

Review

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Antimicrobial Resistance in Poultry in Nigeria: Systematic Review of Reports Within 2005–2020

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Remiero

Antimicrobial Resistance in Poultry in Nigeria: Systematic Review of Reports Within 2005-2020

Short title: Antimicrobial Resistance in Poultry

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Abstract: Antimicrobial resistance (AMR) remains a persistent world-wide health concern. Poultry production in Nigeria has become a large-scale venture and is largely characterised by intensive management system. This intensive management system disposes to more antibiotic use and thus a greater risk of AMR. Previous reviews on AMR data in Nigeria have not considered the accuracy of methods and interpretation criteria. Therefore, in contribution to the "One Health" AMR Surveillance system, which is one of the five focus areas described by the Nigeria action plan on AMR control, this comprehensive review collates, curate, and analyse AMR published data in the Nigeria poultry sector and assessed the AST methods and interpretation criteria. A search of AMR reports from poultry in Nigeria prior to 2020 after screening for relevance yielded 69 studies from 2005-2020 with 91 AMR reports (some studies reported more than one organism). From these studies we reviewed the organisms reported, antibiotics tested, resistance to tested antibiotics, distribution of reports within the country's geopolitical zones, multidrug resistance pattern, the inclusion or absence of control strains in antimicrobial testing, the antimicrobial testing method and interpretative criteria used. There was no AMR study in poultry earlier than 2005 in Nigeria, highest number of study was in 2019 (n=12). South-Western region has the highest number (n = 27) of study and the North-Western region has the lowest (n = 5). Organisms mostly reported were Salmonella, 31%, Escherichia coli, 26% and Staphylococcus aureus, 15%). Other organisms (≤5%) were Campylobacter, Klebsiella, Bacillus, Listeria, Enterobacter, Proteus, Norcardia, Streptococcus, Alkaligenes, Shigella, Micrococcus, Enterococcus, Ochrobacturm, and Yersinia. The antimicrobial susceptibility testing (AST) method mostly used was disc diffusion (80%), also majority (95.7%) of the studies used the CLSI standards for interpretation and 79.7% of the studies did not mention the use of control strains for AST. The 69 studies reviewed tested 69 antimicrobials belonging to 17 classes of Antibiotics; the most tested classes of antimicrobial were β -lactams (33%), quinolones (21%) and aminoglycosides (15.4%). All of the organisms showed varying multi-drug resistance; greater proportion (50-100%) has MAR Index greater than 0.2. The review of the various studies showed very high multi-drug resistance among the various organisms of poultry origin and also a gap in AST carried out in Nigeria. We therefore recommend that AST research methods must be of global standard and additional efforts should be made towards education of the general public on best practices of antibiotics usage and the dangers of AMR.

Keywords: antimicrobial resistance; multiple antibiotic resistance; poultry; interpretation criteria; multiple antibiotic resistance index

Introduction

Importantly, AMR is extensively recognised as a global health problem, yet in many parts of the world its enormity is not well expounded [1]. Rising drug resistance is caused mainly by abuse or misuse of antimicrobials in humans and livestock. The spread of resistant strains in humans and livestock is increasing globally because of greater access to antibiotic drugs in low- and middle-income countries [2]. The necessity to improve human health, animal health and agricultural productivity has led to the extensive use of antimicrobials without respecting therapy guidelines in many places particularly, in Africa.

Current status of AMR

In developing countries like Nigeria, the misuse and abuse of antimicrobials poses serious threats to public health. Also, farmer's ignorance of the hazards related to AMR in food-producing animals and the urgent need to safe their dying animals has escalated the emergence of AMR (AMR) [3–9]. The worldwide public health concerns with antibiotics used in food-producing animals are that they are closely related to those used in human medicine and select for resistance. Cross-species transmission of resistant bacteria or resistance genetic elements from animals or environment to humans has been reported [10,11]. Worldwide, AMR is increasing; if present trends carry on unrestricted, there may possibly be as many as 10 million annual AMR-associated deaths from a widespread array of infections by 2050 [12]. The World Health Organization (WHO) has incorporated AMR as one of the topmost ten threats to global health in 2019 [13]. Thus, the WHO and other international agencies (e.g. FAO, OIE), along with many individual countries, have developed comprehensive action plans to address the AMR crisis and one of the five main pillars of the Global action plan on AMR 2016-2020 is to strengthen the knowledge and evidence base through surveillance and research [14]. The report from the monitoring of the global action plan by FAO has suggested and recommended the need for situation analysis and production of action plans for individual countries [15].

Factors Contributing to AMR in Nigeria

Nigeria is faced with the challenge of poverty and poor environmental conditions such as lack of potable water and sanitation, and poor infection control in farms and hospitals. These factors aid the spread of resistant strains of microbes in the environment. Also, the understanding and awareness of AMR in Nigeria is poor [16]. Antimicrobials, are mainly acquired over-the-counter without proper prescription and disease diagnosis thereby increasing the indiscriminate use of antimicrobials. The repeated use of antimicrobials in small doses in food animals for growth promotion, feed proficiency enhancement and prophylaxis, has been identified as a significant contributing factor to increasing antimicrobial resistance [17].

Implication of AMR for Human and Animal Health

Antimicrobial resistance leads to 'treatment failure', the main consequence of which is Morbidity and mortality in patients. Resistant microbes will increase the chances of developing serious health conditions, longer hospital stays, higher medical costs and even death [18]. The impact of AMR on animal health has been given far less attention than that for human health. Similar to humans Impacts of AMR on livestock is significant. AMR in animals will lead to infections that would not have occurred, increased frequency of treatment failures and increased severity of resulting in increased mortality and morbidity. The increase in resistance to antimicrobials will make treatments on animals ineffective and cause the infections to become more severe.

Antimicrobial Susceptibility Testing

Antimicrobial susceptibility testing (AST) methods are available to determine bacterial susceptibility to antimicrobials. The selection of a method is centred on many factors including practicality, flexibility, automation, cost, reproducibility, accuracy, and individual preference. It is

important that AST methods provide reproducible results in day-to-day laboratory use and that the data be as good as with those results obtained by an acknowledged 'gold standard' reference method. In the absence of standardised methods or reference procedures, susceptibility results from different laboratories cannot be reliably compared [19]. The following three methods have been shown to consistently provide reproducible and repeatable results when followed correctly; disc diffusion, broth dilution and agar dilution [20]. The AST results should be recorded quantitatively; as distribution of MICs in milligrams per litre or µg/ml or as inhibition zone diameters in millimetres. Generally, antimicrobial susceptibility breakpoints are established by national standards organisations, professional societies or regulatory agencies. Only the CLSI has developed protocols for susceptibility testing of bacteria of animal origin and determination of interpretive criteria [20]. However, protocols and guidelines are available from a number of standards organisations and professional societies for susceptibility testing for similar bacterial species that cause infections in humans. It is possible that such guidelines can be adopted for susceptibility testing for bacteria of animal origin. Additionally, efforts focusing on both standardisation and harmonisation of susceptibility/resistance breakpoints on an international scale are progressing. These efforts have primarily focused on the adoption of the standards and guidelines of CLSI and EUCAST, which provide laboratories with methods and quality control values enabling comparisons of AST methods and generated data [20]. The results of AST are reported with interpretative categories; as "susceptible" which indicates that the bacteria was inhibited by the achievable concentrations of antimicrobial agent when the dosage recommended to treat the site of infection is used, "intermediate" which defines the bacteria for which the response rates to usually attainable blood and tissue levels of antimicrobial agent are lower compared to susceptible isolates and "resistant" which defines that the bacteria are not inhibited by the usually achievable concentrations of the agent with normal dose regimens. The AST procedure is sensitive to changes in the environment and test conditions, therefore it is crucial that each variable in the procedure should be standardised and carefully controlled [21]. The continuous monitoring of the performance is best achieved by testing of quality control strains. Quality control strains are well-characterised organisms with defined susceptibility or resistance mechanisms to the antimicrobial agent(s) tested, testing quality control strains help concurrently monitor the performance of the test and ensures that the test is being performed properly [22].

Nigeria Poultry Industry

The Nigeria Poultry industry contributes approximately 25% to agricultural GDP, consumption of chicken meat in Nigeria is growing so fast that capacity is not meeting demand [23]. An estimated 454 billion tonnes of meat and 3.8 million eggs per year is produced in the poultry sector in Nigeria, with a standing population of 180 million birds. Approximately 80 million chicken are reared in extensive systems, 60 million in semi-intensive systems and the remaining 40 million in intensive systems [24]. Poultry production in Nigeria has become increasingly intense consequently playing a key role in food security. There is no religious or cultural restrictions to its consumption therefore its wide consumption is associated with some food safety risks including AMR. Nigeria constitutes a large market for pharmaceuticals; accounting for 60% of volume of health products consumed in ECOWAS sub-region, but has limited regulatory capacity. Furthermore, manufacturers and importers have vertical drug distribution channels for wholesalers, retailers and hospitals with minimal government supervision. Thus, medicines are often sold in unregistered and unlicensed premises and in many cases by non-pharmacists. The Nigeria action plan on AMR control developed in 2017 has five focus areas, which one of it is, building a "One Health" AMR surveillance system [25]. An earlier report of the trend, status and situation of AMR arising from food animals and the environment in Nigeria, Oloso, et al. [26], reviewed 48 AMR reports from different food animals and the environment within 2000-2016. This present study is focused on AMR in poultry and were viewed 91 AMR reports from 69 AMR studies from poultry only within 2005-2020 and we also considered other data not covered in the previous study such as the inclusion or absence of control strains in antimicrobial testing, the antimicrobial testing method and interpretative criteria used.

This study was carried out to collate, curate, and analyse data on AMR in Nigeria related to poultry and the immediate human link as contributors to the AMR burden in Nigeria and used the Preferred Reporting Items for Systematic Review and Meta-Analysis (PRISMA) for retrieval of articles and screening processes. This study serves as a reference source towards the development of good antimicrobial stewardship program by stakeholders through the "One Health Platform" for poultry in Nigeria.

Methods

Search Strategy and Selection Criteria

To obtain relevant articles the Scopus database was searched for English text articles from all years for records relating to AMR in Nigerian poultry using the search terms defined in Figure 1. Additionally, the PubMed database was searched using related terms as defined in Figure 1. The records from the two searches were combined and duplicates removed (Figure 1). The title and abstracts (when available) were then screened for relevance and articles excluded if any of the following criteria were met: not related to AMR/AST or non-bacterial; out of geographic area (*i.e.* not Nigeria); did not concern poultry; publication year earlier than 2005; and reviews. Records that remained were included in the review.

Data Extraction and Analysis

To harmonize the diverse data for analysing the situation of AMR from poultry reports in Nigeria the scale developed by Oloso *et al.* [26] was employed. In this scheme the percentage (proportion) of resistant microbe populations (species) per study was assigned a scale from 1 (no resistance) to 5 (very high resistance), as shown in Table 1.



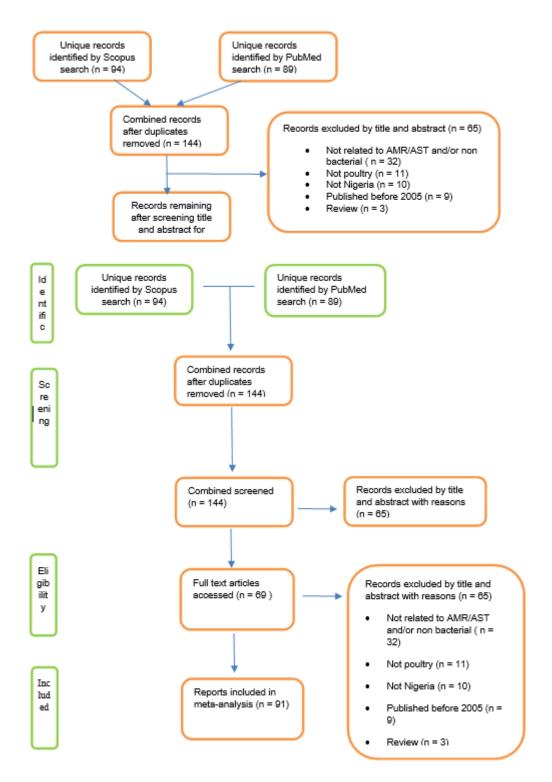


Figure 1. PRISMA diagram of the article selection procedure for records.

Table 1. Scale for the categorisation of resistance proportion. This table shows the scale developed by Oloso *et al.*, 2018, adopted to measure the level of resistance from different diverse measurements from the several studies. Percentage referred to the proportion of resistant microbe populations (species) per study.

Group Scale	Categorization	Antimicrobial Resistance Studies
1	≤1%	Sensitive or No resistance
2	>1 ≤24%	Moderately sensitive or very low resistance
3	>24 ≤50%	Weakly sensitive or low resistance
4	>50 ≤74%	Low sensitive or High resistance
5	>74%	Very low (no) sensitive or Very high resistance

Some studies reported AMR results from more than one bacterial species and in these cases each species was analysed separately. A total of 69 studies were reviewed (Table 2). Given the diversity of antimicrobials employed in the different studies and to further facilitate analysis and comparison, antimicrobials were grouped by class according to OIE, [27] (Table 3). The sulphonamides, trimethoprim and the combination drug trimethoprim-sulfamethoxazole were analysed as a single group named folic acid synthesis inhibitors. A large diversity of drug-bug combinations was observed in the overall data set, and to ensure integrity of interpretation only drug-bug combinations with a clinical breakpoint defined by CLSI or EUCAST were included for analysis. For example, the disc diffusion method for polymyxin susceptibility testing is not recommended by CLSI or EUCAST, and there are currently no established zone diameter breakpoints for colistin or polymyxin B [28]. Therefore, colistin susceptibility data derived from this method were not included in subsequent analysis. The number of reports describing each drug-bug combination and each antimicrobial classbug combination was calculated and these data were summarised by bacterial species. Additional information extracted for analysis from each study included: year of publication, Nigerian State or geopolitical zone, bacterial species tested, antibiotics tested, antimicrobial susceptibility testing (AST) method, AST interpretation standard, and use of quality control strains.

Table 2. Publications included in review by year.

Publication Year	Number of studies Reviewed	Number of AMR Report
2005	2	2
2006	2	2
2007	3	3
2008	2	2
2009	1	1
2010	7	13
2011	7	10
2012	7	9
2013	4	4
2014	3	4
2015	4	4
2016	5	6
2017	5	5
2018	3	7
2019	12	17
2020	2	2
Total	69	91

		Antibiotics in Peer-	u u						
		Reviewed Studies	Generation	Very High	High	Low	Very low	No	Total
Aminocyclitol	s	Spectinomycin	NGC				1		1
Aminoglycosi	des	Amikacin	NGC				1	1	2
		Apramycin	NGC				1	2	3
		Gentamicin	NGC	10	9	20	14	6	59
		Kanamycin	NGC	2	1	1		1	5
		Neomycin	NGC	2	2	1	2		7
		Streptomycin	NGC	10	8	7	8	8	41
		Tobramycin	NGC		1		1		2
Ansamycins		Rifampicin	NGC		1	1		1	3
β-lactams	Cephalosporins	Ceftazolin	1			1		1	2
		Cephalexin	1		4		1	1	6
		Cephalothin	1	1			2	1	4
		Cefoxitin	2	2			1	2	5
		Cefuroxime	2	5	4		3	1	13
		Cefixime	3			1		1	2
		Cefotaxime	3	3	1	2	3		9
		Ceftazidime	3	6	4	2	5		17
		Ceftiofur	3				3		3
		Ceftriaxone	3	4	6	3	3		16
		Cephaximide	3			5			5
		Cefpodoxime	3		1		1		2
		Cefepime	4				1		1
	Carbapenems	Imipenem	NGC				1	2	3
		Ertapenem	NGC				2	1	3
		Floxapenem	NGC			5			5

		Meropenem	NGC		1	1	1		3
	Penicillins	Penicillin	1	8					8
		Penicillin G	1	2	2	1	2		7
		Cloxacillin	2	6	1				7
		Oxacillin	2	2					2
		Amoxicillin	3	13	6	3	2	1	25
		Ampicillin	3	23	5	6	3	3	40
		Amoxicillin/clavulani	4	18	7	5	9	1	40
		c acid							
		Ampicillin/cloxacillin	4	7					7
		Carbenicillin	4	7			1		8
		Piperacillin/tazobacta	4				1		1
		m							
		Ticarcillin	4	1					1
		Ticarcillin/clavulanic	4				1		1
		acid							
		Methicillin	NGC	1	1				2
	Monobactam	Aztreonam	NGC			1	1		2
Glycopeptides	3	Vancomycin	NGC	1	3	6			10
Lincosamides		Clindamycin	NGC	1	1			1	3
		Lincomycin	NGC	1				1	2
Macrolides		Azithromycin	NGC	1	1		1		3
		Clarithromycin	NGC	1					1
		Erythromycin	NGC	10	4	10	3		27
		Tylosin	NGC	2	1				3
Nitrofuran		Nitrofuratoin	NGC	11	6	2	2	2	23
Phenicols		Chloramphenicol	NGC	9	15	4	3	5	36
		Florfenicol	NGC			1			1

Sulphonamide	Sulphamethoxazole	NGC	2	1	4	1	1	9
Diaminopyrimidine inhibitor	Trimethoprim	NGC	3	4	3	2		12
Sulphonamide+Diaminopyrim	trimethoprim-	NGC	18	8	16	3	2	47
idine inhibitor	sulfamethoxazole							
Tetracycline	Doxycycline	NGC		2		1		3
	Oxytetracycline	NGC	1	1				2
	Tetracycline	NGC	25	7	4	2	3	41
Quinolones	Nalidixic acid	1	13	10	4	3	2	32
	Norfloxacin	1	1	1	9	3	1	15
	Enrofloxacin	2	2	2	1	1	2	8
	Ciprofloxacin	2	8	14	14	15	5	56
	Ofloxacin	2	5	7	6	7	6	31
	Pefloxacin	3	4	4		5	4	17
	Levofloxacin	3	1		1	1	1	4
Others	Fosfomycin	NGC					1	1
Combinatio	Gendox	NGC				1		1
n drugs	(Gentamicin/Doxycyc							
	line)							
Total		тота	252	157	158	122	71	760
		L						

NGC: No generation Classification. 1, 2, 3 and 4; first, second, third and fourth generation antibiotics respectively.

Determination of Multiple Antibiotics Resistance (MAR) index followed the procedure described by Ayandele $et\ al.$ [29] in which the number of antibiotics an isolate is resistant to (a) is divided by the total number of the antibiotics used in the study (b), so that the MAR Index = a/b. Resistance to antibiotics was defined by the resistance proportion that fall under scale 4 or 5 (high or very high) and it was calculated for each bacterial species from all 69 reviews containing 91 reports.

Results

Overview of Records Returned

Search of the Scopus database yielded 94 unique records and the PubMed search yielded 89 unique records. In total 144 records were identified following combination of the two searches and removal of duplicates (Figure 1). Of these, 69 met the inclusion criteria to give 91 AMR records (nine studies reported AMR results for more than one organism (range 2-6) and were included in the final

analysis. All 69 records were from surveillance studies and none was a clinical report. These spanned the years 2005 to 2020 with an average of four records per year; the greatest number of records was from the year 2019 (n=12), whereas only a single report was from 2009 (Table 2). All geopolitical zones in Nigeria were represented, and some records covered more than one zone; the highest number of reports was from South West Nigeria (27) and, in descending order, South East (18), South South (10) North Central (8), North West (6), and North East (5) (Supplementary File 3).

The most commonly used antimicrobial susceptibility testing method was disc diffusion (n=60; 87%), broth dilution was used in eight studies (12%) and one study used agar dilution (1%). The majority (n=66, 96%) of the studies used the CLSI standards for interpretation and EUCAST was used in three studies (4%). Most of the studies (n=55; 80%) did not describe the use of quality control strains for antimicrobial susceptibility testing. Eight of the 69 studies reviewed did not provide AMR results as percentage resistance or sensitivity.

Cumulatively the 91 reports tested 69 different antimicrobials or combination antimicrobials belonging to 16 different classes, yielding a total of 760 AMR categorisations using to the scale of Oloso *et al.* [26] (Table 3). The most commonly used antimicrobial groups for susceptibility testing included: quinolones (163 resistance categorisations; 21%), penicillins (153; 20%), aminoglycosides (119; 15%), and cephalosporins (85; 11%). Within these groups the most commonly tested antimicrobials included: ciprofloxacin and nalidixic acid (quinolones); ampicillin and amoxicillin/clavulanic acid (penicillins); gentamicin and streptomycin (aminoglycosides); and ceftazidime and ceftriaxone (cephalosporins) (Table 3). Other antimicrobials groups used in these studies included trimethoprim-sulfamethoxazole, tetracyclines, and phenicols; the full list is given in Table 3. Eight studies reported susceptibilities for polymyxins but in all cases these were determined by disc diffusion and consequently these data were not included for further analysis.

Table 3. List of antibiotics and the number of reports of each antimicrobial resistance.

		Antibiotics in Peer- Reviewed Studies	Generation	Very High	High	Low	Very low	No	Total
Aminocyclito	ls	Spectinomycin	NGC				1		1
Aminoglycos	ides	Amikacin	NGC				1	1	2
		Apramycin	NGC				1	2	3
		Gentamicin	NGC	10	9	20	14	6	59
		Kanamycin	NGC	2	1	1		1	5
		Neomycin	NGC	2	2	1	2		7
		Streptomycin	NGC	10	8	7	8	8	41
		Tobramycin	NGC		1		1		2
Ansamycins		Rifampicin	NGC		1	1		1	3
β-lactams	Cephalospori	Ceftazolin	1			1		1	2
	ns	Cephalexin	1		4		1	1	6
		Cephalothin	1	1			2	1	4

Cefoxitin 2	I	I	1	1					
Cefixime		Cefoxitin	2	2			1	2	5
Cefotaxime		Cefuroxime	2	5	4		3	1	13
Ceftazidime		Cefixime	3			1		1	2
Ceftiofur 3		Cefotaxime	3	3	1	2	3		9
Ceftriaxone		Ceftazidime	3	6	4	2	5		17
Cephaximide 3		Ceftiofur	3				3		3
Cefpodoxime 3 1 1 2 Cefepime 4 1 1 1 Carbapenems Imipenem NGC 1 2 3 Ertapenem NGC 5 2 1 3 Floxapenem NGC 1 1 1 3 Meropenem NGC 1 1 1 3 Penicillins 1 2 2 1 2 7 Cloxacillin 2 6 1 7 7 Oxacillin 2 2 2 2 2 Amoxicillin 3 13 6 3 2 1 25 Ampicillin/clavula 4 18 7 5 9 1 40 n 1 4 7 1 8 Piperacillin/tazobact 4 7 1 1 8		Ceftriaxone	3	4	6	3	3		16
Cefepime		Cephaximide	3			5			5
Carbapenems Imipenem NGC 1 2 3		Cefpodoxime	3		1		1		2
Ertapenem NGC 2 1 3 Floxapenem NGC 5 5 5 Meropenem NGC 1 1 1 1 1 3 Penicillins Penicillin 1 8 8 8 Penicillin G 1 2 2 1 2 7 Cloxacillin 2 6 1 7 Oxacillin 2 6 1 7 Oxacillin 3 13 6 3 2 1 25 Ampicillin 3 23 5 6 3 3 40 Amoxicillin/clavula 4 18 7 5 9 1 40 nic acid Ampicillin/cloxacilli 4 7 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Cefepime	4				1		1
Floxapenem NGC 5 5 5	Carbapenems	Imipenem	NGC				1	2	3
Meropenem NGC 1 1 1 3 3		Ertapenem	NGC				2	1	3
Penicillins Penicillin G 1 8 8 Penicillin G 1 2 2 1 2 7 Cloxacillin 2 6 1 7 7 Oxacillin 2 2 2 2 Amoxicillin 3 13 6 3 2 1 25 Ampicillin 3 23 5 6 3 3 40 Amoxicillin/clavula 4 18 7 5 9 1 40 nic acid Ampicillin/cloxacilli 4 7 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 7 1 1 1		Floxapenem	NGC			5			5
Penicillin G 1 2 2 1 2 7 Cloxacillin 2 6 1 7 Oxacillin 2 2 2 2 2 2 Amoxicillin 3 13 6 3 2 1 25 Ampicillin 3 23 5 6 3 3 40 Amoxicillin/clavula 4 18 7 5 9 1 40 nic acid 7 n 7 Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Meropenem	NGC		1	1	1		3
Cloxacillin 2 6 1 7 Oxacillin 2 2 2 2 Amoxicillin 3 13 6 3 2 1 25 Ampicillin 3 23 5 6 3 3 40 Amoxicillin/clavula 4 18 7 5 9 1 40 nic acid Ampicillin/cloxacilli 4 7 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1	Penicillins	Penicillin	1	8					8
Oxacillin 2 2 2 Amoxicillin 3 13 6 3 2 1 25 Ampicillin 3 23 5 6 3 3 40 Amoxicillin/clavula 4 18 7 5 9 1 40 nic acid Ampicillin/cloxacilli 4 7 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Penicillin G	1	2	2	1	2		7
Amoxicillin 3 13 6 3 2 1 25 Ampicillin 3 23 5 6 3 3 40 Amoxicillin/clavula 4 18 7 5 9 1 40 nic acid 4 7 7 7 7 7 7 7 1 8 Piperacillin/tazobact 4 7 1 1 1 1 1 1 1		Cloxacillin	2	6	1				7
Ampicillin 3 23 5 6 3 3 40 Amoxicillin/clavula nic acid 4 18 7 5 9 1 40 Ampicillin/cloxacilli 4 7 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Oxacillin	2	2					2
Amoxicillin/clavula 4 18 7 5 9 1 40 nic acid Ampicillin/cloxacilli 4 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Amoxicillin	3	13	6	3	2	1	25
nic acid 7 7 Ampicillin/cloxacilli 4 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Ampicillin	3	23	5	6	3	3	40
Ampicillin/cloxacilli 4 7 7 n Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Amoxicillin/clavula	4	18	7	5	9	1	40
n		nic acid							
Carbenicillin 4 7 1 8 Piperacillin/tazobact 4 1 1 1		Ampicillin/cloxacilli	4	7					7
Piperacillin/tazobact 4 1 1 1		n							
		Carbenicillin	4	7			1		8
am		Piperacillin/tazobact	4				1		1
		am							
Ticarcillin 4 1 1		Ticarcillin	4	1					1

		Ticarcillin/clavulani	4				1		1
		c acid							
		Methicillin	NGC	1	1				2
	Monobactam	Aztreonam	NGC			1	1		2
Glycopeptide	s	Vancomycin	NGC	1	3	6			10
Lincosamides		Clindamycin	NGC	1	1			1	3
		Lincomycin	NGC	1				1	2
Macrolides		Azithromycin	NGC	1	1		1		3
		Clarithromycin	NGC	1					1
		Erythromycin	NGC	10	4	10	3		27
		Tylosin	NGC	2	1				3
Nitrofuran		Nitrofuratoin	NGC	11	6	2	2	2	23
Phenicols		Chloramphenicol	NGC	9	15	4	3	5	36
		Florfenicol	NGC			1			1
Sulphonamid	e	Sulphamethoxazole	NGC	2	1	4	1	1	9
Diaminopyrii	nidine	Trimethoprim	NGC	3	4	3	2		12
inhibitor									
Sulphonamid	e+Diaminopyri	trimethoprim-	NGC	18	8	16	3	2	47
midine inhibi	tor	sulfamethoxazole							
Tetracycline		Doxycycline	NGC		2		1		3
		Oxytetracycline	NGC	1	1				2
		Tetracycline	NGC	25	7	4	2	3	41
Quinolones		Nalidixic acid	1	13	10	4	3	2	32
		Norfloxacin	1	1	1	9	3	1	15
		Enrofloxacin	2	2	2	1	1	2	8
		Ciprofloxacin	2	8	14	14	15	5	56
		Ofloxacin	2	5	7	6	7	6	31
		Pefloxacin	3	4	4		5	4	17

	Levofloxacin	3	1		1	1	1	4
Others	Fosfomycin	NGC					1	1
Combinatio	Gendox	NGC				1		1
n drugs	(Gentamicin/Doxyc							
	ycline)							
Total		тот	252	157	158	122	71	760
		AL						

NGC: No generation Classification. 1, 2, 3 and 4; first, second, third and fourth generation antibiotics respectively.

A total of seventeen bacterial species or genera were tested for antimicrobial susceptibility across the 91 reports. The most tested were *Salmonella* spp. (n=28 reports; 31%), *Escherichia coli* (n=24; 26%), *Staphylococcus aureus* (n=14; 15%) and *Campylobacter* spp. (n=5; 5%). *Klebsiella* and *Bacillus* each had three reports, and there were two reports each for *Listeria*, *Enterobacter*, *Staphylococcus* spp. and *Proteus*. There was one report each for *Alkaligenes*, *Enterococcus*, *Micrococcus*, *Norcardia*, *Ochrobacturm*, *Shigella*, *Streptococcus*, and *Yersinia*. The scale developed by Oloso *et al.*, [26] was employed to assess the burden of AMR in the four most reported bacterial species or genera: *Salmonella* spp., *Escherichia coli*, *Staphylococcus aureus*, and *Campylobacter* spp.

Salmonella

There were 28 reports of AMR in *Salmonella* spp. from Nigerian poultry. All geopolitical zones were represented, and studies spanned the years 2005-2019. Data for 29 antibiotics belonging to 10 antimicrobial groups were available, to give a total of 237 resistance categorisations according to the Oloso *et al.* [26] scale (Figure 3a). In aggregate, resistance categorisations of "very high" or "high" were the most abundant (n=117; 49%), and "no" or "very low" categorisations were less common (n=80; 34%). Many studies employed the same antimicrobials; at least ten used ampicillin, amoxicillin/clavulanic acid, chloramphenicol, ciprofloxacin, folic acid inhibitors (sulphonamides, trimethoprim, or trimethoprim-sulfamethoxazole), gentamicin, nalidixic acid, nitrofurantoin, ofloxacin, streptomycin, and tetracycline. The 3rd generation cephalosporins cefotaxime, ceftazidime, and ceftriaxone were employed in five or more studies. However, due to the diversity of antibiotics examined in the 28 reports, the total number of categorisations differed by antimicrobial class (Figure 2a).

The most tested class was the quinolones (n=60), with 26 "very high" or "high" resistance categorisations and 26 categorised as "no" or "very low" (Figure 2a). High proportions of "very high" or "high" resistances were noted for nitrofurantoin (8/10 reports), tetracyclines (16/21 reports), folic acid inhibitors (15/26 reports), penicillins (20/37 reports), cephalosporins (11/24 reports), and chloramphenicol (5/11 reports) (Figure 2a). Three reports investigated susceptibility towards carbapenems and each had "very low" and "no" resistance categorisation. There was a single report for fosfomycin which noted "no" resistance.

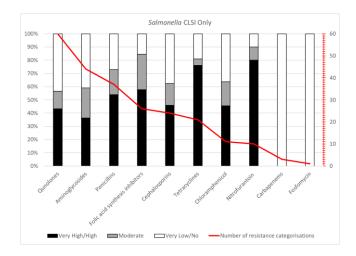


Figure 2. a. Summary of antimicrobial resistances for *Salmonella* **spp..** The data are presented by antimicrobial group, with resistance categorizations expressed as proportions (left side scale) and ordered by the number of categorisations, as indicated by the red line (right-side scale).

Escherichia coli

E. coli was the second most represented bacterium in the papers examined, with 24 AMR reports spanning the years 2007-2020. Data was from all geopolitical zones, with the greatest number from the South West and South East. Data for 34 antibiotics belonging to 10 antimicrobial groups were analysed, to give a total of 134 resistance categorisations according to the Oloso et al. [26] scale (Figure 2b). Overall, resistance categorisations of "very high" or "high" were in the majority (n=76; 57%) whereas "no" or "very low" categorisations were less common (n=42; 31%). As was observed with Salmonella, the number of resistance categorisations differed by antimicrobial class (Figure 2b), again reflecting the diversity of antibiotics employed in the papers included for analysis. Certain antimicrobials were employed in several studies and the following were used in at least ten: ampicillin, ciprofloxacin, gentamicin, trimethoprim-sulfamethoxazole, and tetracycline. High proportions of "very high" or "high" resistances were noted for nitrofurantoin (4/4 reports), tetracycline (9/10 reports), chloramphenicol (6/7 reports), folic acid inhibitors (10/14 reports), penicillins (11/16 reports), and cephalosporins (13/25 reports). Quinolones were the most tested and 47% (15/32 reports) had a "very high" or "high" categorisation. Four studies examined nitrofurantoin resistance, and each reported a "very high" or "high" categorisation. Conversely the four studies (with seven reports) that examined carbapenem resistance all had "very low" (1.2 - 8.0%) or "no" resistance while a study reported 'no' resistance to fasfomycin. The percentage MAR greater than 0.5 for both Salmonella and *E.coli* are 38.9% and 44.8% respectively.

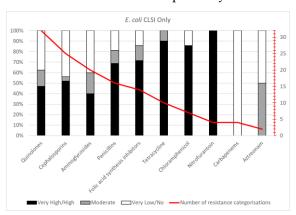


Figure 2. b. Summary of antimicrobial resistances for *E. coli*. The data are presented by antimicrobial group, with resistance categorizations expressed as proportions (left side scale) and the number of categorisations for each group indicated by the red line (right-side scale).

Staphylococcus aureus

Fourteen studies had tested susceptibilities for *Staphylococcus aureus*, from the years 2010-2018, with a least one report from each geopolitical zone of the country. Data for 20 antibiotics belonging to 12 antimicrobial groups were analysed, to yield 79 resistance categorisations (Figure 2 c). The most tested individual antibiotics (≥9 studies) were gentamicin and erythromycin; 5-8 studies employed trimethoprim-sulfamethoxazole, tetracycline, ciprofloxacin, chloramphenicol, and vancomycin; other antibiotics were used in less than five studies. In aggregate, resistance categorisations of "very high" or "high" comprised 52% (n=41), whereas "no" or "very low" categorisations were 25% (n=20). The most tested class was quinolones (n=14), with 36% categorised as "very high" or "high". Macrolides and folic acid synthesis inhibitors were the next most tested (12 reports each) with 75% and 42% categorised as "very high" or "high", respectively. All four reports for penicillins had a "very high" resistance categorisation. The studies on *Staphylococcus aureus isolates reported many to be MDR and* 70-100% resistance to Macrolides, penicillin G1/G2, clindamycin and nitrofuratoin.

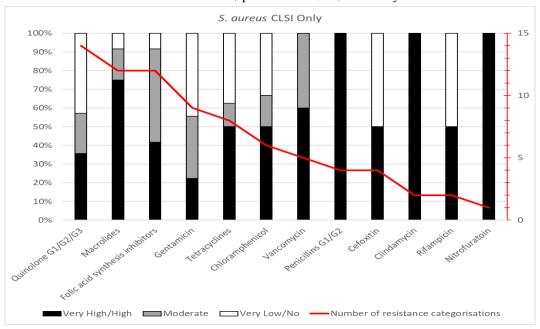


Figure 2. c. Summary of antimicrobial resistances for *Staphylococcus aureus*. The data are presented by antimicrobial group, with resistance categorizations expressed as proportions (left side scale) and the number of categorisations for each group indicated by the red line (right-side scale) .

Campylobacter spp.

Five studies provided susceptibility data for *Campylobacter* spp., limited to the South West, South East and North Central geo-political zones of Nigeria. Although a wide diversity of antimicrobial was employed across these studies (n=20) clinical breakpoints were available for only erythromycin, tetracycline, and ciprofloxacin.

Data for 20 antibiotics belonging to 12 antimicrobial groups were available for analysis. All five studies reported susceptibility data for gentamicin and streptomycin and four studies had used erythromycin, chloramphenicol, trimethoprim-sulfamethoxazole, nalidixic acid and tetracycline. The remaining antimicrobials were used in one or two studies only. Due to the low numbers of studies and susceptibility data available, interpretation of proportions for resistance categorizations was not undertaken.

More studies reported very high/high (50% or above) resistance to aminoglycosides, cephalosporins(G1), penicillins (G2), phenicols, quinolones (G1) and sulphonamide derivatives. However, there was very low or no (50% or above) resistance to penicillins (G3 and G4), lincosamides, quinolone (G3 and G4) and nitrofurans.

16

Other Organisms

All other organisms that made minute contributions (with three or less reports) were pooled together as Gram positive and Gram negative organisms. This organisms include Gram negatives; Klesiella, Proteus, Yersinia, Shigella, Ochrobactrum and Gram positives; Listeria, Streptococcus, Bacillus, Enteroccus, Alkaligenes, Enterobacter, Nocardia and Micrococcus.

These Gram positive organisms tested 32 antibiotics belonging to 16 antimicrobial groups; the penicillins, quinolones and aminoglycosides, were the most tested in descending order. Of the 16 groups, phenicols, had 100% proportion of "very high to high" resistance, followed bytetracycline, 87.5%, penicillins (3rd and 4th generation) and (1st and 2nd), 81.8% and 81.3% each, then nitofurans, 66.7% and quinolones (2nd and 3rd generation) 53.3%. The proportion of "very low to no resistance" was very small or absent for most of the antibiotics groups.

Resistance Pattern of the Organisms Studied

The frequency of report of the various organisms in descending order were; Salmonella, E. coli, S. aureus, Campylobacter, Klebsiella, Bacillus, Listeria, Enterobacter, Proteus, Norcardia, Streptococcus, Alkaligenes, Shigella, Micrococcus, Enterococcus, Ochrobacturm, and Yersinia (Figure 3; Table 6). All AMR reports were from surveillance studies, none was clinical report. All of the organisms showed varying multi-drug resistance. Very high resistance was reported in various studies for Quinolones (n=34), Tetracycline (n=26) and Aminoglycosides (n=24) (Table 4). Greater proportion of the isolates for each organism has MAR greater than 0.2; E. coli (83.3%), Salmonella (71.5%), Staphylococcus (85.7%), Klebsiella and Bacillus, (66.7%), Listeria (50%), while Campylobacter and all others (100%) (Tables 5 & 6, Figure 3). Overall, isolates were multi-drug resistant and more resistance was reported for South west, North central and North west regions in Nigeria (Table 7 and Figure 4).

Table 4. Number of reports for each resistance level category within the classes of antimicrobial in the AMR studies.

Antibiotics class		Number of Reports and Category of Resistance Level							
		Very	High	Low	Very	No	Total (%)		
		High			low				
					T		-		
Aminocyclitols					1		1 (0.1)		
Aminoglycos	ides	24	21	29	27	18	119 (15.7)		
Ansamycin			1	1		1	3 (0.4)		
β-lactam	Cephalosporins	21	20	14	23	7	85 (11.2)		
	Carbapenems	0	1	6	4	3	14 (1.85)		
	Penicillins	87	21	15	20	5	148 (19.5)		
	Monobactam			1	1		2 (0.26)		
Diaminopyrimidine inhibitors		3	4	3	2		12 (1.58)		
Fosfomycin						1	1 (0.1)		

Total	252	157	158	122	71	760
Quinolones	34	38	35	35	21	163 (21.52)
Tetracyclines	26	10	4	3	3	46 (6.1)
inhibitors						
Sulphonamide+Diaminopyrimidine	18	8	16	3	2	47 (6.2)
Sulphonamides	2	1	4	1	1	9 (1.2)
Phenicols	9	15	5	3	5	37 (4.9)
Nitrofuran	11	6	2	2	2	23 (3.0)
Macrolides	14	6	10	4		34 (4.49)
Lincosamides	2	1			2	5 (0.66)
Glycopeptides	1	3	6			10 (1.32)
Gendox (Gentamicin/Doxycycline)				1		1(0.1)

^{*}Note that total report (n=110)for [△]-lactams were 43, 36, 50, and 15 respectively for Very high, high, low, very low and no resistance.

Table 5. Multiple Antibiotics Resistance (MAR) Index for the AMR Reviews Salmonella (n=28).

		E. col	i (n=24)	Staph	ylococcus	Klebsiel	la (n=3)	Listeria (n	=2)
				(n=14)					
S/N	MAR	S/N	MAR	S/N	MAR	S/N	MAR	S/N	MAR
1.	0.46	1.	0.4	1.	1	1	1	1	0.8
2.	0.33	2.	0.7	2.	0.27	2	0.3	2	
3.	0.1	3.	0.5	3.	0.29	3	NA		
4.	0	4.	0.17	4.	0.86				
5.	0.42	5.	0.92	5.	0.64	Proteus ((n=2)	Streptococcus (n=1)	
6.	0.45	6.	0.5	6.	0.82	S/N	MAR	S/N	MAR
7.	0.75	7.	0.4	7.	0	1	0.35	1	0.35
8.	0.4	8.	0.7	8.	0.44	2	0.16		
10.	0.1	10.	0.57	10.	0.33				
11.	0.7	11.	0.86	11.	1				
12.	0.75	12.	0.78	12.	0.38	Yersinia (n=1)		Bacillus (n=3)	

13.	0.71	13.	0.7	13.	0.2	S/N	MAR	S/N	MAR
14.	0.2	14.	0.12	14.	0.8	1	0.4	1	0.35
15.	0.17	15.	0.09					2	0.7
16.	0.15	16.	1					3	NA
17.		17.		Campy	lobacter				
	0.71		0.8	(n=5)					
18.	0.5	18.	1	S/N	MAR	Shigella((n=1)	Enterococo	cus (n = 1)
19.	0.08	19.	0.8	1.	0.27	S/N		S/N	MDR
20.	0.69	20.	NA	2.	0.9	1		1	0.83
21.	0.57	21.	NA	3.	0.51				
22.		22.	NA	4.		Ochroba	ctrum (n		
	0.73				0.62	= 1)		Alkaligen	es (n=1)
23.	0.6	23.	NA	5.	0.89	S/N	MAR	S/N	MAR
24.	0.56	24.	NA			1	0.71	1	0.83
25.	0.7								
26.	0	Lister	ria (n=2)	Norcai	dia (n=1)	Micrococcus (n=1)		Enterobac	ter (n=2)
27.	0.1	S/N	MAR	S/N	MAR	S/N	MAR	S/N	MAR
28	0.73	1	0.8	1	0.35	1	0.43	1	0.3
		2	NA					2	0.5

Key: NA refers to those 8 studies that did not provide AMR results as percentage resistance or sensitivity.

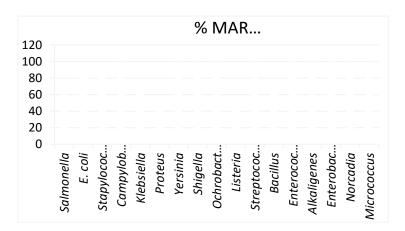


Figure 3. Reported Organisms and the percentage of their MAR Index that is greater than 0.2.

Table 6. Reported Organisms and the percentage of their Multiple Antibiotics Resistance (MAR) Index that is greater than 0.2.

Organisms	Percentage (%) MAR index > 0.2
Salmonella	71.5
E. coli	83.3
Stapylococcus	85.7
Campylobacter	100
Klebsiella	66.7
Proteus	50
Yersinia	100
Shigella	100
Ochrobactrum	100
Listeria	50
Streptococcus	100
Bacillus	66.7
Enterococcus	100
Alkaligenes	100
Enterobacter	100
Norcadia	100
Micrococcus	100

Table 7. Resistant microbes against different antibiotics (2005-2020).

S/	Resistant Microbe	Drug Bug	Ye	AMR Studies/Reference
N	Reported		ar	
1	Bacillus	Amoxicillin, Cloxacillin, Colistin	20	Adelowo et al., 2008
			08	
2	Salmonella	Cephalexin	20	Enabulele et al., 2008
			08	
3	Campylobacter	Cephalothin, Cephalexin, Ampicillin, Cotrimoxazole,	20	Akwobu et al., 2010
		Streptomycin, Amoxicillin/Clavulanic Acid	10	
		Nalidixic Acid, Pefloxacin, Ofloxacin, Ciprofloxacin,		
İ		Gentamicin		

4	Salmonella	Cephalexin	20	Enabulele et al., 2010
			10	
5	Salmonella	Neomycin	20 10	Fashae et al., 2010
6	ESBL E. coli	Penicillin G, Amoxicillin, Cephalexin, Erythromycin,	20	Mamza, 2010
0	ESBL E. COII		10	ivianiza, 2010
_		Tylosin		
7	Staphylococcus	Tylosin	20	Mamza, 2010
			10	
8	Salmonella	Cephalothin, Colistin	20	Muhammed et al., 2010
			10	
9	E. coli	Polymyxin B, Penicillin, Ampliclox, Carbenicillin,	20	Okonko et al., 2010
		Erythromycin	10	
10	Staphylococcus	Polymyxin B, Ampliclox, Streptomycin, Carbenicillin	20	Okonko et al., 2010
			10	
11	Salmonella	Amoxicillin	20	Adeleke E.O. and
			11	Omafuvbe B.O, 2011
12	Salmonella	Amoxicillin	20	Ekundayo and Ezeoke 2011
12	Samionena	Amoachin	11	Ekulidayo alid Ezeoke 2011
10	T 11			F 111 / 1 0044
13	E. coli	Amoxicillin	20	Ezekiel et al., 2011
			11	
14	Salmonella	Amoxicillin	20	Ezekiel et al., 2011
			11	
15	Staphylococcus	Amoxicillin-Clavulanic Acid	20	Otalu <i>et al.,</i> 2011
			11	
16	Salmonella	Amoxicillin	20	Eja et al., 2012
			12	
17	E. coli	Neomycin, Enrofloxacin	20	Ojo et al., 2012
			12	
18	Salmonella	Neomycin, Enrofloxacin	20	Ojo et al., 2012
10	Samorena	recontychy Entonoxuch	12	0,0 0, 41., 2012
19	Salmonella	Oxytetracycline, Enrofloxacin	20	C1: -t -1 2012
19	Samonena	Oxytetracycline, Enrolloxacin		Sasaki et al., 2012
			12	
20	Salmonella	Ceftiofur	20	Raufu et al., 2013
			13	
21	Staphylococcus	Augmentin	20	Suleiman et al., 2013
			13	
	6.1. 11	Oxacillin	20	Agada et al., 2014
22	Salmonella			İ
22	Saimonella		14	
22	Salmonella	Amoxicillin, Ceftiofur	14 20	Raufu <i>et al.,</i> 2014
		Amoxicillin, Ceftiofur		Raufu <i>et al.,</i> 2014
		Amoxicillin, Ceftiofur Cephalothin	20	Raufu <i>et al.,</i> 2014 Eyitayo <i>et al.,</i> 2015

25	Salmonella	Carbenicillin	20 15	Nchawa et al., 2015
26	Staphylococcus	Ampliclox, Amoxicillin	20	Owuna et al., 2015
27	Campylobacter	Azithromycin, Clindamycin, Chloramphenicol Streptomycin, Trimethoprim/Sulpha	20 15	Sati et al., 2015
28	Staphylococcus	Nalidixic Acid, Gentamicin Cloxacillin, Augmentin	20 15	Ugwu <i>et al.,</i> 2015
29	Salmonella	Penicillin G, Amoxicillin,	20	Ifeanyichukwu et al., 2016
30	MRSA	Cloxacillin, Amoxicillin	20 16	Igbinosa et al., 2016
31	ESBL E. coli	Cephalothin	20 16	Ojo et al., 2016
32	Campylobacter	Chloramphenicol Streptomycin, Enrofloxacin, Nalidixic Acid, Gentamicin	20 16	Olatoye and Ogunsemoyin, 2016
33	MRSA	Augmentin	20	Nworie et al., 2017
34	Staphylococcus	Penicillin G, Streptomycin, Imipenem	20 17	Ranjdar et al., 2017
35	Salmonella	Amoxicillin	20 17	Uzeh et al., 2017
36	E. coli	Penicillin G	20 18	Ayadiran et al., 2018
37	Staphylococcus	Penicillin G	20 18	Ayadiran et al., 2018
38	MRSA	Cloxacillin, Ceftazidime, Ceftriaxone, Amoxicillin/Clavunate	20 18	Kwoji et al., 2018
39	Staphylococcus	Streptomycin	20 18	Kwoji et al., 2018
40	ESBL E. coli	Ampliclox	20 18	Okpara et al., 2018
41	Salmonella	Neomycin	20	Ahmed et al., 2019
42	E. coli	Erythromycin, Amoxicillin	20	Bodunde et al., 2019
43	Salmonella	Erythromycin, Amoxicillin	20	Bodunde et al., 2019

44	Staphylococcus	Amoxicillin, Streptomycin, Augmentin	20	Bodunde et al., 2019
			19	
45	Campylobacter	Contrimazole, Streptomycin, Chloramphenicol,	20	Ogbor <i>et al.,</i> 2019
		Cloxacillin	19	82 ** ** ***, = ***
		Nitrofurantoin, Ofloxacin, Gentamycin and Nalidixic		
		Acid		
46	Salmonella	Enrofloxacin, Neomycin, Erythromycin, Colistin,	20	Oloso et al., 2019
		Tylosin	19	
47	Campylobacter	Augmentin, Amoxicillin, Nitrofurantoin, Gentamicin,	20	Oreva et al., 2019
		Ofloxacin	19	
		Clarithromycin, Cotrimoxazole, Cloxacillin,		
		Chloramphenicol, Nalidixic Acid		
48	Salmonella	Erythromycin, Cloxacillin	20	Ugwu et al., 2019
			19	
49	Salmonella	Neomycin	20	Uzeh <i>et al.,</i> 2019
			19	
50	Salmonella	Neomycin	20	Ahmed <i>et al.</i> , 2020
			20	

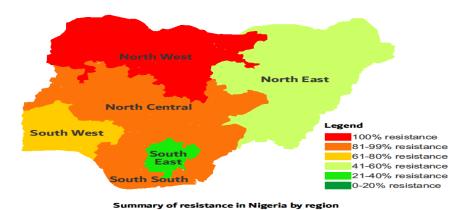


Figure 4. Summary of resistance in Nigeria by region.

Discussion

According to the OIE, [19], the disc diffusion, broth dilution and agar dilution methods have been shown to consistently provide reproducible and repeatable results, when followed correctly. These three methods were the only once used for AMR surveillance in poultry in Nigeria, but the disc diffusion method was mostly used (80.2%). A number of standards and guidelines are currently available for AST for bacteria of human origin and can be adopted for testing bacteria of animal origin but each country must evaluate its own AST guidelines [19]. The disc diffusion and CLSI standard was mostly used (94.5%) for AST in Nigeria (Supplementary figures 1 & 2), which aligns with the guideline of the Nigeria National Action Plan (NAP) for AMR, 2017-2022 that recommends the CLSI standard. It is noteworthy that the AST done for colistin in all papers reviewed is not the appropriate method, thus the colistin resistance may have been reported differently, therefore this data was not suitable for analysis at this time. In conducting microbial analysis, positive control bacteria strains are routinely used to assure the satisfactory performance of various elements in the test procedures [30]. It was observed that only 27.5% of the studies included control strains, failure to include control

strains in AST is not a standard practice. The importance of using approved standard methods for AST and use of proper interpretation criteria from CLSI cannot be over emphasised. The consideration of Epidemiological Cut-Off (ECOFF) is useful in the selection of the Minimal Effective concentration for the antimicrobial in question for effective treatment. Inadequate AST methods will result in inconsistencies and errors in the susceptibility results obtained. These errors may result in the initiation of inadequate antimicrobial therapy which has fatal outcomes in severely ill patients where these antibiotics are common first-line substances. Conversely, errors may deny the patients of treatment with an effective antibiotic and lead to the use of second-choice drugs, usually more recent and expensive, and thus contribute to economic losses and the selection of resistant strains.

It is imperative that studies on antibiogram in Nigeria should consider current susceptibility data being generated by national AMR surveillance projects and AMR surveillance at Veterinary Teaching Hospital sentinel centers nationwide. Furthermore penicillins and tetraxyclines have very resistant profiles across all species studied making them inappropriate for disease treatment in the poultry sector in Nigeria. This study revealed several patterns of multidrug resistance with greater proportion having MAR index above 0.2 as reported in the different studies reviewed and points out the high levels of resistance of the common pathogens in poultry to readily available antimicrobials commonly used and studied in these sectors within Nigeria and this agrees with the findings of Oloso *et al.* [26]. A survey of antibiotics commonly used in Poultry in Nigeria reported trimethoprim-sulfamethoxazole (71.7%), neomycin (66.7%), oxytetracycline (48.3%), chloramphenicol (45%), tetracycline (33.3%) and penicillin (30%) as the most used antibiotics [31].

The broad purview of the reviewed studies indicates that despite the increase surveillance on AMR in the country (across all geopolitical zones), there seem to be low compliance with the standard practice for antimicrobial usage amongst poultry farmers. The overall results in this study, therefore, reflects great AMR risk to humans through the food chain, exposure of farm workers to resistant strains of bacteria and untreatable poultry diseases. Antibiotics used in food-producing animals and in particular the poultry sector are closely related to those used in humans and transmission through the food chain results in the spread of resistance. There would also be increase in microbial risk and associated economic loss to the poultry industry, as poultry organisms are resistant to commonly used antibiotics. The resistance pattern of similar organisms in poultry as reviewed by this study is mostly related. Salmonella had the highest reports followed by E. coli, Staphylococcus and Campylobacter, these organisms were reported across all geopolitical zones of the country which may indicate that these organisms are implicated in diseases in the poultry sector of the nation or are easy to study in every parts of the country. The AMR reports showed that some antimicrobial groups had similar resistance proportion to almost all the organisms tested. Highest proportion of "very high to high" resistance across most of the organisms was noticed among the; penicillins (1st, 2nd, 3rd and 4th generations), tetracyclines, nitrofurans, cephalosporins (particularly the 1st and 2nd generations), sulphonamide derivatives, quinolones (mostly 1st generation) and phenicols in no particular order. While the highest proportion of "very low" to "no" resistance across most of the organisms were observed among; the carbapenems, quinolones (2nd and 3rd generations) and aminoglycosides. This widespread resistance to all generations of penicillins can be explained by the findings of the report that stated that penicillins and its combinations were the most consumed over-the-counter (selfmedicated) drugs by humans and in animals in Nigeria [32,33].

Observations of AMR within the classes of antibiotics reflected higher percentages of resistance in the antibiotics belonging to the macrolides, tetracyclines and β -lactams (mostly penicillins) raising further concerns of the threat posed by AMR. These drugs have been listed by WHO as medically important antimicrobials in food-producing animals as they can result to resistance in human and thus their overall reduction in use in food-producing animals has been recommended [34]. Additionally, the presence low levels of resistance to carbapenems is of public health significance since these drugs are the last-line drugs in therapy of serious infections. The situation of AMR in Nigeria as reflected in this study and other reports from Nigeria is a reason to stimulate and encourage the Nigeria government and other appropriate bodies to embrace and actively practice the "One Health" approach to tackle the nations AMR situation through the establishment and

In addition, studies that tested Polymixin and colistin used the disc diffusion method for analysis, whereas there is no CLSI or EUCAST recommendation for the disc diffusion (DD) method usage for polymyxin and colistin susceptibility testing. Hence, we do not really have a reliable picture of polymyxin and colistin resistance in poultry derived bacteria in Nigeria in this review (Table 8). Also worthy to note, is the low AMR record for *Campylobacter*, an organism of public health importance but hard to grow.

In conclusion, this review revealed very high multi-drug resistance among bacterial of poultry origin, important zoonotic pathogens like *Salmonella*, *E. coli* and *S. aureus* among others showed high MARI >0.2 and also gaps were identified in AST carried out in Nigeria. Studies based on genotypes not addressed in this study were limited. We therefore recommend improvement in research methods to be of global standard and additional efforts to be made towards creating awareness of the general public to the dangers of AMR and also education of farmers on the benefits of animal production with minimal antibiotics usage; the use of preventive measures such as vaccinations, probiotics, hygienic production and biosecurity should be encouraged. As minimal or no usage of antibiotics will result in no antibiotics resistance. Future research should target existing regions where resistance is emerging to understand current trends in the epidemiology of AMR in poultry Nigeria.

Supplementary Materials: The following supporting information can be downloaded at the website of this paper posted on Preprints.org.

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