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## **Original Research Article**

# The study of antibiotic-sensitivity and resistance pattern of bacteria causing catheter associated urinary tract infection

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### ABSTRACT

**Background:** present study is done to study the antibiotic-sensitivity and resistance pattern of bacteria causing catheter associated urinary tract infection. Objectives of this study were to study the bacterial etiology of CAUTI, to study the prevalence of various bacteria causing catheter associated urinary tract infection, the antibiogram (sensitivity and resistance) pattern of isolated bacteria and the percentage of asymptomatic bacteriaria in the study population.

**Methods:** In this prospective observational study, under aseptic precautions, urine sample was taken after 48 hours of catheterization and sent for culture and sensitivity pattern is studied.

**Results:** In this study 500 urine samples were cultured and its antibiotic sensitivity pattern was observed. Out of the 53 culture positive samples most the subjects had asymptomatic bacteriuria. The study gave the incidence of catheter associated urinary tract infection (CAUTI) to be 10.6% and about 9% were polymicrobial. In this study about 7 causative bacteria were isolated. *Escherichia coli* were the most common organism that was isolated. On studying the antibiotic susceptibility pattern of each isolate, it has been observed that all of them are multidrug resistant and the sensitivity pattern is migrating towards higher antibiotics.

**Conclusions:** Empirical use of antibiotics must be avoided and antibiotics must be used only after sensitivity testing. This will help in selection of the appropriate antibiotic for therapeutic use and prevent indiscriminate and irrational use of antibiotics. This will also improve the cost efficiency and decrease the duration of hospital stay.

Keywords: Antibiotic resistance, Antibiotic sensitivity, Catheter associated urinary tract infections

#### **INTRODUCTION**

According to WHO, at any time, up to 7% of patients in developed and 10% in developing countries will acquire at least one hospital acquired infections. Some of these infections include catheter associated bloodstream infection, ventilator associated pneumonia, surgical site infection, catheter associated urinary tract infection (CAUTI). Urinary tract infections (UTIs) are the fourth most common type of healthcare-associated infection, with an estimated 93,300 UTIs in Intensive Care Units in 2011.<sup>1</sup> Among UTIs acquired in the hospital approximately 75% are associated with a urinary catheter.<sup>1</sup>

The risk factors for catheter associated urinary tract infections can be of two types, modifiable and non-modifiable. The total duration of catheterization, taking steps to adhere to aseptic catheter care, inserting catheter outside the operating room, a breach in the closed system of catheter drainage are few of the modifiable risk factors, which if controlled can prevent it from happening.<sup>2</sup> Whenever CAUTI is undiagnosed or untreated, it can lead to complications such as cystitis, pyelonephritis, gram-negative bacteraemia, endocarditis, vertebral osteomyelitis, septic arthritis, endophthalmitis and meningitis in patients. Complications associated with CAUTI cause discomfort to the patient, prolonged hospital stay and increased cost. Bacteriuria also leads

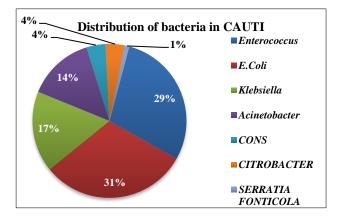
unnecessary antimicrobial use and urinary drainage systems can be reservoirs for multidrug-resistant bacteria and source of transmission to other patients.

#### **METHODS**

After written consent, women who were admitted in obstetric ward and in need of catheterization, were included in the study. Under aseptic precautions, first urine sample was taken at the time of catheterization and sent for culture and sensitivity and to rule out pre-existing urinary tract infections. Women were managed with antibiotics as per standard protocol. Those samples that were negative for urinary tract infections were included in the study. On the third post-operative day (after 48 hours), at the time of removal of catheter, again urine sample was collected for culture and sensitivity. Symptoms of urinary tract infections were noted. Type of bacteria isolated and their antibiotic sensitivity pattern was observed. Asymptomatic bacteriuria was identified if women did not develop symptoms of UTI and the growth of bacteria was more than or equal to 105 CFU/ml.

#### RESULTS

This study included 500 adult patients who received indwelling urethral Foley's catheter and urinary drainage system in wards in DR BRAM Hospital, from April 2018 to March 2019.



# Figure 1: Percentage of bacterial distribution in CAUTI.

Overall incidence of CAUTI was 10.6%. Of 500 urine samples cultured, 53 (10.6%) were culture positive and 447 (89.4%) showed no evidence of growth. Duration of catheterization varied from 48-72 hours. Incidence of CAUTI was directly proportional to duration of catheterization and 4% subjects developed clinical signs or symptoms of UTI after 2 calendar days from the time of insertion of indwelling urinary catheter. Monomicrobial CAUTI was obtained from 48 samples and rest was polymicrobial. The study revealed a total of 7 bacterial isolates, out of which 2 were gram positive and 5 were gram negative bacteria. Overall predominant etiological agent was Escherichia coli (31%), followed by Enterococcus species (29%), Klebsiella species (17%), Acinetobacter species (14%), coagulase-negative Staphylococcus (CoNS) species (4%), Citrobacter species (4%) and Serratia fonticola (1%), as shown in Figure 1.

#### Sensitivity pattern

Antimicrobial sensitivity pattern of bacterial isolates is depicted in Table 1. Among Enterococcus species isolates about 16 (94.1%) were sensitive to vancomycin, followed by 12 (70.5%) to nitrofurantoin. Other sensitive drugs were meropenem, linezolid, amoxyclav. Among E.coli isolates, maximum (72.2%) were sensitive to colistin. followed by meropenem, ceftazidime. cefotaxime and nitrofurantoin. Among CoNS species isolates most effective antibiotics were nitrofurantoin, linezolid, vancomycin (100%). For Acinetobacter species, the most effective drug was colistin (87.5%) followed by ciprofloxacin. Among Klebsiella species, maximum sensitivity was found for colistin (90%), meropenem and ciprofloxacin. Citrobacter species showed maximum sensitivity to cefepime, ampicillin, amoxyclav, meropenem, norfloxacin, cefaperazone, nitrofurantoin, clotrimazole and colistin, 50% each. Serratia fonticola was sensitive to amoxyclay, ciprofloxacin and cefepime-tazobactam (100%).

#### Resistance pattern:

Antimicrobial resistance pattern of bacterial isolates is depicted in Table 2. Among Enterococcus species isolates, maximum was resistant to ceftriaxone (100%), amoxycillin, ampicillin (94.4%), norfloxacin, cefazolin, gentamycin, teicoplanin, tetracycline (88.2%). Among E. coli isolates, maximum (100%) were resistant to ceftriaxone followed by norfloxacin, amoxyclav, cefazolin, cefotaxime-clavulanate, gentamycin, ofloxacin, cefaperazone-sulbactum (94.4% each). Other resistant drugs were linezolid, imipenem, cefepime, ampicillin, clotrimazole. Among CoNS species isolates most resistance was found to ceftriaxone, norfloxacin, ofloxacin, penicillin, colistin (100%) followed by oxacillin, gentamycin, cefoxitin. For Acinetobacter species, the most resistant drug was ceftriaxone (100%) followed by ampicillin, cefotaxime, cefotaximeclavulanate (87.5% each), meropenem, amoxyclav, cefazolin, cefuroxime, gentamicin, ceftriaxone, cefepimetazobactum, nitrofurantoin, teicoplanin (75% each). Among Klebsiella species, maximum resistance was to ceftriaxone (100%) followed by ampicillin, amoxicillin, cefazolin, amoxiclav, piptaz, ceftazidime, cefepime, cotrimoxazole, cefaperazone, cefotaxime-clavulanate (90%). Citrobacter species showed maximum resistance ceftriaxone (100%) followed by ampicillin. to amoxyclav, amoxicillin. cefazolin. ceftazidime. cefuroxime, imipenem, norfloxacin, nitrofurantoin, ciprofloxacin, cefixime, cefaperazone (50% each). Serratia fonticola was resistant to ampicillin, ceftriaxone, cefepime, cefuroxime, norfloxacin, colistin, cefixime.

### Table 1: Antibiotic sensitive/total isolate (%) of uropathogens isolated.

Antibiotic	Enterococcus	E. coli	Cons	Acinetobacter	Klebsiella	Citrobacter	Serratia fonticola
Ampicillin	1/17 (5.8%)	3/18 (16.6%)		1/8 (12.5%)	1/10 (10%)	1/2 (50%)	
Amoxycillin	1/17 (5.8%)					1/2 (50)	
Amoxyclav	6/17 (35.2%)	1/18 (5.5%)		2/8 (25%)	1/10 (10%)	1/2 (50%)	1/1 (100%)
Imipenam		7/18 (38.8%)		3/8 (37.5%)	3/10 (30%)		
Meropenam	8/17 (47.0%)	11/18 (61.1%)		2/8 (25%)	6/10 (60%)	1/2 (50%)	
Piperacillin tazobactum		6/18 (33.3%)			1/10 (10%)		
Cefazolin	2/17 (11.7%)	1/18 (5.5%)		2/8 (25%)	1/10 (10%)		
Cefoxitin	_, _, (, ,,,,,	-,	1/2 (50)				
Ceftazidime		9/18 (50%)			1/10 (10%)		
Cefotaxime		8/18 (44.4%)		1/8 (12.5%)			
Cefotaxime clavulate		1/18 (5.5%)		1/8 (12.5%)	1/10 (10%)		
Cefepime		4/18 (22.2%)		3/8 (37.5%)	3/10 (30%)	1/2 (50%)	
Cefuroxime		9/18 (50%)		2/8 (25%)	2/10 (20%)		
Cefaperazone sulbactum		1/18 (5.5%)					
Cefepime tazobactum		6/18 (33.3%)		2/8 (25%)	5/10 (50%)		1/1 (100%)
Gentamicin	2/17 (11.7%)	1/18 (5.5%)	1/2 (50%)	2/8 (25%)	4/10 (40%)		
High level gentamicin	3/17 (17.6%)						
Amikacin		6/18 (33.3%)		2/8 (25%)	2/10 (20%)		
Netilmycin				2/8 (25%)			
Norfloxacin	2/17 (11.7%)	1/18 (5.5%)			5/10 (50%)	1/2 (50%)	
Ofloxacin		1/18 (5.5%)					
Oxacillin			1/2 (50%)				
Teicoplanin	2/17 (11.7%)			1/8 (12.5%)			
Cotrimazole	1/17 (5.8%)	2/18 (11.1%)			1/10 (10%)	1/2 (50%)	
Nitrofurantoin	12/17 (70.5%)	9/18 (50%)	2/2 (100%)	2/8 (25%)	4/10 (40%)	1/2 (50%)	
Doxycycline	3/17 (17.6%)		1/2 (50%)				
Polymyxin-B	1/17 (5.8%)			2/8 (25%)	1/10 (10%)		
Vancomycin	16/17 (94.1%)		2/2 (100%)				
Linezolid	6/17 (35.2%)	2/18 (11.1%)	2/2 (100%)				
Tetracycline	2/17 (11.7%)						
Colistin		13/18 (72.2%)		7/8 (87.5%)	9/10 (90%)	1/2 (50%)	
Cefaperazone					1/10 (10%)	1/2 (50%)	
Ciprofloxacin	2/17 (11.7%)			4/8 (50%)	6/10 (60%)		1/1 (100%)

### Table 2: Antibiotic resistant/total isolate (%) of uropathogens isolated.

Antibiotic	Enterococcus	E. coli	CoNS	Acinetobacter	Klebsiella	Citrobacter	Serratia fonticola
Ampicillin	16/17 (94.1%)	15/18 (83.3%)		7/8 (87.5%)	9/10 (90%)	1/2 (50%)	1/1 (100%)
Amoxycillin	16/17 (94.1%)					1/2 (50%)	
Amoxyclav	11/17 (64.7%)	17/18 (94.4%)		6/8 (75%)	9/10 (90%)	1/2 (50%)	
Imipenam		11/18 (61.1%)		5/8 (62.5%)	7/10 (70%)		
Meropenam	9/17 (52.9%)	7/18 (38.8%)		6/8 (75%)	4/10 (40%)	1/2 (50%)	
Penicillin			2/2(100%)				
Piperacillin tazobactum		12/18 (66.6%)			9/10 (90%)		
Cefazolin	15/17 (88.2%)	17/18 (94.4%)		6/8 (75%)	9/10 (90%)		
Cefoxitin			1/2 (50%)				
Ceftazidime		9/18 (50%)			9/10 (90%)		
Cefotaxime		10/18 (55.5%)		7/8 (87.5%)			

Antibiotic	Enterococcus	E. coli	CoNS	Acinetobacter	Klebsiella	Citrobacter	Serratia fonticola
Cefotaxime clavulate		17/18 (94.4%)		7/8 (87.5%)	9/10 (90%)		
Cefepime		14/18 (77.7%)		5/8 (62.5%)	7/10 (70%)		1/1 (100%)
Cefuroxime		9/18 (50%)		6/8 (75%)	8/10 (80%)		1/1 (100%)
Cefaperazone sulbactum		17/18 (94.4%)					
Ceftriaxone	17/17 (100%)	18/18 (100%)	2/2 (100%)	8/8 (100%)	10/10 (100%)	2/2 (100%)	1/1 (100%)
Cefepime tazobactum		12/18 (66.6%)		6/8 (75%)	5/10 (50%)		
Gentamicin	15/17 (88.2%)	17/18 (94.4%)	1/2 (50%)	6/8 (75%)	6/10 (60%)		
High level gentamicin	14/17 (82.3%)						
Amikacin		12/18 (66.6%)		6/8 (75%)	8/10 (80%)		
Tobramycin				2/8 (25%)			
Norfloxacin	15/17 (88.2%)	17/18 (94.4%)	2/2 (100%)	1/8 (12.5%)	5/10 (50%)	1/2(50%)	1/1 (100%)
Ofloxacin		17/18 (94.4%)	2/2 (100%)				
Oxacillin			1/2 (50%)				
Teicoplanin	15/17 (88.2%)			6/8 (75%)			
Cotrimazole		16/18 (88.8%)		1/8 (12.5%)	9/10 (90%)		
Nitrofurantoin	5/17 (29.4%)	9/18 (50%)		6/8 (75%)	6/10 (60%)	1/2 (50%)	
Doxycycline	14/17 (82.3%)						
Tetracycline	15/17 (88.2%)						
Ciprofloxacin	10/17 (58.8%)	7/18 (38.8%)		4/8 (50%)	4/10 (40%)	1/2 (50%)	
Colistin			2/2 (100%)				1/1 (100%)
Piperacillin	2/17 (11.7%)	3/18 (16.6%)					
Linezolid	11/17 (64.7%)	16/18 (88.8%)					
Cefixime		4/18 (22.2%)		5/8 (62.5%)	6/10 (60%)	1/2 (50%)	1/1 (100%)
Vancomycin	1/17 (5.8%)						
Cefaperazone					9/10 (90%)	1/2 (50%)	
Levofloxacin				1/8 (12.5%)	1/10 (10%)		
Sulbactum				1/8 (12.5%)			
Moxifloxacin					1/10 (10%)		

#### DISCUSSION

Catheter associated bacteriuria is the most widespread healthcare associated infection in developed as well as in developing countries. In this study, the overall incidence of CAUTI was 10.6%. Study by Verma S et al, found 15.9% incidence of CAUTI.<sup>3</sup> Incidence of CAUTI in this study was found less than other compared studies. This could be due to overall less duration of catheterization compared to other studies. This variation could also be attributed to difference in study protocols, type of patients included, number of centers where the study was performed and duration of study.

In this study, incidence of CAUTI was directly proportional to the duration of catheterization and out of 53 culture positive CAUTI, 98% of patients had asymptomatic UTI. In this study 91% of CAUTI are monomicrobial and most frequent pathogen responsible for CAUTI as *Escherichia coli* (31%). On comparison to other studies, Raminder et al, found maximum (90%) of CAUTI was due to *E. coli.*<sup>4</sup> *E. coli* is the most common

pathogen in the presence of a catheter as well. The persistence of *E. coli* strains is related to the presence of type 1 pili, an adhesin for uroepithelium as well as the Tamm–Horsfall protein.<sup>5</sup>

Other organisms that were isolated in this study includes *Enterococcus* (29%), *Klebsiella species* (17%), *Acinetobacter species* (14%), *Coagulase negative staphylococcus aureus* and *Citrobacter species* (4%) and *Serratia fonticola* (1%).

# Antibiotic susceptibility pattern of isolated gram negative uropathogens

In this study, most of the *E. coli* were sensitive to colistin (72.2%), meropenem (61.1%), ceftazidime, nitrofurantoin (50%), cefotaxime (44.4%). On comparing with other similar studies, one study by Gupta et al, found similar results to that of this study that is, most sensitive antibiotics for gram negative isolates were colistin followed by tigecycline, ciprofloxacin, imipenem.<sup>6</sup>

In this study, among other gram-negative bacteria, most *Acinetobacter species* were sensitive colistin (87.5%). Among *Klebsiella species*, maximum sensitivity was found for colistin (90%), meropenem and ciprofloxacin. For *Citrobacter species*, maximum sensitivity was to cefepime, ampicillin, amoxyclav, meropenem, norfloxacin, cefaperazone, cotrimazole and colistin 50% each. *Serratia fonticola* was sensitive to amoxyclav, ciprofloxacin and cefepime-tazobactam (100%).

This study shows that, 55.5% isolates of *E. coli* were resistant to cefotaxime, 50% to cefuroxime, 100% to ceftriaxone and 50% to ceftazidime. This higher resistance to *cephalosporins* can also be on compared with studies.<sup>3,6-8</sup> Among other organisms, in this study 80% of *Klebsiella species*, 75% of *Acinetobacter species* and 100% *Serratia fonticola* was resistant to ceftazidime. 90% of *Klebsiella species* were resistant to ceftazidime.

High resistance rates to ciprofloxacin - a fluoroquinolone considered highly effective in the treatment of CAUTIs was another finding of this study. In this study 38.8% of E. coli isolates, 50% of Citrobacter and Acinetobacter species isolates and 40% of Klebsiella species isolates were resistant to ciprofloxacin. Similar study found 88.89% of gram-negative bacteria resistant to ciprofloxacin.<sup>4</sup> Another study found 77.7% resistant Enterobacteriaceae to ciprofloxacin.<sup>5</sup> Norfloxacin is another fluoroquinolone, to which 94.4% of E. coli, 50% of Klebsiella and Citrobacter species, 12.5% of Acinetobacter species were resistant. This is in accordance with Bhani et al, where CAUTIs due to gramnegative bacteria were significantly resistant to norfloxacin (81.25%).<sup>9</sup> Cotrimoxazole is another commonly used drug. In present study 88.8% of E. coli, 90% of Klebsiella species, 12.5% of Acinetobacter species were resistant to cotrimazole. Two other similar studies have got similar results, that is about 88.9% of Enterobacteriaceae isolates resistant to Cotrimoxazole.<sup>3,4</sup>

In this study 58.33% of *Enterobacteriaceae* were resistant to nitrofurantoin, another commonly used drug for treatment of catheter associated urinary tract infections.

High incidence of Carbapenem resistance in the isolates was obtained in present study. 61.1% of *E. coli*, 62.5% of *Acinetobacter species* and 70% of *Klebsiella species* were resistant to *imipenam*. 38.8% of *E. coli*, 75% of *Acinetobacter species*, 40% of *Klebsiella species* and 50% of *Citrobacter species* were resistant to meropenem.

Aminoglycoside resistance among the organisms isolated was also observed in this study; with 94.4% *E. coli* were resistant to gentamycin. 66.6% of *E. coli*, 75% *Acinetobacter species*, 80% of *Klebsiella species* were resistant to amikacin. Similar study showed 33.3% of *Enterobacteriaceae* isolates being resistant to both gentamicin and amikacin.<sup>3</sup>

In this study 94.4% of *E. coli*, 90% of *Klebsiella species*, 75% of *Acinetobacter species* and 50% of *Citrobacter species* were resistant to amoxyclav. Mishra et al, found 100% of *E. coli* showing amoxyclav resistance.<sup>6</sup>

# Antibiotic susceptibility pattern of isolated gram positive uropathogens

In this study, gram-positive bacteria- *Enterococcus species* were sensitive to vancomycin (94.1%), followed by nitrofurantoin (70.5%) and they were resistant to ceftriaxone (100%), amoxycillin, ampicillin (94.4%), norfloxacin, cefazolin, gentamycin, teicoplanin, tetracycline (88.2%). CoNS showed 100% sensitivity to vancomycin, nitrofurantoin and linezolid and maximum resistance to ceftriaxone, norfloxacin, ofloxacin and colistin.

Another study shows that gram positive organisms were sensitive to vancomycin which is similar finding of this study.<sup>6</sup> They also have shown maximum sensitivity of these organisms to linezolid which contrasts with this study as about 64.7% of *Enterococci* were resistant to linezolid. They have also found resistance of grampositive organisms to ciprofloxacin.<sup>6</sup>

#### CONCLUSION

The empirical use of antibiotics must be avoided, and antibiotics must be used only after susceptibility testing. This will help in selection of the appropriate antibiotic for therapeutic use, prevent indiscriminate, irrational use of antibiotics and its resistance. This will also improve the cost efficiency and decrease the duration of hospital stay. Implementation of infection control practices must be strictly done and followed for prevention and control of CAUTI.

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