

African Journal of Bacteriology Research

Full Length Research Paper

Prevalence and antimicrobial susceptibility of the borderline oxacillin-resistant *Staphylococcus aureus* (BORSA) strains in Bamako, Mali

Dicko O. A.¹, Maïga A.^{1,2}, Diarra B.^{2,3} and Maïga I. I.^{1,2*}

¹Laboratoire de Biologie Médicale et Hygiène Hospitalière du CHU du Point G, Bamako, Mali.
²Faculté de Médecine et d'Odontostomatologie de Bamako (FMOS), Université des Sciences, des Techniques et des Technologies de Bamako (USTTB), Bamako, Mali.
³Centre Universitaire de Recherche Clinique (UCRC), USTTB, Bamako, Mali.

Received 13 January, 2023; Accepted 24 February, 2023

The borderline oxacillin-resistant *Staphylococcus aureus* (BORSA) strains have a low-level resistance to penicillinase-resistant penicillins. The aim of this study was to study the prevalence and the antimicrobial susceptibility of BORSA strains in Bamako. A retrospective study was conducted at the University Teaching Hospital of the Point G in Bamako. The *S. aureus* strains were isolated on Columbia agar supplemented with 5% sheep blood, nalidixic acid and colistin. Antimicrobial susceptibility testing was performed using the disc diffusion method on Mueller-Hinton agar. β -lactamase production was determined by nitrocephin disc. Among 735 nonrepetitive strains of *S. aureus*, 41 (5.6%) were BORSA and 335 (45.6%) methicillin-resistant *S. aureus* (MRSA). The prevalence of BORSA strains was 25 (4.9%) and 16 (7.2%) in hospital and out-patients areas, respectively. Of the 41 BORSA strains, five were not β -lactamase producers. Cefoxitin (100%), cephalothin (97.6%), gentamycin (90.2%), amikacin (90.2%), netilmicin (87.8%), pristinamycin (87.8%), tobramycin (82.9%), fusidic acid (83%), lincomycin (78%), amoxicillin combined with clavulanic acid (73.2%), sulfonamides (73.2%), kanamycin (73.2%) fosfomycin (73%) and chloramphenicol (70.7%) were the most active against the BORSA strains. The prevalence of BORSA strains was not high in this study, and the strains were susceptible to a great range of antibiotics.

Key words: Antimicrobial susceptibility, borderline oxacillin-resistant *Staphylococcus aureus*, prevalence, Bamako (Mali).

INTRODUCTION

Staphylococcus aureus belongs to the Staphylococcaceae family (Foster and Geoghegan, 2015). The *S. aureus* infections appear under various clinical aspects such as urinary tract infections, abscesses, septicemia, vaginitis, and/or pleuropulmonary infections (Avril et al., 2000). The

borderline oxacillin-resistant *Staphylococcus aureus* (BORSA) has a low level of resistance to penicillin resistant to penicillinases which are oxacillin, cloxacillin and methicillin, that is, antibiotics which are not hydrolyzed by staphylococcal β -lactamases (Hryniewicz

*Corresponding author. E-mail: <u>ibraizet@gmail.com</u>. Tel: +223 76081707.

Author(s) agree that this article remain permanently open access under the terms of the <u>Creative Commons Attribution</u> <u>License 4.0 International License</u> and Garbacz, 2017). β -lactams are the antibiotics of choice in the treatment of staphylococcal infections (Hryniecz and Garbacz, 2017).

The resistance of the BORSA strains is typically linked to hyperproduction of β -lactamases or in some cases to other mechanisms which are the production of an inducible methicillinase with plasmid mediation, the modification of genes of proteins binding penicillin (PBP) (Cros et al., 2010; Hryniecz and Garbacz, 2017; Maalej et al., 2012). The activity of oxacillin on the BORSA strains is restored by β -lactamase inhibitors such as clavulanic acid or sulbactam (Hryniecz and Garbacz, 2017; Maalej et al., 2012). β -lactamase inhibitors do not restore oxacillin activity when altered proteins binding penicillin occurs (Hryniecz and Garbacz, 2017).

The resistance to methicillin, on the contrary, is due to the acquisition of a new PBP, the PBP2a or PBP2' encoded by chromosomal genes *mecA* or *mecC* (mecALGA251) (Hryniecz and Garbacz, 2017; Maïga et al., 2017). In addition, the BORSA strains are not classified as either methicillin-resistant *S. aureus* strains or methicillin-susceptible *S. aureus* strains (Hryniecz and Garbacz, 2017). The methicillin resistance is crossed between penicillins and cephalosporins (Fleurette, 1989). The BORSA strains, also called methicillin-resistant borderline *S. aureus*, do not have PBP2a or PBP2' (Montanari et al., 1990). The BORSA has been described in Tunisia (Maalej et al., 2012).

In Mali, to the best of our knowledge, no study has been focused on the borderline-oxacillin resistant *Staphylococcus aureus* stains. Thus, the aim of our study was to determine the prevalence and susceptibility to antibiotics of the BORSA strains in Bamako both for epidemiological and clinical purposes.

MATERIALS AND METHODS

Study design and settings

A retrospective study was conducted between January 2014 and December 2020 at the medical biology and hospital hygiene laboratory of the University Teaching Hospital (UTH) of the Point G, Bamako Mali. The UTH of the Point G is the third-pyramidal reference in Mali, and has 522 beds divided between the surgical, intensive care and medical departments.

Bacterial strains

Patients admitted to the different departments of the UTH of the Point G, were hospitalized patients (in-patients), those who were not hospitalized and coming directly to the laboratory for different reasons were called out-patients.

The specimens collected were mainly respiratory origin, urine, pus, blood cultures, and/or vaginal secretions (Table 1).

The isolation of *S. aureus* strains was carried out on Columbia agar supplemented with nalidixic acid (15 mg/l), colistin (10 mg/l) and sheep blood (5%) at 37°C at the oven.

Given the retrospective study design, we included in this study, the *S. aureus* strains with complete drug susceptibility results

with additional details of sampling source and date. On the other hand, the *S. aureus* strains without information of the sampling source and/or origin were not included. In this study hospital strains refer to strains isolated from hospitalized patients at the UTH of the Point G, and extra-hospital strains those from outpatients (non-hospitalized patients).

While the molecular methods for identification are rapid and could save more time and labor, they are quite expensive and not implemented in our laboratory yet. Thus, the identification of the strains of *S. aureus* was made by the combination of Gram staining (bioMérieux, France), the presence of a catalase (bioMérieux, France), coagulase (Bio-Rad, France), the Pastorex reagent Staph Plus (Bio-Rad, France) and the API STAPH gallery (bioMérieux, France).

Susceptibility to antibiotics test

The antimicrobial susceptibility testing was carried out on Mueller-Hinton agar (Bio-Rad, France) by the disc method (Agar diffusion method) (Kahlmeter and Turnidge, 2012). The strains of *S. aureus* were classified as "susceptible", "intermediate" or "resistant" according to the recommendations of the Antibiotic Committee of the French Society of Microbiology/European Committee on Antimicrobial Susceptibility Testing in 2015 (CA-SFM/EUCAST) (Amara et al., 2022; Bonnet et al., 2010). For the purpose of analysis, the strains of *S. aureus* classified as <<intermediate>> to the antibiotics tested were considered resistant.

Laboratory procedure

The antimicrobial susceptibility testing was performed on Mueller-Hinton agar poured in a Petri dish. Five to eight colonies identical and isolated from an 18 to 24 h culture of S. aureus were suspended in a 5 ml of sterile saline solution which was calibrated to 0.5 McFarland. One milliliter of this suspension was then added to 9 ml of sterile distilled water (dilution 1:10). This second suspension is poured over the entire surface of the Mueller-Hinton agar poured into a Petri dish. The excess is poured into bleach. The seeded agar is left to dry for 15 min at 37°C inside the incubator. Please kindly note that the incubator is not used to dry seeded agar plate. Instead, the plates are allowed to stand on the bench for a while, before the antibiotics dics are introduced. After the seeded agar was dried, the blotting paper discs impregnated with the antibiotic discs to be tested are placed on the surface of the agar using a disc dispenser according to manufacturer's instructions (Bio-Rad, France).

After a first diffusion of the antibiotics at 30 min at room temperature, the Petri dish is incubated at 37°C for 18 to 24 h, in the inverted position (cover down). The paper disc impregnated with oxacillin is placed on the surface of another seeded agar in Petri dish (diameter 90 mm) and placed in the incubator at 30°C for 24 to 48 h in the inverted position also. The reading is performed in measuring the diameter of inhibition of each antibiotic disc using a caliper in contact of growth.

Antibacterial agents tested

The antibiotics tested are penicillin G (6 μ g), cephalothin (30 μ g), oxacillin (5 μ g), amoxicillin combinated with clavulanic acid (20 μ g + 10 μ g), cefoxitin (30 μ g), gentamycin (10 UI), kanamycin (30 UI), tobramycin (30 μ g), netilmicin (30 μ g), amikacin (30 μ g), streptomycin (10 μ g), erythromycin (15 μ g), lincomycin (15 μ g), pristinamycin (15 μ g), chloramphenicol (30 μ g), tetracycline (30 μ g), sulfonamides (200 μ g), trimethoprim(5 μ g), norfloxacin (5 μ g),

Specimen	Hospital strains (No. of strains)	Extra-hospital strains (No. of strains)	Total (Rate in %)	
Urines	196	165	361 (49.1)	
Pus	186	29	215 (29.2)	
Blood cultures	106	0	106 (14.4)	
Vaginal secretions	0	18	18 (2.5)	
Pleurisy	15	0	15 (2.4)	
Catheters	6	5	11 (1.5)	
Sputum	1	3	4 (<1)	
Ascites	3	0	3 (<1)	
Prostatic fluid	0	2	2 (<1)	
Total	513	222	735 (100)	

Table 1. Distribution of 735 strains of Staphylococcus aureus according to the origin and the specimens.

Source: Authors

fosfomycin (50 µg) and fusidic acid (10 µg) (Bio-Rad, France).

β-lactams resistance phenotypes

While the *S. aureus* strains susceptible to cefoxitin and oxacillin were considered methicillin-susceptible *S. aureus* (MSSA), those resistant to cefoxitin and oxacillin have been considered methicillin-resistant *Staphylococcus aureus* (MRSA) (Cattoir and Leclercq, 2012; Mougeot et al., 2001). In addition, the *S. aureus* strains resistant or intermediate to oxacillin and susceptible to cefoxitin were considered as borderline oxacillin-resistant *Staphylococcus aureus* (BORSA) (Hryniewicz and Garbacz, 2017). Moreover, the BORSA strains resistant or intermediate to penicillin G produced β -lactamase. The production of β -lactamase was confirmed using the nitrocephine (cefinase) disc (bioMérieux, France). The penicillin G-susceptible BORSA strains with a fuzzy border were not β -lactamase producers. We compared the susceptibility of the BORSA strains to that of randomly selected MSSA strains.

Ethical consideration

The use of the results of the susceptibility to antibiotics test of isolated *S. aureus* strains was done after patients signed an informed consent. The results were de-identify with anonymous numbers, and the publication of the results was made with the permission of the leadership of the Point G's UTH.

Statistical analysis

The results of the susceptibility to antibiotics were used to monitor the bacterial resistance's profile at the Point-G's UTH and Bamako.

Data entry and computer processing were done using Epi Info version 7.1 software. The χ^2 test was used to compare our proportions with a significance level of p \leq 0.05.

RESULTS

Global characteristics of patients and isolates

In total, 735 non-repetitive strains of S. aureus were

isolated from 734 persons between January 2014 and December 2020 and included in the study. The mean age of patients was 47.61 ± 19.17 years with extremes of three and 98 years. The sex ratio (male/female) of patients was 0.94. They were in 513 (69.8%) of hospital strains and 222 (30.2%) out-patients' strains.

The annual frequency of the *S. aureus* strains during the study period is presented in Figure 1.

Frequency of BORSA within S. aureus strains

Of the 513 hospital strains of *S. aureus*, 25 (4.9%) were BORSA, 243 (47.4%) MRSA and 245 (47.8%) MSSA. At the same time, of 222 out-patients' strains, 16 (7.2%) were BORSA, 92 (41.4%) MRSA and 114 (51.4%) MSSA. Thus, the BORSA strains prevalence was independent of origin (p= 0.18).

The distribution of the 735 strains according to the phenotype of resistance to β -lactams and the sample is reported in Table 2. The prevalence of the BORSA strains was independent of the sampling sites. Moreover, the prevalence of the MRSA strains was higher in the urine than in the other samples (p < 10⁻⁶).

The bacteriological characteristics of the BORSA hospital and extra-hospital strains are reported in Table 3. Two out-patients' BORSA strains were isolated from urine in the same patient in April 2017: one of them had diminished susceptibility to norfloxacin, chloramphenicol and streptomycin.

Moreover, of the 25 BORSA hospital strains, four did not produce β -lactamase, and of 16 BORSA extra-hospital strains, one did not produce β -lactamase.

Susceptibility profile of S. aureus strains

All our BORSA strains were resistant to oxacillin, but susceptible to cefoxitin. In addition, 40 (97.6%) were

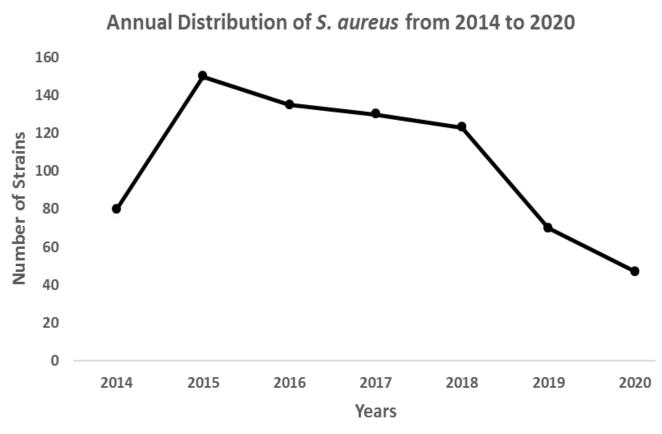


Figure 1. Annual distribution of the *Staphylococcus aureus* strains from 2014 to 2020 at the Point G University Teaching Hospital. Source: Authors

Table 2. Distribution of 735 strains of Staphylococcus aureus according to the phenotype and the specimens.

Specimen	BORSA [n (%)]	MRSA [n (%)]	MSSA [n (%)]	Total [n (%)]
Urines	13 (3.6)	223 (61.8)	125 (34.6)	361 (100)
Pus	17 (7.9)	54 (25.1)	144 (67)	215 (100)
Blood cultures	7 (6.6)	41 (38.7)	58 (54.7)	106 (100)
Vaginal secretions	1	6	11	18
Pleurisy	1	1	13	15
Catheters	2	6	3	11
Sputums	0	3	1	4
Ascites	0	0	3	3
Prostatic fluid	0	1	1	2
Total	41	335	359	735
	5.6	45.6	48.8	100

 χ^2 = 72.76; d.d.l. = 1; P < 10⁻⁶. BORSA = borderline oxacillin-resistant *S. aureus*; M RSA= methicillin-resistant *S. aureus*; M SSA= methicillin-susceptible *S. aureus*.

Source: Authors

susceptible to cephalothin, 30 (73.2%) to the combination of amoxicillin + clavulanic acid, and five (12.2%) to the penicillin G. Among the 36 BORSA strains producing β -lactamase, 25 were susceptible to the combination amoxicillin + clavulanic acid and 11 were resistant. The comparative antibiotic susceptibility of the BORSA and MSSA strains is reported in Table 4. The most active antibiotics on the BORSA strains were cefoxitin, cephalothin, gentamycin, amikacin, netilmicin, amoxicillin + clavulanic acid association, tobramycin, pristinamycin, Table 3. Bacteriological characteristics of 41 borderline-oxacillin resistant *Staphylococcus aureus* hospital and extra-hospital strains production.

Date	Specimens	Departments	Diameter zone inhibition (mm)					
			ΟΧΑ	Р	AMC	CF	FOX	⁻ β-lactamase
1/17/2015	Pus	Surgery B	19	30	35	36	31	Negative
2/8/2015	Pus	Int med.	16	33	30	32	30	Negative
4/27/2015	Pus	Surgery B	0	0	19	21	27	Positive
5/16/2015	Pus	HMO	13	10	17	23	25	Positive
6/14/2015	Pus	Surgery B	18	38	44	36	32	Negative
8/29/2015	Blood culture	HMO	15	21	25	15	26	Positive
10/5/2015	Urine	Int med.	18	12	23	32	27	Positive
12/4/2015	Blood culture	Int med.	16	14	30	28	26	Positive
12/27/2015	Blood culture	Emergencies	15	10	25	27	26	Positive
1/9/2016	Blood culture	Emergencies	15	11	26	27	27	Positive
1/31/2016	Urine	Neurology	15	27	29	33	29	Positive
4/5/2016	Blood culture	Emergencies	18	0	23	25	28	Positive
1/7/2017	Pus	Emergencies	18	21	30	35	28	Positive
1/29/2017	Pus	Surgery A	14	18	34	40	25	Positive
1/9/2017	Pus	Surgery A	0	14	25	28	28	Positive
6/9/2017	Pus	Surgery B	17	20	17	23	25	Positive
3/14/2017	Blood culture	Emergencies	12	13	23	29	30	Positive
3/25/2017	Pus	Int med.	14	16	26	31	32	Positive
9/23/2018	Pus	Surgery A	0	16	23	28	29	Positive
9/29/2018	Pus	Gynecology	0	10	20	27	30	Positive
10/6/2018	Pus	Inf Dis	18	33	40	29	33	Negative
9/17/2019	Pus	Surgery B	0	0	20	25	26	Positive
0/20/2019	Pleurisy	Emergencies	0	0	20	23	25	Positive
9/27/2019	Blood culture	Int med.	0	0	17	27	28	Positive
1/28/2020	Urine	ICU	0	0	23	22	25	Positive
3/20/2015	Urine	EHS	0	0	24	30	28	Positive
6/9/2015	Urines	EHS	18	35	33	36	26	Negative
12/28/2016	Pus	EHS	17	22	35	35	30	Positive
4/2/2017	Pus	EHS	14	24	30	33	28	Positive
4/9/2017	Urine	EHS	0	11	23	29	25	Positive*
4/9/2017	Urine	EHS	0	12	23	29	27	Positive
4/15/2017	VS	EHS	15	15	29	32	30	Positive
7/15/2017	Urine	EHS	0	24	34	20	32	Positive
7/24/2017	Urine	EHS	0	0	21	30	33	Positive
3/12/2017	Urine	EHS	16	19	30	34	26	Positive
9/16/2017	Urine	EHS	17	20	29	19	30	Positive
1/11/2017	Urine	EHS	18	22	30	32	27	Positive
12/19/2017	Urine	EHS	0	0	29	34	30	Positive
6/13/2019	Catheter	EHS	19	0	17	25	28	Positive
6/20/2019	Pus	EHS	18	8	20	27	28	Positive
5/25/2020	Catheter	EHS	0	0	18	30	29	Positive

VS = Vaginal secretions; HMO = hematology-medical oncology; Int med = internal medicine; Inf Dis= Infectious Diseases; ICU = intensive care unit; EHS = extra-hospital strains; OXA = oxacillin; P = penicillin G; AMC = amoxicillin combined with clavulanic acid; CF = cephalothin; FOX = cefoxitin. *Diminished susceptibility to norfloxacin, chloramphenicol and streptomycin. Source: Authors

fusidic acid, lincomycin, chloramphenicol, sulfonamides, kanamycin, and fosfomycin. Moreover, MSSA strains were

more susceptible to amoxicillin + clavulanic acid, kanamycin, tobramycin, erythromycin, lincomycin,

Antibiotics -	BORSA (No. of strains = 41)		MSSA (No. of strains = 82)		
	S [n (%)]	I+R [n (%)]	S [n (%)]	l+R [n (%)]	P-value
Penicillin G	5 (12.2)	36 (87.8)	16 (19.5)	66 (80.5)	0.4
AMC	30 (73.2)	11 (26.8)	82 (100)	0 (0)	0.001
Oxacillin	0 (0)	41 (100)	82 (100)	0 (0)	
Cephalothin	40 (97.6)	1 (2.4)	82 (100)	0 (0)	0.72
Gentamycin	37 (90.2)	4 (9.8)	82 (100)	0 (0)	0.02
Kanamycin	30 (73.2)	11 (26.8)	75 (91.5)	7 (8.5)	0.015
Tobramycin	34 (82.9)	7 (17.1)	81 (98.8)	1 (1.2)	0.0029
Amikacin	37 (90.2)	4 (9.8)	82 (100)	0 (0)	0.019
Netilmicin	36 (87.8)	5 (12.2)	82 (100)	0 (0)	0.006
Streptomycin	25 (61)	16 (39)	56 (68.3)	26 (31.7)	0.54
Erythromycin	24 (58.5)	17 (41.5)	72 (87.8)	10 (12.2)	0.001
Lincomycin	32 (78)	9 (22)	82 (100)	0 (0)	0.001
Pristinamycin	36 (87.8)	5 (12.2)	82 (100)	0 (0)	0.006
Norfloxacin	24 (58.5)	17 (41.5)	63 (76.8)	19 (23.2)	0.058
Chloramphenicol	29 (70.7)	12 (29.3)	75 (91.5)	7 (18.5)	0.006
Tetracycline	16 (39)	25 (61)	35 (42.7)	47 (57.3)	0.846
Sulfonamides	30 (73.2)	11 (26.8)	69 (84.2)	13 (15.8)	0.23
Trimethoprim	25 (61)	16 (39)	60 (73.2)	22 (26.8)	0.24
Fusidic acid	34 (83)	7 (17)	79 (96.3)	3 (3.7)	0.0267
Fosfomycin	30 (73.2)	11 (26.8)	81 (98.8)	1(1.2)	0.001

Table 4. Comparative antimicrobial susceptibility of BORSA and MSSA strains.

BORSA = borderline oxacillin-resistant *S. aureus*; MSSA = methicillin-susceptible *S. aureus*; AMC = amoxicillin combined with clavulanic acid; S = susceptible; I = intermediate; R = resistant; P = probability. Source: Authors

pristinamycin, chloramphenicol, fusidic acid and fosfomycin the BORSA strains: the differences are statistically significant.

DISCUSSION

In this study, we determine the prevalence and susceptibility to antibiotics of the BORSA strains between 2014 and 2020, using the disk method (agar diffusion technique): international reference technique an (Kahlmeter and Turnidge, 2012). To the best of our knowledge, this study is the first study of its kind conducted in Bamako, Mali making difficult comparison of previous data from Mali. The identification of our strains was based on their morphological, physiological and biochemical characteristics (Avril et al., 2000; Fleurette, 1989). The identification of S. aureus was characterized by its susceptibility to lysostaphin, nitrofurantoin and its resistance to the vibriostatic compound O/129 and to bacitracin (Fleurette, 1989).

The results of the antibiogram were interpreted in accordance with the recommendations of the Antibiogram Committee of the French Society of Microbiology and the European Committee on Antimicrobial Susceptibility Testing (Amara et al., 2022; Bonnet et al., 2010).

Other higher sensitivity and specificity methods were used for the detection of resistance to methicillin such as the oxacillin disc placed either on a Petri-dish of Mueller-Hinton agar incubated between 25 and 30°C and observed after 24 and 48 h or on a Petri-dish of hypersalted Mueller-Hinton agar incubated at 37°C, the use of probes recognizing *mec* genes, and the agglutination reactions of latex particles sensitized with anti-PBP2a antibodies (April et al., 2000). But the agglutination reactions of latex particles sensitized with anti-PBP2a antibodies is not routinely used (Maalej et al., 2012).

Currently, the identification of the MRSA and BORSA strains are based on the use of oxacillin and cefoxitin discs on Mueller-Hinton agar incubated at 37° C and examined 24 h later. For oxacillin, the interpretation must be done within 48 h if there is a low amount of culture (Bonnet et al., 2010; Cattoir and Leclercq, 2012). The use of the cefoxitin disc alone could make it possible to wrongly classify the BORSA strains among those with MSSA. The use of cefoxitin (30 µg) and moxalactam (30 µg) discs make it possible to identify MRSA at the Saint-Louis Hospital in France with excellent sensitivity (Felten et al., 2002).

In this study, the BORSA strains prevalence was 5.6%

and independent of the sampling site and the origin of the strains. The MRSA strains prevalence was 46.2% and was higher in the urine than in the other samples (Table 2). As observed in Myanmar (Soe et al., 2021), this high prevalence in urine samples, may have occurred because of urinary catheterization practice and the colonization by MRSA of indwelling urinary catheters. In Tunisia, of the 1,895 strains of S. aureus isolated between 2006 and 2011 at the University Hospital of Sfax, 451 (21.9%) were MRSA and 23 (1.2%) BORSA (Maalej et al., 2012). These rates are lower than ours. In a previous study conducted in 2007 in our department, the frequency of MRSA strains was 50.3 and 50.8% among hospital and out-patients' settings, respectively (Tchougoune, 2007). Those rates are closed to these current findings. Usually, the prevalence of the BORSA varies from 1.4 to 12.5% (Hrynievicz and Garbacz, 2017), and our rate is within the intermediate values (Table 2).

In this study, we isolated the BORSA strains from different collection sites, and we did not find a correlation between the strains and the site of collection (Table 2). This was the same in Sfax (Tunisia), and in Toronto (Canada) (Maalej et al., 2012; Leahy et al., 2011). Moreover, the hospital BORSA strains were coming from different units, and it was the same in Tunisia and Canada. In addition, we did not observe an outbreak of the BORSA strains during the study period, and this was the same in Tunisa as well (Maalej et al., 2012). Moreover, when analysis the 16 BORSA strains isolated from children with cystic fibrosis by pulsed-field restricted DNA electrophoresis in Toronto: no link was observed between the strains (Leahv et al., 2011). No link between the 23 strains was observed in Tunisa (Maalej et al., 2012).

Of the 41 BORSA strains, five did not produce β lactamase (Table 3), and this was the same in Canada, where out of 16, four did not produce β -lactamase (Leahy et al., 2011). At the same time in Tunisia, all the BORSA strains produced β -lactamase (Maalej et al., 2012). This could probably be explained by the sample size which was different in the three countries.

In contrast to Tunisia and Canada (Leahy et al., 2011; Maalej et al., 2012), the combination of amoxicillin + clavulanic acid was not active on 11 BORSA strains out of 41 (Tables 3 and 4). This is probably explained by other mechanisms (Croes et al., 2010; Hryniewicz and Garbacz, 2017; Jupeau-Vessières and Scavizzi, 1994; Maalej et al., 2012). Moreover, both BORSA strains in Mali and Tunisia were susceptible to cefoxitin (Table 3) (Maalej et al., 2012).

In addition, penicillin G, cephalothin, streptomycin, norfloxacin, tetracycline, sulfonamides and trimethoprim were not more active on MSSA strains than on BORSA strains (Table 4). Gentamycin (90.2%), amikacin (90.2%), tobramycin (82.9%), netilmicin (87.8%) and kanamycin (73.2%) were the most active aminoglycosides on our BORSA strains (Table 4). Among the BORSA strains isolated in Sfax, 13 out of 23 strains were susceptible to

kanamycin, 20 to gentamycin and 19 to tobramycin. Fusidic acid (83%), fosfomycin (73%) and chloramphenicol (70.7%) had good activity on our BORSA strains (Table 4). All the BORSA strains isolated in Sfax were susceptible to fosfomycin, chloramphenicol and glycopeptides (Maalej et al., 2012). Among our BORSA strains, one out of two strains was susceptible to erythromycin (Table 4). Eleven out of 23 BORSA strains were susceptible to erythromycin in Sfax, Tunisia (Maalej et al., 2012). Thirteen BORSA strains out of 16 were susceptible to erythromycin in Toronto, Canada (Leahy et al., 2011).

Thus, the search of BORSA and MRSA strains is a priority at the UTH of the Point G in Bamako.

Our study has some limitations including the retrospective approach which may have precluded to firmed conclusion because of the loss of sensitive information. In addition, we have more patients who were not included due to missing data either on sample types and origins or culture results and have used patients who were diagnosed in urban Bamako district only who may have different results from rural areas of Mali.

Despite these limitations, we studied a relatively high sample size to determine the prevalence and the antimicrobial susceptibility of the borderline oxacillinresistant *Staphylococcus aureus* strains in Bamako in sample types. In addition, given that almost all the regions in Mali are getting a bacterial culture system, thus, a larger sample size and sampling from all the different regions of Mali would be essential for obtaining a true country profile of BORSA and MRSA phenotypes.

Conclusion

The BORSA strains prevalence is not high in Bamako and was not associated with the origin and sampling site. The BORSA strains were susceptible to many molecules: cefoxitin, cephalothin, gentamycin, amikacin, netilmicin, tobramycin, the combination amoxicillin + clavulanic acid, pristinamycin, acid fusidic, lincomycin, chloramphenicol, sulfonamides, kanamycin, and fosfomycin. In contrast, antibiotics such as erythromycin, norfloxacin, tetracycline, trimethoprim, and streptomycin should not be used in the treatment of infections caused by strains of the BORSA strains in Bamako.

CONFLICT OF INTERESTS

The authors have not declared any conflict of interests.

REFERENCES

Amara M, Aubin G, Cattoir V, Dortet L, Goutelle S, Jeannot K, Lepeule R, Lina G, Marchandin H, Mérens A, Ploy MC, Schramm F, Varon E (2022). Comité de l'Antibiogramme de la Société Française de Microbiologie/European Committee on Antimicrobial Susceptibility Testing. Recommandations 2022; Avalable on www.sfmmicrobiologie.org : accessed on september 26th 2022.

- Avril JL, Dabernat H, Denis F, Monteil H (2000). Bactériologie clinique, 3rd édition. Paris: Ellipses. 602 p.
- Bonnet R, Cavallo JD, Chardon H, Chidiac C, Courvalin P, Dabernat H, Drugeon H, Dubreuil L, Guery B, Jarlier V, Jehl F, Lambert T, Leclerq R, Nicolas-Chanoine M-H, Plesiat P, Quentin C, Soussy CJ, Varon E, Weber P (2010). Comité de l'Antibiogramme de la Société Française de Microbiologie/European Committee on Antimicrobial Susceptibility Testing. Recommandations 2010 ; Avaliable on http://www.sfm.asso.fr accessed on april 22nd 2010.
- Cattoir V, Leclercq R (2012). β-lactamines et staphylocoques. In. Courvalin P, Leclercq R (eds). Antibiogramme, 3rd édition. Paris: Eska pp.137-145.
- Croes S, Beisser SP, Terporten PH, Neef V, Deurenberg RH, Toberingh EE (2010). Diminished *in vitro* antibacterial activity of oxacillin against clinical isolates of borderline oxacillin-resistant *Staphylococcus aureus*. Clinical Microbiology Infections 16:PMID: 20880412 DOI: 10.1111/j.1469-0691.2010.02956.x
- Felten A, Grandry B, Lagrange PH, Casin I (2002). Evaluation of three techniques for detection of low-level methicillin-resistant *Staphylococcus aureus* (MRSA): a disk diffusion method with cefoxitin and moxalactam, the vitek2 system and the MRSA- screen latex agglutination test. Journal of Clinical Microbiology 40: PMID: 12149327, PMCID: PMC120619.
- Fleurette J (1989). Staphylocoques et microcoques. In. Le Minor L, Véron V (eds).Bactériologie médicale, 2nd édition. Paris: Flammarion pp. 773-794.
- Foster TJ, Geoghegan JA (2015). Staphylococcus aureus. In. Tang YW, Sussman M, Liu D, Poxton I and Schwartzman J (eds). Molecular medical microbiology, Volume II. 2nd edition. New York, NY: Elsevier pp. 655-674.
- Hryniewicz MM, Garbacz K (2017). Borderline oxacillinresistant *Staphylococcus aureus* (BORSA) – amore common problem than expected? Journal of Medical Microbiology DOI 10.1099/jmm.0.000585
- Jupeau-Vessières AM, Scavizzi MR (1994). Evolution de la résistance bactérienne aux antibiotiques. Encyclopédie médico-chirurgicale Maladies Infectieuses 16 p.
- Kahlmeter G, Turnidge J (2012). Techniques phénotypiques. In. Courvalin P, Leclercq R (eds). Antibiogramme, 3rd édition. Paris : Eska pp. 59-69.

- Leahy TR, Yau YCW, Atenafu E, Corey M, Ratjen F, Waters V (2011). Epidemiology of borderline oxacillin-resistant *Staphylococcus aureus* in pediatric cystic fibrosis. Pediatric Pulmonology 46. PMID: 21337531.
- Maalej SM, Rhimi FM, Fines M, Mnif B, Leclercq R, Hammami A (2012). Analysis of borderline oxacillin-resistant *Staphylococcus aureus* (BORSA) strains in Tunisia. Journal of Clinical Microbiology 50. PMID: 22814459 PMCID: PMC3457433.
- Maïga A, Dicko OA, Tchougoune LM, Fofana DB, Coulibaly DM, Maïga II (2017). Haute prévalence des souches de *Staphylococcus aureus* résistantes à la méticilline au centre hospitalier universitaire du Point G à Bamako (Mali). Mali Médical 32(3):1-8.
- Montanari MP, Tonin È, Biavasco F, Varaldo PE (1990). Further characterization of borderline methicillin-resistant *Staphylococcus aureus* and analysis of penicillin- binding proteins. Antimicrobial Agents and Chemotherapy 34. PMID: 2360829 PMCID: PMC171719.
- Mougeot C, Guillaumat-Taillet J, Libert JM (2001). *Staphylococcus aureus* : nouvelle détection de la résistance intrinsèque par la méthode de diffusion. Pathologie Biologie 49. PMID: 11367553.
- Soe PE, Han WW, Sagili KD, Satyanarayana S, Shrestha P, Htoon TT, Tin HH (2021). High Prevalence of Methicillin-Resistant *Staphylococcus aureus* among Healthcare Facilities and Its Related Factors in Myanmar (2018–2019). Tropical Medicine and Infectious Disease 6:70.
- Tchougoune LM (2007). Prévalence des souches de *Staphylococcus aureus* résistantes à la méticilline au CHU du Point G [thèse]. Bamako: Université des Sciences, des Techniques et des Technologies de Bamako 74 p.