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Antibiotic Resistance Pattern of Hospital Isolates of *Staphylococcus aureus* in Mashhad-Iran During 2009 - 2011

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Background: *Staphylococcus aureus* is a major human pathogen. Due to high prevalence of *S. aureus* infections and increasing resistance to antibiotics, physicians have been facing problems in choosing the appropriate empirical antibiotic therapy for such infections. **Objectives:** The aim of this study was to determine the pattern of antibiotic resistance in *S. aureus* clinical isolates of Mashhad Quaem Hospital between 2009 and 2011.

Materials and Methods: In this study 170 isolates of *Staphylococcus aureus* were identified in laboratory. The specimens (including 82 urine, 43 wound, 37blood, 8 stool samples) were collected and examined by standard diagnostic methods. Determination of *S. aureus* sensitivity to antibiotics was performed using standard disc diffusion method. The antibiotic sensitivity of bacteria were reported according to the clinical laboratory standards institute (CLSI) manual as sensitive (S), intermediate (I) and resistant (R).

Results: Among tested antibiotics, *Staphylococci* isolates were highly resistant against Ceftazidime (94%), followed by Penicillin (91%), Ampicillin (82%), Cefotaxime (65%), Erythromycin (60%), and Oxacillin (43%). Nearly all strains were susceptible to tested Vancomycin. **Conclusions:** Our results are similar to the reports from other parts of Iran. According to this study, resistance pattern among *Staphylococcus*

aureus strains were widespread in Quaem hospital. The implication of this high resistance is that *Staphylococcus aureus* infections should be treated more consciously and not with Penicillin and other ineffective antibiotics.

Keywords: Antibiotics, Antineoplastic Agents; Drug Resistance; Staphylococcus aureus

1. Background

Staphylococcus aureus is a bacterium that colonizes both in the community and hospital settings (1). Globally, S. aureus is one of the principal microorganisms involved in nosocomial infections. Indubitably, the fact that makes this pathogen a cure-challenging germ is its innate capacities to endure hard conditions such as high salt concentrations, high temperatures, extremes pH, survival in the presence of many antibiotics and disinfectants and even in macrophages (2). The prevalence of Staphylococcus aureus resistance to conventional antibiotics has been highly increasing in some hospitals. The use of wide spectrum of antibiotics in the hospital environment, leads to the development of increased resistance to these antimicrobial agents. This increasing prevalence of bacterial resistance to antibiotics, are mostly associated with the extensive use of antimicrobial agents, may result in insufficiency of the selected substances to combat some bacterial infections (1, 3).

The feature which makes S. aureus a distinct pathogen is its competence in acquiring resistance against new drugs. This ability is mainly due to its plasmids, which carry an extensive range of drug-resistant genes (4). In Staphylococci, after inactivation of Penicillin by penicillinase, an enzyme that is coded by plasmid, methicillin will be inactivated. Resistanceto this semi-synthetic β-lactamase resistant Penicillin was reported in 1961 in London (5). Many studies have been conducted since then and suggested a steady increase in the prevalence of Methicillin Resistant Staphylococcus aureus (MRSA) from late 1970s until now. They suggested that MRSA is increasingly causing infection in hospitals and community with higher mortality and morbidity than infections due to MSSA (4-6). In 1997, decreased susceptibility to Vancomycin, the drug of last resort against MRSA

Implication for health policy/practice/research/medical education:

This paper recommends physicians to pay close attention to how to prescribe the right antibiotic for the treatment of *Staphylococcus aureus* infection. Copyright © 2012, Shahid Beheshti University of Medical Sciencces; This is an Open Access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/3.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited. infections, was reported followed by full resistance in 2002 (3, 7, 8). In the word of increasing resistance of *Staphylococcus aureus* to antibiotics, especially to routine ones, physicians have faced problems in treating *Staphylococcus aureus* infections.

2. Objectives

The purpose of the study was to determine the antibiotic susceptibility pattern in *Staphylococcus aureus* isolates in Quaem hospital of Mashhad, from 2009 to 2011.

3. Materials and Methods

From September 2009 to March 2011. a total number of 2527 samples were collected from patients attending to Quaem hospital, Mashhad. Among these samples (urine, blood, wound and stool) 170 were identified as Staphylococcus aureus infected samples. For bacterial isolation and identification, the specimens were cultured according to standard methods. Culture media were incubated at 37°C for 18 hours under sterile circumstances. The isolates were identified using motility test, colony morphology, Gram staining, and biochemical tests such as catalase test, coagulase test, sensitivity to novobiocin, mannitol fermentation, and DNase tests. In order to determine the antimicrobial resistance pattern of isolates, we used 0.5 McFarland suspensions of bacteria on Mueller-Hinton's media. Disc diffusion susceptibility test was performed using seven antibiotic discs (i.e. Vancomycin, Ampicillin, Cefotaxime, Erythromycin, Penicillin, Oxacillin and Ceftazidime). The results reported after 18 hours incubation at 37°C according to the Clinical Laboratory Standards Institute (CLSI) (9).

4. Results

In this study *Staphylococcus aureus* was ranked as the second cause of hospital infections. Out Of 2527 clinical samples, 170 (6.7%) were positive for *Staphylococcus aureus*. The total of 2527 specimens was obtained from different parts of the body. Most of isolated *S. aureus* was collected from wound (12.9%) while the least was from urine (5.6%) (Table 1).

Table 1. Occurrence of S. aureus from Different Clinical Samples					
Source	Number	S. aureus Isolated Numbers, No. (%)			
Urine	1457	82 (5.6%)			
Blood	661	37 (5.6%)			
Wound	334	43 (12.9%)			
Stool	75	8 (10.7%)			
Total	2527	170 (6.7%)			

1457 urine, 661 blood, 334 wound and 75 stool specimens were cultured. *Staphylococcus aureus* were isolated from 82 urine specimens (5.6%), 37 blood specimens (5.6%), 43 wound specimens (12.9%) and 8 stool specimens (10.7%). Most of isolates detected as *S. aureus* were collected from urine (n = 82) and the least number were isolated from stool samples (n = 8). The degree of sensitivity of *S. aureus* isolates to antibiotics, following the disc diffusion method was analyzed according to CLSI 2007 (9) (Table 2).

Table 2. Sensitivity of S. aureus Isolates to Different Antibiotics Using Disc Diffusion Method							
Oxacillin	Erythromycin	Cefotaxime	Ampicillin	Penicillin	Ceftazidime		
46%	43%	39%	38%	7%	5%		



Figure 1. Resistance Percentage of S. aureus Isolates Against Various Antibiotics

5. Discussion

S. aureus is a major cause of hospital infections, which has been proved by many other studies (10, 11). Over past

decades, the frequency of antimicrobial resistance and its association with serious infectious disease has been a global public health concern. Antimicrobial resistance resulted in increases in the number of illnesses, death and health-care costs (12). In this study, out of 2527 analyzed samples, 170 (6.7%) were positive for *Staphylococcus aureus* with the most occurrences in wound (12.9%), followed by stool specimens (10.7%) and the least amount were reported in urine samples (5.6%). This is in line with previous reports in which *S. aureus* was the most common organism isolated from wound samples (13-15). The results of this study indicated that *S. aureus* isolates showed the least resistance against Oxacillin (43%) comparing other tested antibiotics. This suggests that Oxacillin can be an appropriate drug for *S. aureus* infections treatment. On the other hand, in another study done in Asia, high levels of Oxacillin resistance (82.1%) were reported (16). Similarly, Pourakbari et al. reported 79% resistance of *S. aureus* against Oxacillin in Tehran, Iran (17).

We expected higher Penicillin resistance rate in our isolates because other studies show more Penicillin resistance compared to our study (14, 18, 19). 60% Erythromycin resistance was reported in this study, which is similar to another study conducted in Mashhad with 52.3% resistance to Erythromycin (19). Furthermore, Pai et al. reported 45.9% Erythromycin resistance (18). This was in accordance with the Indian study in which Ahmed et al. recorded 74% resistance of MRSA isolates to Erythromycin (20). Although disk diffusion method is unable to generate the minimum inhibitory concentration (MIC) value (i.e. not quantitative as is the main disadvantage of this method), this method is a relatively easy and has been currently standardized in clinical settings by CLSI. In this study, we recorded 82% and 65% resistance of S. aureus isolates to Ampicillin and Cefotaxime, respectively. Nevertheless Naderinasab et al. reported 97.7% Ampicillin resistance and 51.2% Cefotaxime resistance in Imam Reza and Quaem hospitals in Mashhad (19). In conclusion, S. arouses is the major cause of nosocomial infections and the prevalence of antibiotic resistance among isolates is very high among patients referred to Quaem hospital, and this issue should be taken into consideration.

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Authors' Contribution

All authors contributed equally to this manuscript.

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The authors declare that there are no conflicts of interest.

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