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Prevalence of Multi-Drug Resistant Bacteria Isolated from Patients Admitted to the Intensive Care Unit in Tehran, Iran: a Five-Year Prospective Study- 2019-2023

Ali Ahmadi¹, Soghra Khani², Abas Ali Imani Fooladi^{1*}

¹ Molecular Biology Research Center, Biomedicine Technologies Institute, Baqiyatallah University of Medical Sciences, Tehran, Iran.

² Department of Biochemistry, Pasteur institute of Iran, Tehran, Iran.

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*Corresponding Authors: Abas Ali Imani Fooladi, Applied Microbiology Research Center, Biomedicine Technologies Institute, Baqiyatallah University of Medical Sciences, Tehran, Iran. Tel: +98-21-88055925, E-mail: imanofouladi.a@gmail.com.

ABSTRACT

Background: The study aims to identify bacterial agents and study the prevalence of pathogenic bacteria isolated from patients admitted to the Intensive Care Unit, as well as recognize factors affecting nosocomial infection that can lead to control and treatment of these infections.

Methods: This study was conducted in the Intensive Care Unit of Baqiyatallah Hospital in Tehran. Totally 738 samples were obtained from patients with a bacterial infection were included in this study. Patient information, including age, gender, type of clinical sample, susceptibility testing, year of hospitalization in to the Intensive Care Unit, and the prevalence of isolated bacteria, was analyzed.

Results: The findings of this study revealed that *Acinetobacter*, *Pseudomonas aeruginosa*, and *Staphylococcus aureus* had the highest and *Escherichia coli*, *Enterobacter*, and *Streptococci* had the lowest prevalence in the to the Intensive Care Unit isolates. Gram-negative bacteria were responsible for most nosocomial infections in the Intensive Care Unit. The rate of nosocomial infections in the Intensive Care Unit had a constant incidence in the recorded years. The rate of nosocomial infections was higher in the male patients than female, with a higher average year of age. Also, the results showed that most isolates were multiple drug resistant.

Conclusion: The high rate of nosocomial infections in to the Intensive Care Unit belonged to respiratory infections. Factors such as intubation and elderly age are associated with the type of infection. Bacterial isolates had a high level of resistance to most antibiotics. Therefore, periodic antibacterial sensitivity assessment could help optimize empirical antibiotic therapy against to the Intensive Care Unit acquired infections.

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Introduction

According to World Health Organization, the rate of Nosocomial infections (NIs) is between 5.7%-19.1% in low- and middle-income countries and 3.5%-12% in 3.5% to 12% in developed countries (1]. Although less than 10% of total hospitalized patients are admitted to the Intensive Care Unit (ICUs), between 20% to 45% of all nosocomial infections occur in the ICU (2, 3). Since patients admitted to ICUs suffer most often from severe underlying diseases, they are more susceptible to NIs caused by ICU- resident opportunistic bacterial pathogens. On the other hand, the prevalence of MDR pathogens such as Extended-spectrum beta-lactamases produced *Acinetobacter*, and *Pseudomonas* in the ICUs and their antibiotic resistance patterns are significantly higher than that of other sections of hospitals (in which there are various resistance reports) (4-6). They are often resistant to most clinically used antibiotics and cause higher treatment failure rates. Therefore, they must pay more attention to the types of ICU resident pathogens and their resistance rates to help physicians to make better therapeutic decisions (7-9). According to the reports, although the overall rate of NIs in Iran is estimated at 15-20 %, there are a few reports on their microbial diversity and resistance rates, the age, gender, and clinical source of the patients, year and ward of isolation, and the frequency of poly bacterial patterns. Considering the importance of microbial resistance in the intensive care unit, it seems necessary to investigate the common causative agents and their microbial resistance. In this study, we aimed to evaluate, in a trendy manner, the prevalence of ICU pathogens, NI and antibiotic resistance patterns in ICU-isolated

nosocomial pathogens at Baqiyatallah Hospital in Iran.

Materials and Methods

Patients and clinical samples

This study was done in Baqiyatallah hospital ICUs in Tehran and assessed the prevalence of nosocomial infections and pattern of antibiotic resistance in all patients admitted to the hospital ICUs between 2019 and 2023. The protocol of this study was approved by the ethics committee of the Baqiyatallah University. According to the standard definition, nosocomial infection is an infection that involves patients at least 48 hours after hospitalization (10). For ICU patients, clinical samples were regularly obtained, including associate, bronchial aspirate, tracheal aspirate, pleural fluid, blood, CSF, and wound. The patients whose collected samples yielded the growth of organisms were then identified. The following variable, such as age, sex, time of admission, ward, and site of infection were investigated. Procedures performed according to the ethical standards of the responsible committee on human experimentation (institutional and national) and with the Helsinki Declaration of 1975, as revised in 2000. Informed consent was obtained from all patients for being included in the research.

Bacterial isolation and identification

All collected samples were inoculated on the various microbiological media and incubated at 37°C for 24-48 hours. The identification of the isolates was performed by colony characteristics and biochemical tests.

Antibacterial susceptibility testing

Antibacterial susceptibility testing was performed by Kirby-Bauer disk diffusion of the Muller Hinton Agar medium according to CLSI guidelines (11).

Statistical analysis

Data have been analyzed by SPSS version 17. Data were presented in number and percentage of each variant using tables and charts.

Results

During the study period, from 738 hospitalized patients in the ICUs, a total of 914 non-duplicated pathogenic bacteria were isolated. Overall, the prevalence of pathogenic bacteria was slightly higher among males (52%) than females. The prevalence had a stable rate among males and females during the study period and the average age of the patients was 53 years old. More than 50% of the patients were over 50 years old. Most of the NI patients in ICUs were elderly males. The rate of nosocomial infections in the ICUs had a constant incidence in the recorded years. The respiratory system was the most common site of nosocomial infections, the bronchoalveolar lavage was the most frequent sample, and the blood was the least (Table 1).

Table 2 reveals the prevalence of bacteria responsible for nosocomial infections. *Acinetobacter* spp (50%) and *P. aeruginosa* (18.9%) had the highest frequency rates, followed by *S. aureus* (9.8%), *Enterococcus* (6.8%), *Klebsiella pneumoniae* (4.6%) and *Enterobacter* (4.1%). The relationship between the isolates and different samples was investigated. According to the results presented in Table 2, the distribution of isolates varies among different samples. Because of the complexity, multiple pathogens were also isolated from some ICU patients. Among the total cases of nosocomial infections, 63.4% were poly microbial, and 36.5% were monobacterial (Table 3). The results showed that although Gram-negative bacteria were predominant only for polybacterial cases, Streptococci ssp. was most frequent (77.7%). The most common bacteria found in poly bacterial infections were included *Acinetobacter*, *P. aeruginosa*, and *S. aureus*. Also,

the most abundant strain isolated each year is *Acinetobacter*, followed by *P. aeruginosa*.

The most predominant polybacterial rate was seen for Ascites (53.3%), followed by pleural fluid (44.4%), wound (36.6%) and bronchoalveolar fluid (33.9%) specimens. The Ascite and bronchial samples were found to have most poly bacterial infections rates. The rate of NI of the total ICU admissions in the ICU patients was 25%. The results showed that most of the isolates were Multiple drug resistant, and the resistance pattern is seen for each year in tables 4. The average rates of antibiotic resistance during 2019-2023 were 80, 93, 117, 139, and 192, respectively. Based on the obtained findings of this study, a trend of decreasing antibiotic susceptibility was shown in isolated strains during the five years.

Discussion

This study presents a profile of pathogenic bacterial strains isolated from patients in the ICUs of one hospital over five years. Our research aimed at determining the prevalence rate of different bacterial isolates, the frequent resistance patterns, and evaluating polybacterial infections. According to WHO, the rate of NI in developed, and developing countries is 5-15 and >25%, respectively. In addition, the prevalence of NI caused by MDR bacteria is 1-5% (12). In this study, the rate of NI in ICUs was 25% without any remarkable change during the five years. Consistent with studies by Shehabi et al., Amini et al., and Jean et al., *Pseudomonas*, *Acinetobacter*, and *S. aureus* were the most prevalent bacteria causing NIs in this study. (13-15). Unlike our study, in another study in ShahreKord, *Klebsiella* and coagulase-negative Staphylococci were more prevalent in ICU patients (16).

Table 1. Frequency (%) of clinical specimens in ICU wards during 2019-2023.

Year Specimen	2019	2020	2021	2022	2023	Total
Ascite	1(0.10)	0	5(0.54)	6(0.66)	3(0.32)	15(1.6)
BAL	134(14.6)	96(10.50)	129(14.1)	30(3.3)	24(2.6)	413(45.2)
Pleural fluid	2(0.22)	0	5(0.54)	0	2(0.22)	9(0.98)
Bronchial	16(1.7)	5(0.54)	5(0.54)	1(0.10)	0	27(2.9)
Trachea	45(4.9)	13(1.43)	44(4.8)	96(10.50)	137(14.98)	335(36.6)
Blood	2(0.22)	0	3(0.32)	0	0	5(0.54)
wound	17(1.8)	9(0.98)	32(3.51)	18(1.98)	25(2.74)	101(11)
CSF	3(0.32)	0	3(0.32)	1(0.10)	2(0.22)	9(0.98)
Total	220(24.07)	123(13.4)	226(24.7)	152(16.6)	193(21.1)	914(100)

Table 2. Frequency (%) of isolates according to the specimen types.

Specimen Bacteria isolated	Ascites	BAL	Pleural fluid	Bronchial	Trachea	Blood	Wound	CSF
<i>Acinetobacter</i>	0	221(24.1)	4(0.4)	12(1.3)	182(19.9)	1(0.1)	30(3.28)	9(0.98)
<i>S. aureus</i>	0	52(5.68)	1(0.1)	2(0.22)	27(2.95)	1(0.1)	7(0.77)	0
<i>Klebsiella</i>	0	8(0.87)	0	1(0.1)	23(2.5)	0	10(1.1)	0
<i>P. aeruginosa</i>	2(0.22)	89(9.73)	1(0.1)	8(0.87)	65(7.1)	0	13(1.4)	0
<i>Streptococcus</i>	4(0.44)	7(0.77)	1(0.1)	0	9(0.98)	0	15(1.6)	0
<i>Enterococcus</i>	8(0.87)	15(1.6)	2(0.2)	2(0.22)	18(1.97)	2(0.22)	16(1.7)	0
<i>E. coli</i>	0	3(0.3)	0	0	1(0.10)	0	4(0.44)	0
<i>Enterobacter</i>	1(0.10)	18(1.9)	0	2(0.22)	10(1.09)	1(0.1)	6(0.66)	9(0.98)

Table 3. Poly bacterial pattern frequency (%) according to the year.

Year Bacterial pattern	2019	2020	2021	Total	2022	2023	Total
One bacterial	129(14.1%)	77(8.4%)	149(16.3%)	580(63.4%)	97(10.6%)	128(14%)	580(63.4%)
Two bacterial	82(8.9%)	34(3.7%)	74(8.09%)	280(30.6%)	40(4.38%)	50(5.5%)	280(30.6%)
Three bacterial	9(0.98%)	12(1.3%)	3(0.32%)	54(5.9%)	15(1.62%)	15(1.6%)	54(5.9%)

Table 4. Resistance Rate (%) of bacterial isolates.

	DC ¹	E ²	CAZ ³	C ⁴	TE ⁵	D ⁶	AT ⁷	CZ ⁸	CRO ⁹	AMX ¹⁰	P ¹¹	SXT ¹²	V ¹³	AK ¹⁴	IMP ¹⁵	TZP ¹⁶	CP ¹⁷	MRO ¹⁸	GM ¹⁹
<i>Acinetobacter</i>	70	70	70	70	70	70	70	70	70	70	70	69	70	68	69	70	70	69	69
<i>P. aeruginosa</i>	70	70	70	70	70	70	70	70	70	70	70	70	68	67	68	66	68	70	70
<i>S. aureus</i>	70	70	70	70	70	70	70	70	70	70	70	53	10	66	68	62	70	70	70
<i>Klebsiella</i>	70	70	70	70	70	70	70	70	70	70	70	70	70	70	60	70	65	65	70
<i>Enterococcus</i>	70	70	70	70	70	70	70	70	70	70	70	70	55	70	70	55	70	70	70
<i>Enterobacter</i>	70	70	70	70	70	70	70	70	70	70	70	70	64	70	64	64	70	66	70
<i>Streptococcus</i>	70	70	70	70	70	70	70	70	70	70	65	70	50	70	70	60	70	70	70
<i>E. coli</i>	70	70	70	70	70	70	70	70	70	70	70	70	70	53	53	30	70	70	70

Although *E. coli* is an important NI pathogen in ICU, the prevalence of *E. coli* in this study was low, possibly because there were no urine samples in our research and the fact that *E. coli* are especially important for urinary tract infections.

Other studies have reported different prevalence rates for *P. aeruginosa* (36% in India; 33% in Brazil), *Klebsiella* (20% in India; 12% in Brazil), and *Acinetobacter* (17% in Brazil) (17-20). Polybacterial infections are more challenging to treat and impose heavier costs on the healthcare system and patients (21). We found a high rate of polybacterial infections in our study. The most prevalent bacteria, including *Acinetobacter* and *Pseudomonas*, were MDR as being resistant to different classes of antibiotics. As other studies have implied, *Acinetobacter*, *P. aeruginosa*, *Enterobacter*, *Klebsiella*, and *E. coli* were the most prevalent Gram-negatives, and *S. aureus*, Streptococci, and Enterococci were the most common Gram-positive bacteria isolated from poly bacterial infections (22). Similar studies have

shown that polybacterial infections predominantly involve Gram-negative bacilli rather than Gram-positive or anaerobic bacteria (23-26). In this study, most pathogens were isolated from BAL, trachea, and wound specimens. Instead, plural aspirate, sputum, and ascites were less frequent. In a study by Harbarth and colleagues, surgical wound infections were the most common nosocomial infections (30%), followed by urinary tract infections, pneumonia, and bloodstream infection (27). In a study by Poor and colleagues, the highest incidence of nosocomial infections belonged mainly to urinary tract infections and hospital pneumonia (28). In our study, the incidence of nosocomial infections was higher at extreme ages, similar to two other domestic studies in Tehran (29) and Sanandaj (30). Also, similar to other studies, most MDR in this study was seen in patients older than 50 years old (31). Continuous monitoring of antibiotic resistance in bacteria is a routine surveillance program in many countries (32). In a study between 2002 upto 2007 in the US,

out of 29723 Gram-negative bacteria, 6644 MDRs were isolated (33). Also, a study of ICU patients in China indicated a high rate of antibiotic resistance in Gram-negative bacteria. They did not find a significant relationship between sex and the rate of nosocomial infections in ICU patients (34). In the present study, there was no relationship between sex and MDR infections. Antimicrobial administration in ICUs is ten times higher than other wards of the hospital (35-37), demonstrating a major reason for such a high level of resistance in ICUs. Our results indicated that there was no constant or certain time trend in the resistance rate of the ICU bacteria in our hospital from 2019 to 2024.

Conclusion

In conclusion, the etiology of ICU infections is polymicrobial, and MDR Gram-negative bacteria are the most predominant isolates of ICU patients. There was not a constant time trend in the resistance rate of the bacteria during five years. The rate of NI in the present study was almost the same as in other developing countries.

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Ethics approval and consent to participate

Not needed.

Conflict of interest

All of authors declare that they have no conflict of interest.

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