (N=188)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
		R	R				R	R	1	0.4	0.3	0	0	0	1
R						R	R		1	0.4	0.3	1	0	0	0
R						R	R	R	1	0.4	0.3	0	0	1	0
R			R		R	R		R	1	0.4	0.3	0	0	1	0
R			R	R	R	R		R	1	0.4	0.3	0	1	0	0
R		R	R	R		R	R		1	0.4	0.3	0	0	0	1
R		R	R	R	R	R	R	R	1	0.4	0.3	0	1	0	0
R	R		R						1	0.4	0.3	0	1	0	0
	Total							•	257	100	64.4	16	61	16	164

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).



Appendix Table MDRP2. Multi-resistance patterns of interest in Salmonella spp. from meat from pigs in MSs reporting isolate-based data, 2012

		Mult	i-resi	stand	ce pat	ttern				MS gro (N=538	up 3)	Czech Republic (N=33)	Denmark (N=41)	Estonia (N=22)	Germany (N=163)	Ireland (N=69)	Italy (N=85)	Romania (N=125)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
R					R	R	R		106	38.7	19.7	5	14	0	37	19	19	12
					R	R	R		38	13.9	7.1	1	2	2	3	5	8	17
R					R	R	R	R	18	6.6	3.3	2	0	0	7	2	4	3
R		R			R	R	R		18	6.6	3.3	3	0	0	5	2	2	6
R		R			R	R	R	R	16	5.8	3.0	0	1	0	4	5	5	1
R					R	R			12	4.4	2.2	1	7	0	3	0	0	1
R		R			R	R			6	2.2	1.1	0	0	1	5	0	0	0
R		R	R		R	R	R		6	2.2	1.1	1	0	0	5	0	0	0
			R			R	R		5	1.8	0.9	0	0	0	0	0	0	5
R		R R R R R R R R R R R R							5	1.8	0.9	0	0	0	0	0	0	5
R		R		R	R				5	1.8	0.9	0	0	0	0	0	0	5
		R			R	R	R	R	3	1.1	0.6	0	0	0	1	2	0	0
R			R		R	R	R		3	1.1	0.6	0	0	0	2	0	0	1
R		R	R			R	R	R	3	1.1	0.6	0	0	0	0	0	0	3
			R		R	R	R		2	0.7	0.4	1	0	0	0	0	0	1
R					R		R		2	0.7	0.4	0	0	0	1	0	0	1
R				R	R	R	R		2	0.7	0.4	0	0	0	1	0	0	1
R		R	R		R	R	R	R	2	0.7	0.4	0	0	0	0	1	1	0
R						R	R	R	2	0.7	0.4	0	0	0	0	0	2	0
R				R	R	R	R	R	2	0.7	0.4	0	0	0	0	2	0	0
R		R					R		2	0.7	0.4	0	0	0	0	0	0	2
R		R				R	R	R	2	0.7	0.4	0	0	0	0	0	0	2
						R	R	R	1	0.4	0.2	0	0	0	1	0	0	0
					R	R		R	1	0.4	0.2	0	0	0	0	0	1	0
					R	R	R	R	1	0.4	0.2	0	0	0	0	1	0	0
		R		R	R	R	R		1	0.4	0.2	0	0	0	0	0	1	0



Appendix Table MDRP2 (continued). Multi-resistance patterns of interest in Salmonella spp. from meat from pigs in MSs reporting isolate-based data, 2012

		Mult	i-resi	stanc	ce pat	ttern				MS gro (N=538	up 3)	Czech Republic (N=33)	Denmark (N=41)	Estonia (N=22)	Germany (N=163)	Ireland (N=69)	Italy (N=85)	Romania (N=125)
Amp	Ctx	ChI	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
R							R	R	1	0.4	0.2	0	0	0	0	0	0	1
R						R	R		1	0.4	0.2	0	0	0	0	0	0	1
R					R	R		R	1	0.4	0.2	0	0	0	1	0	0	0
R		R				R			1	0.4	0.2	0	0	0	1	0	0	0
R		R				R		R	1	0.4	0.2	0	0	0	0	1	0	0
R		R				R	R		1	0.4	0.2	0	0	0	1	0	0	0
R		R		R	R	R	R	R	1	0.4	0.2	0	0	0	0	1	0	0
R	R					R	R	R	1	0.4	0.2	0	0	0	0	0	0	1
R	R	R			R	R			1	0.4	0.2	0	0	0	1	0	0	0
R	R	R	R			R	R		1	0.4	0.2	0	0	0	0	0	0	1
				Total					274	100	50.9	14	24	3	79	41	43	70

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP3. Multi-resistance patterns of interest in Salmonella spp. from broilers in MSs reporting isolate-based data, 2012

	Mu	lti-r	esis	stan	ice	patt	ern			MS gro (N=1,63	up 33)	Austria (N=113)	Czech Republic (N=351)	Denmark (N=24)	Hungary (N=175)	Ireland (N=38)	ltaly (N=105)	Romania (N=781)	Spain (N=29)	United Kingdom (N=17)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n	n	n
			R			R	R		200	26.4	12.2	16	35	0	86	0	0	63	0	0
			R		R	R	R		83	11.0	5.1	5	19	0	28	0	0	31	0	0
			R			R	R	R	47	6.2	2.9	0	0	0	0	0	2	45	0	0
R			R	R	R	R	R		47	6.2	2.9	0	4	0	0	0	0	43	0	0
R		R	R			R			30	4.0	1.8	0	0	0	0	0	0	30	0	0
R			R	R		R	R		28	3.7	1.7	0	1	0	1	0	0	26	0	0
			R		R	R	R	R	26	3.4	1.6	0	0	0	0	0	0	26	0	0
R			R			R	R		24	3.2	1.5	0	5	0	5	0	2	12	0	0
R			R			R	R	R	18	2.4	1.1	0	0	0	0	0	1	16	1	0
R			R			R			16	2.1	1.0	0	0	0	0	0	0	16	0	0
R			R		R	R	R		15	2.0	0.9	0	3	0	2	0	0	10	0	0
R			R		R		R		12	1.6	0.7	2	0	0	0	0	5	5	0	0
R	R		R			R	R	R	11	1.5	0.7	0	0	0	0	0	10	1	0	0
			R		R		R		10	1.3	0.6	0	0	0	1	0	2	6	1	0
R					R	R	R		10	1.3	0.6	0	0	6	0	0	1	1	2	0
		R	R		R	R	R	R	9	1.2	0.6	0	0	0	0	0	0	9	0	0
R					R	R			8	1.1	0.5	0	0	0	0	0	0	8	0	0
R			R		R	R	R	R	8	1.1	0.5	0	0	0	0	0	2	6	0	0
R	R	R	R	R	R	R	R	R	8	1.1	0.5	0	0	0	0	0	0	8	0	0
R			R				R		7	0.9	0.4	0	0	0	1	0	2	4	0	0
R		R	R	R	R	R	R	R	6	0.8	0.4	0	0	0	0	0	2	4	0	0
R						R	R	R	5	0.7	0.3	0	0	1	0	2	1	1	0	0
R		L	R		R				5	0.7	0.3	0	0	0	0	0	1	4	0	0
R			R	R	R	R	R	R	5	0.7	0.3	0	0	0	0	0	1	4	0	0
R		R	R			R	R		5	0.7	0.3	0	0	0	0	0	0	5	0	0
R		R	R		R	R	R	R	5	0.7	0.3	0	0	0	0	0	0	5	0	0



Appendix Table MDRP3 (continued). Multi-resistance patterns of interest in Salmonella spp. from broilers in MSs reporting isolate-based data, 2012

	Mu	lti-r	esis	stan	ice	patt	ern			MS gro (N=1,63		Austria (N=113)	Czech Republic (N=351)	Denmark (N=24)	Hungary (N=175)	Ireland (N=38)	ltaly (N=105)	Romania (N=781)	Spain (N=29)	United Kingdom (N=17)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n	n	n
R	R		R			R	R		5	0.7	0.3	0	0	0	0	0	0	5	0	0
R	R		R	R	R	R	R		5	0.7	0.3	0	0	0	0	0	0	5	0	0
R	R	R	R		R	R		R	5	0.7	0.3	0	0	0	0	0	0	5	0	0
				R	R	R			4	0.5	0.2	0	0	0	0	0	0	3	0	1
R			R	R		R			4	0.5	0.2	0	0	0	0	0	0	4	0	0
R	R	R	R	R		R			4	0.5	0.2	0	0	0	0	0	0	4	0	0
R			R	R	R	R			3	0.4	0.2	0	0	0	0	0	0	1	2	0
R	R		R						3	0.4	0.2	0	0	0	0	0	1	2	0	0
R			R					R	3	0.4	0.2	0	0	0	0	0	0	3	0	0
R			R	R	R				3	0.4	0.2	0	0	0	0	0	0	3	0	0
			R	R	R	R	R		2	0.3	0.1	0	0	0	0	0	1	1	0	0
			R	R		R	R	R	2	0.3	0.1	0	0	0	0	0	0	2	0	0
			R	R	R	R	R	R	2	0.3	0.1	0	0	0	0	0	0	2	0	0
		R	R			R	R		2	0.3	0.1	0	0	0	2	0	0	0	0	0
		R	R			R	R	R	2	0.3	0.1	0	0	0	0	0	0	2	0	0
R						R		R	2	0.3	0.1	0	0	0	0	0	2	0	0	0
R			R	R		R	R	R	2	0.3	0.1	0	0	0	0	0	0	2	0	0
R		R			R	R	R		2	0.3	0.1	0	0	0	0	0	2	0	0	0
R		R	R						2	0.3	0.1	0	0	0	0	0	0	2	0	0
R		R	R			R		R	2	0.3	0.1	0	0	0	0	0	0	2	0	0
R		R	R			R	R	R	2	0.3	0.1	0	0	0	0	0	0	2	0	0
R		R	R		R	R	R		2	0.3	0.1	0	0	0	0	0	0	2	0	0
R		R	R	R		R	R		2	0.3	0.1	0	0	0	0	0	0	2	0	0
R	R	R	R		R	R	R		2	0.3	0.1	0	0	0	0	0	0	2	0	0
R	R	R	R		R	R	R	R	2	0.3	0.1	0	0	0	0	0	0	2	0	0
						R	R	R	1	0.1	0.1	0	0	0	0	1	0	0	0	0
					R	R		R	1	0.1	0.1	0	0	0	0	0	0	0	0	1
					R	R	R		1	0.1	0.1	0	0	0	0	0	1	0	0	0



Appendix Table MDRP3 (continued). Multi-resistance patterns of interest in Salmonella spp. from broilers in MSs reporting isolate-based data, 2012

	Mu	lti-r	esis	stan	ce	patt	ern			MS gro (N=1,6	oup 33)	Austria (N=113)	Czech Republic (N=351)	Denmark (N=24)	Hungary (N=175)	Ireland (N=38)	Italy (N=105)	Romania (N=781)	Spain (N=29)	United Kingdom (N=17)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n	n	n
			R				R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
			R		R	R			1	0.1	0.1	0	0	0	1	0	0	0	0	0
			R	R			R		1	0.1	0.1	0	0	0	1	0	0	0	0	0
			R	R		R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
			R	R		R	R		1	0.1	0.1	0	0	0	0	0	0	0	1	0
			R	R	R	R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
		R				R	R		1	0.1	0.1	0	0	0	0	0	1	0	0	0
		R	R		R	R		R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
		R	R	R	R	R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
R					R		R		1	0.1	0.1	0	0	0	0	0	0	1	0	0
R					R		R	R	1	0.1	0.1	0	0	0	0	0	0	0	1	0
R				R	R				1	0.1	0.1	0	0	0	0	0	0	1	0	0
R				R	R		R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R				R	R	R	R		1	0.1	0.1	0	0	0	0	0	1	0	0	0
R			R				R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R			R	R	R		R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R			R	R	R	R		R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R		R				R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
R		R	R	R			R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R		R	R	R		R	R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R		R	R	R	R		R		1	0.1	0.1	0	0	0	0	0	0	0	1	0



Appendix Table MDRP3 (continued). Multi-resistance patterns of interest in Salmonella spp. from broilers in MSs reporting isolate-based data, 2012

	Mu	lti-r	esis	stan	ice	oatt	ern			MS gro (N=1,63	up 33)	Austria (N=113)	Czech Republic (N=351)	Denmark (N=24)	Hungary (N=175)	lreland (N=38)	ltaly (N=105)	Romania (N=781)	Spain (N=29)	United Kingdom (N=17)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n	n	n
R		R	R	R	R	R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
R		R	R	R	R	R	R		1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R						R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R					R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R				R			R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R			R	R	R	R		1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R		R			R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R		R		R		R		1	0.1	0.1	0	0	0	0	0	0	0	1	0
R	R		R		R	R	R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R		R	R		R	R		1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R		R	R		R	R	R	1	0.1	0.1	0	0	0	0	0	1	0	0	0
R	R		R	R	R	R			1	0.1	0.1	0	0	0	0	0	1	0	0	0
R	R		R	R	R	R		R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R	R	R			R			1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R	R	R			R	R	R	1	0.1	0.1	0	0	0	0	0	0	1	0	0
R	R	R	R	R	R	R	R		1	0.1	0.1	0	0	0	0	0	0	1	0	0
			1	lota	ıl				757	100	46.4	23	67	7	128	3	43	474	10	2

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP4. Multi-resistance patterns of interest in Salmonella spp. from laying hens in MSs reporting isolate-based data, 2012

	Μι	ulti-r	esis	stan	ce p	oatte	ern			MS gro (N=667	up 7)	Austria (N=63)	Germany (N=51)	Hungary (N=86)	ltaly (N=161)	Romania (N=145)	Spain (N=150)	United Kingdom (N=11)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
			R			R	R		20	17.7	3.0	3	0	8	0	9	0	0
R			R				R		11	9.7	1.6	0	0	3	6	2	0	0
			R			R	R	R	11	9.7	1.6	3	0	0	0	8	0	0
R			R		R		R		8	7.1	1.2	0	0	0	0	8	0	0
R					R	R	R		7	6.1	1.0	0	0	0	4	0	2	0
R			R		R	R	R		5	4.4	0.8	0	0	0	2	2	1	0
			R		R	R	R		5	4.4	0.8	0	0	1	0	4	0	0
R			R	R		R	R		5	4.4	0.8	0	0	0	0	5	0	0
R			R	R	R	R	R		4	3.5	0.6	0	0	2	0	2	0	0
R			R			R	R		4	3.5	0.6	0	0	0	0	4	0	0
R		R			R	R	R		3	2.6	0.4	2	0	1	0	0	0	0
			R		R		R		3	2.6	0.4	0	0	0	0	3	0	0
R		R				R	R		3	2.6	0.4	0	0	0	3	0	0	0
R		R	R			R	R		3	2.6	0.4	0	0	0	0	3	0	0
					R	R	R		2	1.8	0.3	0	0	0	1	0	0	1
R			R		R	R	R	R	2	1.8	0.3	0	0	1	1	0	0	0
R			R			R			2	1.8	0.3	0	0	0	0	2	0	0
R		R	R			R	R	R	2	1.8	0.3	0	0	0	2	0	0	0
R	R		R						2	1.8	0.3	0	0	0	2	0	0	0
						R	R	R	1	0.9	0.2	0	0	0	0	0	1	0
				R	R	R	R		1	0.9	0.2	1	0	0	0	0	0	0
			R	R		R	R		1	0.9	0.2	0	0	0	0	1	0	0
			R	R	R	R	R		1	0.9	0.2	0	0	0	0	0	1	0
R						R		R	1	0.9	0.2	0	0	0	1	0	0	0
R					R	R	R	R	1	0.9	0.2	0	0	0	0	0	1	0
R			R			R		R	1	0.9	0.2	0	0	0	1	0	0	0
R			R			R	R	R	1	0.9	0.2	0	0	0	0	0	1	0
R			R	R	R		R	R	1	0.9	0.2	0	0	0	0	1	0	0
R		R			R	R	R	R	1	0.9	0.2	0	1	0	0	0	0	0
R		R	R		R	R	R		1	0.9	0.2	0	0	0	0	1	0	0
R		R	R	R	R	R	R	R	5	4.4	0.8	0	0	0	0	0	1	0
				Γota	l				113	100	16.9	9	1	16	23	55	8	1

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP5. Multi-resistance patterns of interest in Salmonella spp. from turkeys in MSs reporting isolate-based data, 2012

	М	ulti-	resi	stan	ce p	oatte	rn			MS gro (N=55		Austria (N=38)	Czech Republic (N=27)	Germany (N=87)	Hungary (N=174)	Ireland (N=14)	ltaly (N=48)	Spain (N=169)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
R		R	R			R	R	R	69	18.6	12.4	0	0	0	0	0	0	69
R			R			R	R	R	41	11.1	7.4	0	0	2	0	0	0	39
R			R				R		39	10.5	7.0	0	2	0	37	0	0	0
R			R	R	R	R	R		30	8.1	5.4	0	6	0	21	0	1	2
R					R	R	R		20	5.4	3.6	0	0	17	0	0	2	1
			R			R	R		19	5.1	3.4	1	0	3	15	0	0	0
R			R		R		R		19	5.1	3.4	0	1	0	0	0	1	17
R			R	R		R	R		18	4.9	3.2	0	2	1	15	0	0	0
R						R	R	R	15	4.1	2.7	0	0	2	0	0	11	2
R			R	R	R	R			13	3.5	2.3	3	0	10	0	0	0	0
R		R	R		R	R	R	R	13	3.5	2.3	0	0	0	0	0	0	13
R				R	R		R		8	2.2	1.4	0	0	5	0	0	3	0
			R		R		R		6	1.6	1.1	0	0	0	0	0	5	1
			R		R	R	R		5	1.4	0.9	1	0	1	3	0	0	0
R		R				R			5	1.4	0.9	0	0	1	0	0	0	4
R			R		R	R		R	5	1.4	0.9	5	0	0	0	0	0	0
					R	R	R		3	0.8	0.5	0	0	0	0	2	1	0
R				R	R	R	R		3	0.8	0.5	0	0	0	0	0	3	0
R		_	R	R		R			3	0.8	0.5	0	0	3	0	0	0	0
R		R	R		R	R	R		3	0.8	0.5	0	0	0	0	0	0	3
R			R		R	R	R		2	0.5	0.4	0	0	1	1	0	0	0
		R	<u> </u>		R	R	R	R	2	0.5	0.4	0	0	0	0	0	2	0
R					R	R		R	2	0.5	0.4	0	0	2	0	0	0	0
R		<u> </u>	R			R	R		2	0.5	0.4	0	0	0	2	0	0	0
R			R	R	R		R		2	0.5	0.4	0	0	0	0	0	2	0
R		R			R	R			2	0.5	0.4	0	0	0	0	0	0	2
R			R		R	R			2	0.5	0.4	0	0	1	0	0	1	0



Appendix Table MDRP5 (continued). Multi-resistance patterns of interest in Salmonella spp. from turkeys in MSs reporting isolate-based data, 2012

	M	ulti-	resi	stan	ce p	oatte	rn			MS grou (N=557)	ip)	Austria (N=38)	Czech Republic (N=27)	Germany (N=87)	Hungary (N=174)	Ireland (N=14)	Italy (N=48)	Spain (N=169)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
R		R	R	R	R	R	R	R	2	0.5	0.4	0	0	0	0	0	2	0
						R	R	R	1	0.3	0.2	0	0	1	0	0	0	0
			R		R		R	R	1	0.3	0.2	0	0	0	0	0	1	0
			R		R	R	R	R	1	0.3	0.2	0	0	0	1	0	0	0
R					R	R	R	R	1	0.3	0.2	0	0	0	0	0	1	0
R			R			R		R	1	0.3	0.2	0	0	0	1	0	0	0
R			R		R		R	R	1	0.3	0.2	0	0	0	0	0	0	1
R			R		R	R	R	R	1	0.3	0.2	0	0	0	0	0	1	0
R			R	R		R	R	R	1	0.3	0.2	0	0	0	0	0	0	1
R		R				R	R		1	0.3	0.2	0	0	0	0	0	0	1
R		R				R	R	R	1	0.3	0.2	0	0	0	0	0	0	1
R		R			R	R	R		1	0.3	0.2	0	0	1	0	0	0	0
R		R			R	R	R	R	1	0.3	0.2	0	0	0	0	0	0	1
R		R	R			R	R		1	0.3	0.2	0	0	0	0	0	0	1
R	R		R	R		R	R	R	1	0.3	0.2	0	0	0	0	0	0	1
R	R	R				R	R		1	0.3	0.2	0	0	0	0	0	1	0
R	R	R	R		R	R	R	R	1	0.3	0.2	0	0	0	0	0	1	0
R	R	R	R	R	R	R	R	R	1	0.3	0.2	0	0	0	0	0	0	1
	Mare		•	Tota	I				370	100	66.4	10	11	51	96	2	39	161

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP6. Multi-resistance patterns in Salmonella spp. from fattening pigs in MSs reporting isolate-based data, 2012

	Mu	lti-r	esis	tan	ce I	patt	ern			MS grou (N=1,15		Denmark (N=374)	Estonia (N=14)	Germany (N=627)	Hungary (N=38)	Ireland (N=24)	ltaly (N=25)	Spain (N=48)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group%	n	n	n	n	n	n	n
R					R	R	R		280	40.6	24.3	60	0	198	8	4	2	8
R					R	R	R	R	90	13.0	7.8	4	0	81	1	0	1	3
R		R			R	R	R		75	10.9	6.5	9	0	60	3	3	0	0
R		R			R	R	R	R	29	4.2	2.5	2	0	25	0	1	0	1
R					R	R			24	3.5	2.1	13	0	7	4	0	0	0
R		R	R		R	R	R	R	17	2.5	1.5	3	1	10	1	0	1	1
R						R	R	R	17	2.5	1.5	0	0	14	0	3	0	0
					R	R	R		12	1.7	1.0	3	1	3	0	2	3	0
R			R		R	R	R		11	1.6	1.0	7	0	3	1	0	0	0
R		R		R	R	R	R		11	1.6	1.0	1	0	9	0	0	1	0
R		R		R	R	R	R	R	11	1.6	1.0	0	0	11	0	0	0	0
R			R		R	R	R	R	10	1.4	0.9	0	0	9	0	1	0	0
			R			R	R		8	1.2	0.7	0	0	5	0	0	1	2
R		R	R		R	R	R		8	1.2	0.7	0	0	2	6	0	0	0
R		R			R	R		R	7	1.0	0.6	4	0	2	0	0	0	1
R					R	R		R	7	1.0	0.6	0	0	6	1	0	0	0
R				R	R	R	R		6	0.9	0.5	3	0	2	0	0	1	0
						R	R	R	6	0.9	0.5	1	0	5	0	0	0	0
					R	R		R	4	0.6	0.3	2	0	2	0	0	0	0
					R	R	R	R	4	0.6	0.3	1	3	0	0	0	0	0
R						R		R	4	0.6	0.3	0	0	4	0	0	0	0
R	R		R		R	R		R	3	0.4	0.3	2	0	1	0	0	0	0
R						R	R		3	0.4	0.3	2	0	1	0	0	0	0
R					R		R		3	0.4	0.3	1	0	2	0	0	0	0
R	R					R		R	3	0.4	0.3	1	0	2	0	0	0	0
R		R				R	R	R	3	0.4	0.3	0	0	3	0	0	0	0
R		R	R	R		R	R	R	3	0.4	0.3	0	0	3	0	0	0	0



Appendix Table MDRP6 (continued). Multi-resistance patterns of interest in Salmonella spp. from fattening pigs in MSs reporting isolate-based data, 2012

	R	ChI	Cip	_						(N=1,1	roup 50)	Denmark (N=374)	Estonia (N=14)	Germany (N=627)	Hungary (N=38)	Ireland (N=24)	ltaly (N=25)	Spain (N=48)
R			0	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
					R	R	R		2	0.3	0.2	0	0	1	0	0	0	1
R				R	R	R	R	R	2	0.3	0.2	2	0	0	0	0	0	0
		R				R	R		2	0.3	0.2	2	0	0	0	0	0	0
				R	R	R			1	0.1	0.1	1	0	0	0	0	0	0
			R			R		R	1	0.1	0.1	1	0	0	0	0	0	0
			R		R		R		1	0.1	0.1	1	0	0	0	0	0	0
			R		R	R	R		1	0.1	0.1	0	0	0	0	1	0	0
		R				R	R	R	1	0.1	0.1	0	0	0	0	0	0	1
	_	R			R	R			1	0.1	0.1	0	0	0	1	0	0	0
	_	R			R	R	R		1	0.1	0.1	0	0	0	0	0	0	1
		R			R	R	R	R	1	0.1	0.1	1	0	0	0	0	0	0
		R		R	R		R		1	0.1	0.1	0	0	1	0	0	0	0
R				R		R	R		1	0.1	0.1	0	0	1	0	0	0	0
R				R		R	R	R	1	0.1	0.1	0	0	0	0	0	0	1
R				R	R	R			1	0.1	0.1	0	0	1	0	0	0	0
R		_	R				R		1	0.1	0.1	0	0	0	0	0	0	1
R		_	R			R			1	0.1	0.1	0	0	1	0	0	0	0
R		_	R		R		R		1	0.1	0.1	0	0	1	0	0	0	0
R			R	R	R		R		1	0.1	0.1	0	0	0	1	0	0	0
R	_	R				R			1	0.1	0.1	0	0	1	0	0	0	0
R	_	R		_	R				1	0.1	0.1	0	0	0	0	0	1	0
R	_	R		R	R	R		R	1	0.1	0.1	0	0	0	0	0	0	1
R		R	R			R	R		1	0.1	0.1	0	0	1	0	0	0	0
	R				D	R	D	D.	1	0.1	0.1	0	0	0	0	0	0	1
	R	_	D.		R	R	R	R	1	0.1	0.1	0	0	0	0	0	0	1
	R	_	R R		R	R	R	D.	1	0.1 0.1	0.1	0	0	1	0	0	0	0
	R R	R	R R	R	R R	R R	R R	R R	1	0.1	0.1	0	0	1 0	0	0	0	0
K	ĸ	ĸ	ĸ	К	ĸ	ĸ	ĸ	ĸ	I	100	60.0	127	5	480	27	15	U	25

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP7. Multi-resistance patterns of interest in Salmonella spp. from cattle in MSs reporting isolate-based data, 2012

	N	∕lulti∙	-resi	stan	ce pa	atter	n			MS gro (N=20	up 5)	Belgium (N=42)	Finland (N=19)	Germany (N=68)	Ireland (N=36)	ltaly (N=14)	Spain (N=9)	Sweden (N=17)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
R		R			R	R	R		25	33.8	12.2	4	0	2	15	2	0	2
R					R	R	R		16	21.6	7.8	2	0	5	5	2	2	0
					R	R	R		5	6.8	2.4	3	0	0	0	0	2	0
		R	R		R	R			3	4.1	1.5	3	0	0	0	0	0	0
R		R			R	R	R	R	2	2.7	1.0	0	0	0	1	0	1	0
			R			R	R	R	2	2.7	1.0	0	0	2	0	0	0	0
		R	R			R			2	2.7	1.0	2	0	0	0	0	0	0
R		R				R	R	R	2	2.7	1.0	0	0	0	0	0	2	0
			R		R			R	1	1.4	0.5	0	0	1	0	0	0	0
			R		R		R		1	1.4	0.5	0	0	1	0	0	0	0
			R		R	R		R	1	1.4	0.5	0	0	1	0	0	0	0
		R			R	R			1	1.4	0.5	0	0	1	0	0	0	0
		R	R	R	R		R		1	1.4	0.5	0	0	0	0	1	0	0
R						R	R		1	1.4	0.5	1	0	0	0	0	0	0
R					R	R			1	1.4	0.5	0	0	0	0	0	0	1
R					R	R		R	1	1.4	0.5	0	1	0	0	0	0	0
R					R	R	R	R	1	1.4	0.5	0	1	0	0	0	0	0
R			R			R			1	1.4	0.5	1	0	0	0	0	0	0
R			R		R	R	R	R	1	1.4	0.5	1	0	0	0	0	0	0
R			R	R		R	R	R	1	1.4	0.5	0	0	0	0	1	0	0
R		R		R	R	R	R	R	1	1.4	0.5	0	0	1	0	0	0	0
R		R	R		R	R			1	1.4	0.5	1	0	0	0	0	0	0
R		R	R		R	R		R	1	1.4	0.5	1	0	0	0	0	0	0
R		R	R		R	R	R	R	1	1.4	0.5	1	0	0	0	0	0	0
R	R		R		R	R	R	R	1	1.4	0.5	1	0	0	0	0	0	0
				Tota					74	100	36.1	21	2	14	21	6	7	3

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP8. Multi-resistance patterns of interest in Salmonella Enteritidis from broilers in MS reporting isolate-based data, 2012

		N	/lulti-res	sistance	e patter	n				MS (N=10)		Romania (N=10)
Amp	Ctx	сы	Cip	Gen	Str	Su	Tmp	n	%	MS	n	
			R			R	R		2	40.0	20.0	2
R			R			R			1	20.0	10.0	1
R		R				R			1	20.0	10.0	1
R		R	R	R	R	R	R	R	1	20.0	10.0	1
				Total					5	100	50.0	5

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP9. Multi-resistance patterns of interest in Salmonella Enteritidis from laying hens in MSs reporting isolate-based data, 2012

		N	lulti-res	sistance	e patter	n				MS grou (N=109	ıp)	Spain (N=43)	Romania (N=66)
Amp	Ctx	ChI	Cip	Gen	Str	Su	Tmp	n	%	MS group %	n	n	
			R			R	R		4	33.3	3.7	0	4
			R			R	R	R	4	33.3	3.7	0	4
R			R			R			1	8.3	0.9	0	1
R			R			R	R		1	8.3	0.9	0	1
R			R			R	R	R	1	8.3	0.9	1	0
R			R	R		R	R		1	8.3	0.9	0	1
				Total					12	100	11.0	1	11

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP10. Multi-resistance patterns of interest in Salmonella Typhimurium from broilers in MSs reporting isolate-based data, 2012

		Mu	lti-res	istanc	e patt	ern				MS group (N=8)		Romania (N=1)	ltaly (N=3)	Denmark (N=4)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n
R					R	R	R		4	57.1	50.0	1	0	3
R		R			R	R	R		2	28.6	25.0	0	2	0
R		R	R	R	R	R	R	R	1	14.5	12.5	0	1	0
				Total					7	100	87.5	1	3	3

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimetoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP11. Multi-resistance patterns of interest in Salmonella Typhimurium from laying hens in MSs reporting isolate-based data, 2012

		Mu	lti-res	istanc	e patte	ern				MS group (N=38))	Germany (N=13)	Romania (N=4)	ltaly (N=5)
Amp	Ctx	сы	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n
R		R				R	R		3	42.9	7.9	0	0	3
R					R	R	R		1	14.3	2.6	0	0	1
R			R		R	R	R		1	14.3	2.6	0	1	0
R			R		R	R	R	R	1	14.3	2.6	0	0	1
R		R			R	R	R	R	1	14.3	2.6	1	0	0
				Total					7	100	18.4	1	1	5

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimetoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP12. Multi-resistance patterns of interest in Salmonella Typhimurium from fattening pigs in MSs reporting isolate-based data, 2012

	N	lulti-	resi	stan	ce p	atter	'n			MS gr (N=37	oup 74)	Denmark (N=63)	Estonia (N=6)	Germany (N=273)	Hungary (N=8)	Ireland (N=15)	Spain (N=6)	Sweden (N=3)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
R		R			R	R	R		73	26.5	19.5	9	0	58	2	3	0	1
R					R	R	R	R	62	22.5	16.6	0	0	62	0	0	0	0
R					R	R	R		45	16.4	12.0	7	0	33	3	1	1	0
R		R			R	R	R	R	16	5.8	4.3	0	0	15	0	1	0	0
R		R	R		R	R	R	R	9	3.3	2.4	0	0	6	0	3	0	0
R						R	R	R	9	3.3	2.4	0	0	9	0	0	0	0
					R	R	R		9	3.3	2.4	2	6	0	0	1	0	0
R					R	R			8	2.9	2.1	6	0	1	1	0	0	0
R		R	R		R	R	R		7	2.5	1.9	0	0	6	1	0	0	0
R		R		R	R	R	R		7	2.5	1.9	1	0	6	0	0	0	0
R		R		R	R	R	R	R	5	1.8	1.3	0	0	4	0	1	0	0
R		R			R	R		R	4	1.5	1.1	0	0	4	0	0	0	0
R			R		R	R	R	R	3	1.1	0.8	0	0	3	0	0	0	0
					R	R		R	2	0.7	0.5	1	0	1	0	0	0	0
R		R				R	R		2	0.7	0.5	1	0	1	0	0	0	0
R					R	R		R	2	0.7	0.5	0	0	2	0	0	0	0
R				R	R	R	R		2	0.7	0.5	0	0	2	0	0	0	0
						R	R	R	1	0.4	0.3	0	0	1	0	0	0	0
					R	R	R	R	1	0.4	0.3	0	0	1	0	0	0	0
		R		R	R		R		1	0.4	0.3	0	0	1	0	0	0	0
R						R	R		1	0.4	0.3	0	0	1	0	0	0	0
R				R	R	R	R	R	1	0.4	0.3	0	0	1	0	0	0	0
R			R			R			1	0.4	0.3	0	0	1	0	0	0	0
R		R	R			R	R		1	0.4	0.3	0	0	0	0	0	1	0
R	R					R			1	0.4	0.3	0	0	1	0	0	0	0
R	R				R	R	R	R	1	0.4	0.3	0	0	1	0	0	0	0
R	R		R		R	R	R		1	0.4	0.3	0	0	0	0	0	1	0
				Tota	I				275	100	73.5	27	6	221	7	10	3	1

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP13. Multi-resistance patterns of interest in Salmonella Typhimurium from cattle in MSs reporting isolate-based data, 2012

	Μ	lulti-	resis	stan	ce p	atter	'n			MS gro (N=11	up 7)	Belgium (N=25)	Finland (N=16)	Germany (N=35)	Ireland (N=24)	Italy (N=4)	Spain (N=1)	Sweden (N=12)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n	n
R		R			R	R	R		25	64.1	21.4	4	0	2	15	2	0	2
R					R	R	R		3	7.7	2.6	2	0	0	0	0	1	0
					R	R	R		3	7.7	2.6	3	0	0	0	0	0	0
R						R	R		1	2.6	0.9	1	0	0	0	0	0	0
R					R	R			1	2.6	0.9	0	0	0	0	0	0	1
R					R	R		R	1	2.6	0.9	0	1	0	0	0	0	0
R					R	R	R	R	1	2.6	0.9	0	1	0	0	0	0	0
R			R		R	R	R	R	1	2.6	0.9	1	0	0	0	0	0	0
R		R			R	R	R	R	1	2.6	0.9	0	0	0	1	0	0	0
R		R		R	R	R	R	R	1	2.6	0.9	0	0	1	0	0	0	0
R		R	R		R	R	R	R	1	2.6	0.9	1	0	0	0	0	0	0
			-	Tota					39	100	33.3	12	2	3	16	2	1	3

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP14. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from turkey meat in MS reporting isolate-based data, 2012

		N	lulti-res	sistance	e patter	n				MS (N=22))	Germany (N=22)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS	n
R			R		R	R	R		12	63.2	54.5	12
R					R	R	R		7	36.8	31.8	7
				Total					19	100	86.4	19

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP15. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from pig meat in MSs and one non-MS reporting isolate-based data, 2012

		Mul	ti-resi	stanc	e pat	tern				Group reportir countrie (N=98)	ng es	Czech Republic (N=8)	Denmark (N=22)	Germany (N=37)	Ireland (N=23)	ltaly (N=8)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	group %	n	n		n	n
R					R	R	R		55	66.3	56.1	4	13	20	14	4
					R	R	R		6	7.2	6.1	1	1	0	4	0
R		R			R	R			5	6.0	5.1	0	0	5	0	0
R					R	R			4	4.8	4.1	1	2	1	0	0
R		R			R	R	R	R	3	3.6	3.1	0	1	1	1	0
		R			R	R	R	R	2	2.4	2.0	0	0	1	1	0
R		R			R	R	R		2	2.4	2.0	0	0	1	0	1
R				R	R	R	R	R	2	2.4	2.0	0	0	0	2	0
R					R	R	R	R	1	1.2	1.0	0	0	1	0	0
R				R	R	R	R		1	1.2	1.0	0	0	1	0	0
R			R		R	R	R		1	1.2	1.0	0	0	1	0	0
R		R				R			1	1.2	1.0	0	0	1	0	0
				Total					83	100	84.7	6	17	33	22	5

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP16. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from bovine animals meat in MSs reporting isolate-based data, 2012

		Mu	lti-res	istanc	e patt	ern				MS gro (N=4)	up)	Germany (N=2)	Ireland (N=2)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n
R					R	R	R		2	50.0	50.0	2	0
					R	R	R		1	25.0	25.0	0	1
		R			R	R	R	R	1	25.0	25.0	0	1
				Total					4	100	100	2	2

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimetoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP17. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from broilers in MSs reporting isolate-based data, 2012

		M	ulti-res	istanc	e patte	rn				MS group (N=8)		Denmark (N=4)	Spain (N=4)
Amp	Ctx	ChI	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n
R					R	R	R		5	71.4	62.5	3	2
R					R		R	R	1	14.3	12.5	0	1
R		R	R	R	R		R		1	14.3	12.5	0	1
				Total					7	100	87.5	3	4

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP18. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from laying hens in MSs reporting isolate-based data, 2012

		Mul	ti-res	istanc	e pat	tern				MS grou (N=11)	p	Denmark (N=1)	Italy (N=5)	Spain (N=4)	United Kingdom (N=1)
Amp	Ctx	chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
R					R	R	R		6	60.0	54.5	1	3	2	0
R			R		R	R	R		2	20.0	18.2	0	1	1	0
					R	R	R		1	10.0	9.1	0	0	0	1
R					R	R	R	R	1	10.0	9.1	0	0	1	0
				Total					10	100	90.9	1	4	4	1

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP19. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from turkeys in MSs reporting isolate-based data, 2012

		Mul	ti-resi	istanc	e pat	tern				MS grouj (N=28)	D	Germany (N=22)	ltaly (N=4)	Spain (N=1)	United Kingdom (N=1)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
R					R	R	R		17	85.0	60.7	15	0	1	1
R			R		R	R	R		1	5.0	3.6	1	0	0	0
R			R	R	R	R	R		1	5.0	3.6	0	1	0	0
R		R			R	R	R		1	5.0	3.6	1	0	0	0
				Total					20	100	71.4	17	1	1	1

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP20. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from fattening pigs in MSs reporting isolate-based data, 2012

		Mult	i-resi	stand	ce pat	ttern				MS gro (N=27	oup 9)	Denmark (N=29)	Germany (N=228)	lreland (N=6)	ltaly (N=2)	Spain (N=14)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n
R					R	R	R		192	74.4	68.8	18	163	3	1	7
R					R	R	R	R	11	4.3	3.9	1	10	0	0	0
R		R			R	R	R	R	10	3.9	3.6	0	9	0	0	1
R			R		R	R	R		10	3.9	3.6	0	10	0	0	0
R	R R R R								7	2.7	2.5	2	5	0	0	0
R	R R R R R								4	1.6	1.4	0	3	0	1	0
R	R R R R R Image: Constraint of the second sec								3	1.2	1.1	2	0	0	0	1
R					R		R		3	1.2	1.1	0	3	0	0	0
R		R		R	R	R	R	R	3	1.2	1.1	0	3	0	0	0
R		R	R		R	R	R	R	3	1.2	1.1	0	3	0	0	0
			R			R	R		2	0.8	0.7	0	2	0	0	0
					R	R	R		1	0.4	0.4	0	0	1	0	0
		R			R	R	R	R	1	0.4	0.4	0	0	0	0	1
R				R		R	R		1	0.4	0.4	0	0	0	0	1
R			R		R	R	R	R	1	0.4	0.4	0	1	0	0	0
R		R				R	R	R	1	0.4	0.4	0	1	0	0	0
R		R			R	R	R		1	0.4	0.4	0	1	0	0	0
R		R	R	R	R	R	R	R	1	0.4	0.4	0	0	0	0	1
R	R				R	R	R		1	0.4	0.4	0	1	0	0	0
R	R		R		R	R	R	R	1	0.4	0.4	0	0	0	0	1
R	R	R	R	R	R	R	R	R	1	0.4	0.4	0	1	0	0	0
				Total					258	100	92.5	23	216	4	2	13

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP21. Multi-resistance patterns of interest in monophasic Salmonella Typhimurium from cattle in MSs reporting isolate-based data, 2012

		Mul	ti-res	istanc	e pati	tern				MS grou (N=15)	ip	Germany (N=7)	lreland (N=5)	ltaly (N=2)	Spain (N=1)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
R					R	R	R		13	92.9	86.7	5	5	2	1
		R			R	R			1	7.1	6.7	1	0	0	0
				Total					14	100	93.3	6	5	2	1

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP22. Multi-resistance patterns of interest in Salmonella Kentucky from broiler meat in MSs reporting isolate-based data, 2012

		М	ulti-res	sistanc	e patte	rn				MS gro (N=65	oup 5)	Ireland (N=60)	Romania (N=5)
Amp	Ctx	сы	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n
R			R	R	R	R	R		6	50.0	9.2	1	5
R		R				R			4	33.3	6.2	4	0
R						R		R	1	8.3	1.5	1	0
R						R	R	R	1	8.3	1.5	1	0
				Total					12	100	18.5	7	5

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP23. Multi-resistance patterns of interest in Salmonella Kentucky from broilers in MSs reporting isolate-based data, 2012

	N	∕lulti∙	-resi	stan	ce pa	atter	n			MS gro (N=12	oup 0)	Czech Republic (N=9)	Hungary (N=1)	Ireland (N=31)	ltaly (N=2)	Romania (N=68)	Spain (N=6)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n	n
R			R	R	R	R	R		45	57.0	37.5	4	0	0	0	41	0
R			R	R		R	R		17	21.5	14.2	1	1	0	0	15	0
R	R		R	R	R	R	R		4	5.1	3.3	0	0	0	0	4	0
R			R	R	R	R			2	2.5	1.7	0	0	0	0	0	2
R	R	R	R	R	R	R	R	R	2	2.5	1.7	0	0	0	0	2	0
			R	R		R	R		1	1.3	0.8	0	0	0	0	0	1
R						R	R	R	1	1.3	0.8	0	0	1	0	0	0
R			R				R		1	1.3	0.8	0	0	0	1	0	0
R			R			R	R		1	1.3	0.8	0	0	0	0	1	0
R			R	R		R			1	1.3	0.8	0	0	0	0	1	0
R			R	R	R	R	R	R	1	1.3	0.8	0	0	0	0	1	0
R		R	R	R	R	R	R		1	1.3	0.8	0	0	0	0	1	0
R	R		R						1	1.3	0.8	0	0	0	1	0	0
R	R	R	R	R	R	R	R		1	1.3	0.8	0	0	0	0	1	0
				Tota					79	100	65.8	5	1	1	2	67	3

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella* and multi-resistant; n: number of multi-resistant isolates; R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP24.	Multi-resistance pa	atterns of	interest i	n Salmonella	Kentucky from	m laying
hens in MSs reporting iso	late-based data, 201	12			-	

		Mul	lti-res	istanc	e pati	tern				MS gro (N=29	oup))	Hungary (N=2)	ltaly (N=19)	Romania (N=4)	Spain (N=4)
Amp	Ctx	сы	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
R			R				R		6	35.3	20.7	0	6	0	0
R			R	R	R	R	R		3	17.7	10.3	2	0	1	0
R	R		R						2	11.8	6.9	0	2	0	0
			R	R	R	R	R		1	5.9	3.4	0	0	0	1
R			R			R		R	1	5.9	3.4	0	1	0	0
R			R		R	R	R		1	5.9	3.4	0	1	0	0
R			R	R		R	R		1	5.9	3.4	0	0	1	0
R		R	R			R	R	R	1	5.9	3.4	0	1	0	0
R		R	R	R	R	R	R	R	1	5.9	3.4	0	0	0	1
				Total					17	100	58.6	2	11	2	2

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim; N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *Salmonella* and multi-resistant; n: number of multi-resistant isolates; R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).



Appendix Table MDRP25. Multi-resistance patterns of interest in Salmonella Derby from pig meat in MSs reporting isolate-based data, 2012

		Mu	lti-resi	istanc	e patt	ern				MS gro (N=8		Germany (N=28)	lreland (N=5)	ltaly (N=26)	Romania (N=21)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
					R	R	R		16	64.0	20.0	1	1	7	7
R					R	R	R	R	3	12.0	3.8	1	0	1	1
						R	R	R	1	4.0	1.3	1	0	0	0
					R	R		R	1	4.0	1.3	0	0	1	0
					R	R	R	R	1	4.0	1.3	0	1	0	0
		R		R	R	R	R		1	4.0	1.3	0	0	1	0
R					R		R		1	4.0	1.3	0	0	0	1
R					R	R	R		1	4.0	1.3	0	0	1	0
				Total					25	100	31.3	3	2	11	9

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP26. Multi-resistance patterns of interest in Salmonella Derby from bovine animals meat in MSs reporting isolate-based data, 2012

		Mu	lti-res	istanc	e patte	ern				MS grou (N=8)	ıp	Czech Republic (N=3)	ltaly (N=5)	Ireland (N=2)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n
					R	R	R		1	33.3	12.5	0	1	0
		R			R	R	R		1	33.3	12.5	1	0	0
R		R				R	R	R	1	33.3	12.5	0	1	0
				Total					3	100	37.5	1	2	0

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP27. Multi-resistance patterns of interest in Salmonella Derby from broilers in MS reporting isolate-based data, 2012

		N	/lulti-res	sistance	e patteri	n				MS (N=2))	Romania (N=2)
Amp	Ctx	ChI	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS	n
R					R	R			1	50.0	50.0	1
R	R		R			R			1	50.0	50.0	1
				Total					2	100	100	2

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP28. Multi-resistance patterns of interest in Salmonella Derby from turkeys in MSs reporting isolate-based data, 2012

		Mul	lti-res	istanc	e patt	ern				MS gro (N=124	up 4)	lreland (N=2)	ltaly (N=3)	Spain (N=118)	United Kingdom (N=1)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
R		R	R			R	R	R	62	50.0	50.0	0	0	62	0
R			R			R	R	R	39	31.5	31.5	0	0	39	0
R		R	R		R	R	R	R	10	8.1	8.1	0	0	10	0
					R	R	R		3	2.4	2.4	2	0	0	1
R						R	R	R	3	2.4	2.4	0	1	2	0
R			R		R		R		1	0.8	0.8	0	0	1	0
R			R	R		R	R	R	1	0.8	0.8	0	0	1	0
R		R				R	R	R	1	0.8	0.8	0	0	1	0
R		R			R	R	R	R	1	0.8	0.8	0	0	1	0
R		R	R			R	R		1	0.8	0.8	0	0	1	0
R		R	R	R	R	R	R	R	1	0.8	0.8	0	1	0	0
R	R	R	R		R	R	R	R	1	0.8	0.8	0	1	0	0
				Total					124	100	100	2	3	118	1

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP29. Multi-resistance patterns of interest in Salmonella Derby from fattening pigs in MSs reporting isolate-based data, 2012

		Mult	i-resi	stanc	ce pa	ttern				MS gro (N=17	up 0)	Denmark (N=111)	Germany (N=42)	Hungary (N=4)	ltaly (N=6)	Spain (N=7)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n
					R	R	R		5	16.7	2.9	0	3	0	2	0
R	R R								4	13.3	2.4	2	1	1	0	0
R	R		R		R	R		R	4	13.3	2.4	0	4	0	0	0
R					R	R	R	R	3	10.0	1.8	1	2	0	0	0
					R	R		R	2	6.7	1.2	1	1	0	0	0
R					R	R	R		2	6.7	1.2	1	0	0	1	0
R						R	R		2	6.7	1.2	2	0	0	0	0
						R	R	R	1	3.3	0.6	1	0	0	0	0
		R				R	R	R	1	3.3	0.6	0	0	0	0	1
		R			R	R	R		1	3.3	0.6	0	1	0	0	0
R					R	R		R	1	3.3	0.6	1	0	0	0	0
R				R	R	R			1	3.3	0.6	0	1	0	0	0
R		R				R			1	3.3	0.6	1	0	0	0	0
R		R			R	R		R	1	3.3	0.6	1	0	0	0	0
R	R				R	R	R		1	3.3	0.6	0	0	0	0	1
				Total					30	100	17.6	11	13	1	3	2

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP30. Multi-resistance patterns of interest in Salmonella Derby from cattle in MS reporting isolate-based data, 2012

		r	Multi-res	sistance	e patteri	n				MS (N=2		Spain (N=2)
Amp	Ctx	ChI	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS	n
					R	R	R		2	100	100	2
		-	-	Total					2	100	100	2

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP31. Multi-resistance patterns of interest in Salmonella Infantis from broiler meat in MSs reporting isolate-based data, 2012

		Mult	ti-resi	istano	ce pat	tern				MS grou (N=162	up 2)	Czech Republic (N=10)	Estonia (N=2)	Germany (N=24)	ltaly (N=3)	Romania (N=123)
Amp	Ctx	ChI	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n
			R			R	R		50	33.3	30.9	4	0	13	3	30
			R			R	R	R	43	28.7	26.5	0	0	2	0	41
			R		R	R	R	R	26	17.3	16.0	0	0	0	0	26
			R		R	R	R		14	9.3	8.6	5	1	2	0	6
	R		R			R	R	R	3	2.0	1.9	0	0	0	0	3
			R	R		R	R	R	2	1.3	1.2	0	0	0	0	2
R		R	R			R	R	R	2	1.3	1.2	0	0	0	0	2
			R		R	R			1	0.7	0.6	0	1	0	0	0
		R	R				R	R	1	0.7	0.6	0	0	0	0	1
		R	R			R	R	R	1	0.7	0.6	0	0	0	0	1
		R	R		R	R	R		1	0.7	0.6	0	0	0	0	1
		R	R		R	R	R	R	1	0.7	0.6	0	0	0	0	1
R						R	R		1	0.7	0.6	1	0	0	0	0
R			R			R	R	R	1	0.7	0.6	0	0	0	0	1
R			R		R	R	R	R	1	0.7	0.6	0	0	0	0	1
R		R	R		R	R	R	R	1	0.7	0.6	0	0	0	0	1
R	R	R	R			R	R	R	1	0.7	0.6	0	0	0	0	1
				Total					150	100	92.6	10	2	17	3	118

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP32. Multi-resistance patterns of interest in Salmonella Infantis from pig meat in MSs reporting isolate-based data, 2012

		Mul	ti-resi	istanc	e pati	ern				MS grou (N=23)	up)	Czech Republic (N=6)	Germany (N=5)	Ireland (N=2)	Romania (N=10)
Amp	Ctx	сы	Cip	Gen	Str	ns	Tet	Tmp	n	%	MS group %	n	n	n	n
R						R		R	5	38.5	21.7	0	0	0	5
			R		R	R	R		2	15.4	8.7	1	0	0	1
R					R	R	R	R	2	15.4	8.7	2	0	0	0
			R			R	R		1	7.7	4.3	0	0	0	1
R					R		R		1	7.7	4.3	0	1	0	0
R		R				R	R		1	7.7	4.3	0	1	0	0
R		R		R	R	R	R	R	1	7.7	4.3	0	0	1	0
				Total					13	100	56.5	3	2	1	7

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP33. Multi-resistance patterns of interest in Salmonella Infantis from broilers in MSs reporting isolate-based data, 2012

		Mul	ti-resi	stanc	e pat	tern				MS gro (N=52	oup 9)	Austria (N=21)	Czech Republic (N=51)	Hungary (N=157)	ltaly (N=17)	Romania (N=283)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n
			R			R	R		180	40.4	34.0	16	25	86	0	53
			R		R	R	R		78	17.5	14.7	5	16	28	0	29
			R			R	R	R	41	9.2	7.8	0	0	0	2	39
			R		R	R	R	R	26	5.8	4.9	0	0	0	0	26
R			R			R	R		18	4.0	3.4	0	5	5	2	6
R			R		R	R	R		12	2.7	2.3	0	3	2	0	7
R			R			R	R	R	12	2.7	2.3	0	0	0	0	12
		R	R		R	R	R	R	9	2.0	1.7	0	0	0	0	9
	R		R		R	R	R		8	1.8	1.5	0	0	0	0	8
R			R		R	R	R	R	7	1.6	1.3	0	0	0	1	6
R	R		R			R	R	R	7	1.6	1.3	0	0	0	7	0
R			R	R		R	R		3	0.7	0.6	0	0	0	0	3
R		R	R			R	R		3	0.7	0.6	0	0	0	0	3
R	R		R			R	R		3	0.7	0.6	0	0	0	0	3
			R	R		R	R	R	2	0.5	0.4	0	0	0	0	2
		R	R			R	R		2	0.5	0.4	0	0	2	0	0
R			R	R	_	R	R	R	2	0.5	0.4	0	0	0	0	2
R			R	R	R	R	R	R	2	0.5	0.4	0	0	0	0	2
R		R	R		R	R	R		2	0.5	0.4	0	0	0	0	2
R		R	R		R	R	R	R	2	0.5	0.4	0	0	0	0	2
R	R		R						2	0.5	0.4	0	0	0	0	2
R	R	R	R		R	R	R	R	2	0.5	0.4	0	0	0	0	2
R	R	R	R	R	R	R	R	R	2	0.5	0.4	0	0	0	0	2
			R				R	R	1	0.2	0.2	0	0	0	0	1
			R		R		R		1	0.2	0.2	0	0	1	0	0
			R		R	R			1	0.2	0.2	0	0	1	0	0
			R	R			R		1	0.2	0.2	0	0	1	0	0



Appendix Table MDRP33 (continued). Multi-resistance patterns of interest in Salmonella Infantis from broilers in MSs reporting isolate-based data, 2012

		Mult	ti-resi	istano	ce pat	tern				MS gro (N=52		Austria (N=21)	Czech Republic (N=51)	Hungary (N=157)	ltaly (N=17)	Romania (N=283)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n	n
			R	R	R	R	R		1	0.2	0.2	0	0	0	0	1
			R	R	R	R	R	R	1	0.2	0.2	0	0	0	0	1
		R	R			R	R	R	1	0.2	0.2	0	0	0	0	1
	R		R			R	R		1	0.2	0.2	0	0	0	0	1
	R		R	R	R	R	R	R	1	0.2	0.2	0	0	0	0	1
R			R		R				1	0.2	0.2	0	0	0	0	1
R			R		R		R		1	0.2	0.2	0	0	0	1	0
R			R	R	R	R	R		1	0.2	0.2	0	0	0	0	1
R		R	R			R		R	1	0.2	0.2	0	0	0	0	1
R		R	R			R	R	R	1	0.2	0.2	0	0	0	0	1
R		R	R	R	R	R	R	R	1	0.2	0.2	0	0	0	0	1
R	R		R		R	R	R	R	1	0.2	0.2	0	0	0	0	1
R	R		R	R		R	R		1	0.2	0.2	0	0	0	0	1
R	R		R	R		R	R	R	1	0.2	0.2	0	0	0	1	0
R	R		R	R	R	R	R		1	0.2	0.2	0	0	0	0	1
R	R	R	R		R	R	R		1	0.2	0.2	0	0	0	0	1
				Total					445	100	84.1	21	49	126	14	235

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP34. Multi-resistance patterns of interest in Salmonella Infantis from laying hens in MSs reporting isolate-based data, 2012

		Mul	ti-resi	istanc	e pati	tern				MS grou (N=36)) dr	Austria (N=4)	Denmark (N=1)	Hungary (N=15)	Romania (N=16)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n	n
			R			R	R		16	55.2	44.4	3	0	8	5
			R		R	R	R		5	17.2	13.9	0	0	1	4
			R			R	R	R	2	6.9	5.6	0	0	0	2
R		R	R			R	R		2	6.9	5.6	0	0	0	2
R					R		R		1	3.5	2.8	0	1	0	0
R			R			R	R		1	3.5	2.8	0	0	0	1
R			R		R	R	R		1	3.5	2.8	0	0	0	1
R		R	R		R	R	R		1	3.5	2.8	0	0	0	1
				Total					29	100	80.6	3	1	9	16

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP35. Multi-resistance patterns of interest in Salmonella Infantis from turkeys in MSs reporting isolate-based data, 2012

		Mu	lti-res	istanc	e patt	ern				MS group (N=27)		Austria (N=2)	Germany (N=4)	Hungary (N=21)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n	n
			R			R	R		19	76.0	70.4	1	3	15
			R		R	R	R		5	20.0	18.5	1	1	3
R			R			R	R		1	4.0	3.7	0	0	1
				Total					25	100	92.6	2	4	19

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP36. Multi-resistance patterns of interest in Salmonella Infantis from fattening pigs in MSs reporting isolate-based data, 2012

			Resis	tance p	attern					MS group (N=18))	Germany (N=10)	Hungary (N=8)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	MS group %	n	n
			R			R	R		6	50.0	33.3	0	6
R					R	R	R	R	2	16.7	11.1	2	0
R	R					R		R	2	16.7	11.1	2	0
						R	R	R	1	8.3	5.6	1	0
			R		R	R	R		1	8.3	5.6	0	1
				Total					12	100	66.7	5	7

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for Salmonella and multi-resistant;

n: number of multi-resistant isolates;

R: minimum inhibitory concentration (MIC) above epidemiological cut-off values (ECOFFs) from the European Committee on Antimicrobial Susceptibility Testing (EUCAST).

Appendix Table MDRP37. Multi-resistance patterns of interest in Escherichia coli from broilers in MSs and one non-MS reporting isolate-based data, 2012

		Mu	lti-resi	stance	patter	'n				Group reporti countr (N=53	ng ies	Austria (N=130)	Denmark (N=115)	Hungary (N=104)	Switzerland (N=185)
Amp	Ctx	chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	group %	n	n	n	n
R			R				R		13	7.8	2.4	0	0	5	8
R			R		R	R	R	R	11	6.6	2.1	6	0	2	3
			R		R	R	R		9	5.4	1.7	7	0	1	1
			R		R	R		R	9	5.4	1.7	8	0	1	0
R						R	R	R	8	4.8	1.5	0	3	1	4
			R		R		R		6	3.6	1.1	1	0	2	3
R						R		R	6	3.6	1.1	3	2	1	0
R					R		R		6	3.6	1.1	1	3	0	2
R			R		R	R	R		5	3.0	0.9	3	1	1	0
R		R	R		R	R	R	R	5	3.0	0.9	2	0	1	2
						R	R	R	5	3.0	0.9	1	0	0	4
R			R			R		R	5	3.0	0.9	3	0	2	0
R			R			R	R	R	5	3.0	0.9	0	0	1	4
			R			R	R	R	4	2.4	0.8	2	0	1	1
R					R	R	R	R	4	2.4	0.8	1	0	1	2
R			R			R			4	2.4	0.8	1	0	0	3
R	R		R						4	2.4	0.8	0	0	3	1
			R			R	R		3	1.8	0.6	1	0	1	1
R					R	R		R	3	1.8	0.6	0	2	1	0
R					R	R	R		3	1.8	0.6	0	0	1	2
R		R	R			R	R	R	3	1.8	0.6	1	0	2	0
					R	R		R	3	1.8	0.6	0	3	0	0



Appendix Table MDRP37 (continued). Multi-resistance patterns of interest in Escherichia coli from broilers in MSs and one non-MS reporting isolate-based data, 2012

		Mu	lti-resi	stance	patter	n				Group reporti countri (N=53	ng ies	Austria (N=130)	Denmark (N=115)	Hungary (N=104)	Switzerland (N=185)
Amp	Ctx	ChI	Cip	Gen	Str	Su	Tet	Tmp	n	%	group %	n	n	n	n
			R		R	R			2	1.2	0.4	1	0	0	1
R			R			R	R		2	1.2	0.4	1	0	1	0
R		R	R		R	R	R		2	1.2	0.4	1	0	1	0
R	R		R			R	R	R	2	1.2	0.4	0	0	1	1
					R	R	R	R	2	1.2	0.4	0	0	2	0
R			R		R		R		2	1.2	0.4	0	0	2	0
R	R		R			R	R		2	1.2	0.4	0	0	0	2
			R			R		R	1	0.6	0.2	1	0	0	0
			R		R	R	R	R	1	0.6	0.2	0	0	1	0
		R	R		R	R			1	0.6	0.2	1	0	0	0
		R	R		R	R		R	1	0.6	0.2	1	0	0	0
		R	R		R	R	R	_	1	0.6	0.2	1	0	0	0
		R	R		R	R	R	R	1	0.6	0.2	1	0	0	0
		R	R	R		R			1	0.6	0.2	0	0	1	0
R						R	R		1	0.6	0.2	0	1	0	0
R			_		R			R	1	0.6	0.2	1	0	0	0
R			R				_	R	1	0.6	0.2	0	0	1	0
R			R				R	R	1	0.6	0.2	0	0	1	0
R			R		R	6			1	0.6	0.2	1	0	0	0
R			R R		R R	R R		R	1	0.6	0.2	0	0	1	0
R R			R R	R	R R	R R		R	1	0.6 0.6	0.2	0	0 0	1 0	0
R		R	R R	ĸ	ĸ	ĸ	R	R	1	0.6	0.2	0	0	1	1 0
R		R	R			R	R	R	1	0.6	0.2	0	0	1	0
R		R	R		R	R			1	0.6	0.2	0	0	1	0
R		R	R	R		R		R	1	0.6	0.2	1	0	0	0
R		R	R	R	R	R	R	R	1	0.6	0.2	0	0	1	0
R	R					R			1	0.6	0.2	1	0	0	0
R	R				R			R	1	0.6	0.2	0	0	1	0
R	R				R	R	R		1	0.6	0.2	1	0	0	0
R	R		R		R	R		R	1	0.6	0.2	0	0	1	0
R	R		R		R	R	R	R	1	0.6	0.2	0	0	1	0
R	R	R			R	R	R		1	0.6	0.2	1	0	0	0
R	R	R	R			R		R	1	0.6	0.2	0	0	1	0
R	R	R	R			R	R		1	0.6	0.2	1	0	0	0
				Total				•	166	100	31.1	56	15	49	46

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for *E. coli* and multi-resistant;

n: number of multi-resistant isolates;



Appendix Table MDRP38. Multi-resistance patterns of interest in Escherichia coli from fattening pigs in MSs and one non-MS reporting isolate-based data, 2012

		Mu	lti-res	istanc	e patt	ern				Group reporti countri (N=54	ng ies	Austria (N=140)	Denmark (N=152)	Hungary (N=68)	Switzerland (N=185)
Amp	Ctx	сы	Cip	Gen	Str	Su	Tet	Tmp	n	%	group %	n	n	n	n
					R	R	R		29	16.0	5.3	13	4	2	10
R					R	R	R	R	29	16.0	5.3	3	16	2	8
R					R	R		R	16	8.8	2.9	2	8	0	6
					R	R	R	R	13	7.2	2.4	2	2	1	8
R					R	R			13	7.2	2.4	1	8	0	4
R					R	R	R		7	3.9	1.3	1	3	1	2
R					R		R		6	3.3	1.1	1	0	4	1
R						R	R	R	5	2.8	0.9	0	1	3	1
R			R		R	R	R	R	5	2.8	0.9	3	1	1	0
					R	R		R	5	2.8	0.9	0	1	0	4
						R	R	R	4	2.2	0.7	1	1	0	2
R			R				R		4	2.2	0.7	1	0	3	0
R		R				R	R	R	4	2.2	0.7	0	0	3	1
R		R			R	R	R	R	4	2.2	0.7	1	0	3	0
R		R			R	R	R		3	1.7	0.6	1	0	1	1
		R				R	R	R	3	1.7	0.6	0	1	0	2
		R			R	R	R	R	3	1.7	0.6	1	0	0	2
			R		R	R		R	2	1.1	0.4	0	0	2	0
R					R			R	2	1.1	0.4	0	0	0	2
					R		R	R	1	0.6	0.2	0	0	0	1
			R		R		R		1	0.6	0.2	1	0	0	0
			R		R		R	R	1	0.6	0.2	1	0	0	0
			R		R	R	R		1	0.6	0.2	0	0	1	0
		R			R	R			1	0.6	0.2	0	0	1	0
		R			R	R	R		1	0.6	0.2	0	1	0	0
		R	R				R		1	0.6	0.2	0	0	0	1
R						R		R	1	0.6	0.2	0	0	0	1
R					R		R	R	1	0.6	0.2	0	0	0	1
R				R	R				1	0.6	0.2	0	0	0	1
R				R	R	R	R		1	0.6	0.2	0	0	1	0
R			R		R		R		1	0.6	0.2	0	0	1	0
R			R		R	R	R		1	0.6	0.2	1	0	0	0
R			R	R	R	R		R	1	0.6	0.2	0	0	0	1
R			R	R	R	R	R	R	1	0.6	0.2	0	0	1	0
R		R					R		1	0.6	0.2	0	0	1	0



Appendix Table MDRP38 (continued). Multi-resistance patterns of interest in Escherichia coli from fattening pigs in MSs and one non-MS reporting isolate-based data, 2012

		Mu	lti-resi	istanc	e patt	ern				Group reporti countri (N=54	ng es	Austria (N=140)	Denmark (N=152)	Hungary (N=68)	Switzerland (N=185)
Amp	Ctx	Chl	Cip	Gen	Str	Su	Tet	Tmp	n	%	group %	n	n	n	n
R		R					R	R	1	0.6	0.2	0	0	0	1
R		R				R	R		1	0.6	0.2	0	0	1	0
R		R			R		R		1	0.6	0.2	0	1	0	0
R		R			R	R		R	1	0.6	0.2	0	0	0	1
R		R		R	R	R		R	1	0.6	0.2	0	1	0	0
R	R						R		1	0.6	0.2	0	0	1	0
R	R				R	R	R	R	1	0.6	0.2	0	0	0	1
R	R		R						1	0.6	0.2	0	0	0	1
				Total					181	100	33.2	34	49	34	64

MS: Member State; Amp: ampicillin; Ctx: cefotaxime; Chl: chloramphenicol; Cip: ciprofloxacin; Gen: gentamicin; Str: streptomycin; Su: sulfonamides; Tet: tetracyclines; Tmp: trimethoprim.

N: total number of isolates tested for susceptibility against the whole common antimicrobial set for E. coli and multi-resistant;

n: number of multi-resistant isolates;



APPENDIX 5. High-level resistance to ciprofloxacin

Appendix Table ULCD1	h loval ainroflavaain ragistanaa in Salmanalla caravara from brailar m	at in MSs reporting isolate based data 2012
	h-level ciprofloxacin resistance in Salmonella serovars from broiler me	

Country	Isolates resistant at > 0.06 mg/L ¹ Cip n (%)	Isolates resistant at > 1 mg/L ² Cip n (%)	Isolates resistant at > 2 mg/L ³ Cip n (%)	Isolates resistant at > 4 mg/L ⁴ Cip n (%)	Resistance patterns of isolates resistant at > 4 mg/L Cip	Salmonella serovars
Czech Republic (N=47)	16 (34.0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Estonia (N=3)	2 (66.7 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Germany (N=94)	66 (70.2 %)	1 (1.1 %)	0 (0 %)	0 (0 %)	NA	NA
Ireland (N=70)	8 (11.4 %)	1 (1.4 %)	1 (1.4 %)	1 (1.4 %)	AmpCipGenNalStrSuTet (1)	S. Kentucky
Italy (N=8)	4 (50.0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Romania (N=188)	168 (89.4%)	14 (7.4 %)	3 (1.6 %)	1 (0.5 %)	CipNalSuTetTmp(1)	S. Infantis
Total (6 MSs) (N=410)	264 (64.4 %)	16 (3.9 %)	4 (1.0 %)	2 (0.5 %)		

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of resistant isolates; NA: not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.

4: Very high breakpoint.



Country	Isolates resistant at > 0.06 mg/L ¹ Cip n (%)	Isolates resistant at > 1 mg/L ² Cip n (%)	Isolates resistant at > 2 mg/L ³ Cip n (%)	Isolates resistant at > 4 mg/L ⁴ Cip n (%)	Resistance patterns of isolates resistant at > 4 mg/L Cip	Salmonella serovars
Czech Republic (N=10)	5 (50.0 %)	1 (10.0 %)	1 (10.0 %)	1 (10.0 %)	AmpCipGenNalSuTet (1)	S. Kentucky
Estonia (N=2)	1 (50.0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Germany (N=126)	80 (63.5 %)	10 (7.9 %)	7 (5.6 %)	7 (5.6 %)	AmpCipGenNalSuTet (5)	Other serovars
					AmpCipNal (1)	Other serovars
					AmpCipGenNalStrSuTet (1)	Other serovars
Ireland (N=1)	1 (100 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Italy (N=6)	2 (33.3 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Total (5 MSs) (N=145)	89 (61.4 %)	11 (7.6 %)	8 (5.5 %)	8 (5.5 %)		

Appendix Table HLCR2. High-level ciprofloxacin resistance in Salmonella serovars from turkey meat in MSs reporting isolate-based data, 2012

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of resistant isolates; NA: not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.

4: Very high breakpoint.



Appendix Table HLCR3. High-level ciprofloxacin resistance in Salmonella serovars from pig meat in MSs reporting isolate-based data, 2012

Country	Isolates resistant at > 0.06 mg/L ¹ Cip n (%)	Isolates resistant at > 1 mg/L ² Cip n (%)	Isolates resistant at > 2 mg/L ³ Cip n (%)	Isolates resistant at > 4 mg/L ⁴ Cip n (%)	Resistance patterns of isolates resistant at > 4 mg/L Cip	Salmonella serovars
Czech Republic (N=33)	2 (6.1 %)	1 (3.0 %)	1 (3.0 %)	1 (3.0 %)	CipNalStrSuTet (1)	S. Infantis
Denmark (N=41)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Estonia (N=22)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Germany (N=163)	10 (6.1 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Ireland (N=69)	1 (1.4 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Italy (N=85)	1 (1.2 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Romania (N=125)	13 (10.4 %)	1(0.8 %)	1 (0.8 %)	1 (0.8 %)	AmpChlCipSuTetTmp(1)	S. Typhimurium
Total (7 MSs) (N=538)	27 (5.0 %)	2 (0.4 %)	2 (0.4 %)	2 (0.4 %)		

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of resistant isolates; NA: not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.

4: Very high breakpoint.



Isolates resistant Isolates resistant Isolates resistant Isolates Resistance patterns of isolates resistant at at at at Country resistant at Salmonella serovars $> 0.06 \text{ mg/L}^1 \text{ Cip}$ $> 1 \text{ mg/L}^2 \text{Cip}$ $> 2 \text{ mg/L}^3 \text{ Cip}$ $> 4 \text{ mg/L}^4 \text{ Cip}$ > 4 mg/L Cipn (%) n (%) n (%) n (%) Austria (N=113) 25 (22.1 %) 0 (0 %) NA 0 (0 %) 0 (0 %) NA AmpCipGenNalStrSuTet (4) S. Kentucky Czech Republic (N=351) 73 (20.8 %) 14 (4.0 %) 14 (4.0 %) 5 (1.4 %) AmpCipGenNalSuTet (1) S. Kentucky Denmark (N=24) 0 (0 %) 0 (0 %) 0 (0 %) 0 (0 %) NA NA Germany (N=8) 2 (25.0 %) 0 (0 %) 0 (0 %) 0 (0 %) NA NA AmpCipGenNalSuTet (1) S. Kentucky 2 (1.1 %) Hungary (N=175) 158 (90.3 %) 5 (2.9 %) 4 (2.3 %) AmpCipNalSuTet (1) S. Infantis Ireland (N=38) 0 (0 %) 0 (0 %) 0 (0 %) 1 (2.6 %) NA NA Italy (N=105) 34 (32.4 %) 1 (1.0 %) 1 (1.0 %) 0 (0 %) NA NA AmpCipGenNalStrSu (2) S. Kentucky 4 (13.8 %) CipGenNalSuTet (1) S. Kentucky Spain (N=29) 19 (65.5 %) 6 (20.7 %) 5 (17.2 %) AmpCipNal (1) S. Kentucky Sweden (N=1) 0 (0 %) 0 (0 %) 0 (0 %) 0 (0 %) NA NA

Appendix Table HLCR4. High-level ciprofloxacin resistance in Salmonella serovars from broilers in MSs reporting isolate-based data, 2012



Appendix Table HLCR4 (continued). High-level ciprofloxacin resistance in Salmonella serovars from broilers in MSs reporting isolate-based data, 2012

Country	Isolates resistant at > 0.06 mg/L ¹ Cip	Isolates resistant at > 1 mg/L ² Cip	Isolates resistant at > 2 mg/L ³ Cip	Isolates resistant at > 4 mg/L ⁴ Cip	Resistance patterns of isolates resistant at	Salmonella serovars
	n (%)	n (%)	n (%)	n (%)	> 4 mg/∟ Cip	
Romania (N=781)					> 4 mg/L Cip AmpCipGenNalStrSuTet (39) AmpCipGenNalSuTet (13) AmpCtxCipGenNalStrSuTet (4) AmpCtxChlCipGenNalStrSuTetTmp (2) AmpCipNal (1) AmpCipOenNalSuTet (1) AmpCipGenNalSuTet (1) AmpCipGenNalStrSuTetTmp(1) AmpChlCipGenNalStrSuTet (1) AmpCipGenNalStrSuTet (1) AmpCipGenNalStrSuTet (1) AmpCipGenNalStrSuTet (1) AmpCipGenNalStrSuTet (1) AmpCipGenNalStrSuTet (1) AmpCipGenNalStrSuTetTmp (2) AmpCipGenNalSuTetTmp (2) AmpCipGenNalSuTetTmp (2) AmpCipGenNalStrSuTetTmp (2) AmpCipGenNalStrSuTetTmp (1) CipGenNalStrSuTetTmp (2) AmpCipGenNalStrSuTetTmp (1) AmpCipGenNalStrSuTetTmp (1) AmpCipSuTetTmp (1) AmpChlCipNalStrSuTet (1)	S. Kentucky S. Agona S. Agona S. Infantis S. Infantis S. Infantis S. Infantis S. Infantis S. Infantis S. Infantis S. Infantis
					AmpChlCipSu (1) AmpChlCipSuTmp (1)	S. Liverpool S. Liverpool
					AmpCipGenNalSuTet (3) AmpCipTet (1)	S. Tennessee S. Tennessee
					AmpCipGenNalStrSuTetTmp(1)	S. Tennessee
					AmpCipNalSuTet (1)	S. Livingstone
					CipNalSuTet (1)	S. Rissen
United Kingdom (N=17)	1 (5.9 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Total (11 MSs) (N=1,641)	885 (54.2 %)	164 (10.0 %)	130 (8.0 %)	97(5.9 %)		

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of resistant isolates; NA: not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.



Appendix Table HLCR5. High-level ciprofloxacin resistance in Salmonella serovars from laying hens in MSs reporting isolate-based data, 2012

Country	Isolates resistant at > 0.06 mg/L ¹ Cip n (%)	Isolates resistant at > 1 mg/L ² Cip n (%)	Isolates resistant at > 2 mg/L ³ Cip n (%)	Isolates resistant at > 4 mg/L ⁴ Cip n (%)	Resistance patterns of isolates resistant at > 4 mg/L Cip	Salmonella serovars
Austria (N=63)	7 (11.1 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Czech Republic (N=8)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Denmark(N=4)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Finland (N=5)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Germany (N=51)	1 (2 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Live serve (NL OC)					AmpCipGenNalStrSuTet (2)	S. Kentucky
Hungary (N=86)	22 (25.6 %)	3 (3.5 %)	3 (3.5 %)	3 (3.5 %)	AmpCipNalStrSuTetTmp (1)	S. Saintpaul
Italy (N=161)	29 (18.0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
		13 (9.0 %)	10 (6.9%)	9 (6.2 %)	Cip (2)	S. Kentucky
					AmpCipGenNalSuTet (1)	S. Kentucky
					AmpCipGenNalStrSuTet (1)	S. Kentucky
Demonia $(N + 4.45)$					CipNalTmp (1)	S. Thompson
Romania (N=145)	67 (46.2 %)				AmpCipGenNalStrSuTet (1)	S. Hadar
					AmpCipNalSu (1)	S. Albany
					Cip (1)	S. Corvallis
					AmpCipGenNalSuTet (1)	S. Corvallis
On size (NL 450)	04 (44 0.00)		0 (4 0 0()	0 (4 0 0()	CipGenNalStrSuTet (1)	S. Kentucky
Spain (N=150)	21 (14.0 %)	3 (2.0 %)	2 (1.3 %)	2 (1.3 %)	AmpChlCipGenNalStrSuTetTmp (1)	S. Kentucky
Sweden (N=2)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
United Kingdom (N=11)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Total (11 MSs) (N=686)	147 (21.4 %)	19 (2.8 %)	15 (2.2 %)	14 (2 %)		

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of the resistant isolates; NA: Not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.



Appendix Table HLCR6. High-level ciprofloxacin resistance in Salmonella serovars from turkeys in MSs reporting isolate-based data, 2012

Country	Isolates resistant at > 0.06 mg/L ¹ Cip	Isolates resistant at > 1 mg/L ² Cip	Isolates resistant at > 2 mg/L ³ Cip	Isolates resistant at > 4 mg/L ⁴ Cip	Resistance patterns of isolates resistant at > 4 mg/L Cip	Salmonella serovars
	n (%)	n (%)	n (%)	n (%)	· · ···9· - ···P	
Austria (N=38)	30 (78.9 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Belgium (N=2)	1 (50.0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
					AmpCipGenNalStrSuTet (6)	S. Kentucky
Czech Republic (N=27)	22 (81.5 %)	9 (33.3 %)	9 (33.3 %)	9 (33.3 %)	AmpCipGenNalSuTet (2)	S. Kentucky
					AmpCipNal (1)	S. Kentucky
Finland (N=1)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Germany (N=87)	28 (32.2 %)	3 (3.4 %)	1 (1.1 %)	1 (1.1 %)	AmpCipGenNalSuTet (1)	Other serovars
	159 (91.4 %)	38 (21.8 %)	37 (21.3 %)	35 (20.1 %)	AmpCipGenNalStrSuTet (18)	S. Kentucky
					AmpCipGenNalSuTet (14)	S. Kentucky
Hungary (N=174)					AmpCipNal (1)	S. Kentucky
					AmpCipGenNalSuTet (1)	S. Stanley
					AmpCipGenNalStrSuTet (1)	S. Stanley
Ireland (N=14)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Italy (N=48)	18 (37.5 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Spain (N=169)	152 (89.9 %)	2 (1.2 %)	2 (1.2 %)	2 (1.2 %)	AmpCipGenNalStrSuTet (2)	S. Kentucky
United Kingdom (N=7)	1 (14.3 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Total (10 MSs) (N=567)	411 (72.5 %)	52 (9.2 %)	49 (8.6 %)	47 (8.3 %)		

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of the resistant isolates; NA: Not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.



Appendix Table HLCR7. High-level ciprofloxacin resistance in Salmonella serovars from fattening pigs in MSs reporting isolate-based data, 2012

Country	lsolates resistant at > 0.06 mg/L ¹ Cip	at a	Isolates resistant at > 2 mg/L ³ Cip	at	Resistance patterns of isolates resistant at	Salmonella serovars
	n (%)	n (%)	n (%)	n (%)	> 4 mg/L Cip	
Denmark (N=374)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Estonia (N=14)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Finland (N=2)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Germany (N=627)	47 (7.5 %)	1 (0.2 %)	0 (0 %)	0 (0 %)	NA	NA
Hungary(N= 38)	9 (23.7 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Ireland (N=24)	4 (16.7 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Italy (N=25)	3 (12.0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Spain (N=48)	10 (20.8 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Sweden (N=3)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Total (9 MSs) (N=1,155)	73 (6.3 %)	1 (0.1 %)	0 (0 %)	0 (0 %)		

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of the resistant isolates; NA: Not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.



Appendix Table HLCR8. High-level ciprofloxacin resistance in Salmonella serovars from cattle in MSs reporting isolate-based data, 2012

Country	Isolates resistant at > 0.06 mg/L ¹ Cip n (%)	Isolates resistant at > 1 mg/L ² Cip n (%)	Isolates resistant at > 2 mg/L ³ Cip n (%)	Isolates resistant at > 4 mg/L ⁴ Cip n (%)	Resistance patterns of isolates resistant at > 4 mg/L Cip	Salmonella serovars
Belgium (N=42)	12 (28.6 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Estonia (N=7)	1 (14.3 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Finland (N=19)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Germany (N=68)	6 (8.8 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Ireland (N=36)	2 (5.6 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Italy (N=14)	2 (14.3 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Spain (N=9)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Sweden (N=17)	0 (0 %)	0 (0 %)	0 (0 %)	0 (0 %)	NA	NA
Total (8 MSs) (N=212)	23 (10.8 %)	0 (0 %)	0 (0 %)	0 (0 %)		

MS: Member State; Cip: ciprofloxacin; N: Total number isolates for which relevant data are available; n: number of the resistant isolates; NA: Not applicable.

1: European Committee on Antimicrobial Susceptibility Testing (EUCAST) epidemiological cut-off values (ECOFFs).

2: EUCAST clinical breakpoint.

3: High breakpoint.



APPENDIX 6. Level 3 Tables

Level 3 Tables, containing information on reported MIC distributions and data on the number of resistant isolates, are available on the EFSA website.

APPENDIX 7. List of abbreviations, Antimicrobial substances, Member States and other reporting countries, definitions

List of abbreviations

Abbreviation	Definition
%	Percentage of resistant isolates per category of susceptibility or multiple resistance
% f	Percentage frequency of isolates tested
% Res	Percentage of resistant isolates
-	No data reported
AHVLA	Animal Health and Veterinary Laboratories Agency
AMB	Antimicrobial substance
AMR	Antimicrobial resistance
AST	Antimicrobial susceptibility testing
BIOHAZ	EFSA Panel on Biological Hazards
BSAC	British Society for Antimicrobial Chemotherapy
CA-SFM	French Society for Microbiology
CC	Clonal complex
CLSI	Clinical and Laboratory Standards Institute
CBP	Clinical breakpoints
CTX-M	Cefotaximase
DIN	Deutsches Institut für Normung
DNA	desoxyribonucleic acid
DTU	Technical University of Denmark
EARS-Net	European Antimicrobial Resistance Surveillance Network
EC	European Commission
ECDC	European Control
ECOFF	epidemiological cut-off value
EEA	European Economic Area
EFSA	European Food Safety Authority
EMA	European Medicines Agency
ESBL	Extended Spectrum Beta-Lactamase
ETEC	Enterotoxigenic <i>E. coli</i>
EU EUCAST	European Union
	European Committee on Antimicrobial Susceptibility Testing
EU-RL	European Union Reference Laboratory
FWD	Food- and Waterborne Diseases and Zoonoses
HACCP	Hazard Analysis and Critical Control Point
HPA	Health Protection Agency (UK)
	Intermediate
IZD	Inhibition Zone Diameter
LA-MRSA	Livestock- associated methicillin-resistant Staphylococcus aureus
MDR	Multiple Drug Resistance
MIC	Minimum Inhibitory Concentration
MLST	Nulti-locus Sequence Typing
MRSA	Methicillin-resistant Staphylococcus aureus
MSSA	Methicillin-susceptible Staphylococcus aureus
MS	Member State
NA	Not applicable
NCP	National Control Programme
NRL	National Reference Laboratory
R	Resistant
res1–res9	Resistance to one antimicrobial substance/resistance to nine antimicrobial substance
	of the common set for Salmonella
S	Susceptible
SGI	Salmonella genomic island
spp.	species
ST	Sequence type
TESSy	The European Surveillance System



VTEC	Vero(cyto)toxigenic <i>E. coli</i>
WHO	World Health Organization

Antimicrobial substances

Abbreviation	Antimicrobials
Amp	Ampicillin
Caz	Ceftazidime
Chl	Chloramphenicol
Cip	Ciprofloxacin
Ctx	Cefotaxime
Ery	Erythromycin
Gen	Gentamicin
Nal	Nalidixic acid
Su	Sulfonamides
Str	Streptomycin
Tet	Tetracycline
Tmp	Trimethoprim



Member States of the European Union and other reporting countries in 2012

Member States of the European Union, 2012

Member State	Country abbreviations
Austria	AT
Belgium	BE
Bulgaria	BG
Cyprus	CY
Czech Republic	CZ*
Denmark	DK
Estonia	EE
Finland	FI
France	FR
Germany	DE
Greece	GR
Hungary	HU
Ireland	IE
Italy	IT
Latvia	LV
Lithuania	LT
Luxembourg	LU
Malta	МТ
Netherlands	NL*
Poland	PL
Portugal	PT
Romania	RO
Slovakia	SK
Slovenia	SI
Spain	ES
Sweden	SE
United Kingdom	UK*

* In text, referred to as the Czech Republic, the Netherlands and the United Kingdom

Non-Member States reporting, 2012

Country	Country abbreviations
Iceland	IS
Norway	NO
Switzerland	СН



Definitions

Term	Definition and description				
'Antimicrobial resistant isolate'.	In the case of quantitative data, an isolate was defined as 'resistant' to a selected antimicrobial when its minimum inhibitory concentration (MIC) value (in mg/L) was above the cut-off value or the disc diffusion diameter (in mm) was below the cut-off value. The cut-off values, used to interpret MIC distributions (mg/L) for bacteria from animals and food, are shown in Table MM11.				
		alitative data, an isolate was regarded resistant when the it as resistant using its own cut-off value or break point.			
'Level of antimicrobial resistance':	The percentage of resistant isolates among the tested isolates.				
'Reporting MS group':	table for antimicr	MSs) that provided data and were included in the relevant obial resistance data for the bacteria-food/animal crobial combination.			
Terms used to describe the	Rare:	< 0.1 %			
antimicrobial resistance levels	Very low:	0.1 % to 1 %			
	Low:	>1 % to 10 %			
	Moderate:	>10 % to 20 %			
	High:	>20 % to 50 %			
	Very high:	>50 % to 70 %			
	Extremely high:	>70 %			



APPENDIX 8. List of institutions contributing to antimicrobial resistance monitoring in animals and food

List of institutions contributing to antimicrobial resistance monitoring in animals and food

Member State	Institution
Austria	Federal Ministry for Health
	Austrian Agency for Health and Food Safety (AGES)
	Veterinary and Agrochemical Research Centre (CODA-CERVA), Uccle
Belgium	Institute of Public Health, Brussels
	Federal Agency for the Safety of the Food Chain, Brussels
Bulgaria	National Diagnostic and Research Veterinary Institute, Sofia
	Bulgarian Food Safety Agency, Sofia
Cyprus	 Veterinary Services, Nicosia Ministry of Agriculture, Nicosia
Czech Republic	 State Veterinary Institute, Prague and Olomouc State Veterinary Administration of the Czech Republic, Prague
	National Food Institute, Technical University of Denmark
Denmark	 Danish Veterinary and Food Administration
	Estonian Veterinary and Food Laboratory, Tartu
Estonia	 Veterinary and Food Board, Tallinn
Finland	EVIRA, Finnish Food Safety Authority, Helsinki
	ANSES, French Agency for Food, Environmental Occupational Health and Safety:
France	Fougères Laboratory, Maisons-Alfort Laboratory, Ploufragan/Plouzané Laboratory
	 Ministère de l'agriculture, de l'alimentation, de la pêche, de la ruralité et de l'aménagement du terriroire, Direction Générale de l'Alimentation, Paris
Germany	Federal Institute for Risk Assessment (BfR), Berlin
Crosso	Veterinary Laboratory, Chalkis
Greece	Ministry of Rural Development and Food, Athens
Hungary	Central Agricultural Office, Veterinary Diagnostical Directorate, Budapest
Tungary	Ministry of Rural Agriculture, Budapest
Ireland	Central Veterinary Research Laboratory, Celbridge
	Food Safety Authority of Ireland, Dublin
Italy	Istituto Zooprofilattico Sperimentale delle Regioni Lazio e Toscana, Rome
	Ministry of Health, Rome
Latvia	 Institute of Food Safety, Animal Health and Environment "BIOR", Animal Disease Diagnostic Laboratory, Riga
	 Food and Veterinary Service of Latvia, Riga
Lithur '	National Food and Veterinary Risk Assessment Institute, Vilnius
Lithuania	State Food and Veterinary Service, Vilnius
Luxembourg	Laboratoire de Médecine Vétérinaire, Luxembourg
Malta	Ministry for Resources and Rural Affairs

Table continued overleaf.



List of institutions contributing to antimicrobial resistance monitoring in animals and food (continued)

Member State	Institution
Netherlands	 Central Veterinary Institute, part of Wageningen UR (CVI), Lelystad National Institute of Public Health and the Environment (RIVM), Bilthoven Ministry of Agriculture, Nature and Food Quality Animal Health Service, Deventer
Poland	 National Veterinary Research Institute, Pulawy General Veterinary Inspectorate, WARSAW
Portugal	 Laboratório Nacional de Investigação Veterinária, Lisbon Direcção Geral de Veterinária, Lisbon
Romania	 Institute for Diagnostic and Animal Heath, Bucharest Institute for Hygiene and Veterinary Public Heath, Bucharest National Sanitary Veterinary and Food Safety Authority, Bucharest
Slovakia	 State Veterinary and Food Institute, Dolny Kubin and Bratislava State Veterinary and Food Administration of the Slovak Republic
Slovenia	 National Veterinary Institute, Veterinary Faculty, Ljubljana Ministry for Agriculture and Environment, Veterinary Administration, Ljubljana
Spain	 Laboratorio Central de Sanidad Animal de Santa Fe, Granada Laboratorio Central de Veterinaria de Algete, Madrid VISAVET Health Surveillance Center, Complutense University, Madrid Ministerio de Agricultura, Alimentación y Medio Ambiente Agencia Española de Seguridad Alimentaria y Nutrición
Sweden	 National Veterinary Institute (SVA), Department of Animal Health and Antimicrobial Strategies, Uppsala National Food Administration, Uppsala
United Kingdom	Animal Health and Veterinary Laboratories Agency (AHVLA)

Other reporting country	Institution
Norway	Norwegian Veterinary Institute
Switzerland	 ZOBA–Centre for Zoonoses, Bacterial Animal Diseases and Antimicrobial Resistance– Institute of Veterinary Bacteriology, Vetsuisse Faculty, University of Bern Swiss Federal Veterinary Office