# Exploring the prevalence of antibiotic resistance patterns and drivers of antibiotics resistance of *Salmonella* in livestock and poultry-derived foods: a systematic review and meta-analysis in Bangladesh from 2000 to 2022

Rezaul Karim Ripon<sup>1</sup>\*, Umma Motahara<sup>1</sup>, Ayesha Ahmed<sup>1</sup>, Nishrita Devnath<sup>1</sup>, Fatema Akter Mahua<sup>1</sup>, Rubaiya Binthe Hashem<sup>1</sup>, Kifayat Sadmam Ishadi<sup>1</sup>, Adiba Alam<sup>1</sup>, Md. Safaet Hossain Sujan<sup>1</sup> and Md Samun Sarker<sup>2</sup>

<sup>1</sup>Department of Public Health and Informatics, Jahangirnagar University, Savar, Dhaka, Bangladesh; <sup>2</sup>Antimicrobial Resistance Action Center (ARAC), Bangladesh Livestock Research Institute (BLRI), Savar, Dhaka, Bangladesh

\*Corresponding author. E-mail: riponrezaul5@gmail.com

#### Received 28 October 2022; accepted 19 April 2023

**Background:** Antimicrobial resistance (AMR) is a severe public health problem that Bangladeshis are dealing with nowadays. However, we wanted to investigate the pooled prevalence of *Salmonella* and AMR in *Salmonella* strains isolated from livestock- and poultry-derived foods between 1 January 2000 and 31 August 2022.

**Methods:** The metafor and metareg packages in the R programming language were used to conduct all analyses. We used a random-effect or fixed-effect model for pooled prevalence of *Salmonella* and AMR to *Salmonella*, depending on the heterogeneity test for each antibiotic. The heterogeneity was examined using stratified analyses, the meta-regression approach and sensitivity analysis.

**Results:** The combined prevalence of *Salmonella* in livestock and poultry-derived food in Bangladesh is 37%, according to the 12-research considered (95% CI: 23%–52%). According to subgroup analysis, neomycin had the lowest prevalence of resistance (4%, 95% CI: 1%–13%), whereas tetracycline had the highest prevalence of resistance (81%, 95% CI: 53%–98%). According to univariate meta-analysis and correlation analysis, the prevalence of *Salmonella* increased with the study period ( $\beta$ =0.0179; 95% CI: 0.0059–0.0298, *P*=0.0034;  $R^2$  = 46.11%) and without this, none of aforementioned variables was significantly associated with the detected heterogeneity and there was a positive relationship (*r*=0.692, *P*=0.001) between the *Salmonella* prevalence and study period.

**Conclusions:** AMR is rising alarmingly in Bangladesh by livestock-derived food consumption. However, monitoring and evaluating antibiotic sensitivity trends and developing effective antibiotic regimens may improve *Salmonella* infection inhibition and control in Bangladesh. Policymakers should be concerned about food handling practices. Doctors should be concerned when using prescribing antibiotics.

# Introduction

Salmonella is one of the most commonly recognized pathogens that cause gastroenteritis,<sup>1,2</sup> which results in significant morbidity, mortality and economic loss.<sup>3,4</sup> In 2010, the World Health Organization (WHO) reported 153 million cases of non-typhoidal *Salmonella* (NTS) enteric infections worldwide, of which 56 969 were fatal and 50% were foodborne.<sup>5</sup> *Salmonella* was the second foodborne epidemic in some regions' illness monitoring reports from 2006 to 2010.<sup>6</sup> Among the 2600 *Salmonella* serotypes discovered, NTS serovars such as Typhimurium and Enteritidis are the most common worldwide.<sup>7,8</sup> Poultry has been identified as the one cause of human salmonellosis, and avian salmonellosis affects the poultry business and can infect humans when infected poultry meat and eggs are consumed.<sup>9</sup> Eggs are the principal source of salmonellosis and other foodborne illnesses<sup>10-13</sup> *Salmonella* that grows in animal farms may contaminate eggs and meat during the slaughtering process before being

© The Author(s) 2023. Published by Oxford University Press on behalf of British Society for Antimicrobial Chemotherapy. This is an Open Access article distributed under the terms of the Creative Commons Attribution-NonCommercial License (https:// creativecommons.org/licenses/by-nc/4.0/), which permits non-commercial re-use, distribution, and reproduction in any medium, provided the original work is properly cited. For commercial re-use, please contact journals.permissions@oup.com transmitted to people via the food chain. Indeed, multiple earlier investigations have reported the isolation of *Salmonella* from animal and human diets.<sup>14–17</sup> Human *S*. Enteritidis is commonly associated with the intake of infected eggs and chicken meat, whereas *S*. Typhimurium is commonly associated with the consumption of pork, poultry and beef.<sup>18,19</sup> In addition, *Salmonella* enterica serovars have been found in varying concentrations in animal products and by-products worldwide.<sup>18,20,21</sup> The most often-reported serovars connected with human foodborne diseases are *Salmonella* Typhimurium and Enteritidis.<sup>22</sup> However, untyped *Salmonella* of animal origin is becoming more common in Bangladesh.<sup>23,24</sup>

However, domestic chickens in developing countries live close to humans in urban and rural communities and are frequently housed overnight in the family home.<sup>25</sup> As soon as a chicken becomes infected, it sheds faeces into the environment. Additionally, interacting with employees in poultry farms and slaughterhouses, the main route of human *Salmonella* infections, involves contaminated meat and eggs.<sup>26</sup> One of the primary reasons for animal management is the risk of antimicrobial resistance (AMR) in humans and animals.<sup>27</sup>

AMR is a worldwide public health issue.<sup>28,29</sup> AMR can be caused by one of three fundamental processes: (i) antibiotic modification by lowering absorption or enhancing efflux of the antibiotic via their enzymes; (ii) alteration in the antibiotic's target site and (iii) gaining the capacity to break or change the



Figure 1. PRISMA flow diagram for this study.

antibiotic.<sup>30</sup> Several lines of evidence showed that using antimicrobial agents in food animals contributes to the emergence and spread of AMR in foodborne Salmonella.<sup>31</sup> AMR has recently been a significant issue in treating Salmonella infections.<sup>32,</sup> Salmonella infections in food animals are essential in public health and, in particular, food safety because food products of animal origin are thought to be the most common source of human Salmonella infections.<sup>34</sup> Contamination by healthy food handlers is also assessed during food processing. In recent years, it has been estimated that animals and their products can account for up to 96% of all Salmonella infections in humans.<sup>35,36</sup> AMR is expected to increase by 70% in Asia, posing a national and alobal threat.<sup>37</sup> The WHO estimates that Salmonella infections cause 93.8 million instances of gastroenteritis worldwide each year and that Salmonella infections cause 155000 fatalities.<sup>38</sup>

Moreover, according to a recent Shanghai study, just 1.1% of strains were responsive to all 16 medications, and AMR rates for third and fourth-generation cephalosporins (cefotaxime and cefepime) were 10% and 8.1%, respectively.<sup>39</sup> Furthermore, according to a study conducted in Guanazhou, annual resistance rates of ampicillin are reasonably consistent. However, resistance rates of NTS to ceftazidime in 2015 (31.43%) were significantly areater than in 2014 (16%). Furthermore, AMR to ampicillin was considerably higher in serotype Typhimurium and Enteritidis isolates than in other serotypes.<sup>40</sup> Salmonella drug resistance rates to cephalosporin and cefepime were 22.3% and 13.1%, respectively, in four hospitals in Shenzhen,<sup>41</sup> which were all higher than the results of earlier Chinese investigations.<sup>42,43</sup> This phenomenon demonstrates that the outlook for AMR is bleak. Regarding AMR mechanisms, the corresponding resistance genes are usually found on plasmids, transposons, gene cassettes or variants of the Salmonella genomic islands SGI1 and SGI2.44

However, this study revealed a prevalence of *Salmonella* in poultry and livestock-derived foods. In this regard, this meta-analytical study will be evidential for assessing the prevalence of *Salmonella* in livestock and poultry-derived food and AMR in livestock and poultry-derived foods and comprehensively investigate the whole scenario of Bangladesh. We hypothesize that *Salmonella* pooled prevalence and antibiotic resistance are increasing in Bangladesh.

# Materials and methods

#### Data sources and systematic search strategy

We looked for research focused on investigating the prevalence of *Salmonella* and AMR in livestock and poultry-derived food in both English and Bangla. In English and Bangla, an attempt was made to locate grey and published scientific literature. From 2000 to August 2022, search engines such as Google Scholar, Scopus, ISI Web of Knowledge and PubMed were used to identify published literature, with reference lists of pertinent articles manually searched. We did not find any article on Bangla. The screening of titles and abstracts to find relevant publications was followed by full-text scanning of the relevant articles in a two-step approach.

Additionally, we were concerned with resolving any conflicts that developed during the data extraction to eliminate selection bias. *'Salmonella'* AND 'antibiotic resistance OR antimicrobial resistance OR

Resistance antibiotics	Ampicillin, Tetracycline, Ialidixic acid, Ciprofloxacin Sentamicin, Erythromycin, Rifampicin, Streptomycin, Kanamycin, zithromycin	Enrofloxacin	Ampicillin, Cloxacillin, Nalidixic acid, Ervthromvcin	Ampicillin, Amoxicillin, Tetracycline, hloramphenicol, Nalidixic acid, ientamicin, Erythromycin, Kanamcin	Ampicillin, Penicillin, Tetracycline, Inloramphenicol, Nalidixic acid, Ciprofloxacin, Erythromycin, Norfloxacin, Rifampicin, Cenhalexin	Ampicillin, Nalidixic acid, Gentamicin, Azithromvcin. Cebhalexin	Amoxicillin, Oxytetracycline, Doxycycline, Sulpha/Trimetho/Cotri, Ciprofloxacin, ientamicin, Erythromycin, Neomycin, Amikacin Axithromycin	Ampicillin, Amoxicillin, Tetracycline, 
Quality of articles	4 7 0 1	٢	4	4	4	4	ى ب	~ 0
AST standard	CLSI	CLSI	CLSI	CLSI	CLSI	CLSI	CLSI	NCCLS
AST method	Disc diffusion	Disc diffusion, Broth	Disc diffusion	Disc diffusion	Agar	Disc diffusion	Disc diffusion	Disc
Sample type	Meat	egg shell surface, egg content	Meat	Clocal swab, Foot pad Faeces	Cloacal swab, Intestinal fluid, Egg surface	Cloacal swab, Pharyngeal swab	Chicken meat, Milk, Beef	Egg
Positive no.	10	71	24	40	71	17	37	148
Sample no.	20	240	24	112	210	50	169	200
Host	Chicken	Chicken	Chicken	pigeon	Chicken	Pigeon	Chicken, Cow	Chicken
Location	Dhaka	Chittagong	Mymensingh, Feni, Dhaka	Mymensingh	Dhaka	Mymensingh	Mymensingh, Gazipur	Dhaka
Study year	2013	2013	2004	2012	2010	2015	2017	2017
Author, year	Fatema et al., 2014	Mahmud et al., 2016	Khan <i>et al.</i> , 2000	Hosain et <i>a</i> l., 2012	Akond et al., 2012	Saifullah et al., 2016	Rahman et al., 2018	Islam et al., 2018

Review

# JAR

Resistance antibiotics	acid, Ciprofloxacin, Gentamicin, Kanamycin, Azithromycin, Ceftriaxone Amoxicillin, Tetracycline, Sulpha/Trimetho/Cotri, Erythromycin,	Tetracycline, Tetracycline, Oxytetracycline, Chloramphenicol, Ciprofloxacin, Gentamicin, Erythromycin, Neomycin, Kanamycin, Ertapenem, Meropenem, Tminenem	Ampicillin, Amoxicillin, Tetracycline, Ciprofloxacin, Ervthromvcin Azithromvcin	Ampicillin, Tetracycline, Gentamicin	Amoxicillin, Tetracycline, Doxycycline, Colistin, Ciprofloxacin, Pefloxacin, Kanamycin, Enrofloxacin	Ampicillin, Tetracycline, Chloramphenicol, Ciprofloxacin, Strentomycin, Ertanenem	Amoxicillin, Tetracycline, Colistin, Erythromycin, Neomycin	Ampicillin, Tetracycline, Sulpha/Trimetho/Cotri, Ciprofloxacin, Erythromycin, Neomycin
Quality of articles	Ч	4	Ø	4	σ	4	4	4
AST standard	CLSI	CLSI	CLSI	CLSI	NCCLS		CLSI	CLSI
AST method	Agar	Disc	Disc	Disc	Disc	Disc	Disc	Disc
Sample type	Aii	Cow dung, Milk	Meat	Egg	Liver, Spleen	Cloacal swab	Cloacal swab	Gut materials
Positive no.	24	90	14	50	13	35	10	46
Sample no.	200	100	20	50	30	100	75	320
Host	Cattle	Cattle	Chicken	Chicken	Chicken	Chicken	Quails	Chicken
Location	Sylhet	Mymensingh	Jamalpur, Netrokona, Tangail, Kishoreaani	Dhaka	Chittagong	Mymensingh	Mymensingh	Sylhet
Study year	2018	2018	2017	2017	2012	2017	2017	2017
Author, year	Khan et al., 2019	Sobur et al., 2019	Islam <i>et al.</i> , 2018a	Rahman et al 2019	Hassan et al., 2014	Alam <i>et al.</i> , 2020	Jahan <i>et al.</i> , 2018	Joy et al., 2017

Table 1. Continued

Not clear	Erythromycin; Cloxacillin; Ampicillin; Trimethoprim; Nitrofurantoin; Ciprofloxacin; Cefotaxime; Ceftriaxone; Levofloxacin; Co-Trimoxazole	Amikacin, Streptomycin, Gentamicin, Azithromycin, Oxytetracycline, Amoxicillin, Oxacillin, Sulfamethoxazole- Trimethoprim, Ceftriaxone, Ciprofloxacin	Amoxicillin, Penicillin, Chloramphenicol, Erythromycin, Cefradine, Oxytetracycline, Enrofloxacin, Penicillin, Erythromycin, Cotrimoxazole, Cefradine, Amoxicillin, Penicillin, Cefradine	Amoxicillin, Penicillin, Chloramphenicol, Erythromycin, Cefradine, Oxvtetracvcline.	Penicillin, Chloramphenicol, Erythromycin, Cefradine, Oxytetracycline
6	7	∞	٥	2	7
CLSI	CLSI	CLSI	CLSI	CLSI	CLSI
Disc	Disc	Disc	Disc	Disc	Disc
Chicken meat, Liver	Meat	Chicken meat, Frozen milk	Chicken meat, Cattle meat, Buffalo meat and Goat meat	Meat	Meat
8		13	19	82	35
16	52	40	205	189	50
Poultry Chickens	Poultry	Poultry	Chicken, Cattle, Goat	Chicken	Chicken
Chattogram	Dhaka	Barishal	Barishal, Pirojpur and Bhola	Dhaka	Dhaka
2019- 2020	2020	2020- 2021	Not clear	2017	2006
Islam <i>et al.</i> , 2022	Rabby et al., 2021	Rahman et al., 2022	Matubber et al., 2021	Rahman et al., 2018a	Begum et al., 2007



Figure 2. Sensitivity analysis for individual studies.



Figure 3. Funnel plot for study bias.

drug resistance OR AMR' AND 'prevalence OR incidence OR morbidity OR odd ratio OR risk ratio OR confidence interval OR *P* value OR rate' AND 'antibiotic resistance OR antimicrobial resistance OR drug resistance' AND 'Bangladesh'. Studies conducted on livestock and poultry were chosen using the Species filters in PubMed and Google Scholar. Because there was no sorting filter for species in the Web of Science and Cochrane Library, new search phrases were added to select species. The articles found during the search were exported to EndNote to be checked for duplicates. The unique hits were then uploaded to the Rayyan QCRI website for data extraction and screening. The title and abstract were screened first, followed by the complete content of the article.

#### Data collection process and data items

As a result of our search approach, all items were exported to the EndNote program. Duplicated articles were removed from the database. One neutral reviewer (A.A.) examined the titles and abstracts of the discovered papers. The whole texts of qualifying papers were obtained and appropriately evaluated for eligibility. The records were examined for legitimacy by the second examiner (M.S.S.). In the event of a disagreement between the two reviewers (A.A. and M.S.S.), the reviewer was then consulted by F.A.M. and A.M.S. Then, the article information was included by the three reviewers (N.D., K.S.I. and R.B.G.).

Antibiotic name	Tau <sup>2</sup>	$I^2$	$\chi^2$	Random/fix effect of resistance prevalence, % [95% CI]	P for H	P for Egger
Ervthromvcin	0.13	94%	27.40	84[53–99]	<0.001	0.424
Ciprofloxacin	0.01	95%	143.4	20[4-42]	< 0.001	0.523
Gentamicin	0.05	90%	71.82	11[2-24]	<0.001	0.242
Oxytetracycline	0.0097	65%	5.57	52[39–66]	0.06	0.654
Sulpha/Trimetho/Cotri	0.15	97%	144.29	42[11-70]	< 0.001	0.524
Tetracycline	0.1	93%	60.48	80[53–98]	<0.001	0.525
Neomycin	0.1	60%	7.45	4[1-13]	0.06	0.131
Doxycycline	0.122	95%	35.63	51[12-89]	<0.001	0.354
Overall Salmonella prevalence	0.105	97%	558.13	37[23-52]	<0.001	0.636

Table 2. Overall result in meta-analysis

 Table 3. Univariate meta-regression analysis

Covariates	Beta [β] [95%CI]	P value	R <sup>2</sup> [%]
Studv vear	0.0179 [0.0059 -0.0298]	0.0034	46.11%
Location	-0.0073 [-0.0199 -0.0053]	0.2557	Nil
Host	-0.0085 [-0.0363 -0.0193]	0.5497	Nil
Sample number	0.0024 [-0.0017 -0.0064]	0.2623	20.98%
Positive sample	-0.0069 [-0.0230 -0.0093]	0.4037	Nil
AST method	-0.0106 [-0.0274 -0.0061]	0.2125	Nil
AST standard	0.0028 [-0.0016 -0.0072]	0.2140	15.22%
Quality of articles	-0.0384 [-0.1333 -0.0565]	0.4280	Nil
Resistance antibiotic	0.0097 [-0.0313 -0.0119]	0.3796	Nil

Furthermore, the Final was checked by one reviewer (R.K.R.). Next, statistical analysis was performed by R.K.R. and U.M. The search strategy was depicted in a PRISMA flow chart, which showed which studies were included and excluded and the reasons for exclusion (Figure 1).

#### Data extraction and quality assessment

We choose studies on the following criteria: (i) reported AMR in livestock and poultry-derived foods in Bangladesh; (ii) published between 2000 and August 2022; (iiii) samples indicated poultry and livestock-derived foods and (iv) were scientifically reported on location, number of samples and outcomes. The analysis did not cover the detection of *Salmonella* and the resistance pattern of sample types of faeces, faecal, cloacal, intestinal and rectal. In Bangladesh, there were no restrictions on when the study might be published.

#### Reporting bias assessment

Data quality was assessed according to the Joanna Briggs Institute.<sup>45–47</sup> Appropriate sample frame, study participants sampled, sample size, description of study subjects and setting, sample size justification, power description or variance and effect estimates, valid methods for condition identification, a standard and reliable condition measured appropriate statistical analysis, and adequate response rate were among the nine factors used to assess the risk of bias. The phrases 'yes', 'no', 'unclear' and 'not available' were used to convey the risk assessment criteria. They received a one-point score, while the others received a zero. The total score was 0 to 9. The likelihood of bias was considered low when the overall score was greater than 70%, moderate when it was 50%–69% and high when it was 0%–49% (Table 1).<sup>45,47</sup>

#### Statistical analysis

After reviewing the entire article for quality, the authors independently extracted data using purpose-built forms that specified the pertinent factors. Disagreements were settled by debating the articles and coming to an agreement. The significant dependent variable was a binary categorization of study results on the basis of whether the article supported the prevalence of Salmonella and antibiotic resistance to Salmonella or not. The proportion for the study based on the standardized effect size was the dependent variable in the meta-analysis. The statistical analysis was carried out with the help of the R programming language. The metafor and metareg in the R programming language were used to determine the pooled prevalence after excluding low-quality articles.<sup>40,45,46</sup> Metafor uses inverse-variance weights from a random-effects model to pool proportions and offers a weighted subgroup and overall pooled estimate. It entailed a meta-analysis of the prevalence values of each publication, weighted by sample size and allowing for potential heterogeneity across studies in this context. Before the random-effects model, the Freeman-Tukey transformation produced a proportional meta-analysis. Prevalence estimates were merged using a random-effects metaanalysis for accounting for between-study heterogeneity. The statistical heterogeneity was assessed using the chi-square test on the  $Q(H, I^2)$  statistic, which was quantified by the  $I^2$  values under the assumption that I-square values of 25%, 50% and 75% were nominally assigned as low, moderate and high estimates, respectively.<sup>40,45</sup> The following grouping variables were used in stratified analyses and meta-regression for covariates to examine potential sources of heterogeneity. To establish whether one or more studies had an impact on the outcomes by being excluded one at a time, a sensitivity analysis was also performed. The distribution of the observed studies was visually inspected on a funnel plot to determine publication bias. Egger's linear regression and Begg's rank correlation test were then used to measure the level of bias.40,45,46 The significance threshold was kept at <0.05.

# Results

#### Search result and eligible studies

PRISMA was used to outline the specific steps of the systematic review and meta-analysis method, and Figure 1 outlines the process of selecting relevant papers. We identified 463 new studies via PubMed, Google Scholar, Scopus, ISI and Web of Knowledge databases. Owing to duplicate records, ineligible by automation tools, titles, sample types and study had mixing with other bacteria, 419 articles were removed. Twenty-two studies were removed for not indicating prevalence, being systematic and not being peer-reviewed. According to the quality assessment standards, two studies received a score of 9, two got an 8, six got a 7, two got a 6 and 10 got a 4. According to the quality scores, the studies were generally deemed to be of acceptable quality (scores more than or equal to  $6)^{47,48}$ . Ten articles were also removed for methodological quality. For that, we included 12 studies in this meta-analysis.

#### Characteristics of eligible studies

Table 1 included the characteristics of eligible studies The study sample included Chittagong, Mymensingh, Sylhet, Chittagong and Barisal divisions. The hosts were chickens, cattle and goats. These three are the most commonly used livestock for a portion of food in Bangladesh. According to the included study, 1411 livestock-derived food were used as a study sample, and the review found 471 samples to be *Salmonella* positive and resistant to some antibiotics. Most samples were identified by the disc diffusion method, and the CLSI method was used as an AST method. Tetracycline, oxytetracycline, doxycycline, sulpha/trimetho/ cotri, ciprofloxacin, gentamycin and neomycin have frequently shown resistance against *Salmonella* in many reviewed studies.

#### Systematic review and meta-analysis results

A higher level of multi-drug resistance was present in Bangladesh. In Bangladesh, the combined prevalence of Salmonella isolated from livestock and poultry-derived foods (sample types faeces, faecal, cloacal and rectal excluded) was 37% of the sample. Overall, 37% of the sample had Salmonella in livestock and poultry-derived foods [95% CI: (23%–52%),  $P \leq$ 0.001,  $R^2 = 97\%$ ]. A high prevalence was seen for some of the commonly prescribed antibiotics like tetracycline 81% [(95% CI:53-98),  $P \le 0.001$ ,  $R^2 = 93\%$ ], oxytetracycline 52% [(95% CI: 38%-66%), P = 0.06,  $R^2 = 65\%$ ], doxycycline 51% [(95% CI: 12%-86%),  $P \le 0.001$ ,  $R^2 = 95\%$ ], sulpha/trimetho/cotri 42% [(95% CI: 11%-77%),  $P \le 0.001$ ,  $R^2 = 97\%$ ], ciprofloxacin 20% [(95% CI: 4-43),  $P \le 0.001$ ,  $R^2 = 95\%$ ], gentamycin 11% [(95% CI: 2%–25%),  $P \le 0.001$ ,  $R^2 = 90\%$ ] and neomycin 4% [(95% CI:1%-13%), P = 0.06,  $R^2 = 60\%$ ] (Table 2 and Figures S1-S9 (available as Supplementary data at JAC-AMR Online)). Among the included covariate in this study, only the study period showed significance in the univariate meta-analysis ( $\beta = 0.0179$ ; 95% CI: 0.0059–0.0298, P = 0.0034;  $R^2 = 46.11\%$ ) (Table 3). Sensitivity analysis was carried out by calculating pooled Salmonella prevalence once more when any single study was eliminated to verify the meta-analysis's stability and liability. The related pooled Salmonella prevalence was shown range from 40.2% (33.0%-48.5%) to 33.8% (30.0%-35.6%) without significantly changing (Figure 2). The statistically identical findings indicated that no single study had an impact on the stability

of the overall estimate of the prevalence of *Salmonella* in this meta-analysis. Even though the funnel plot's visual inspection exhibited asymmetry (Figure 3), Egg's test demonstrated substantial value, and there was no chance of publication bias (Table 2).

There was considerable variation between studies. To investigate potential sources of heterogeneity, meta-regression (univariate) was used. The meta-regression method was used to examine the study period, location, host, sample number, positive sample, AST method, AST standard, quality of articles and resistance antibiotic as potential sources of heterogeneity. Table 3 contains the meta-finding on regressions. None of those previously mentioned factors were found to be substantially correlated with the identified heterogeneity through the regression model, except for research study time (P = 0.004). Therefore, we conducted additional research to examine the relationship between *Salmonella* prevalence and the study period. The prevalence of *Salmonella* was found to be positively correlated with the research study period (r = 0.692, P = 0.001) (Table 4).

#### Evidence-based antibiotic resistance drivers

AMR is an upcoming predicted pandemic for a low-income country such as Bangladesh.<sup>29-33</sup> Bangladesh has limited information on antibiotic use and resistance.<sup>34</sup> Antibiotics were considered able to treat both the physical and social elements of infection, which has severe implications for AMR.<sup>22</sup> AMR has mainly spread throughout the country's hospitals and unregulated pharmacy stores for their widely available antibiotics, overprescribing and selling antibiotics without prescription.<sup>32</sup> People in hospitals are predisposed to developing nosocomial and other infections due to the abuse of antibiotics and a lack of adherence to isolation techniques in these hospitals.<sup>27</sup> In Bangladesh, most antibiotic prescriptions are written by unqualified practitioners.<sup>28,30</sup> Owing to poor health systems, reported paucity of testing facilities, prescribing antibiotics without laboratory tests, using antibiotics for common infections and not finishing the entire course of antibiotics were causes of widespread resistance.<sup>33</sup> Unsafe drinking water, inadequate sanitation, lack of diagnostic facilities growing private practice and excessive demand for antibiotics are all contributing causes of AMR.<sup>30,33</sup> Antibiotic usage before seeking medical help may impair the sensitivity of blood cultures, which has ramifications for patients and doctors.<sup>29</sup> Antibiotics are used in various food animal production and meat systems, such as commercial poultry and aquaculture.<sup>36</sup> They have been discovered to be frequently used to boost food animal production, leading to the growth of antibiotic-resistant bacteria.<sup>33</sup>

Moreover, its use in aquaculture and aquatic ecology has been linked to AMR development.<sup>28</sup> Antimicrobial drugs used in aquaculture have established AMR bacteria reservoirs in fish and other aquatic and non-aquatic animals.<sup>29-32</sup> The long-term use of

Table 4. The correlation between the Salmonella prevalence and potential sources

Covariate	Study year	Location	Host	Sample no.	Positive no.	Sample type	AST method	AST standard	Quality of articles	Resistance antibiotics
P	0.001	0.744	0.563	0.785	0.325	0.325	0.563	0.566	0.743	0.774
r	0.692	0.764	0.346	0.633	0.764	0.763	0.567	0.568	0.366	0.852

these antibiotics indiscriminately and unnecessarily increases the prevalence of AMR.<sup>28-31</sup> This also leads to antibiotic residue. Antibiotic residues, other drugs and pollutants are found in high concentrations in Bangladesh's ponds, canals, lakes, rivers and other bodies of water.<sup>36</sup> Higher residues are caused by poor sanitation, hygiene, antibiotic misuse on the farm and inappropriate use.<sup>33</sup> As part of the food production cycle, food animals, seafood and vegetables are regarded significant reservoirs of AMR bacteria by these residues.<sup>26</sup> Some antibiotics may have lost their effectiveness against specific microorganisms due to the misuse of those antibiotics.<sup>30</sup> Antimicrobials are only available on prescription and to pharmacists who devote more time to patients. Antibiotics are sought and used during an emergency health problem.<sup>24</sup> Policy measures such as restrictions on licensing specific antibiotics are vital to human health.<sup>25</sup> In Bangladesh, specific and targeted measures to combat AMR should include teaching about the proper use of antibiotics.<sup>31</sup> There is a significant frequency of AMR in Bangladesh, and major gaps in surveillance and information, and there has been a drop in the rate of new antibiotic development.<sup>23</sup>

# Discussion

The present study found a significant prevalence of Salmonella from livestock and poultry-derived foods in Bangladesh (sample type faeces, faecal, collocal and rectal exclude). We found that 37% of livestock and poultry food had Salmonella. Consequently, livestock and poultry-derived foods appear to be one of the essential Salmonella reservoirs in Bangladesh. A meta-analysis of Salmonella in Ethiopia indicates that the prevalence of Salmonella in slaughterhouse may vary from 7% to 43%,<sup>46</sup> and a worldwide meta-analysis of *Salmonella* prevalence in food indicate that it was less than 1%.<sup>45,49</sup> Therefore, researchers have investigated Salmonella prevalence for many years, and different rates have been found in different studies. The reported prevalence of Salmonella found in this study matched and differed from the findings of other researchers studying the prevalence of Salmonella in poultry.<sup>50-57</sup> Thus, it can be mentioned that the significant amount of Salmonella indicates a severe problem for both livestock and poultry-derived food and ensures public health.

Moreover, globally AMR is becoming a significant health issue.<sup>53</sup> Resistant to many drugs, *Salmonella* has become a considerable public health concern worldwide.<sup>57</sup> Our present study findings also revealed that a high percentage of the *Salmonella* isolates were resistant to routinely used antibiotics such as tetracycline, oxytetracycline, doxycycline, sulpha/trimetho/cotri, ciprofloxacin, gentamycin and neomycin and so on. Almost all the isolates tested in this investigation were determined to be multi-drug resistant, which is a concerning finding (Table 1).

However, among these antibiotics, the highest prevalence rate was seen for tetracycline (81%, 95% CI: 53–98). Some studies showed that isolated *Salmonella* was highly resistant to tetracycline in the antimicrobial investigation.<sup>50–52</sup> The present study found the next high prevalence rate for resistance to oxytetracycline that shows 52% from the isolates (95% CI: 38–66), but some studies found a higher prevalence rate of oxytetracycline rather than our findings.<sup>45,46,48,49</sup> According to a couple of studies, *Salmonella* resistance to tetracycline and oxytetracycline was found in layers and broilers in Bangladesh.<sup>58–60</sup> Doxycycline is another antibiotic used in humans and animals to treat various diseases. Investigating multi-drug resistance due to *Salmonella*, our findings showed that a high prevalence for this commonly prescribed antibiotic is 51% (95% CI: 12–86). In addition, a recent study showed that *Salmonella* in poultry in Bangladesh has also been found to have a considerable number of isolates resistant to doxycycline.<sup>46,48,60</sup> Moreover, according to univariate analysis and correlation in this meta-analysis, the prevalence of *Salmonella* increased with every year. Antibiotic resistance creates a severe problem in the treatment of *Salmonella*.

Furthermore, *Salmonella* has been proven to be a significant cause of creating resistance against several commonly prescribed antibiotics. Resistant to many drugs, *Salmonella* isolates were discovered in various food samples, and the gene responsible for multi-drug resistance could be passed on to consumers through food and unhealthy, unhygienic poultry handling systems, posing a severe public health risk. These findings also revealed that multi-drug resistance in *Salmonella* is rising due to the indiscriminate use of antibiotics in the dairy and poultry industries, pet animal usage and human practice in Bangladesh. In the future, rational usage of this antibiotic may help to prevent the formation of *Salmonella*-resistant isolates.

The main strength of this article was the pooled prevalence of *Salmonella* and antibiotic resistance to *Salmonella* from livestock and poultry-derived foods. We solely refer to the antibiotic subgroup. We looked through numerous databases and websites to discover all relevant and grey publications to eliminate database bias, but some databases or websites may be massing. The limitations mentioned before, as well as publication bias and heterogeneity for some of the pooled results, must be considered when interpreting the results.

#### Conclusions

Antimicrobial agents used in food animal production may lead to the emergence of multi-drug-resistant *Salmonella* strains. The irrational and excessive use of antibiotics in humans and foodproducing animals raises the risk of AMR worldwide. To reduce the risk of pathogenic AMR bacteria originating from animal origin foods, raising awareness about the rational use of antibiotics in food animals, safe food handling and safe cooking practices is obligatory. Doctors should take concern when treating with an antibiotic.

## Acknowledgements

We are thankful to Sojib Bhuiyan for his technical support, Department of Accounting, Government Bangla College, Bangladesh.

# Funding

Self-funded.

#### **Transparency declarations**

None to declare.

# Supplementary data

Figures S1 to S9 are available as Supplementary data at JAC-AMR Online.

## References

**1** Hawker J, Begg N, Reintjes R *et al. Communicable Disease Control and Health Protection Handbook.* John Wiley & Sons, 2018.

**2** Jain P, Chowdhury G, Samajpati S *et al.* Characterization of nontyphoidal *Salmonella* isolates from children with acute gastroenteritis, Kolkata, India, during 2000–2016. *Braz J Microbiol* 2020; **51**: 613–27. https://doi. org/10.1007/s42770-019-00213-z

**3** Lin D, Yan M, Lin S *et al.* Increasing prevalence of hydrogen sulfide negative *Salmonella* in retail meats. *Food Microbiol* 2014; **43**: 1–4. https://doi. org/10.1016/j.fm.2014.04.010

**4** Sallam KI, Mohammed MA, Hassan MA *et al.* Prevalence, molecular identification and antimicrobial resistance profile of *Salmonella* serovars isolated from retail beef products in Mansoura, Egypt. *Food Control* 2014; **38**: 209–14. https://doi.org/10.1016/j.foodcont.2013.10.027

**5** Kirk MD, Pires SM, Black RE *et al.* World Health Organization estimates of the global and regional disease burden of 22 foodborne bacterial, protozoal, and viral diseases, 2010: a data synthesis. *PLoS Med* 2015; **12**: e1001921. https://doi.org/10.1371/journal.pmed.1001921

**6** Pang L, Zhang Z, Xu J. Surveillance of foodborne disease outbreaks in China in 2006–2010. *Chin J Food Hyg* 2011; **23**: 560–3.

**7** Issenhuth-Jeanjean S, Roggentin P, Mikoleit M *et al*. Supplement 2008–2010 (no. 48) to the White-Kauffmann-Le minor scheme. *Res Microbiol* 2014; **165**: 526–30. https://doi.org/10.1016/j.resmic.2014.07.004

**8** Takaya A, Yamamoto T, Tokoyoda K. Humoral immunity vs. *Salmonella*. *Front Immunol* 2020; **10**: 3155. https://doi.org/10.3389/fimmu.2019.03155

**9** Behravesh CB, Brinson D, Hopkins BA *et al*. Backyard poultry flocks and salmonellosis: a recurring, yet preventable public health challenge. *Clin Infect Dis* 2014; **58**: 1432–8. https://doi.org/10.1093/cid/ciu067

**10** Gieraltowski L, Higa J, Peralta V *et al.* National outbreak of multidrug resistant *Salmonella* Heidelberg infections linked to a single poultry company. *PLoS ONE* 2016; **11**: e0162369. https://doi.org/10.1371/journal. pone.0162369

**11** Keerthirathne TP, Ross K, Fallowfield H *et al.* Reducing risk of salmonellosis through egg decontamination processes. *Int J Environ Res Public Health* 2017; **14**: 335. https://doi.org/10.3390/ijerph14030335

**12** Biswas S, Li Y, Elbediwi M *et al.* Emergence and dissemination of MCR-carrying clinically relevant *Salmonella* Typhimurium monophasic clone ST34. *Microorganisms* 2019; **7**: 298. https://doi.org/10.3390/microorganisms7090298

**13** Yu H, Elbediwi M, Zhou X *et al*. Epidemiological and genomic characterization of *Campylobacter jejuni* isolates from a foodborne outbreak at Hangzhou, China. *Int J Mol Sci* 2020; **21**: 3001. https://doi.org/10. 3390/ijms21083001

**14** Ed-Dra A, Karraouan B, El Allaoui A *et al*. Antimicrobial resistance and genetic diversity of *Salmonella* Infantis isolated from foods and human samples in Morocco. *J Glob Antimicrob Resist* 2018; **14**: 297–301. https://doi.org/10.1016/j.jgar.2018.05.019

**15** Paudyal N, Pan H, Wu B *et al.* Persistent asymptomatic human infections by *Salmonella enterica* serovar Newport in China. *mSphere* 2020; **5**: e00163-20. https://doi.org/10.1128/mSphere.00163-20

**16** Jiang Z, Paudyal N, Xu Y *et al.* Antibiotic resistance profiles of *Salmonella* recovered from finishing pigs and slaughter facilities in Henan, China. *Front Microbiol* 2019; **10**: 1513. https://doi.org/10.3389/fmicb.2019.01513

**17** Elbediwi M, Pan H, Biswas S *et al*. Emerging colistin resistance in *Salmonella enterica* serovar Newport isolates from human infections. *Emerg Microbes Infect* 2020; **9**: 535–8. https://doi.org/10.1080/22221751.2020.1733439

**18** Park HC, Baig IA, Lee SC *et al.* Development of ssDNA aptamers for the sensitive detection of *Salmonella* Typhimurium and *Salmonella* Enteritidis. *Appl Biochem Biotechnol* 2014; **174**: 793–802. https://doi.org/10.1007/s12010-014-1103-z

**19** Spector MP, Kenyon WJ. Resistance and survival strategies of *Salmonella* Enterica to environmental stresses. *Food Res Int* 2012; **45**: 455-81. https://doi.org/10.1016/j.foodres.2011.06.056

**20** de Freitas CG, Santana AP, da Silva PH *et al*. PCR multiplex for detection of *Salmonella* Enteritidis, Typhi and Typhimurium and occurrence in poult-ry meat. *Int J Food Microbiol* 2010; **139**: 15–22. https://doi.org/10.1016/j. ijfoodmicro.2010.02.007

**21** Shah AH, Korejo NA. Antimicrobial resistance profile of *Salmonella* serovars isolated from chicken meat. *J Vet Anim Sci* 2012; **2**: 40–6.

**22** Suresh T, Hatha AA, Sreenivasan D *et al*. Prevalence and antimicrobial resistance of *Salmonella* Enteritidis and other salmonellas in the eggs and egg-storing trays from retails markets of Coimbatore, South India. *Food Microbiol* 2006; **23**: 294–9. https://doi.org/10.1016/j.fm.2005.04.001

**23** Momtaz S, Saha O, Usha MK *et al.* Occurrence of pathogenic and multidrug resistant *Salmonella* spp. in poultry slaughter-house in Bangladesh. *Bioresearch Communications (BRC)* 2018; **4**: 506–15. https://www.bioresearchcommunications.com/index.php/brc/article/view/80

**24** Sultana M, Bilkis R, Diba F *et al.* Predominance of multidrug resistant zoonotic *Salmonella* Enteritidis genotypes in poultry of Bangladesh. *J Poult Sci* 2014; **51**: 424–34. https://doi.org/10.2141/jpsa.0130222

**25** Dana N, Van der Waaij LH, Dessie T *et al.* Production objectives and trait preferences of village poultry producers of Ethiopia: implications for designing breeding schemes utilizing indigenous chicken genetic resources. *Trop Anim Health Prod* 2010; **42**: 1519–29. https://doi.org/10. 1007/s11250-010-9602-6

**26** Barrow PA, Jones MA, Smith AL *et al*. The long view: *Salmonella*-the last forty years. *Avian Pathol* 2012; **41**: 413–20. https://doi.org/10.1080/03079457.2012.718071

**27** WHO. Drug Resistant *Salmonella*. 2014. http://www.who.int/ mediacentre/factsheets/fs139/en/.

**28** Su LH, Chu C, Cloeckaert A *et al.* An epidemic of plasmids? Dissemination of extended-spectrum cephalosporinases among *Salmonella* and other Enterobacteriaceae. *FEMS Immunol Med Microbiol* 2008; **52**: 155–68. https://doi.org/10.1111/j.1574-695X.2007.00360.x

**29** García-Fernández A, Fortini D, Veldman K *et al.* Characterization of plasmids harbouring qnrS1, qnrB2 and qnrB19 genes in *Salmonella. J Antimicrob Chemother* 2009; **63**: 274–81. https://doi.org/10.1093/jac/dkn470

**30** Strohl WA, Rouse H, Fisher BD. *Microbiologia Ilustrada*. Artmed, Porto Alegre, 2004.

**31** Angulo FJ, Johnson KR, Tauxe RV *et al*. Origins and consequences of antimicrobial-resistant nontyphoidal *Salmonella*: implications for the use of fluoroquinolones in food animals. *Microb Drug Resist* 2000; **6**: 77–83. https://doi.org/10.1089/mdr.2000.6.77

**32** Murugkar HV, Rahman H, Kumar A *et al.* Isolation, phage typing and antibiogram of *Salmonella* from man & animals in northeastern India. *Indian J Med Res* 2005; **122**: 237–42.

**33** Aragaw K, Molla B, Muckle A *et al.* The characterization of *Salmonella* serovars isolated from apparently healthy slaughtered pigs at Addis Ababa abattoir, Ethiopia. *Prev Vet Med* 2007; **82**: 252–61. https://doi.org/10.1016/j.prevetmed.2007.05.022

34 Zewdu E, Cornelius P. Antimicrobial resistance pattern of Salmonella serotypes isolated from food items and personnel in Addis Ababa,

Ethiopia. *Trop Anim Health Prod* 2009; **41**: 241–9. https://doi.org/10.1007/s11250-008-9181-y

**35** Bhan MK, Bahl R, Bhatnagar S. Typhoid and paratyphoid fever. *Lancet* 2005; **366**: 749–62. https://doi.org/10.1016/S0140-6736(05)67181-4

**36** WHO. Food Safety and Food Borne Illness. 2014. http://www.who.int/ mediacentre/factsheets/fs237/edu.

**37** Kang CI, Song JH. Antimicrobial resistance in Asia: current epidemiology and clinical implications. *Infect Chemother* 2013; **45**: 22–31. https://doi.org/10.3947/ic.2013.45.1.22

**38** Majowicz SE, Musto J, Scallan E *et al*. The global burden of nontyphoidal *Salmonella* gastroenteritis. *Clin Infect Dis* 2010; **50**: 882–9. https://doi.org/10.1086/650733

**39** Wei ZQ, Chang HL, Li YF *et al.* Clinical epidemiology and antimicrobial resistance of nontyphoidal *Salmonella* enteric infections in children: 2012–2014. *Chin J Pediatr* 2016; **54**: 489–95. https://doi.org/10.3760/cma.j.issn.0578-1310.2016.07.003

**40** Liang B, Xie Y, He S *et al.* Prevalence, serotypes, and drug resistance of nontyphoidal *Salmonella* among paediatric patients in a tertiary hospital in Guangzhou, China, 2014–2016. *J Infect Public Heal* 2019; **12**: 252–7. https://doi.org/10.1016/j.jiph.2018.10.012

**41** Dong-Ling L, Jie-hong W, Jing-Song WU *et al.* Characteristics of drug resistance and molecular typing for *Salmonella* in diarrhea patients from four hospitals in Shenzhen. *Chin J Zoonoses* 2017; **33**: 897–902.

**42** Hopkins KL, Kirchner M, Guerra B *et al.* Multi resistant *Salmonella enterica* serovar 4,[5],12:i- in Europe: a new pandemic strain? *Euro Surveill* 2010; **15**: 19580. https://doi.org/10.2807/ese.15.22.19580-en

**43** Liang DW, Lu JH, Jiang LX. Pattern of antimicrobial resistance and molecular typing for *Salmonella* isolated from diarrhea cases in Guangzhou. *Mod Prev Med* 2016; **43**: 696–9.

**44** Michael GB, Schwarz S. Antimicrobial resistance in zoonotic nontyphoidal *Salmonella*: an alarming trend? *Clin Microbiol Infect* 2016; **22**: 968–74. https://doi.org/10.1016/j.cmi.2016.07.033

**45** Lash TL, Fox MP, MacLehose RF *et al.* Good practices for quantitative bias analysis. *Int J Epidemiol* 2014; **43**: 1969–85. https://doi.org/10. 1093/ije/dyu149

**46** Nik Hazlina NH, Norhayati MN, Shaiful Bahari I *et al.* Worldwide prevalence, risk factors and psychological impact of infertility among women: a systematic review and meta-analysis. *BMJ Open* 2022; **12**: e057132. https://doi.org/10.1136/bmjopen-2021-057132

**47** Abd-Elghany SM, Sallam KI, Abd-Elkhalek A *et al.* Occurrence, genetic characterization and antimicrobial resistance of *Salmonella* isolated from chicken meat and giblets. *Epidemiol Infect* 2015; **143**: 997–1003. https://doi.org/10.1017/S0950268814001708

**48** Ahmed MM, Rahman MM, Mahbub KR *et al.* Characterization of antibiotic resistant *Salmonella* spp isolated from chicken eggs of Dhaka city. *J Sci Res* 2010; **3**: 191–6. https://doi.org/10.3329/jsr.v3i1.6109

**49** Tadesse G, Tessema TS. A meta-analysis of the prevalence of *Salmonella* in food animals in Ethiopia. *BMC Microbiol* 2014; **14**: 270. https://doi.org/10.1186/s12866-014-0270-y

**50** Akond MA, Shirin M, Alam S *et al.* Frequency of drug resistant *Salmonella* spp. isolated from poultry samples in Bangladesh. S *J Microbiol* 2013; **2**: 15–9. https://doi.org/10.3329/sjm.v2i1.15207

**51** Alam SB, Mahmud M, Akter R *et al.* Molecular detection of multidrug resistant *Salmonella* species isolated from broiler farm in Bangladesh. *Pathogens* 2020; **9**: 201. https://doi.org/10.3390/pathogens9030201

**52** Nahar A, Islam MA, Sobur MA *et al.* Detection of tetracycline resistant *Salmonella* and *Salmonella* spp. in sewage, river, pond and swimming pool in Mymensingh, Bangladesh. *Afr J Microbiol Res* 2019; **13**: 382–7. https://doi.org/10.5897/AJMR2019.9156

**53** Dahlén G, Blomqvist S, Almståhl A *et al.* Virulence factors and antibiotic susceptibility in enterococci isolated from oral mucosal and deep infections. *J Oral Microbiol* 2012; **4**: 10855. https://doi.org/10.3402/jom.v4i0.10855

**54** Iwabuchi E, Yamamoto S, Endo Y *et al*. Prevalence of *Salmonella* isolates and antimicrobial resistance patterns in chicken meat throughout Japan. *J Food Prot* 2011; **74**: 270–3. https://doi.org/10.4315/0362-028X. JFP-10-215

**55** Karim MR, Giasuddin M, Samad MA *et al*. Prevalence of *Salmonella* spp. in poultry and poultry products in Dhaka, Bangladesh. *Int J Anim Biol* 2017; **3**: 18–22.

**56** Mahmud MS, Kabir ML, Alam SS *et al.* Prevalence of *Salmonella* spp. in poultry eggs from different retail markets at Savar area, Bangladesh. *Am J Food Sci Health* 2015; **1**: 27–31.

**57** Marshall BM, Levy SB. Food animals and antimicrobials: impacts on human health. *Clin Microbiol Rev* 2011; **24**: 718–33. https://doi.org/10. 1128/CMR.00002-11

**58** Paul P, Akther S, Zulfekar Ali M *et al.* Isolation, identification and antibiogram study of *Salmonella* spp. from poultry farm environment. *Int J Animal Biol* 2017; **3**: 5–11.

**59** Talukder M, Islam MS, Ievy S *et al.* Detection of multidrug resistant *Salmonella* spp. from healthy and diseased broilers having potential public health significance. *J Adv Biotechnol Exp Ther* 2021; **4**: 248–55. https://doi.org/10.5455/jabet.2021.d125

**60** Haque AKMZ, Akter MR, Islam SKS et al. Salmonella gallinarum in small-scale commercial layer flocks: occurrence, molecular diversity and antibiogram. Vet Sci 2021; **8**: 71. https://doi.org/10.3390/vetsci8050071