

Prevalence of microorganism and emergence of bacterial resistance in ICU of Bangabandhu Sheikh Mujib Medical University of Bangladesh

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Abstract

Background Antibiotic resistant bacterial nosocomial infections are a leading problem in intensive care units (ICU).

Objective To study the pattern of microorganism and bacterial resistant to antibiotic in ICU of Bangabandhu sheikh Mujib Medical University of Bangladesh.

Methods This retrospective study was conducted in ICU of Bangabandhu Sheikh Mujib Medical University, Bangladesh from January 2010 to December 2012. Total number of samples were 448. The samples of tracheal aspirate, blood and urine for culture and sensitivity was collected from the patient admitted in ICU. Analysis of tracheal aspirate, blood and urine culture was done from hospital record. All bacteria was identified by standard microbiological methods, and their antibiotic sensitivity was performed using disk diffusion method.

Results Total number of samples 448. Samples of tracheal aspirate was 159, positive culture 121(76%), most frequent identified organism was acinetobacter 45.45%, followed by pseudomonas 32.23%, proteus 11%, klebsiella 10% and E.coli 3%, samples of blood culture was 148, positive culture 22(14.86%), most frequent identified organism was pseudomonas 63.63%, followed by acinetobacter 22.72%, salmonella 4.54% and E.coli 4.54% and samples of urine culture was 141, positive culture 36 (25.53%) most frequent identified organism was enterococcus 22.22%, followed by acinetobacter 19.44%, candida 16.66%, klebsiella 13.88% and E.coli 13.88%. Drug resistant organism of tracheal aspirate was 12(20.33%) in 2010, 2(20%) in 2011 and 13(25%) in 2012. only collistin sensitive organism identified was 28(23.14%).

Conclusion From this study we concluded that most common site of infection was respiratory tract and most prevailing organism was acinetobacter & pseudomonas and antibiotic resistant infection is increasing and at present around one fourth organisms were resistant to all antibiotics.

Keywords Microorganism, drug resistance, respiratory tract, acinetobacter and pseudomonas aeruginosa.

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Introduction

Increasingly rapid emergence and dissemination of antimicrobial-resistant bacteria has become a world wide problem during the last decades. The intensive care unit often is called the epicenter of infections, due to its extremely vulnerable population of reduced host defences deregulating the immune responses and increased risk of

becoming infected through multiple procedures and use of invasive devices distorting the anatomical integrity-protective barriers of patients like-intubation, mechanical ventilation, vascular access, etc. In addition, several drugs may be administered, which also predispose for infections, such as pneumonia, e.g., by reducing the cough and swallow reflexes (sedatives, muscle relaxants)

or by distorting the normal nonpathogenic bacterial flora (e.g., stress ulcer prophylaxis)¹.

Consequently, the ICU population has one of the highest occurrence rates of nosocomial infections (20-30% of all ICU-admissions)^{2,3}, leading to an enormous impact on morbidity, hospital costs, and often, survival⁴⁻⁶. Along with the problem of nosocomial infection goes the burden of “multidrug” antimicrobial resistance (MDR). The ongoing emergence of resistance in the community and hospital is considered a major threat for public health. Antibiotic resistance increases the morbidity and mortality associated with infections and contributes substantially to rising costs of care, resulting from prolonged hospital stays and the need for more expensive drugs.

Both infection and MDR result in a considerable clinical and economic burden. As such, the presence of MDR boosts the deleterious impact of nosocomial infection⁷. Compared with infections not caused by MDR microorganisms, the additional cost of multidrug resistance in hospitalized patients with infections has been estimated at \$6,000 to \$30,000 (per patient) in Belgium⁸. This burden of resistance, however, is probably more due to the higher rate of inappropriate empiric antimicrobial treatment associated with infections caused by MDR pathogens than with the virulence of particular MDR strains⁹.

We are currently faced with (multi drug) resistant bacteria that are difficult and sometimes impossible to treat¹⁰. The tremendous therapeutic advantage afforded by antibiotics is being threatened by the emergence of increasingly resistant strains of microbes¹¹. The problem has recently been worsened by the steady increase in multi-resistant strains and by the restriction of antibiotic discovery and development programs¹⁰.

In hospital settings world-wide Intensive care Units are faced with rapid and increasing resistance of bacteria. ICUs are the source and site of development of multidrug resistant (MDR) bacteria.

The widespread use of antibiotics both inside and outside of medicine is playing a significant role in the emergence of resistant bacteria¹². Antimicrobials have transformed our ability to treat many infectious diseases that were killers only a few decades ago. The increasing use of antimicrobials in humans, animals, and agriculture has resulted in many pathogens developing

resistance to these powerful drugs¹³. A number of factors contribute to the emergence of antimicrobial resistance in ICUs, including extended length of hospital stay, and the widespread use of prophylactic and therapeutic anti-infective agents. Bacteria have developed resistance to all different classes of antibiotics discovered to date. The most frequent type of resistance is acquired and transmitted horizontally via the conjugation of a plasmid¹⁴. In recent times new mechanisms of resistance have resulted in the simultaneous development of resistance to several antibiotic classes creating very dangerous multidrugresistant (MDR) bacterial strains, some also known as “superbugs”¹⁵. The need for new antimicrobial agents is greater than ever because of the emergence of multidrug resistance in common pathogens, the rapid emergence of new infections, and the potential for use of multidrug-resistant agents in bioweapons¹⁶.

Controlling the spread of resistance requires the collaboration of several participants such as Medical, Veterinary and Public Health Communities¹⁷. Multidrug resistant organisms (MDROs) are resistant to one or more classes of antimicrobial agents and the knowledge of susceptibility pattern is helpful in selecting the empirical therapy and improving the likelihood of a satisfactory outcome for patient¹⁸. The objective of this study was to determine bacterial pathogens prevalence and to assess the multi-drug resistant (MDR) strains to different antibiotics in the intensive care unit of Bangabandhu Sheikh Mujib Medical University.

Materials

This retrospective study was carried out in the intensive care unit of Bangabandhu Sheikh Mujib Medical University, Dhaka, Bangladesh from 2010-2012. The approval of the University ethical committee was taken before conducting the study. Data of total 448 samples (tracheal aspirate 159, blood culture 148 and urine culture 141) for culture and sensitivity were collected from intensive care unit record. The pattern of microorganism and bacterial resistant to antibiotic was recorded. All bacteria was identified by standard microbiological methods, and their antibiotic sensitivity was performed using disk diffusion method.

Sample Collection: Tracheal aspirate, blood and urine specimens were collected aseptically for bacteriological examination from the patient of Intensive care unit of BSMMU.

Isolation and Identification

Tracheal aspirate, blood and urine specimens were cultured into blood agar and MacConkey agar media. Bacteriological smears were prepared from the growing colonies then stained with gram stain for morphological identification. All the bacterial isolates were preserved on nutrient agar slants at 4°C and subcultured periodically. The obtained pure cultures were identified biochemically.

Antimicrobial Susceptibility Test

Antimicrobial susceptibility pattern was performed using disk diffusion method on Muller Hinton agar plate. The isolates were tested against amoxicillin, cotrimoxazole, ceftazidime, Cefotaxime, Cefuroxime, Aztreonam, Netilmicin, Ticarcillin, Cephradine, gentamicin, imipenem, ciprofloxacin, ceftriaxone, amikacin, Colistin Sulphate, Tazobactam+ piperacillin.

The proportion of susceptible organisms was calculated as the sum of susceptible isolates relative to the total number of organisms tested. The organism considered as multidrug resistant if it is resistant to three or more antimicrobials.

Results

Table I Tracheal aspirate-2010, prevalence & pattern of organism, n=74

Positive c/s	Name of organism	Frequency	Percentage
59(79.7%)	Acenatobacter	32	54.2
	Pseudomonas	24	40.7
	Proteus	2	3.3
	Klebsiella	1	1.69

Table II Tracheal aspirate-2010, pattern of antibiotic sensitivity, n=59

Sensitive to none	Sensitive to one		Sensitive to many
Total	Antibiotic	Number	Total
12(20.3%)	Colistin	12(66.6%)	29(49.1%)
18(30.5%)	Tazobactam	2(11.1%)	
	Imepenem	3(16.6%)	
	Netilmicin	1(5.5%)	

Table III Blood-2010, prevalence & pattern of organism, n=61

Positive c/s	Name of organism	Frequency	Percentage
13(21.3%)	Pseudomonas	8	61.5
	Acenatobacter	3	23
	E coli	1	7.6
	Salmonella	1	7.6

Table IV Blood -2010, pattern of antibiotic sensitivity, n=13

Sensitive to none	Sensitive to one		Sensitive to many
Total	Antibiotic	Number	Total
1(7.6%)	Colistin	3(50%)	6(46.15%)
6	Ceftazidim	3(50%)	

Table V Urine -2010, pattern of antibiotic sensitivity, n=54

Positive c/s	Name of organism	Frequency	Percentage
19(35.1%)	Acenatobacter	6	31.5
	Enterococcus	5	26.3
	E coli	3	15.7
	Pseudomonas	2	10.5
	Klebsiella	2	10.5
	Candida	1	5.2

Table-VI Urine -2010. pattern of antibiotic sensitivity, n=19

Sensitive to none	Sensitive to one		Sensitive to many
Total	Antibiotic	Number	Total
5(26.3%)	Colistin	3(60%)	9(47.3%)
5(26.3%)	Imipenem	1(20%)	
	Ciprofloxacin	1(20%)	

Table-VII Tracheal aspirate-2011, prevalence & Pattern of Organism n=14

Positive c/s	Name of organism	Frequency	Percentage
10(71.4%)	Acenetobacter	5	50
	Proteus	2	20
	Pseudomonas	1	10
	Klebsiella	1	10
	Streptococcus	1	10

Table VIII Tracheal aspirate-2011, Pattern of antibiotic sensitivity, n=10

Sensitive to none	Sensitive to one		Sensitive to many
Total	Antibiotic	Number	Total
2(20%)	Colistin	4(100%)	4(40%)
4(40%)			

Table IX Blood-2011, prevalence & pattern of organism n=16

Positive c/s	Name of organism	Frequency	Percentage
3(18.7%)	Pseudomonas	2	66.6
	Salmonella	1	33.3

Table X Urine-2011, prevalence & pattern of organism n=14

Positive c/s	Name of organism	Frequency	Percentage
4(28.5%)	Candida	2	50
	Enterococcus	1	25
	Acinetobacter	1	25

Table XI Tracheal aspirate-2012, prevalence & pattern of organism n=71

Positive c/s	Name of organism	Frequency	Percentage
52(73.23%)	Acinetobacter	18	34.61
	Pseudomonas	14	26.92
	Klebsiella	8	15.38
	Proteus	7	13.46
	Enterococcus	1	1.92
	E. coli	3	5.76
	Gram +ve cocci	1	1.92

Table XII Tracheal aspirate-2012, Pattern of antibiotic sensitivity, n=52

Sensitive to none	Sensitive to one	Sensitive to many
13(25%)	Total Antibiotic	26(50%)
	13(25%) Colistin	12(92.7%)
	Tazobactam	1(7.7%)

Table XIII Blood-2012, prevalence & pattern of organism n=71

Positive c/s	Name of organism	Frequency	Percentage
6(8.45%)	Pseudomonas	4	66.6
	Acinetobacter	2	33.3

Table XIV Urine-2012, prevalence & pattern of organism n=73

Positive c/s	Name of organism	Frequency	Percentage
13(17.80%)	Pseudomonas	2	15.38
	Enterococcus	2	15.38
	E.coli	2	15.38
	Klebsiella	3	23.07
	Candida	3	23.07
	Proteus	1	7.06

Table XV Prevalence of positive culture of trachea, blood and urine- comparison during 2010-2012

Tracheal aspirate		Blood		Urine	
Total Sample	Positive c/s	Total Sample	Positive c/s	Total Sample	Positive c/s
159	121(80.66%)	148	22(14.86%)	141	36(25.53%)

Discussion

Surveys of the prevalence and antibacterial susceptibility patterns of bacterial isolates are important for determining appropriate empirical therapy for infections in critically-ill patients. Also, epidemiological analysis of patient data can be informative for appropriate management of patients in ICUs.

In our study most common site of infection is respiratory tract. Culture of 159 sample of tracheal aspirate was done in 2010 to 2012. Among them

121 culture was positive and most frequent organism *Acinetobacter* was isolated in maximum 55(45.45%) samples followed by *Pseudomonas* in 39 (32.23%) samples, *Proteus* in 11 (9%) samples, *Klebsiella* in 10(8.26%) samples, *E.Coli* in 3 (2.47%) samples and *Streptococcus* in 1(0.82%) sample. A very high rate 27(22.31%) of resistant organism was identified, sensitive to only one antibiotic was 35(28.92%) and sensitive to many antibiotic was 59(48.76%). Among the antibiotic collistin (polymyxin E) was only sensitive for 23.14%

organism, carbapenem 2.47% and tazobactam-piperacillin combination 2.47%.

In blood culture, among 148 samples 22(14.86%) samples were positive. Among identified organism pseudomonas was 14(63.63%), acenatobacter 5(22.72%), E. coli 1(4.54%) and salmonella 1(4.54%). In urine culture, among 141 samples- 36(25.53%) samples were positive. Among identified organism enterococcus was 8(22.22%), acenatobacter 7(19.44%), Candida 6(16.66%), E.coli 5(13.88%), klebsiella 5(13.88%) and pseudomonas 4(11.11%).

In Asian countries including Indonesia, the most frequent pathogen isolated from infections in the ICU are *P. aeruginosa*, *Klebsiella* spp., *E. coli*, *Enterococcus*, and *Staphylococcus aureus*. For example, in 12 ICUs in seven Indian cities, overall 87.5% of all *Staphylococcus aureus* health care associated infections were caused by methicillin-resistant strains, 71.4% of *Enterobacteriaceae* were resistant to ceftriaxone and 26.1% to piperacillintazobactam; 28.6% of the *P. aeruginosa* strains were resistant to ciprofloxacin, 64.9% to ceftazidime and 42.0% to imipenem¹⁹.

In Thailand the predominance causative pathogens in ICU, were the imipenem resistant *P. aeruginosa*, ceftazidime-resistant *Acinetobacter baumannii*, third-generation-cephalosporin-resistant *K. pneumoniae*, and quinolone-resistant *E. coli*²⁰. Another study performed at ICU of a tertiary care center in Saudi Arabia showed that the most frequent pathogens are *Acinetobacter baumannii*, *P. aeruginosa*, *E. coli*, *K. pneumoniae*²¹. Recently, similar studies were conducted in hospitals and several ICUs in Asian countries including Philippine²², India^{23,24,25,26} Iran^{27,28}, China²⁹, Malaysia³⁰, Singapore³¹, and Nepal³², demonstrated that the most frequent microorganism derived from ICU samples were *P. aeruginosa*, *Klebsiella* spp. and *Staphylococcus aureus*. In Canada, the Canadian National Intensive Care Unit study conducted during 2005-2006, showed that *P. aeruginosa*, *Staphylococcus aureus*, *Haemophilus influenzae*, *Enterococcus* spp., *Staphylococcus pneumoniae*, and *K. pneumoniae* are the most common isolates recovered from clinical specimens in Canadian ICUs. Moreover, *P. aeruginosa* is the most frequent multi drug-resistant phenotype, which is resistance to three or more of the antibiotics

including cefepime, piperacillin-tazobactam, meropenem, amikacin or gentamicin, and ciprofloxacin³³.

In our study showed that the most common pathogens identified from clinical specimen were *Acinetobacter baumannii*, *P. aeruginosa*, *E. coli*, *Klebsiella pneumoniae* and *acenetobacter* is the most frequent multidrug resistant phenotype and about 22.31% organisms were resistant to all available antibiotics in market. Ultimate fate of these patients is no need to mentioned. It's a grave situation. As we are being unable to treat many of these infections, so due important is to be given on prevention of infection and prevention of development antimicrobial resistance. Most sensitive antibiotic is polymixin- colistin. About one fourth organisms were resistant to carbanem which is also an alarming sign for us.

The prescribing of antibiotics in the ICU is usually empiric. Therefore, the ongoing surveillance of antibiotic susceptibility patterns of predominant bacteria is a fundamental effort to monitor changes in susceptibility patterns and to guide the clinician in choosing empirical or directed therapy appropriately, especially in ICU setting. Appropriate antibiotic utilization in ICU is crucial not only in ensuring an optimal outcome, but also in preventing the emergence of multi drug resistance bacteria. Therefore developing nation wide antibiotic policy and guidelines is essential to limit multidrug resistance and to maintain low level of resistance to newer antibiotics. Several societies have published guidelines for optimizing antibiotic use and curtailing antibiotic resistance in hospitals. Key components of these guidelines include multidisciplinary coordination between hospital administrators, clinicians, infectious disease specialists, infection control teams, microbiologist and hospital pharmacists.

In order to prevent the emergence and spread of antimicrobial resistance pathogens in ICU, the pattern of antimicrobial use has to be determined. A multidisciplinary approach is required to succeed in combating the problem. Hospital should monitor antimicrobial use to determine whether in ICUs or the entire hospital is overusing antimicrobials.

Introduction of strict and mandatory infection control program in hospital setting like proper hand washing before and after contact with

patients, appropriate isolation of patients, standard guideline for disinfection and sterilization of medical equipment should be introduced.

This study conclude that most common site of infection was respiratory tract. Most prevailing organism was acinetobacter & pseudomonas. Around one fourth organisms were resistant to all antibiotics. Around one fourth organisms are sensitive to only one antibiotic. Most sensitive antibiotic is polymixin- colistin. Around half of the organisms are sensitive to many antibiotics. Positive blood culture reports are lowest in comparison to others.

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