

Emerging Trends of Methicillin-Resistant Staphylococcus Aureus: A Tertiary Care Hospital-Based Study

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Author's Contribution

^{1,3,5}Substantial contributions to the conception or design of the work or the acquisition, Final approval of the version to be published. ^{2,4}Drafting the work or revising it critically for important intellectual content, ⁶Active participation in active methodology

Funding Source: None

Conflict of Interest: None

Received: Jan 17, 2024

Accepted: May 22, 2024

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ABSTRACT

Objective: To analyze the frequency and antimicrobial susceptibility patterns of Methicillin-resistant Staphylococcus aureus (MRSA) in a tertiary care hospital in Lahore, Pakistan.

Methodology: A retrospective study was conducted from October 2020 to December 2021 in a tertiary care hospital in Lahore, Pakistan. The clinical specimens meeting the inclusion criteria underwent processing in the microbiology section of a tertiary care hospital in Lahore, Pakistan, following the CLSI 2021 standards. Identification of all Methicillin-resistant Staphylococcus aureus (MRSA) isolates was conducted using standard laboratory procedures. Antibiotic susceptibility testing was performed utilizing the modified Kirby-Bauer disc diffusion method, with E-strips employed for vancomycin assessment.

Results: A total of 65 Staphylococcus aureus specimens were isolated. Among these, 49% were identified as Methicillin-resistant Staphylococcus aureus (MRSA). The majority of MRSA isolates (56%) were found in males, and a significant proportion (59%) fell within the age range of 21-60 years. Pus specimens accounted for the highest frequency (78%) among all samples. Notably, all MRSA isolates exhibited 100% resistance to beta-lactam drugs and combination therapies such as penicillin, cephalosporin, carbapenem, and amoxicillin-clavulanic acid. Conversely, Linezolid demonstrated complete sensitivity, while Chloramphenicol showed a sensitivity rate of 93%. Fluoroquinolones and aminoglycosides displayed a sensitivity of up to 50%. Additionally, the minimum inhibitory concentration (MIC) for vancomycin indicated sensitivity across all study isolates.

Conclusion: The study concluded that MRSA is continuously on the rise, with increasing resistance and limiting the antibiotic options for the treatment of these superbugs.

Keywords: Beta-Lactam drugs, Cephalosporin, MRSA, Tertiary care Hospitals, Lahore.

Cite this article as: Ikram S, Inam S, Shahid F, Aslam A, Inam A, Ashraf M. Emerging Trends of Methicillin-Resistant Staphylococcus Aureus: A Tertiary Care Hospital-Based Study. Ann Pak Inst Med Sci. 2024; 20(3):272-276. doi: 10.48036/apims.v20i3.1098.

Introduction

Staphylococcal infections are amongst the top causes of global mortality and morbidity, leading to hospital-acquired as well as community-acquired infections.⁽¹⁾ These include bloodstream infections, skin and soft tissue

infections, pneumonia, and post-operative wound infections.⁽²⁾ Treatment of staphylococcal infections is becoming difficult due to the drastic changes in the drug susceptibility profiles worldwide⁽³⁻⁵⁾ and rising resistance against beta-lactam antibiotics.⁽⁶⁾ As a consequence, the treatment of serious life-threatening infections caused by

Methicillin-resistant *Staphylococcus aureus* (MRSA), leads to prolonged hospital stay and increased cost of care. Once established, MRSA is very difficult to eradicate as it spreads more readily than other strains once introduced into hospitals. In many countries, 75 % of all *S. aureus* isolates in hospitals are MRSA, and infected or colonized patients are the major source of transmission. ^(7, 8)

MRSA is difficult to control and eradicate and has emerged due to injudicious antimicrobial chemotherapy. Hence, the knowledge of their antibiotic sensitivity pattern and prevalence is necessary for the selection of appropriate treatment. With this objective, the current study was conducted in a tertiary care hospital in Lahore to analyze the antimicrobial susceptibility patterns of MRSA.

Methodology

The study was initiated in a tertiary care hospitals of Lahore during the period of October 2020 and December 2021. All clinical samples obtained during that time frame that met the inclusion criteria were processed in the microbiological department of the pathology laboratory and included in the study. After inoculation on blood and MacConkey agar, specimens were incubated at 35-37 °C for 18 to 24 hours. Biochemical tests were conducted on gram-stained bacterial colonies that had beta hemolysis on blood agar or colonies that were tinted a golden yellow. *Staphylococcus aureus* was confirmed by performing DNase and coagulase assays on Gram-positive, catalase-positive bacteria that showed bunches on Gram stain. ⁽⁹⁾

Kirby Bauer disc diffusion method was used to assess the antimicrobial susceptibility patterns, as per Clinical Laboratory Standards Institute (CLSI, 2021) guidelines. Following antibiotics were analyzed: Oxacillin, penicillin, gentamicin, amikacin, trimethoprim/sulfamethoxazole, ciprofloxacin, levofloxacin, and erythromycin and clindamycin chloramphenicol, linezolid. ⁽¹⁰⁾ Cefoxitin was used to interpret the sensitivity of cephalosporin's, carbapenem and combination drugs. The zone of inhibition was measured and interpreted according to CLSI 2021 guidelines. ⁽¹⁰⁾ Vancomycin susceptibility was done by E test (Epsilometer) techniques and the results were interpreted based on determination of MIC of vancomycin in µg/ml after 48 h of incubation usually 2 µg/ml to 16 µg/ml concentration. Data was compiled on Microsoft Excel.

Inclusion Criteria

Patients with a history of no antibiotic use and symptoms of illness, as well as specimens yielding Cefoxitin-

resistant *Staphylococcus aureus* (MRSA) were included in the study.

All the coagulase negative *staphylococcus* species and Cefoxitin sensitive *staphylococcus aureus* (MRSA) were excluded from the study. The study was approved from Institutional Ethical Review Board, Azra Naheed Medical College with reference no IRB/ANMC/2020/008.

Results

A total of 65 *staphylococcus aureus* were isolated from the specimens, out of which 32 were confirmed as Methicillin resistant *staphylococcus aureus* (MRSA), making a percentage of 49%.

Males (56%) were affected more by Methicillin resistant staph aureus infections as compared to females. (Figure: 1). 59% of the MRSA isolates (n=19) obtained were in the age range of 21-60 years, as shown in figure 2.

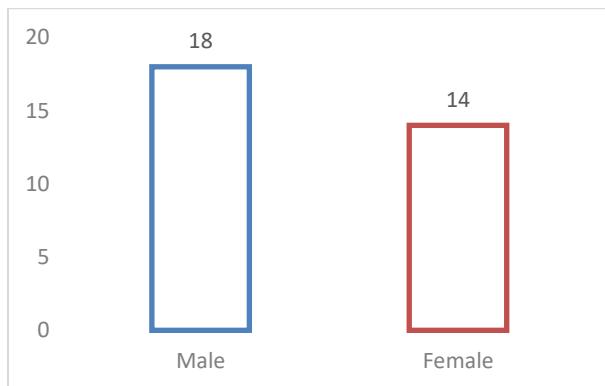


Figure 1. The demographic distribution, predominantly in terms of gender of patients affected by Methicillin-resistant *Staphylococcus aureus* (MRSA) infections.

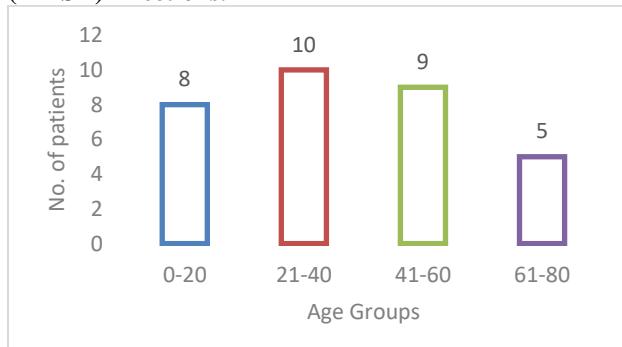


Figure 2: The demographic breakdown specifically in terms of age of patients with Methicillin-resistant *Staphylococcus aureus* (MRSA) infections.

Staphylococcus was identified in diverse specimens, with 78% of Methicillin-resistant *Staphylococcus aureus* (MRSA) being isolated from pus samples.

Based on the sensitivity pattern observed, all isolates obtained exhibited 100% resistance to beta-lactam drugs and combination drugs (including penicillin, cephalosporin, carbapenem, and amoxicillin-clavulanic acid). Conversely, Linezolid demonstrated 100% sensitivity, while Chloramphenicol showed 93% sensitivity. Fluoroquinolones and aminoglycosides displayed 50% sensitivity. Moreover, the minimum inhibitory concentration (MIC) of vancomycin for all isolates fell within the sensitive range refer to Figure 4.

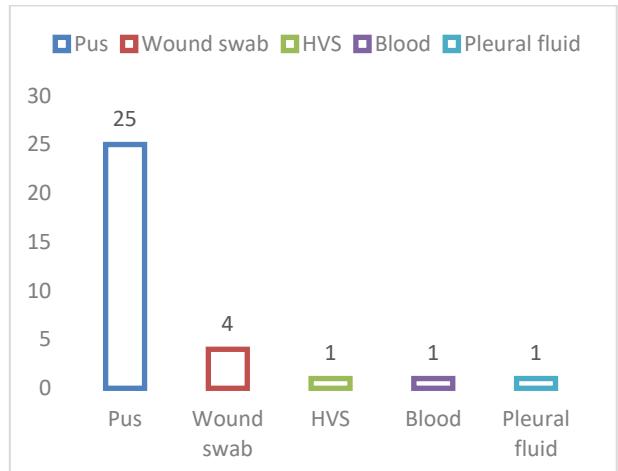


Figure 3. The distribution of Methicillin-resistant Staphylococcus aureus (MRSA) varies according to specimen type.

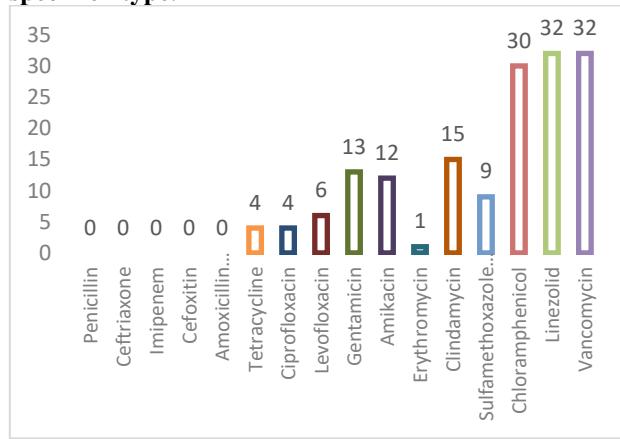


Figure 4: The susceptibility of Methicillin-resistant Staphylococcus aureus (MRSA) isolates to various antibiotics

Discussion

In this study, out of 65 Staphylococcus aureus isolates, almost half (49%) turned out to be MRSA. A study from Debre Markos Hospital reported similar magnitude of MRSA (49.7%), which is consistent with our study findings,⁽¹¹⁾ whereas, slightly lower prevalence was observed in studies from Moi Teaching and Referral

Hospital of Kenya (37%)⁽¹²⁾ South Africa (30.9%).⁽¹³⁾ Bangladesh (72%).⁽¹⁴⁾ Differences in the age groups, study subjects and specimen type could be attributed to the difference amongst various studies. Antibiotic misuse can also be associated with this substantial magnitude of MRSA.^(15, 16)

Males (56% vs 44%) were affected more by Methicillin resistant staph aureus infections as compared to females. This subgroup analysis is comparable with a meta-analysis reported from India (60.4% vs 39.6%), thereby representing MRSA infections more common in male gender.⁽¹⁷⁾ This is very similar to the findings of a work done by Harbath and colleagues and Kupfer et al.^(18, 19) Behavioral and physiological factors, (including hand hygiene, nose picking and nail-biting) and working with livestock industry are considered as the major contributory factors to the unusual occurrence of MRSA in male population.⁽²⁰⁾

Most of the MRSA isolates obtained (59%) in our study were falling in the age range of 21-60 years, consistent with a study conducted in Ethiopia. However, a rising trend has been observed in the prevalence of MRSA (25%) amongst population group of less than 20 years, as seen in a study by Wu et al, reporting MRSA in young Chinese population.⁽²¹⁾ The major source of MRSA in the current study was pus specimen (78%), followed by 13% of wound swabs as observed in a study by Tebelay Dilnessa et al.⁽²¹⁾

Looking at the susceptibility profile of the study isolates against anti-staphylococcal drugs, Penicillin, Cephalosporin, Carbapenem and Amoxicillin clavulanic acid were found to be 100% resistant. A similar picture was witnessed in India,⁽⁸⁾ Trinidad & Tobago.⁽²²⁾ Apart from high resistance to the reported β -lactams, trimethoprim/sulfa-methoxazole, clindamycin, erythromycin and gentamicin showed resistance of 72%, 53%, 97% and 59% respectively, which is higher in comparison to a systematic review from Tanzania showing 50% resistance in co-trimoxazole, 45% in clindamycin, 37% in erythromycin and 32% in gentamicin. Considering the fluoroquinolones, 81% resistance was observed against levofloxacin as compared to ciprofloxacin that showed 87% resistance. This is in concordance with a study from Assam, India showing 87.5% resistance to ciprofloxacin, and a study from Nigera showing 78.6% of the isolates showing resistance to the fluoroquinolone antibiotic ciprofloxacin.⁽²³⁾

Almost negligible resistance was observed against chloramphenicol (6.2%). Linezolid and vancomycin showed 0% resistance amongst the MRSA study isolates, similar to the study conducted in Nepal.⁽²⁴⁾ This is in contrast to a study from Ethiopia showing 29.4 % vancomycin resistant in MRSA. Similar trends were observed in studies from Trinidad⁽²⁵⁾ Argentina, Brazil and Mexico.⁽²⁶⁾

Variable trend against vancomycin have been reported in previous studies, ranging from 0% in Ethiopia, Karachi and Uganda^(8,27) to 8 % in Iran, Malaysia and Nigeria,^(28,29) not comparable to our study. Studies from United States and Latin America reported increased resistance to other classes of antibiotics, thereby necessitating the use of vancomycin for these infectious agents.⁽³⁰⁾ Prudent use of drugs and constant surveillance susceptibility testing of MRSA against vancomycin has been stated as a remedy to deal with these organisms.⁽³¹⁻³³⁾

Conclusion

The study concluded that the prevalence of MRSA is on a rise. The most significant remedy is the wise use of anti-staphylococcal drugs. Prompt diagnosis and appropriate management of infected patients is the key to contain MRSA infections. Along with this adherence to the infection control practices also play a substantial role in this regard.

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