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Research Article

Prevalence and Potential Risk Factors for the Acquisition of Antibiotic-Resistant *Staphylococcus* spp. Bacteria Among Pastoralist Farmers in Kajiado Central Subcounty, Kenya

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Antimicrobial resistance (AMR) is a growing health problem globally. To address this challenge, there is a need to generate baseline data on the prevalence and AMR profile of the main disease-causing bacteria. Here, we interrogated the prevalence of bacteria in the nasal cavity of healthy pastoralists in Kajiado Central Subcounty, Kenya, and the occurrence of AMR in Staphylococcus isolates among the study subjects. Nasal swabs from 176 pastoralists were cultured, and the bacteria isolates identified using standard phenotypic and biochemical bacteriological methods. Among the obtained 195 isolates, the most prevalent isolates were coagulase-negative Staphylococcus (CoNS) (44.9%), followed by Enterococci spp. (43.2%) while Staphylococcus aureus prevalence was 8%. Antimicrobial sensitivity of the Staphylococcus spp. isolates to 14 antibiotics representing six antibiotic groups was undertaken using the Kirby-Bauer disk diffusion method. Among the CoNS, the highest resistance was reported in amoxicillin (78.7%) and ceftazidime (76%), while the most resistance for S. aureus was reported in ceftazidime (100%), amoxicillin (71.4%), and streptomycin (71.4%). From an administered questionnaire looking at gender, animal contact frequency, history of hospital visitation and antibiotic usage, and habitual intake of raw milk, the study showed that male participants had a higher risk of carrying multiple drug resistant (MDR) bacteria than females (p = 0.02, OR = 1.3). Likewise, habitual intake of raw milk was significantly associated MDR acquisition (p = 0.02, OR = 1.82). This study reveals a high prevalence of AMR Staphylococcus isolates in the study area laying a foundation for further analysis of molecular characterization of the observed resistance as well as the development of interventions that can reduce the occurrence of AMR in the study area.

1. Introduction

The upper respiratory tract harbors commensal and potentially pathogenic bacteria, i.e., *Staphylococcus* spp., *Moraxella* spp., *Corynebacterium* spp., *Streptococcus* spp., and *Haemophilus* spp. [1]. The commensal bacteria function as competitors to pathogenic bacteria and continuously prime the immune system against the pathogenic bacteria [2]. The *Staphylococcus* spp. colonizing the nasal cavity are

divided into coagulase-positive (*S. aureus*) and coagulase-negative (CoNS) and account for a variety of infections and diseases in both humans and livestock [3]. In African countries, Staphylococcal carriage ranges from 8.0 to 57% in both patients and healthy populations [4–6].

The first-line treatment for Staphylococcal infection is penicillinase-stable penicillin [7, 8]. However, resistance to this class of antibiotics was first identified in1960s in *Staphylococcus aureus* [9]. AMR in Staphylococcus is highest in

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low-income countries and low middle-income countries [10-12]. Currently, methicillin-resistant Staphylococcus (MRS) pathogens are among the leading causes of death with a prevalence of 1-80% worldwide [13, 14]. MRS is majorly caused by the acquisition of the staphylococcal cassette chromosome mec (SCCmec) comprised mainly of mecA gene or its homologues (mecB or mecC) which encodes for a novel penicillin-binding protein (PBP2a) with a lower affinity to the beta-lactam ring and the site-specific recombinase gene (ccrA/B/C). After its first recovery in the 1980s, the diversity of the SCCmec was noted first in the 1990s, and afterward, more diversity was detected between 2001 and 2008 [15]. To date, fourteen types of SCCmec (I-XIV) have been reported [16]. The first community-associated (CA-MRS) was detected in the 1980s [17]. There is a steady increase of CA-MRS in the 1990s and early 2000s [7, 15, 18-20]. CA-MRSA has majorly been associated with SCCmec types IV, V, and VI [16, 21-23].

The integrons have been reported to foster the spread of AMR in Staphylococcus spp. given their ability to naturally clone and express resistant genes. Integrons contain up to 40 genes associated with resistance to beta-lactams, sulfonamides, aminoglycosides, chloramphenicol, and macrolides [24, 25]. The occurrence of integrons in HA-MRS isolates was first reported in 2007 in South China [26]. Since then, integrons have been identified in antibiotic-resistant staphylococcal isolates largely in the Asian continent [27-31]. However, data on integron associated Staphylococcus AMR in other parts of the world especially in the developed countries and Africa are lacking. Coexistence of class 1/2 integron gene cassettes and SCCmec in MRS has been reported occasionally [28, 32-34]. The simultaneous existence of both mechanisms is anticipated to increase the adaptability of microbes to antibiotic selective pressure speeding up the spread and survival fitness of resistant Staphylococcus while fostering MDR evolution [32–35].

MRS infections are treated using a combination of penicillin and vancomycin [36, 37]. However, suboptimal responses to vancomycin treatment, vancomycin intermediate susceptible Staphylococcus (VIS) minimum inhibitory concentration (MIC 4-8 µg/ml), were first detected in Japan in 1997 whereas total resistance to vancomycin, vancomycin-resistant Staphylococcus (VRSA) MIC $\geq 16 \,\mu \text{g/}$ ml, was detected in the USA in 2002 [38, 39], whereas the VIS is thought to be because, by cell wall thickening, VRS is caused by the acquisition of the Van operon from Enterococcus faecalis bacteria [40, 41]. Since their emergence, VIS and VRS isolates have been detected in different parts of the world albeit at low percentages [42-45]. Penicillin-resistant Staphylococcus on the other hand is caused by a blaz gene carried in the R plasmids which encodes for the penicillinase enzyme that hydrolyses the β -lactam ring [46, 47]. On the other hand, accumulation of mutations on the drug target genes has been identified as a main cause of resistance for protein and DNA replication inhibitors [48, 49].

The emergence of antimicrobial resistance is mainly caused by the indiscriminate use of antibiotics for treatment/prophylaxis in both human and animal ecologies [50]. Globalization and lifestyle factors such as the intake

of raw/undercooked animal products, low hygiene level, and high population density as well foster the spread of antimicrobial resistance [5, 51]. In Africa, AMR in Staphylococcus spp. ranges from 12 to 80% [52-54]. In Kenya, Staphylococcus spp. AMR ranges between 20 and 84.1% [54-57] with a steady increase from <50% in the 2000s to >50% by 2020 in Staphylococcus antimicrobial resistance [55, 57-63], whereas most Staphylococcus spp. AMR studies in Kenya have been conducted in the urban hospital setting (Nairobi, Kiambu, Kilifi, Kisumu, and Kericho); majority of the studies [60, 64, 65] investigating antibiotic resistance in rural areas, especially among pastoralist communities such as the Maasai, who use large quantities of antibiotics for veterinary care with >95% being self-administered and up to 75% carried out without professional consultation, are lacking or poorly documented [51]. The current study is aimed at determining the prevalence, risk factors, and detection of mecA or mecC antibiotic-resistant marker in Staphylococcus isolates obtained from the anterior nares of livestock farmers in Kajiado Central Subcounty in Kenya.

2. Material and Methods

- 2.1. Study Area. The study was undertaken in the Kajiado Central Subcounty region in Kajiado County, Kenya, located in the southern part of Kenya, approximately 80 km from Nairobi City and borders Northern Tanzania: 1.83874 latitudes and 36.79135 longitudes. The subcounty is divided into five wards: Matapato South, Matapato North, Purko, Ilidamati, and Dalalekutuk, which were included in this study. Kajiado Central experiences bimodal rainfall ranging from 500 to 1200 mm, with short rains occurring from October to December and long seasons between March and May, and an average temperature of 25°C. The total human population of the subcounty is 161,862, most of which are of the Maasai community and are pastoralists. The total livestock population in the subcounty as per the 2019 census was 802,898 [66].
- 2.2. Study Design. A descriptive cross-sectional study design was used. The study sample size was determined using the Thrusfield method [67], which showed that a minimum of 150 participants were sufficient for the current study. A total of 176 participants were sampled. The sampling unit of interest was household members, which were selected randomly from a sampling frame provided by the county extension and public health officers. Based on the human population in specific wards, the study participants were from Matapato South (n = 54), Matapato North (n = 51), Purko (n = 25), Ilidamati (n = 30), and Dalalekutuk (n = 16) wards (Table 1).
- 2.3. Nasal Swab Collection and Administration of the Questionnaire. Nasal swabs were collected, as described previously [68]. Briefly, the swabs were collected using a sterile nasal swab (BD BBL™ CultureSwab™) dipped in normal saline by slightly swabbing both anterior nares of the participants and transferred to the transport media (Scharlab S.L, Spain). The sample was labelled and transported back to the microbiology laboratory based at the Institute of Primate

TABLE 1: The number of study participants as distributed per ward
in Kajiado Central Subcounty, Kenya.

Wards	Total respondents sampled	Number of homesteads sampled
Matapato North	54	12
Matapato South	51	8
Purko	25	5
Ilidamati	30	4
Dalalekutuk	16	5
Total	176	36

Research, Kenya, in a cool box. The data on the risk factors, age, gender, history of antibiotic use, hospital visitation < 3 months prior to the study, intake of raw milk, and animal contact, were obtained through a structured questionnaire administered in local Maasai language to the study participants by the investigator.

2.4. Culture and Identification of Bacteria. To identify the bacteria in the collected samples, the nasal swabs were first inoculated on MacConkey agar (Scharlab S.L, Spain) and sheep blood agar (Scharlab S.L, Spain) and incubated at 37°C for 18-24 hrs. The inoculated sheep blood agar plates were incubated in an environment enriched with 5% CO₂. Colonies were then identified based on morphological appearance on the respective agar plates, Gram stain reaction, and biochemical tests [69]. The catalase test was used to distinguish staphylococci from streptococci. Catalase-positive staphylococci were further subjected to a coagulase test to distinguish S. aureus from coagulase-negative Staphylococcus. Pure isolates of S. aureus and coagulase-negative Staphylococcus were then subjected to antimicrobial susceptibility testing [69].

2.5. Antibiotic Susceptibility Testing. To determine the antibiotic susceptibility profiles of the isolated S. aureus and coagulase-negative Staphylococcus, a standardized Kirby-Bauer disk diffusion method utilizing the Mueller-Hinton agar (Scharlab S.L, Spain) plate technique was performed as per the Clinical and Laboratory Standards Institute guidelines [70]. Briefly, a uniform suspension of each isolate was prepared and adjusted to a concentration equivalent to 0.5 McFarland with sterile normal saline. Each suspension was evenly spread on two Mueller-Hinton agar (Scharlab S.L, Spain) plates followed by placing seven antibiotics disks on every plate. A total of fourteen antibiotic disks representing six antibiotic groups, beta-lactams (amoxicillin $30 \,\mu g$, oxacillin $1 \,\mu g$, Augmentin $20/10 \,\mu g$, ceftriaxone 30 µg, ceftazidime 30 µg, cefoxitin 30 µg, and cefotaxime 30 µg), tetracyclines (tetracycline 30 µg), glycopeptides (vancomycin 30 µg), aminoglycopeptides (gentamicin $10 \,\mu g$ and streptomycin $10 \,\mu g$), macrolides (erythromycin 15 μ g and clindamycin 2 μ g), and fluoroquinolones (ciprofloxacin $30 \mu g$), were placed on the plates as per the CLSI guidelines. Measurements of the inhibitory

zones were done, and the results were interpreted according to the CLSI guidelines. The results were reported as resistant, intermediate, or susceptible to specific antibiotics [70].

2.6. Detection of the mecA and mecC Genes Using Singleplex PCR. DNA from the resistant bacteria was extracted using the heat lysis method. Briefly, one bacterial colony was suspended in 500 µl of nuclease-free water, heated at 95°C for 10 minutes followed by freezing at -80°C for 5 minutes. The DNA was harvested by centrifugation at 1200 rpm for 10 min, the supernatant was picked, and pellets were discarded. The extracted DNA was subjected to PCR for detection of mecA and mecC genes using primers mecA 513 bp C05/C06 (F-AAA ATC GAT GGT AAA GGT TGG C) and D09/D10 (R-AGT TCT GCA GTA CCG GAT TTG C) and mecC 718 bp (GAAAAAAAGGCTTAGAACGCC TC CCTGAATCTGCTAATAATATTTC). A PCR protocol of 25 μ l final volume containing 2x Taq polymerase master mix (NEBNext, New England Biolabs, Inc.) was adopted to run 94°C for 5 min followed by 35 cycles of 94°C for 30 seconds, 58°C for 40 seconds, 72°C for 40 seconds, and a final extension at 72°C for 5 minutes. The products of PCR amplification were analyzed using 2% agarose gel electrophoresis 1x tris acetate ethylene (TAE) diamine tetra acetic acid buffer stained with SYBR Sate TM fluorescent dye. Positive Staphylococcus reference strains S. aureus ATCC 25923 and S. epidermidis ATCC 12228 were used as controls. A template control was added as a negative control in both assays.

2.7. Ethical Approval. The ethical clearance was granted by the JKUAT Ethics Review Committee (JKU/2/4/896B) and Kajiado County Medical and Public Health Service Department. The livestock farmers were informed about the study, and written consent was subsequently obtained from those more than 18 years while assent was sort from the parent/guardian for the minors.

2.8. Data Analysis. The obtained data was entered and analyzed using Microsoft Excel. The proportion of the identified bacteria species was determined by dividing the total number of the specific bacteria by the sample size. Analysis of the odds ratio (OR) and p value was done to evaluate the association between risk factors (age, history of hospital visitation, contact with cattle, history of antibiotic usage, intake of raw milk, and gender factors) and presence of MDR Staphylococcus bacteria for a potential link to acquisition. A p value of < 0.05 was considered statistically significant.

3. Results

3.1. Characteristics of Study Participants. A total of 176 study participants were sampled from the study area. Majority of the respondents were males (61.4%) with 82.4% of them reporting frequent contact with livestock. Of the study participants, only 26.7% recorded having visited a hospital within 3 months prior to this study, while 22.7% of the participants consumed raw milk habitually with the majority (40%) being <20 years. Other investigated predisposing factors are outlined in Table 2.

Table 2: Demographic characteristics of the study participants (n = 176) from Kajiado Central Subcounty.

Characteristics			Frequency (%) bas	sed on origin of w	ards	
Characteristics	M. South	M. North	Purko	Ilidamati	Dalalekutuk	Total
Gender						
Male	31 (57.4)	31 (60.8)	18 (72.0)	13 (43.3)	15 (93.8)	108 (61.4)
Female	23 (42.6)	20 (39.2)	7 (28.0)	17 (56.7)	1 (6.3)	68 (38.6)
Age						
<20 years	26 (48.1)	11 (21.6)	9 (36.0)	10 (33.3)	1 (6.3)	57 (32.4)
21-30 years	8 (14.8)	14 (27.5)	8 (32.0)	8 (26.7)	9 (56.3)	47 (26.7)
31-40 years	7 (13.0)	5 (9.8)	4 (16.0)	4 (13.3)	4 (25.0)	24 (13.6)
41-50 years	4 (7.4)	4 (7.8)	1 (4.0)	5 (16.7)	2 (12.5)	16 (9.1)
>51 years	2 (3.7)	11 (21.6)	3 (12.0)	2 (6.7)	0 (0.0)	18 (10.2)
Could not remember	7 (13.0)	6 (11.8)	0 (0.0)	1 (3.3)	0 (0.0)	14 (8.0)
Contact with livestock						
Frequent	46 (85.2)	45 (88.2)	25 (100)	18 (60.0)	11 (68.8)	145 (82.4)
Infrequent	8 (14.8)	6 (11.8)	0 (0.0)	12 (40.0)	5 (31.3)	31 (17.6)
History of hospital visitation	n					
Yes	19 (35.2)	15 (29.4)	6 (24.0)	3 (10.0)	4 (25.0)	47 (36.4)
No	35 (64.8)	36 (70.6)	19 (76.0)	27 (90.0)	12 (75.0)	129 (73.3)
Milk consumption						
Boiled	17 (31.5)	20 (39.2)	3 (12.0)	30 (100)	0 (0.0)	70 (39.8)
Raw	17 (31.5)	26 (51.0)	22 (88.0)	0 (0.0)	16 (100)	81 (46.0)
Not sure	20 (37.0)	5 (9.8)	0 (0.0)	0 (0.0)	0 (0.0)	25 (14.2)
Antibiotic use						
Yes	9 (5.6)	3 (5.8)	1 (4.0)	3 (10.0)	0 (100)	16 (9.1)
No	45 (83.3)	48 (94.1)	24 (96.0)	27 (90.0)	16 (100)	160 (90.9)

M: Matapato.

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3.2. Prevalence of Nasal Bacteria. A total of 195 bacterial isolates were obtained from the collected 176 nasal swabs. All (100%) participants' nares were colonized by various bacteria. The most common isolates were coagulase-negative Staphylococcus (CoNS) (44.9%) and Enterococci spp. (43.2%) (Table 3). The Staphylococcus aureus prevalence was 8%. Other isolates obtained from the study are outlined in Table 3. Respondents from Matapato South and Matapato North had the highest (50%) prevalence of CoNS, while Enterococcus spp. were more common in respondents from Matapato South ward.

3.3. Antibiotic Resistance. The obtained Staphylococcus spp. were further investigated for antimicrobial susceptibility. Among the CoNS, the highest resistance was amoxicillin (78.7%), followed by ceftazidime (76%) and oxacillin (49.3%). Majority of the CoNS isolates, however, were sensitive to Augmentin (88.0%) and ciprofloxacin (84%). Resistance to other tested antibiotics is shown in Table 4.

For *S. aureus*, all (100%) isolates were resistant to ceftazidime. The percentage of *S. aureus* resistant to amoxicillin (71.4%), streptomycin (71.4%), and vancomycin (64.3%) was also high. The *S. aureus* isolates resistant to oxacillin (28.6%) and cefotaxime (21.4%) resistance were lower compared to the CoNS isolates. In contrast, *S. aureus* isolates were highly (100%) susceptible to Augmentin. The summary

Table 3: Prevalence of bacteria isolated from nasal cavities of 176 nasal swabs of participants in Kajiado Central Subcounty, Kenya.

Туре	Total isolates (N)	Prevalence (%)
CoNS	79	44.9
Enterococci spp.	76	43.2
Bacillus spp.	14	8
Staphylococcus aureus	14	8
Corynebacterium spp.	5	2.8
Morganella spp.	2	1.1
Micrococcus spp.	1	0.6
Lactococcus spp.	1	0.6
Proteus mirabilis	1	0.6
Pantoea spp.	1	0.6
Aeromonas hydrophila	1	0.6

CoNS = coagulase-negative Staphylococcus.

of the sensitivities of *S. aureus* to individual antibiotics is shown in Table 4.

From the study findings, majority of the CoNS isolates were resistant to amoxicillin in combination with oxacillin 61/79 (77.2%) and ceftazidime 56/79 (70.9%) while *S. aureus* major resistance combination was observed in the amoxicillin and ceftazidime 10/14 (71.4%), streptomycin (42.9%),

Table 4: Antibiotic sensitivity profile of Staphylococcus isolates from nasal swabs of participants in Kajiado Central Subcounty, Kenya.

Antibiotic	Isolate		ivity levels for Stap ant $(n, \%)$		ates liate (n, %)	Susceptible (n, %)	
	CoNS	37	49.3	0	0	38	50.7
Oxacillin	S. aureus	4	28.6	1	7.14	9	64.3
A 4:	CoNS	9	12.0	0	0	66	88.0
Augmentin	S. aureus	0	0.0	0	0.0	14	100.0
Ceftazidime	CoNS	57	76.0	14	18.7	4	5.3
Certazidilile	S. aureus	14	100.0	0	0.0	0	0.0
Amoxicillin	CoNS	59	78.7	5	6.7	11	14.7
Tilloxiciiiii	S. aureus	10	71.4	2	14.3	2	14.3
Emythromycin	CoNS	28	37.3	14	18.7	33	44.7
Erythromycin	S. aureus	3	21.4	3	21.4	8	57.1
Tatua avalina	CoNS	26	34.7	16	21.3	33	44.7
Tetracycline	S. aureus	2	14.3	2	14.3	10	71.4
Cefoxitin	CoNS	21	28.0	0	0	54	72.0
Celoxitiii	S. aureus	2	14.2	0	0	11	78.6
C	CoNS	17	22.7	21	28.0	37	49.3
Streptomycin	S. aureus	10	71.4	4	28.6	0	0.0
Vancomycin	CoNS	19	25.3	0	0	56	74.7
v ancomychi	S. aureus	5	35.7	0	0	9	64.3
Clindomycin	CoNS	11	14.7	30	40.0	34	45.3
Clindamycin	S. aureus	3	21.4	7	50.0	4	28.5
Gentamicin	CoNS	13	17.3	13	17.3	49	65.3
Gentamicii	S. aureus	0	0.0	11	78.6	3	21.4
Cefotaxime	CoNS	5	6.7	16	21.3	54	72.0
	S. aureus	0	0.0	2	14.3	12	85.7
Ceftriaxone	CoNS	3	4.0	25	33.3	47	62.7
Certifiaxone	S. aureus	2	14.3	1	7.1	11	78.6
Ciprofloxacin	CoNS	4	5.3	8	10.7	63	84.0
Сіртопохасні	S. aureus	0	0.0	3	21.4	11	78.6

ceftazidime in combination with streptomycin 7/14 (50.0%), and vancomycin 5/14 (35.7%). Other major observed resistance combination is shown in Table 5.

Occurrence of MDR was observed across both species of Staphylococcus. In this study, most of the MDR involved beta-lactams and cephalosporins. For CoNS, the MDR involved beta-lactams and other classes of antibiotics: cephalosporins (57/79, 72.2% isolates), macrolides (37/79, 46.8% isolates), oxytetracycline (27/79, 34.2% isolates), aminoglycosides (16/79, 20.3%), and glycosides (15/79, 19%). For the *S. aureus*, MDR was observed majorly in beta-lactams and cephalosporins (13/14, 92.9%), aminoglycosides (7/14, 50% isolates), oxytetracycline (2/14, 14.29% isolates), and glycosides (3/14, 21.4% isolates). Resistance for the beta-lactam class was consistently high across all wards.

The prevalence of resistant *Staphylococcus* spp. strains varied across the study wards. Study participants from Mata-

pato South ward had bacteria resistant to all classes of antibiotics. Details of the distribution of resistance to the six classes of antibiotics tested are shown in Table 6.

3.4. Relationship between Risk Factors and Prevalence of Multidrug Resistance in Bacteria Isolated from the Study Participants. The antibiotic susceptibility results were used to evaluate the potential association between various factors: gender, age, consumption of raw milk, animal contact, antibiotic usage, hospital visits, and the occurrence of MDR bacteria in nares (Table 7. From the analysis, male participants had a higher risk of carrying MDR bacteria than females (p = 0.02, OR = 1.3), while habitual intake of raw milk was significantly associated with a higher prevalence of MDR bacteria (p = 0.02, OR = 1.82). Majority of the study participants (82.4%) were frequently in contact with livestock; however, this factor was not associated with the occurrence

Table 5: Antibiotic resistance combination patterns of CoNS and S. aureus isolated from nasal swabs of participants in Kajiado Central Subcounty, Kenya.

Bacteria	Antibiotic combination	No. of isolates (n)	%
	AMX and OX	61	77.2
	AMX and E	27	34.2
	AMX and VAN	20	25.3
	AMX and TE	24	30.4
C-NC	AMX and CTX	41	51.9
CoNS	E and CTX	23	29.1
	OX and CTX	56	70.9
	OX and VAN	20	25.3
	OX and TE	26	32.9
	OX and E	30	38.0
S. aureus	AMX and CTX	10	71.4
	AMX and STR	6	42.9
	CTX and STR	7	50.0
	CTX and VAN	5	35.7

Key: AMX: amoxicillin; CTX: cefotaxime; E: erythromycin; OX: oxacillin; STR: streptomycin; TE: tetracycline; VAN: vancomycin.

Table 6: Distribution of Staphylococcus spp. antibiotic resistance in participants from Kajiado Central Subcounty, Kenya.

	M. South (<i>n</i> , %)	M. North (<i>n</i> , %)	Wards Ilidamati (n, %)	Purko (<i>n</i> , %)	Dalalekutuk (n, %)	Total (n, %)
Beta-lactams	44 (100)	31 (100)	7 (100)	12 (92.3)	4 (100)	98 (99.0)
Macrolides	19 (43.2)	15 (48.4)	2 (28.6)	1 (7.7)	0 (0.0)	37 (37.7)
Glycopeptide	15 (34.1)	6 (19.4)	1 (14.3)	0 (0.0)	0 (0.0)	22 (22.2)
Aminoglycosides	11 (25.0)	6 (19.4)	1 (14.3)	4 (30.8)	1 (25.0)	23 (23.2)
Tetracycline	10 (22.7)	8 (25.8)	3 (42.9)	6 (46.2)	0 (0.0)	27 (23.3)
Quinolones	2 (4.5)	0 (0.0)	0 (0)	0 (0.0)	0 (0.0)	2 (2.0)

Key: M = Matapato.

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Table 7: Relationship between various factors and the occurrence of MDR bacteria in participants from Kajiado Central Subcounty, Kenya.

Factor	Number	MDR prevalence	p value, odds ratio
Gender	Male	98.1	p = 0.02, OR = 0.63
Gender	Female	100	p = 0.02, OK = 0.03
	<40	96.4	6 > 0.05 OD 1.61
Age	>40	100	p > 0.05, OR = 1.61
7 . 1 . 6	Yes	100	. 0.02 OD 1.02
Intake of raw milk	No	98.3	p = 0.02, OR = 1.82
	Frequently	98.8	5 1 OD 0.04
Animal contact	Infrequently	100	p = 1, OR = 0.94
	Yes	15.6	5 0.5 OD 0.72
Hospital visitation (≤3 months)	No	84.4	p = 0.5, OR = 0.72
A (11 (1 (1 (1 (1 (1 (1 (1 (1 (1 (1 (1 (1	Yes	15.6	5 0.07 OD 1.0
Antibiotic usage (≤3 months)	No	84.4	p = 0.07, OR = 1.0

of drug resistance bacteria (p = 1, OR = 9.7). The relationship between the prevalence of various risk factors and the occurrence of MDR bacteria is outlined in Table 7.

3.5. Prevalence of mecA and mecC Genes among Methicillin-Resistant Staphylococcus Isolates. The phenotypically methicillin-resistant isolates were further investigated for

the genotypic presence of *mecA* and *mecC* genes. The study showed that 16% of the CoNS isolates were mecA positive. Eight percent (8%) of the CoNS isolates harbored the *mecC* while the *S. aureus* isolates lacked both the *mecA* and *mecC* genes.

4. Discussion

The current study was geared at determining the spectrum of bacteria colonizing the nasal cavity of a pastoralist community in Kajiado Central Subcounty in Kenya as well as determine the antibiotic sensitivity of the obtained Staphylococcus isolates. First, CoNS, Enterococci spp., and S. aureus were the dominant bacteria in the nasal cavity of the study subjects with CoNS contributing close to half of the isolates. The dominance of these bacteria is not unique to these settings. Different studies across the world (Brazil, Saudi Arabia, Kuwait, Middle East, and Egypt) have reported similar trends [71-73]. The dominance of these bacteria is largely due to their commensal nature in humans, mainly on the skin, nasal cavity, and gut [2, 74]. The prevalence of S. aureus is however lower compared to that recorded elsewhere in hospital settings and among healthcare workers in Kenya [4, 56] and across communities in East Africa and sub-Saharan regions in Africa [55, 75, 76]. The difference may have been influenced by the difference in study, laboratory, experimental setting, and climate factors where the high temperature in the study area could have reduced S. aureus colonization [77, 78]. Moreover, the high rates of CoNS isolated in this study might have lowered S. aureus colonization by secreting molecules that negatively affect the virulence of S. aureus [79, 80].

Secondly, we show that most of the circulating Staphylococcus isolates in the study population are resistant to at least one antibiotic tested in our study. The highest resistance was recorded in the penicillin which is in line with other studies across the world [54, 56, 60, 81-85]. High penicillin resistance is linked with overuse of these antibiotics in treatment of bacterial infections especially in livestock [84-86]. In our opinion, HGT and interspecies transfer within the livestock and human ecology might have caused the observed resistance in the human population. Methicillin resistance among the CONS isolates was close to that reported in Nigeria (65%) [87] and Ghana (60-93%) [88, 89] but slightly lower than that recorded in some sub-Saharan countries (<90%) [90-92]. This resistance is however higher than that recorded in hospital settings in some parts of Kenya (21-30%) [93, 94] suggesting a higher circulation of MRS isolates in the study population compared to the other places which is alarming given the weak association of the history of antibiotic usage and hospital visitations to emergence and spread of MDR in this community. However, methicillin resistance for S. aureus in our study is comparable with that recorded elsewhere in the country in hospital, livestock, and community settings (1-90%) [4, 60, 83, 95].

Interestingly, resistance to vancomycin in the study area was higher than that recorded elsewhere in the world: 0-11% in Kenyan hospital setting [54, 82, 96] and 0-16% in the rest of the world [53, 97, 98]. The resistance is however lower

than that identified among camel slaughterhouse workers in Ethiopia (54%) [99]. Vancomycin resistance has been linked with presence of vanA operon especially in enteric bacteria. Molecular investigation of the causes of the observed vancomycin resistance will be beneficial especially in establishing possibility of HGT of vancomycin resistance gene to the Staphylococcus isolates given the codominance of the two bacteria in the study area. Importantly, the high vancomycin resistance in the current population compared to other studies might be partly due to low sample size especially among the S. aureus which limits the statistical power of the study. Of note, resistance of the Staphylococcus isolates across all six antibiotic classes tested (glycopeptide, aminoglycosides, macrolides, tetracycline, and quinolones) was noted which is alarming. Given the high susceptibility of the Staphylococcus isolates to Augmentin compared to the rest of the tested antibiotics, this drug can serve as an alternative drug for the treatment of staphylococcal infections in this population.

The presence of mecA and mecC genes in some of the CoNS isolates confirms the presence of SCCmec-associated MRS in the study area. The mecA gene is the gold standard for MRS; thus, occurrence of mecA gene and mecC gene in our study was expected. Although the mecC gene has been detected in various clinical studies across the world [100-103], this is the first study to report mecC gene resistance in Kenya. The presence of mecA and mecC genes in the methicillin-resistant isolates forms a basis for further molecular work especially SCCmec typing in the current study population. The absence of both the mecA and mecC genes in some of the MRS isolates is not unique to this setting; some studies have reported similar findings or lack of one of these genes in methicillin-resistant isolates [101, 104-106]. Thus, the possibility of a different MRS mechanism in the study area is highly probable which needs further investigation. Some studies that have explored further mecA and mecC negative isolates have identified the presence of the mecB homologue, hyperproduction of β -lactamase, and production of modified PBPs [17, 107-109]. Further, class 1 and 2 integrons containing dfrA12-orfFaadA2, dfrA17-aadA5, aadA2, and aadB, oxa2, aacA4, orfD-aacA4-catB8, aadB-catB3, orfD-aacA4 and aadBaadA1-cmlA6, dhfrA1-sat2-aadA1, dhfrA11, dhfrA1-sat2 gene cassettes, respectively, have been detected in resistant Staphylococcus isolates in the past [32–34]. Given the ability of the integrons to carry resistant genes for β -lactams, further investigation of the presence of these classes of integrons in the current study population will be necessary.

Multidrug resistance (MDR) was recorded in both CoNS and *S. aureus* majorly to the beta-lactams, macrolides, and aminoglycosides. Although most studies in Kenya and East Africa have focused on multiple drug resistance in livestock, food, and nosocomial infections, our findings are comparable to these studies [84, 110–112]. MDR resistance has been attributed accumulation of multiple drug resistant genes or overexpression of resistance genes encoding for efflux pumps and acquisition of less susceptible genes [113–117]. Investigation of the risk factors associated with MDR linked livelihood factors to the acquisition of MDR Staphylococcus

isolates. Habitual intake of raw milk had a significant impact on the acquisition of resistant MDR Staphylococcus bacteria which is in accord with other studies conducted elsewhere in the world [118–120]. *Staphylococcus* spp. bacteria have been one of the foodborne pathogens occurring in raw milk; thus, the link between raw milk intake and occurrence of MDR in the study participants is not surprising. Also, from the study, the males had higher odds of acquiring resistant bacteria compared to female which is in line with other studies [121–123]. The likelihood of the males to acquire MDR bacteria may be linked to the occupational differences between the two genders in the Maasai community where the males are more involved in animal husbandry compared to the females in this community, as well as differences between hygiene levels in the two genders.

Although antibiotic usage in the current study was not significantly associated with the acquisition of antibioticresistant strain which differs with other studies [51, 124, 125], the study result relied heavily on study participant honesty which was dependent on their memory, a factor that might have limited our study findings. Most pastoralists are known to extensively use antibiotics in the management of livestock diseases with lack of withdrawal period during the medication period. This may have led to consumption of antimicrobial drugs eventually through milk and meat products. In such scenarios, HGT between livestock ecology and human is possible. The link of livelihood factors to acquisition of MDR acquisition necessitates proper understanding of the diverse beliefs in this community, and how they affect AMR acquisition and spread that will be essential in designing a proper AMR intervention to mitigate antibiotic resistance spread.

5. Conclusions

We outline three major findings from this study. First, most of the circulating nasal bacteria in the Kajiado Central were CoNS with some containing the mecA and/or mecC antibiotic-resistant markers, suggesting the possibility of factors promoting high nasal colonization of Staphylococcal spp. in humans that need further investigation. Second, we record a high prevalence of MDR for most of the antibiotics currently used for staphylococcal infection treatment. Lastly, livelihood factors (intake of raw milk and gender) were the major contributors to the acquisition of multidrug resistance strains, while antibiotic pressure and history of hospital visitation < 3 months prior to this study did not significantly impact the acquisition of resistant bacteria strains in the human population. These findings provide a basis to integrate social studies into antimicrobial resistance in this community to understand the cultural, social, and economic factors influencing the acquisition of antimicrobial resistance in this study area. The study also gives insight on the presence of *mecA/mecC* methicillin-resistant Staphylococcus strains laying a foundation for further investigation of molecular mechanisms in the observed resistance as well as building up on efforts to mitigate antimicrobial resistance at both the national and global scales. This will aid in the development of appropriate antimicrobial resistant mitigation strategies.

Data Availability

The datasets generated are available from the corresponding author on reasonable request.

Disclosure

The views expressed herein are those of the authors and not necessary that of the funding agency.

Conflicts of Interest

The authors declare that there is no conflict of interest.

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References

- [1] T. M. Bosch, G. Biesbroek, K. Trzcinski, E. A. M. Sanders, and D. Bogaert, "Viral and bacterial interactions in the upper respiratory tract," *PLoS Pathogens*, vol. 9, no. 1, article e1003057, 2013.
- [2] D. N. Frank, L. M. Feazel, M. T. Bessesen, C. S. Price, E. N. Janoff, and N. R. Pace, "The human nasal microbiota and *Staphylococcus aureus* carriage," *PLoS One*, vol. 5, no. 5, article e10598, 2010.
- [3] S. de Benito, L. Alou, R. B. B. Vallejo et al., "Prevalence of Staphylococcus spp. nasal colonization among doctors of podiatric medicine and associated risk factors in Spain," Antimicrobial Resistance and Infection Control, vol. 7, no. 1, p. 24, 2018.
- [4] A. M. Aiken, M. M. Irene, S. J. Artur, A. Victoria, and M. Jonah, "Carriage of Staphylococcus aureus in Thika Level 5 Hospital, Kenya: a cross-sectional study," Antimicrobial Resistance and Infection Control, vol. 3, no. 1, 2014.
- [5] S. Joachim, J. Moyo, L. Nkinda et al., "Prevalence of methicillin-resistant *Staphylococcus aureus* carriage on admission among patients attending regional hospitals in Dar es Salaam, Tanzania," *BMC Research Notes*, vol. 10, no. 1, p. 417, 2017.
- [6] O. O. Ayepola, S. O. Taiwo, A. Anifowose, and O. Onile-ere, "Nasal carriage of *Staphylococcus aureus* and associated risk factors among students in a Nigerian University," *Acta Scientific Microbiology*, vol. 1, no. 2, pp. 6–8, 2018.
- [7] M. Z. David and R. S. Daum, "Community-associated methicillin-resistant *Staphylococcus aureus*: epidemiology and clinical consequences of an emerging epidemic," *Clinical Microbiology Reviews*, vol. 23, no. 3, pp. 616–687, 2010.

- [8] J. D. Bard, J. A. Hindler, H. S. Gold, and B. Limbago, "Rationale for eliminating *Staphylococcus* breakpoints for β -lactam agents other than penicillin, oxacillin or cefoxitin, and ceftaroline," *Clinical Infectious Diseases*, vol. 58, no. 9, pp. 1287–1296, 2014.
- [9] H. F. Chambers, "Methicillin resistance in staphylococci: molecular and biochemical basis and clinical implications," *Clinical Microbiology Reviews*, vol. 10, no. 4, pp. 781–791, 1997
- [10] E. Y. Garoy, Y. B. Gebreab, O. O. Achila et al., "Methicillin-Resistant Staphylococcus aureus (MRSA): Prevalence and Antimicrobial Sensitivity Pattern among Patients—A Multicenter Study in Asmara, Eritrea," Canadian Journal of Infectious Diseases Medical Microbiolology, vol. 2019, article 8321834, pp. 1–9, 2019.
- [11] S. Kariuki, K. Kering, C. Wairimu, R. Onsare, and C. Mbae, "Antimicrobial resistance rates and surveillance in sub-Saharan Africa: where are we now?," *Infection and Drug Resistance*, vol. Volume 15, pp. 3589–3609, 2022.
- [12] Antimicrobial Resistance Collaborators, "Global burden of bacterial antimicrobial resistance in 2019: a systematic analysis," *The Lancet*, vol. 399, no. 10325, pp. 629–655, 2022.
- [13] G. Marincola, W. Ziebuhr, F. D. R. Wencker et al., "Antimicrobial resistance profiles of coagulase-negative staphylococci in community-based healthy individuals in Germany," Frontiers in Public Health, vol. 9, article 684456, 2021.
- [14] J. B. Owolabi and S. K. Olatunde, "A review of prevalence, antimicrobial susceptibility patterns and molecular characteristics of methicillin-resistant *Staphylococcus aureus* (MRSA) in the Caribbean," *Advances in Applied Microbiol*ogy, vol. 12, no. 8, pp. 459–480, 2022.
- [15] J. Liu, D. Chen, B. M. Peters et al., "Staphylococcal chromosomal cassettes mec (SCCmec): a mobile genetic element in methicillin-resistant *Staphylococcus aureus*," *Microbial Pathogenesis*, vol. 101, pp. 56–67, 2016.
- [16] Y. Uehara, "Current status of staphylococcal cassette chromosome mec (SCCmec)," Antibiotics, vol. 11, no. 1, p. 86, 2022
- [17] H. Liu, G. Buescher, N. Lewis, S. Snyder, and D. Jungkind, "Detection of borderline oxacillin-resistant Staphylococcus aureus and differentiation from methicillin-resistant strains," European Journal of Clinical Microbiology & Infectious Diseases, vol. 9, no. 10, pp. 717–724, 1990.
- [18] K. Loewen, Y. Schreiber, M. Kirlew, N. Bocking, and L. Kelly, "Community-associated methicillin-resistant Staphylococcus aureus infection: literature review and clinical update," Canadian Family Physician, vol. 63, no. 7, pp. 512–520, 2017.
- [19] R. C. Moellering, "MRSA: the first half century," *Journal of Antimicrobial Chemotherapy*, vol. 67, no. 1, pp. 4–11, 2012.
- [20] T. J. Ochoa, J. Mohr, A. Wanger, J. R. Murphy, and G. P. Heresi, "Community-associated methicillin-resistant Staphylococcus aureus in pediatric patients," Emerging Infectious Diseases, vol. 11, no. 6, pp. 966–968, 2005.
- [21] L. Challagundla, J. Reyes, I. Rafiqullah et al., "Phylogenomic classification and the evolution of clonal complex 5 methicillin-resistant *Staphylococcus aureus* in the Western hemisphere," *Frontiers in Microbiology*, vol. 9, p. 1901, 2018.
- [22] A. Hassoun, P. K. Linden, and B. Friedman, "Incidence, prevalence, and management of MRSA bacteremia across patient populations—a review of recent developments in MRSA

- management and treatment," Critical Care, vol. 21, no. 1, p. 211, 2017.
- [23] S. Lakhundi and K. Zhang, "Methicillin-resistant Staphylo-coccus aureus: molecular characterization, evolution, and epidemiology," Clinical Microbiology Reviews, vol. 31, no. 4, 2018.
- [24] N. P. Marathe, S. S. Nagarkar, A. A. Vaishampayan et al., "High prevalence of class 1 integrons in clinical isolates of methicillin-resistant *Staphylococcus aureus* from India," *Indian Journal of Medical Microbiology*, vol. 33, no. 2, pp. 231–236, 2015.
- [25] P. Sabbagh, M. Rajabnia, A. Maali, and E. Ferdosi-Shahandashti, "Integron and its role in antimicrobial resistance: a literature review on some bacterial pathogens," *Iranian Journal of Basic Medical Sciences*, vol. 24, no. 2, pp. 136–142, 2021.
- [26] Z. Xu, L. Shi, C. Zhang et al., "Nosocomial infection caused by class 1 integron-carrying *Staphylococcus aureus* in a hospital in South China," *Clinical Microbiology and Infection*, vol. 13, no. 10, pp. 980–984, 2007.
- [27] M. Afsharian, M. Hemmati, F. Mansouri et al., "Frequency of class I and II Integrons in methicillin-resistant and methicillin-sensitive *Staphylococcus aureus* isolates in the city of Kermanshah," *Archives of Clinical Infectious Diseases*, vol. 14, no. 4, article 86688, 2019.
- [28] F. Hajiahmadi, E. S. Ghale, M. Y. Alikhani, A. Mordadi, and M. R. Arabestani, "Detection of integrons and staphylococcal cassette chromosome mec types in clinical methicillin-resistant coagulase negative staphylococci strains," *Osong Public Health and Research Perspectives*, vol. 8, no. 1, pp. 47–53, 2017.
- [29] L. Li and X. Zhao, "Characterization of the resistance class 1 integrons in *Staphylococcus aureus* isolates from milk of lactating dairy cattle in northwestern China," *BMC Veterinary Research*, vol. 14, no. 1, p. 59, 2018.
- [30] M. Mohammadi, N. Bahrami, M. Khajavian, and J. Faghri, "The occurrence of type I, II, and III integrons in multidrug resistance and methicillin-resistant *Staphylococcus aureus* isolates in Iran," *Current Microbiology*, vol. 77, no. 8, pp. 1653–1659, 2020.
- [31] M. Mostafa, S. D. Siadat, F. Shahcheraghi et al., "Variability in gene cassette patterns of class 1 and 2 integrons associated with multi drug resistance patterns in *Staphylococcus aureus* clinical isolates in Tehran-Iran," *BMC Microbiology*, vol. 15, no. 1, pp. 1–9, 2015.
- [32] Z. Xu, L. Shi, M. J. Alam, L. Li, and S. Yamasaki, "Integron-bearing methicillin-resistant coagulase-negative staphylococci in South China, 2001-2004," FEMS Microbiology Letters, vol. 278, no. 2, pp. 223–230, 2008.
- [33] Y. Deng, J. Liu, B. M. Peters et al., "Antimicrobial resistance investigation on Staphylococcus strains in a local Hospital in Guangzhou, China, 2001–2010," Microbial Drug Resistance, vol. 21, no. 1, pp. 102–104, 2015.
- [34] H. Goudarzi, S. S. Seyedjavadi, E. E. Udo, E. Beiranvand, M. Fazeli, and M. Goudarzi, "Molecular characterization and distribution of class 1 integron-bearing methicillin resistant Staphylococcus aureus strains in burn patients, Tehran, Iran," *Jundishapur Journal of Microbiology*, vol. 10, no. 2, article e40592, 2017.
- [35] M. E. S. Zaki, O. A. Faried, and K. Montaseer, "Molecular study of class 1 and 2 integrons in methicillin-resistant *Staphylococcus aureus*," *The Open Microbiology Journal*, vol. 15, no. 1, 2021.

[36] R. O. Darouiche, "Treatment of infections associated with surgical implants," *New England Journal of Medicine*, vol. 350, no. 14, pp. 1422–1429, 2004.

- [37] L. A. Mermel, M. Allon, E. Bouza et al., "Clinical practice guidelines for the diagnosis and management of intravascular catheter-related infection," *Clinical Infectious Diseases*, vol. 49, no. 1, pp. 1–45, 2009.
- [38] S. Gardete and A. Tomasz, "Mechanisms of vancomycin resistance in *Staphylococcus aureus*," *Journal of Clinical Investigation*, vol. 124, no. 7, pp. 2836–2840, 2014.
- [39] H. Kest and A. Kaushik, "Vancomycin-resistant *Staphylococcus aureus*: formidable threat or silence before the storm?," *Journal of Infectious Disease and Epidemiology*, vol. 5, no. 5, p. 93, 2019.
- [40] P. J. Stogios and A. Savchenko, "Molecular mechanisms of vancomycin resistance," *Protein Science*, vol. 29, no. 3, pp. 654–669, 2020.
- [41] S. Unni, T. J. Siddiqui, and S. Bidaisee, "Reduced susceptibility and resistance to vancomycin of *Staphylococcus aureus*: a review of global incidence patterns and related genetic mechanisms," *Cureus*, vol. 13, no. 10, article e18925, 2021.
- [42] R. Hasan, M. Acharjee, and R. Noor, "Prevalence of vancomycin resistant *Staphylococcus aureus* (VRSA) in methicillin resistant *S. aureus* (MRSA) strains isolated from burn wound infections," *Tzu Chi Medical Journal*, vol. 28, no. 2, pp. 49–53, 2016.
- [43] W. A. Mcguinness, N. Malachowa, and F. R. Deleo, "Vancomycin resistance in *Staphylococcus aureus*," *Yale Journal of Biology and Medicine*, vol. 90, no. 2, pp. 269–281, 2017.
- [44] T. Saber, M. Samir, R. M. El-Mekkawy et al., "Methicillinand vancomycin-resistant *Staphylococcus aureus* from humans and ready-to-eat meat: characterization of antimicrobial resistance and biofilm formation ability," *Frontiers* in *Microbiology*, vol. 12, article 735494, 2022.
- [45] S. Selim, O. A. Faried, M. S. Almuhayawi et al., "Incidence of vancomycin-resistant *Staphylococcus aureus* strains among patients with urinary tract infections," *Antibiotics*, vol. 1, no. 3, p. 408, 2022.
- [46] M. P. Gheorghe and L. G. Măruțescu, Molecular features of virulence and resistance mechanisms in nosocomial and community-acquired Staphylococcus aureus, Intech Open, 2019.
- [47] Y. Guo, G. Song, M. Sun, J. Wang, and Y. Wang, "Prevalence and therapies of antibiotic-resistance in *Staphylococcus aureus*," *Frontiers in Cellular and Infectious Microbiology*, vol. 10, p. 107, 2020.
- [48] R. Leclercq, "Mechanisms of resistance to macrolides and lincosamides: nature of the resistance elements and their clinical implications," *Clinical Infectious Diseases*, vol. 34, no. 4, pp. 482–492, 2002.
- [49] J. M. Munita and C. A. Arias, "Mechanisms of antibiotic resistance," *Microbiology Spectrum*, vol. 4, no. 2, pp. 10-11, 2016.
- [50] S. Deyno, A. Toma, M. Worku, and M. Bekele, "Antimicrobial resistance profile of *Staphylococcus aureus* isolates isolated from ear discharges of patients at University of Hawassa comprehensive specialized hospital," *BMC Pharmacology and Toxicology*, vol. 18, no. 1, 2017.
- [51] M. A. Caudell, A. Dorado-Garcia, S. Eckford et al., "Towards a bottom-up understanding of antimicrobial use and resistance on the farm: a knowledge, attitudes, and practices sur-

- vey across livestock systems in five African countries," *PLoS One*, vol. 5, no. 1, article e0220274, 2020.
- [52] J. Asante, B. A. Hetsa, D. G. Amoako, A. L. Abia, L. A. Bester, and S. Y. Essack, "Multidrug-resistant coagulase-negative staphylococci isolated from bloodstream in the uMgungundlovu district of KwaZulu-Natal province in South Africa: emerging pathogens," *Antibiotics*, vol. 10, no. 2, p. 198, 2021.
- [53] G. E.-S. Mashaly and R. H. El-Mahdy, "Vancomycin heteroresistance in coagulase negative Staphylococcus blood stream infections from patients of intensive care units in Mansoura University Hospitals, Egypt," *Annals of Clinical Microbiology* and Antimicrobials, vol. 16, no. 1, p. 63, 2017.
- [54] F. K. Wangai, M. M. Masika, M. C. Maritim, and R. A. Seaton, "Methicillin-resistant *Staphylococcus aureus* (MRSA) in East Africa: red alert or red herring?," *BMC Infectious Diseases*, vol. 19, no. 1, p. 596, 2019.
- [55] S. Kariuki and G. Dougan, "Antibacterial resistance in sub-Saharan Africa: an underestimated emergency," *Annals of the New York Academy of Sciences*, vol. 1323, no. 1, pp. 43–55, 2014.
- [56] G. Omuse, S. Kariuki, and G. Revathi, "Unexpected absence of meticillin-resistant *Staphylococcus aureus* nasal carriage by healthcare workers in a tertiary hospital in Kenya," *Journal* of *Hospital Infection*, vol. 80, no. 1, pp. 71–73, 2012.
- [57] F. K. Wangai, M. M. Moses, G. N. Lule et al., "Bridging antimicrobial resistance knowledge gaps: the East African perspective on a global problem," *PLoS One*, vol. 14, no. 2, article e0212131, 2019.
- [58] R. Kohli, G. Omuse, and G. Revathi, "Antibacterial susceptibility patterns of blood stream isolates in patients investigated at the Aga Khan University Hospital, Nairobi," *East African Medical Journal*, vol. 87, no. 2, pp. 74–80, 2010.
- [59] S. Ladhani, O. S. Konana, S. Mwarumba, and M. C. English, "Bacteraemia due to *Staphylococcus aureus*," *Archives of Disease in Childhood*, vol. 89, no. 6, pp. 568–571, 2004.
- [60] E. K. Maina, C. Kiiyukia, C. N. Wamae, P. G. Waiyaki, and S. Kariuki, "Characterization of methicillin-resistant Staphylococcus aureus from skin and soft tissue infections in patients in Nairobi, Kenya," International Journal of Infectious Diseases, vol. 7, no. 2, pp. e115–e119, 2013.
- [61] A. W. Mogere, "Carriage rate of methicillin resistant Staphylococcus aureus among health care workers at the Kenyatta National Hospital, [Ph.D. thesis]," University of Nairobi, 2015, http://erepository.uonbi.ac.ke/handle/11295/ 94041.
- [62] J. Nyasinga, G. Omuse, N. John et al., "Epidemiology of Staphylococcus aureus infections in Kenya: current state, gaps and opportunities," Open Journal of Medical Microbiology, vol. 10, no. 4, pp. 204–221, 2020.
- [63] S. Rutare, "Prevalence of Methicillin Resistant Staphylococcus aureus (MRSA) among Paediatric Patients Admitted in Intensive Care Unit and Neonatal Intensive Care Unit at Kenyatta National Hospital-Nairobi, Kenya," 2013, http://erepository.uonbi.ac.ke:8080/xmlui/handle/11295/58806.
- [64] J. M. Hassell, M. J. Ward, D. Muloi et al., "Clinically relevant antimicrobial resistance at the wildlife-livestock-human interface in Nairobi: an epidemiological study," *The Lancet Planetary Health*, vol. 3, no. 6, pp. e259–e269, 2019.
- [65] L. M. Langata, J. M. Maingi, H. A. Musonye, J. Kiiru, and A. K. Nyamache, "Antimicrobial resistance genes in Salmonella and Escherichia coli isolates from chicken droppings in

- Nairobi, Kenya," BMC Research Notes, vol. 12, article 22, 2019
- [66] KNBS, "2019 Kenya population and housing census, vol. II," 2019, http://www.knbs.or.ke.
- [67] M. Thrusfield, Veterinary Epidemiology Third Edition, Blackwell Science Ltd., Oxford, 2007.
- [68] P. Warnke, T. Harnack, P. Ottl, G. Kundt, and A. Podbielski, "Nasal screening for *Staphylococcus aureus* - daily routine with improvement potentials," *PLoS One*, vol. 9, no. 2, article e89667, 2014.
- [69] M. Cheesbrough, District Laboratory Practice in Tropical Countries, Cambridge University Press, Cambridge, 2nd ed. edition, 2009.
- [70] CLSI, CLSI Performance Standards for Antimicrobial M100-S23, Clinical and Laboratory Standards Institute, Payne, 2015.
- [71] G. C. M. De Almeida, N. G. M. Lima, M. M. Dos Santos, M. C. N. De Melo, and K. C. De Lima, "Colonização nasal por Staphylococcus sp. em pacientes internados," Acta Paulista de Enfermagem, vol. 27, no. 3, pp. 273–279, 2014.
- [72] Y. M. Ternes, J. Lamaro-Cardoso, M. C. P. Andre et al., "Molecular epidemiology of coagulase-negative Staphylococcus carriage in neonates admitted to an intensive care unit in Brazil," BMC Infectious Diseases, vol. 13, no. 1, 2013.
- [73] N. S. Alharbi, "Screening of antibiotic-resistant staphylococci in the nasal cavity of patients and healthy individuals," *Saudi Journal of Biological Sciences*, vol. 27, no. 1, pp. 100–105, 2019.
- [74] M. Yan, S. J. Pamp, J. Fukuyama et al., "Nasal microenvironments and interspecific interactions influence nasal microbiota complexity and S. aureus carriage," Cell Host & Microbe, vol. 14, no. 6, pp. 631–640, 2013.
- [75] G. Omuse, B. Kabera, and G. Revathi, "Low prevalence of methicillin resistant as determined by an automated identification system in two private hospitals in Nairobi, Kenya: a cross sectional study," *BMC Infectious Diseases*, vol. 14, no. 669, 2014.
- [76] T. P. Robinson, D. P. Bu, J. Carrique-Mas et al., "Antibiotic resistance is the quintessential One Health issue," *Transactions of the Royal Society of Tropical Medicine and Hygiene*, vol. 110, no. 7, pp. 377–380, 2016.
- [77] H. Wang, W. Dai, X. Feng et al., "Microbiota composition in upper respiratory tracts of healthy children in Shenzhen, China, differed with respiratory sites and ages," *BioMed Research International*, vol. 2018, Article ID 6515670, 8 pages, 2018.
- [78] F. Schwab, P. Gastmeier, and E. Meyer, "The warmer the weather, the more gram-negative bacteria - impact of temperature on clinical isolates in intensive care units," *PLoS One*, vol. 9, no. 3, article e91105, 2014.
- [79] M. Michalik, A. Samet, A. Podbielska-Kubera, V. Savini, J. Międzobrodzki, and M. Kosecka-Strojek, "Coagulase-negative staphylococci (CoNS) as a significant etiological factor of laryngological infections: a review," *Annals of Clinical Micro*biology and Antimicrobials, vol. 19, no. 1, p. 26, 2020.
- [80] P. Peng, M. Baldry, B. H. Gless et al., "Effect of co-inhabiting coagulase negative staphylococci on *S. aureus* agr quorum sensing, host factor binding, and biofilm formation," *Frontiers in Microbiology*, vol. 10, 2019.
- [81] E. Ş. Yılmaz and Ö. Aslantaş, "Antimicrobial resistance and underlying mechanisms in Staphylococcus aureus isolates,"

- Asian Pacific Journal of Tropical Medicine, vol. 10, no. 11, pp. 1059–1064, 2017.
- [82] W. Gitau, M. Masika, M. Musyoki, B. Museve, and T. Mutwiri, "Antimicrobial susceptibility pattern of *Staphylo-coccus aureus* isolates from clinical specimens at Kenyatta National Hospital," *BMC Research Notes*, vol. 11, no. 1, p. 226, 2018.
- [83] C. Kyany'a, J. Nyasinga, D. Matano et al., "Phenotypic and genotypic characterization of clinical *Staphylococcus aureus* isolates from Kenya," *BMC Microbiology*, vol. 19, no. 1, 2019.
- [84] M. Mbindyo, G. C. Gitao, and C. M. Mulei, "Prevalence, etiology, and risk factors of mastitis in dairy cattle in Embu and Kajiado Counties, Kenya," *Veterinary Medicine Interna*tional, vol. 2020, Article ID 8831172, 12 pages, 2020.
- [85] M. Mutonga, M. W. Mureithi, N. N. Ngugi, and F. C. F. Otieno, "Bacterial isolation and antibiotic susceptibility from diabetic foot ulcers in Kenya using microbiological tests and comparison with RT-PCR in detection of S. aureus and MRSA," BMC Research Notes, vol. 12, no. 1, p. 244, 2019.
- [86] M. Virto, G. Santamarina-García, G. Amores, and I. Hernández, "Antibiotics in dairy production: where is the problem?," *Dairy*, vol. 3, no. 3, pp. 541–564, 2022.
- [87] C. Heilmann, W. Ziebuhr, and K. Becker, "Are coagulasenegative staphylococci virulent?," *Clinical Microbiology and Infection*, vol. 25, no. 9, pp. P1071–P1080, 2019.
- [88] M. J. Newman, E. Frimpong, E. S. Donkor, J. A. Opintan, and A. Asamoah-Adu, "Resistance to antimicrobial drugs in Ghana," *Infection and Drug Resistance*, vol. 4, pp. 215–220, 2011.
- [89] B. Egyir, L. Guardabassi, S. S. Nielsen et al., "Prevalence of nasal carriage and diversity of *Staphylococcus aureus* among inpatients and hospital staff at Korle Bu Teaching Hospital, Ghana," *Journal of Global Antimicrobial Resistance*, vol. 1, no. 4, pp. 189–193, 2013.
- [90] E. Udobi, A. F. Obajuluwa, and J. A. Onaolapo, "Prevalence and antibiotic resistance pattern of methicillin-resistant *Staphylococcus aureus* from an orthopaedic hospital in Nigeria," *BioMed Research International*, vol. 2013, Article ID 860467, 4 pages, 2013.
- [91] C. Ntirenganya, O. Manzi, C. M. Muvunyi, and O. Ogbuagu, "High prevalence of antimicrobial resistance among common bacterial isolates in a tertiary healthcare facility in Rwanda," *American Journal of Tropical Medicine and Hygiene*, vol. 92, no. 4, pp. 865–870, 2015.
- [92] S. A. Rebiahi, D. E. Abdelouahid, M. Rahmoun, S. Abdelali, and H. Azzaoui, "Emergence de souches de Staphylococcus aureus r esistantes a la vancomycine isolees du centre hospitalo-universitaire de Tlemcen (Algerie Nord-ouest)," Médecine et Maladies Infectieuses, vol. 41, no. 12, pp. 646– 651, 2011.
- [93] C. Kesah, S. Ben Redjeb, T. O. Odugbemi et al., "Prevalence of methicillin-resistant Staphylococcus aureus in eight African hospitals and Malta," *Clinical Microbiology and Infection*, vol. 9, no. 2, pp. 153–156, 2003.
- [94] G. Omuse, B. Kabera, and G. Revathi, "Low prevalence of methicillin resistant Staphylococcus aureus as determined by an automated identification system in two private hospitals in Nairobi, Kenya: a cross sectional study," *BMC Infec*tious Diseases, vol. 14, no. 1, p. 669, 2014.
- [95] C. K. Mbogori, A. Muigai, and S. Kariuki, "Detection and Characterization of Methicillin resistant *Staphylococcus*

aureus from toilet and classroom door handles in selected secondary schools in Nairobi County," *Open Journal of Medical Microbiology*, vol. 3, no. 4, pp. 248–252, 2013.

- [96] C. Akoru, R. T. Kuremu, S. K. Ndege, A. Obala, J. W. Smith, and M. Bartlett, "Prevalence and anti-microbial susceptibility of methicillin resistant and Staphylococcus aureus; at Moi Teaching and Referral Hospital Eldoret," *Open Journal of Medical Microbiology*, vol. 6, no. 1, 2016.
- [97] L. Pinheiro, C. I. Brito, V. C. Pereira, A. D. Oliveira, C. H. Camargo, and M. D. Cunha, "Reduced susceptibility to vancomycin and biofilm formation in methicillin-resistant Staphylococcus epidermidis isolated from blood cultures," Memórias do Instituto Oswaldo Cruz, vol. 109, no. 7, pp. 871–878, 2014.
- [98] M. Al-Tamimi, J. Abu-Raideh, N. Himsawi, A. Khasawneh, and H. Hawamdeh, "Methicillin and vancomycin resistance in coagulase-negative staphylococci isolated from the nostrils of hospitalized patients," *Journal of Infection in Developing Countries*, vol. 14, no. 1, pp. 28–35, 2020.
- [99] K. Al-Amery, M. Elhariri, A. Elsayed et al., "Vancomycinresistant Staphylococcus aureus isolated from camel meat and slaughterhouse workers in Egypt," Antimicrobial Resistance and Infection Control, vol. 8, no. 1, 2019.
- [100] K. Becker, O. Denis, S. Roisin et al., "Detection of mecA-and mecC-positive methicillin-resistant Staphylococcus aureus (MRSA) isolates by the new Xpert MRSA Gen 3 PCR assay," Journal of Clinical Microbiology, vol. 54, no. 1, pp. 180–184, 2016
- [101] P. Basset, G. Prod'hom, L. Senn, G. Greub, and D. S. Blanc, "Very low prevalence of meticillin-resistant *Staphylococcus aureus* carrying the mecC gene in western Switzerland," *Journal of Hospital Infection*, vol. 83, no. 3, pp. 257–259, 2013.
- [102] A.-K. Lindgren, E. Gustafsson, A. C. Petersson, and E. Melander, "Methicillin-resistant *Staphylococcus aureus* with mecC: a description of 45 human cases in southern Sweden," *European Journal of Clinical Microbiology & Infectious Diseases*, vol. 35, no. 6, pp. 971–975, 2016.
- [103] M. Stegger, T. Wirth, P. S. Andersen et al., "Origin and evolution of European community-acquired methicillin-resistant Staphylococcus aureus," MBio, vol. 5, no. 5, article e01044, 2014.
- [104] A. Cikman, M. Aydin, B. Gulhan et al., "Absence of the mecC gene in methicillin-resistant *Staphylococcus aureus* isolated from various clinical samples: the first multi-centered study in Turkey," *Journal of Infection and Public Health*, vol. 12, no. 4, pp. 528–533, 2019.
- [105] M. M. Elhassan, H. A. Ozbak, H. A. Hemeg, M. A. Elmekki, and L. M. Ahmed, "Absence of the mecA gene in methicillin resistant *Staphylococcus aureus* isolated from different clinical specimens in Shendi City, Sudan," *BioMed Research International*, vol. 2015, Article ID 895860, 5 pages, 2015.
- [106] K. Becker, B. Ballhausen, R. Köck, and A. Kriegeskorte, "Methicillin resistance in Staphylococcus isolates: the "mec alphabet" with specific consideration of mecC, a mec homolog associated with zoonotic S. aureus lineages," International Journal of Medical Microbiology, vol. 304, no. 7, pp. 794–804, 2014
- [107] K. Becker, S. van Alen, E. A. Idelevich et al., "Plasmidencoded transferable mecb-mediated methicillin resistance in *Staphylococcus aureus*," *Emerging Infectious Diseases*, vol. 24, no. 2, pp. 242–248, 2018.

- [108] P. O. Olorunfemi, N. C. Ngwuluka, J. A. Onaolapo, and Y. K. E. Ibrahim, "Susceptibility and molecular characterization of mec A- and mec B-positive community acquired methicillin-resistant *Staphylococcus aureus* isolates from students," *Journal of Pharmacy & Bioresources*, vol. 18, no. 2, pp. 155–171, 2021.
- [109] A. Tomasz, H. B. Drugeon, H. M. De Lencastre, D. Jabes, L. Mcdougall, and J. Bille, "New mechanism for methicillin resistance in *Staphylococcus aureus*: clinical isolates that lack the PBP 2a gene and contain normal penicillin-binding proteins with modified penicillin-binding capacity," *Antimicro*bial Agents and Chemotherapy, vol. 33, no. 11, pp. 1869– 1874, 1989.
- [110] V. S. Sonola, G. Misinzo, and M. I. Matee, "Occurrence of multidrug-resistant *Staphylococcus aureus* among humans, rodents, chickens, and household soils in Karatu, Northern Tanzania," *International Journal of Environmental Research* and Public Health, vol. 33, no. 11, pp. 1869–1874, 2021.
- [111] R. Murugesan, R. Rituparna, S. Ojha et al., A cross sectional study on the prevalence of MDR Staphylococci and E. coli from livestock in Karnataka, India, Research Square, 2022.
- [112] G. O. Omwenga, E. S. Aboge, G. Mitema et al., "Antimicrobial usage and detection of multidrug-resistant *Staphylococcus aureus*, including methicillin-resistant strains in raw milk of livestock from northern Kenya," *Microbial Drug Resistance*, vol. 27, no. 6, pp. 843–854, 2021.
- [113] C. E. DeMarco, L. A. Cushing, E. Frempong-Manso, S. M. Seo, T. A. A. Jaravaza, and G. W. Kaatz, "Efflux-related resistance to norfloxacin, dyes, and biocides in bloodstream isolates of *Staphylococcus aureus*," *Antimicrobial Agents and Chemotherapy*, vol. 51, no. 9, pp. 3235–3239, 2007.
- [114] M. Kuroda, T. Ohta, I. Uchiyama et al., "Whole genome sequencing of meticillin-resistant *Staphylococcus aureus*," *The Lancet*, vol. 357, no. 9264, pp. 1225–1240, 2001.
- [115] P. Mazurkiewicz, A. J. M. Driessen, and W. N. Konings, "Energetics of wild-type and mutant multidrug resistance secondary transporter LmrP of *Lactococcus lactis*," *Biochi*mica et Biophysica Acta, vol. 1658, no. 3, pp. 252–261, 2004.
- [116] H. Nikaido, "Multidrug resistance in bacteria," *Annual Review of Biochemistry*, vol. 78, pp. 119–146, 2009.
- [117] S. M. Li, Y. F. Zhou, L. Li et al., "Characterization of the multi-drug characterization of the multi-drug resistance gene cfr in methicillin-resistant *Staphylococcus aureus* (MRSA) strains isolated from animals and humans in China," *Fron*tiers in Microbiology, vol. 9, p. 2925, 2011.
- [118] A. Schnitt and B. A. Tenhagen, "Risk factors for the occurrence of methicillin-resistant Staphylococcus aureus in dairy herds: an update," Foodborne Pathogens and Disease, vol. 17, no. 10, pp. 585–596, 2020.
- [119] S. Regasa, S. Mengistu, and A. Abraha, "Milk safety assessment, isolation, and antimicrobial susceptibility profile of *Staphylococcus aureus* in selected dairy farms of Mukaturi and Sululta Town, Oromia Region, Ethiopia," *Veterinary Medicine International*, vol. 2019, Article ID 3063185, 11 pages, 2019.
- [120] M. Jamali, B. Paydar, S. I. Radmehr, and A. Dadrasnia, "Prevalence and antimicrobial resistance of *Staphylococcus aureus* isolated from raw milk and dairy products," *Food Control*, vol. 54, pp. 383–388, 2015.
- [121] V. Barasa and L. Waldman, Exploring the Intersection of Sanitation, Hygiene, Water, and Health in Pastoralist

- Communities in Northern Tanzania, Institute of Development Studies, 2022.
- [122] J. P. Cruz, C. P. Cruz, and A. S. Al-Otaibi, "Gender differences in hand hygiene among Saudi nursing students," *Interna*tional Journal of Infection Control, vol. 11, no. 4, 2015.
- [123] K. Eriksson, T. E. Dickins, and P. Strimling, "Global sex differences in hygiene norms and their relation to sex equality," PLOS Global Public Health, vol. 2, no. 6, article e0000591, 2022.
- [124] C. M. Wolfe, B. Cohen, and E. Larson, "Prevalence and risk factors for antibiotic-resistant community-associated blood-stream infections," *Journal of Infection and Public Health*, vol. 7, no. 3, pp. 224–232, 2014.
- [125] S. Sakr, A. Ghaddar, B. Hamam, and I. Sheet, "Antibiotic use and resistance: an unprecedented assessment of university students' knowledge, attitude and practices (KAP) in Lebanon," *BMC Public Health*, vol. 20, no. 1, p. 535, 2020.