

Original Article

Antimicrobial Resistance Pattern of Bacterial Isolates from Intensive Care Unit of a Tertiary Care Hospital in Bangladesh

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Abstract

Infections with resistant strain are one of the leading causes of morbidity and mortality in hospitalized patients especially the critically ill patients in intensive care unit (ICU). The present study was conducted to know the changing pattern of antibiotic resistance of the common isolates from blood, urine, respiratory secretions and pus/ wound swab of patients admitted in ICU at Holy Family Red Crescent Medical College hospital over a two years period from January 2012 to December 2013. A total of 1282 samples were analyzed. Out of which 301 (23.5%) samples were positive for growth of organisms. The predominant isolates were E.coli (28%), Klebsiella (27%), Acinetobacter (17.3%), Pseudomonas (9.6%), Staphylococcus aureus (5.3%) & candida (6%). Majority isolates were resistant to Cephalosporin and Amoxicillin, (>70-90%). Carbapenem were still highly active against E.coli. However, Klebsiella, Pseudomonas and Acinetobacter were showing alarming resistance (55-60%) towards those drugs. Aminoglycosides resistance has also increased for Pseudomonas and Klebsiella. The percentage of Cloxacillin resistant for staphylococcus was near about 40%. The most active drug against Pseudomonas was piperacillin/Tazobactam. The positive result from this study was that organisms showed decreased resistance towards Tetracycline, Cotrimoxazole and Ciprofloxacin. The sensitivity of Nitrofurantoin for urinary isolates was good. Regular surveillance of antibiotic susceptibility pattern & judicious use of antibiotics is very important for reducing the infection rate and antimicrobial resistance.

Key Words: Intensive care units, antimicrobial resistance pattern.

Introduction:

Infections with resistant strain are one of the leading cause of morbidity and mortality in hospitalized patients especially the critically ill patients in intensive care unit (ICU)¹. Patients admitted in ICU have an increased susceptibility to infection because of increased morbidity, decreased mobility and increased use of invasive devices². The therapeutic interventions which are associated with infectious complications include indwelling catheters, sophisticated life support, intravenous fluid therapy, prosthetic devices, immunosuppressive therapy and use of broad spectrum antibiotics leading to a spectrum of multi-drug resistant

pathogens, which contributed to the evolution of the problem of infections³.

The patterns of organisms causing infections and their antibiotic resistance pattern vary widely from one country to another, as well as from one hospital to other⁴. In addition, there is also variation in the frequency and types of infections among different subsets of patients within the same ICU⁵. In a study in an Indian ICU, the most common organisms were *Escherichia coli*, *Proteus sp.*, *Pseudomonas aeruginosa*, *Candida albicans*, *Staphylococcus aureus*, *Klebsiella sp.*, etc⁶. However, in an European ICU, *Staphylococcus aureus* was found as the most frequently isolated organism (30.1%) followed by *Pseudomonas aeruginosa* (28.7%), coagulase negative *Staphylococcus* (19.1%) and yeast (17.1%)⁷.

To formulate an antibiotic policy for the ICU, knowledge of the antibiotic susceptibility of the organisms isolated in the ICU is essential. This also avoids unnecessary use of broad spectrum antibiotics and prevents emergence of drug resistant

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bacterial strains. The data on the changing antibiotic susceptibility trends is important for infection control activities in ICU settings. Therefore, the present study was undertaken to determine the antimicrobial resistance pattern of bacterial isolates from Holy Family Red Crescent Medical College & Hospital over a 2 year period. The results of this study will help clinicians to plan the antibiotic guidelines as well as antibiotic cycling in ICU settings.

Material and Methods

A retrospective record based study was carried out based on data of bacterial isolates from the ICU of Holy Family Red Crescent Medical College & Hospital.

Study period: All the samples obtained for culture and sensitivity tests from patients admitted in the ICU during January, 2012 to December, 2013 were included in the study.

Study samples: Samples included blood, urine, sputum/tracheal aspirate (respiratory secretions) and pus/wound swabs. The samples were cultured in standard media for isolation of potential pathogen. Isolates were identified by standard microbiologic methods and then susceptibility testing was performed by Kirby Bauer disk diffusion method. Isolates with intermediate susceptibility were considered resistant.

Result:

In the 2 year study period, total 1282 samples were analyzed. The cumulative isolates of all samples together were 301 (23.5%). Out of these, 83 isolates (27.6%) were from respiratory tract; 83 (27.6%) isolates from blood; 129 (42.9%) isolates from urine; 6 (2%) isolates from pus/ wound swab as shown in Table-I.

The major organism isolated in a 2 year study period were *Escherichia coli* 84(28%) followed by *Klebsiella sp.* 81(27%), *Acinetobacter sp.* 52(17.3%), *Pseudomonas sp.* 29(9.6%), *Candida* 18(6%) and *Staphylococcus aureus* 16(5.3%). The numbers of percentages of *Klebsiella* and *Staphylococcus aureus* have markedly increased from 17.8% & 1.9% in 2012 to 37.1% & 7.2% in 2013 respectively. *Acinetobacter* was the most commonly isolated organism from blood 30 (36.1%), while *E. coli* was the most frequently isolated organism from urine, 57 (44.1%). The most common organism from respiratory tract was *Klebsiella*, 44 (53%). (Table-II)

Antibiotic resistance pattern of major five bacterial isolates is shown in Table-III. Majority isolates were resistant to Cephalosporin and Amoxicillin, (>70-90%). Carbapenems are still highly active against *E.coli*. However, *Klebsiella*, *Pseudomonas* and *Acinetobacter* are showing alarming resistance (55-60%) towards those drugs. The resistance

patterns of Gentamycin & Amikacin have increased for *Pseudomonas* (37% to 67% & 25% to 52% respectively) and for *Klebsiella* (52% to 62% & 26% to 58% respectively) during this study period. For *E. coli* & *Acinetobacter*; Gentamycin resistance was stable; however, Amikacin resistance has slightly increased from 32% to 45% & 48% to 60% respectively over the 2 years time. The percentage of Cloxacillin resistant for staphylococcus was near about 40%. As per sensitivity analysis, the most active drug against *Pseudomonas* was piperacillin/Tazobactam. Organisms were showing decreased resistance towards Tetracycline, Cotrimoxazole and Ciprofloxacin. The sensitivity of Nitrofurantoin for urinary isolates was good in this 2 year period.

Table-I: Distribution of isolates among samples

Sample	2012 N(%)	2013 N(%)	Total N(%)
Blood	27(25.2)	56(28.9)	83(27.6)
Respiratory tract	30(28)	53(27.3)	83(27.6)
Urine	49(45.8)	80(41.2)	129(42.8)
Pus/Wound swab	01(1)	05(2.6)	06(2)
Total	107	194	301

Figure within parentheses indicate percentage

Table-II: Pattern of organisms isolated from different samples between 2012 & 2013.

Organism most Frequently isolated	Samples	2012 N=107	2013 N=194	Total N=301
<i>E. coli</i> □	Respiratory tract □	5 □	2 □	7
	Blood □	3 □	15 □	18
	Urine	28	29	57
	Pus/Wound	1	1	2
<i>Klebsiella</i>	Respiratory tract	9	35	44
	Blood	3	16	19
	Urine	7	19	26
	Pus/Wound	-	2	2
<i>Acinetobacter</i>	Respiratory tract	12	13	15
	Blood	15	15	30
	Urine	-	7	7
	Pus/Wound	-	-	-
<i>Pseudomonas</i>	Respiratory tract	2	6	8
	Blood	2	7	9
	Urine	4	8	12
	Pus/Wound	-	-	-
<i>Staph aureus</i>	Respiratory tract	-	6	6
	Blood	-	-	-
	Urine	2	6	8
	Pus/Wound	-	-	-
<i>Candida</i>	Respiratory tract	-	-	-
	Blood	-	-	-
	Urine	8	10	18
	Pus/Wound	-	-	-
Others	Respiratory tract	2	1	3
	Blood	4	3	7
	Urine	-	1	1
	Pus/Wound	-	-	-

Table-III: Antibiotic resistance pattern of major organisms isolated from ICU in the year 2012 & 2013 (Numbers are percent resistant).

Antibiotics	E. coli		Klebsiella		Acinetobacter		Pseudomonas		Staph Aureus
	2012 N=37	2013 N=47	2012 N=19	2013 N=72	2012 N=27	2013 N=25	2012 N=8	2013 N=21	2013 N=14
Ceftriaxone	89.2	83	89.5	82	81.5	80	-	-	64.3
Ceftazidime	-	-	-	-	77.8		75	85.7	-
Cefuroxime	94.6	82.97	78.9	84.72	-	-	-	-	
Cephalexin	94.6	93.6	94.7	91.7	-	-	-	-	71.4
Ciprofloxacin	78.4	55.3	52.6	73.6	44.4	44	50	47.6	50
Cotrimoxazole	64.9	66	78.9	70.8	59.1	28	-	-	50
Amikacin	32.4	44.7	31.6	47.2	48.1	60	25	52.4	35.7
Gentamycin	59.5	57.4	52.6	62.5	66.4	68	37.5	66.7	42.8
Imipenem	13.6	21.3	26.3	58.3	40.7	56	12.5	62	43
Meropenem	13.6	19.1	21.1	56.9	37	40	12.5	60	-
Tetracycline	73	61.7	78.9	50	77.8	76	-	-	71.4
Amoxicillin	91.9	100	-	-	100	100	-	-	-
Amoxiclave	83.8	70.2	89.5	75	-	-	-	-	64.3
Cloxacillin	-	-	-	-	-	-	-	-	42.2
Nitrofurantoin (in urine)	21.4	34	14.3	26.4	-	-	-	-	28.6
Piper-Tazobac	-	-	-	-	-	-	-	24	-
Azithromycin	56.8	48.9	68.4	72.2	51.9	44	-	-	57.14

Discussion

During the study period, organisms were mostly isolated from Urinary tract 129 (42.9%) followed by respiratory tract and blood 83 (27.6%). These finding corroborated the results reported in a study from India⁶. Gram negative bacteria were isolated at a significant higher rate (98%) than Gram positive bacteria and candida. This observation is agreed with findings of Barai *et al* (2010)⁸.

In total, predominant organisms isolated from our ICU were *E.coli* 84 (28%), *Klebsiella sp.* 81 (27%), *Acinetobacter sp.* 52. (17.3%), *Pseudomonas sp.* 29 (9.6%), *Candida* 18 (6%) and *Staphylococcus aureus* 16(5.3%). These finding were comparable to the finding of Pattanayak *et al*(2013), Barai *et al* (2010) & Shalinis *et al*(2010)^{6,8,9}. The spectrum of pathogens in ICU may change from country to country with time and by hospital, type of ICU, and specific patient population^{10,11}. *Klebsiella pneumoniae* is one of the common causative organisms for respiratory and urinary tract infection¹². The microorganism can rapidly spread from the gastrointestinal tract of patient and via the hands of the hospital personal to other patients, leading to nosocomial outbreak¹³. In our study, the percentage of *Klebsiella* isolates

markedly increased from 17.8% to 37.11%. Moreover it was the most common isolates in respiratory tract. Similar finding was observed by Jain & Khety (2012)¹².

Acinetobacter sp. has recently advanced to one of the most common pathogens isolated from ICUs. In our study it was the most common isolates in blood and 2nd most common isolates from respiratory tract. A study conducted in ICU of Dhaka city by Haque *et al* (2013)¹⁴ reported that the most common pathogen from tracheal aspirates was *Acinetobacter*. Another study from Barai *et al*(2010)⁸ also found, *Acinetobacter* as the 2nd most common organism from all samples. In our study, the most common isolates from urinary samples were *E. coli* which corresponds to the finding of Pattanayak *et al* (2013)⁶.

Prolonged use of Carbapenem in the treatment of nosocomial infection can favor the development of resistance to those antimicrobial agents. In our study, Carbapenem resistant rate has alarmingly increased for *Klebsiella*, *Acinetobacter* and *Pseudomonas*. However, they are still highly active against *E.coli*. A study from Birdem hospital in 2010 also reported imipenem resistant *Acinetobacter* (>70%) and *Klebsiella* (>20%)⁸. Jain & Khety (2012)¹² also reported carbapenem

resistant pseudomonas. Emergence of Imipenem resistant strains in many parts of the world is alarming and a threat to the effective management of those organisms. Majority isolates were resistant to cephalosporin and amoxicillin (>70-90%). This might be due to the selective influence of extensive usage of cephalosporins and amoxicillin/ clavulanic acid. *Ganneja et al* (2011)¹⁵ also reported the similar findings.

According to *Sheth et al* (2012)¹⁶ piperacillin/ Tazobactam and Ceftazidime are ineffective for *Pseudomonas*. They discourage these drugs for use. However, in our study, as per sensitivity analysis the most active drugs against *Pseudomonas* were piperacillin/ Tazobactam. We can not compare it with previous year because it was not available. The positive result from this study showed that the resistance of Tetracycline, Cotrimoxazole and Ciprofloxacin was decreased. These drugs were not much used in the ICU settings that may be the reason of this decreased resistance. In our study the resistance patterns of Gentamycin & Amikacin have increased for *Pseudomonas* (37% to 67% & 25% to 52% respectively) and for *Klebsiella* (52% to 62% & 26% to 58% respectively) over the 2 years study period. For *E. coli* & *Acinetobacter*; Gentamycin resistance was stable; however Amikacin resistance has slightly increased from 32% to 45% & 48% to 60% respectively. A study from Ibrahim Medical College and Birdem; ICU resistance pattern of these organism to Aminoglycosides showed marked variability ranging from 48.3% to 83.3%⁸.

Nitrofurantoin is an urinary antiseptic. The sensitivity of Nitrofurantoin for urinary isolates was good and stable over this 2 year study period. Therefore, this may be the drug of choice for treatment of urinary isolates from ICU. Hooper explained this consistent and high level susceptibility of urinary isolates to nitrofurantoin may be due to nitrofurantoin's narrow spectrum of activity, limited indication (urinary tract infection), narrow tissue distribution (low or undetectable serum concentration) and limited contact with bacteria outside the urinary tract¹⁷.

The rate of isolation of *Staphylococcus aureus* markedly increased from 1.9% in 2012 to 7.2% in 2013. However, number of *Staphylococcus aureus* was decreased from 25% to 12% reported by Jain & Khety (2012)¹². The reason behind this was the good infection control practices. We can not compare the antibiotic resistance of *Staphylococcus aureus* between 2012 and 2013 as the number of *Staphylococcus aureus* isolates in 2012 was very few. Amikacin showed good susceptibility for *Staphylococcus aureus*. The percentage of Cloxacillin resistant for staphylococcus in our study was near

about 40%. Similar finding was observed by a tertiary care hospital in Bangladesh¹⁸.

Candida was isolated from 14% of urine culture, which may be due to the presence of underlying conditions like poor nutritional status, diabetes mellitus, use of steroid and broad spectrum antibiotics & presence of catheters. This may, however, also indicate an overuse of antibiotics and need for strict control measures.

Conclusion:

Antibiotic resistance continues to rise and complicates empirical selection of antibiotics in the ICU. Our data indicate an alarming pattern of antibiotic resistance in the majority of ICU isolates. Thus, strategies to prevent the emergence of multidrug resistant bacteria in ICU must be implemented. These include: knowledge of the infection rates and common pathogens and judicious use of older and newer antimicrobial agent according to antibiogram. This will assist clinicians in choosing appropriate empiric antibiotics to maximize the patient's chances of receiving early and effective therapy. It is important to use direct discussion and communication between clinicians and microbiologists for appropriate antibiotics based on antibiogram in every 6 month interval in the ICU. Strict infection control measures like good aseptic precautions, isolation of patients infected with resistant isolates and hand washing practices need to be reinforced for the same to prevent the occurrence of infection.

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