

## Original Research

### Bacteriological Profile and Antibiotic Sensitivity Patterns of Catheter Associated Urinary Tract Infection Isolates in a North- Indian Tertiary Care Hospital

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#### ABSTRACT:

**Background:** Catheter associated urinary tract infection (CAUTI) usually resulting from wide spread inappropriate urinary catheterization is a major concern due to its nosocomial potential and association with antibiotic resistance both of which make its management difficult, consequently increasing morbidity, mortality, hospital stay, and economic burden. This study was therefore undertaken to find bacteriological profile and antibiotic sensitivity pattern of the isolates of catheter associated urinary tract infections (CAUTI). **Materials and method:** A total of 115 non-repetitive urine samples were collected aseptically from all catheterised patients in various Intensive Care units (ICUs) of SGRD Charitable Hospital as per standard protocol. Semi-quantitative bacterial cultures were performed without delay by inoculating samples on Blood agar and MacConkey agar and incubating them aerobically at 37°C overnight. Isolate identification and antibiotic sensitivity testing was done by both conventional bacteriological techniques and automated Vitek-2 as per CLSI guidelines. **Results:** Of the 115 samples we processed CAUTI was diagnosed in 100 (86.9%). Overall incidence of CAUTI was 30.94. The average age of the patient was 55±15 years. CAUTI developed more frequently in (95.7%) female patients compared to male patients (80.9%). The most common uropathogen was *E.coli* 42/100 (42%) followed by *Klebsiella pneumoniae* 30/100 (30%), *Acinetobacter baumannii* and *Enterococcus faecium* 4/100 (4%) each, *Klebsiella oxytoca*, *Staph.hemolyticus* and *Pseudomonas aeruginosa* 2 (2%) each and *Enterobacter cloacae* 1 (1%). However, yeast like organisms 13/100 (13%), were a conspicuous observation denoting that immunocompromised status of ICU patients may increase susceptibility to infection. Also high microbial resistance was found in the first two organisms in listed above in the new treatment options. **Conclusion:** Stringent infection control interventions including hand hygiene, closed drainage system aseptic method of insertion and catheter care along with daily need assessment with evidence-based observation can go a long way to keep CAUTI rates in check.

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#### INTRODUCTION

CAUTI is a UTI where an indwelling urinary catheter was in place for >2 calendar days on the date of event.<sup>1</sup> Approximately 12%-16% of adult hospital in patients will have an indwelling urinary catheter (IUC) at some time during their hospitalization, and each day the indwelling urinary catheter remains, a patient has a 3%-7% increased risk of acquiring a catheter-associated urinary tract infection (CAUTI).<sup>2</sup> CAUTI can lead to such complications as prostatitis, epididymitis, and orchitis in males, and cystitis

pyelonephritis, gram-negative bacteremia, endocarditis, vertebral osteomyelitis, septic arthritis, endophthalmitis, and meningitis in either gender. Complications associated with CAUTI cause discomfort to the patient, prolonged hospital stay, and increased cost and mortality.<sup>3</sup> However, reduction of the rate and duration of unnecessary catheterization may have an impact in reduction of HA CAUTI. Bacteria may enter the bladder through contamination of the tip during insertion with the flora of the distal urethra or from bacteria ascending the outside or the

inside of the catheter. Residual urine in the bladder of catheterised patients increases the risk of bacteriuria. During the process of infection, bacteria need first to adhere to the epithelial cells of the urinary tract and/or the surface of the catheter. They will then develop into biofilms on the catheter surface which are resistant to the immune system and antibiotics. Catheters by themselves may cause immediate physical damage to the bladder epithelium; they may be toxic and also cause inflammation. Bacteria can also damage the epithelium and cause inflammation and the combination of both may be synergistic in producing symptoms in the patient.<sup>4</sup>

The development of bacteriuria is mainly attributed to the duration of catheterization in the inpatient that in turn increases the length of hospital stay, cost and morbidity. So it is mandatory to know the prevailing bacterial strains and its antimicrobial susceptibility pattern to start targeted therapy to prevent such sequel. Additionally, a major concern is the growing antibiotic resistance of the uropathogens isolated that makes management difficult, consequently increasing morbidity, mortality, hospital stay and economic burden. Knowledge of antibiotic resistance trends is important for improving evidence based recommendations for empirical treatment of CAUTI. This study was therefore undertaken to find bacteriological profile and antibiotic sensitivity pattern of the isolates of catheter associated urinary tract infections (CAUTI) in SGRD Hospital.

#### MATERIAL AND METHODS

A total of 115 patients catheterized for more than 48 hours and admitted in various ICUs of SGRD Charitable Hospital were studied for development of

CAUTI. Patients who didn't give consent or catheterized prior to admission or having any sub acute/chronic urinary tract infections prior to admission were excluded.

Upon enrollment, and thereafter on days 3<sup>rd</sup>/5<sup>th</sup>/ 7<sup>th</sup> and removal / change of first catheter approximately 3ml of urine was aspirated from the sampling port of the catheter with a sterile syringe, first disinfecting the port with 10% povidone-iodine. The catheter tubing was then clamped just above the port to allow collection of freshly voided urine. The collected urine was then transported in a sterile, wide mouthed, screw capped universal container. Diagnosis of CAUTI was made as per the Centers for Disease Control and Prevention (CDC) guidelines published in 2009.<sup>1</sup> Urine samples were transported immediately in a sterile container to Bacteriology section of Microbiology department of SGRD hospital. In case of delay in processing beyond 2 hours they were refrigerated overnight. Uncentrifuged urine samples were subjected to microscopic examination of wet mount and gram staining for the presence of pus cells and or organisms. With a calibrated platinum loop of 3.26 mm internal diameter, 0.001ml urine was inoculated on Blood agar and MacConkey agar media for semi-quantitative analysis.

#### INCIDENCE OF CAUTI WAS CALCULATED AS FOLLOWS

$$\frac{\text{Total number of CAUTIs in one month} \times 1000}{\text{Number of CAUTIs per Total catheter days}^* \times 1000 \text{ catheter days}}$$

\*Number of patients catheterized in that month × Total number of days of that month

#### RESULTS

**Table 1: Association of age group with culture results**

Age Group	Positive		Negative		Total	
	Number	%	Number	%	Number	%
<=40	15	78.90%	4	21.10%	19	100.00%
41-60	57	91.90%	5	8.10%	62	100.00%
61-80	23	82.10%	5	17.90%	28	100.00%
>80	5	83.30%	1	16.70%	6	100.00%
Total	100	87.00%	15	13.00%	115	100.00%
P value=.381						

Table 1 show that no significant association was present between culture results and age group as depicted

**Table 2: Association of gender with culture results**

Gender	Positive		Negative		Total	
	Number	%	Number	%	Number	%
Male	55	80.9%	13	19.1%	68	100.00%
Female	45	95.7%	2	4.3%	47	100.00%
Total	100	87.00%	15	13.00%	115	100.00%
P value=.020						

In this study, 55/68(80.9%) males and 45/47(95.7%) females were diagnosed with CAUTI. A significant association (p=.012) was seen in gender and culture positivity was found. (Table 2)

**Table 3: Distribution of positive cultures according to day of sampling**

Day	CAUTI Present	
	No. of patients	% age
Day 0	0	0
Day 3	14	12.2
Day 5	25	21.7
Day 7	61	53.0
Culture negative	15	13
Total	115	100

Table 3 shows that 14(12.2%), 25 (21.7%) and 61 (53%) of 115 samples were positive on Day 3, 5 and 7 respectively indicating that CAUTI was commonest after a week of catheterization.

Table 4 shows similar results by both conventional and automated (vitek-2) methods. But 2 additional were detected as *E.coli*, beyond 42 detected by conventional technique.

**Table 4: Comparative analysis of organisms isolated by conventional and automated vitek-2 method**

Bacteria	Conventional Method		Automated vitek-2 method	
	No. of isolates	Percentage (%)	No. of patients	Percentage (%)
<i>E.coli</i>	42	42	44	44
<i>Klebsiella pneumoniae</i>	30	30	28	28
Yeast like organisms	13	13	13	13
<i>Acinetobacter baumannii</i>	4	4	4	4
<i>Enterococcus faecium</i>	4	4	4	4
<i>Klebsiella oxytoca</i>	2	2	2	2
<i>Pseudomonas aeruginosa</i>	2	2	2	2
<i>Staph.hemolyticus</i>	2	2	2	2
<i>Enterobacter cloacae</i>	1	1	1	1
Total	100	100.0	100	100.0

**Table 5: Antibiotic resistance profile of gram negative bacteria**

	Sensitive strains		Resistant strains		Intermediate	
<b>Ampicillin</b>	2 (2.4)	2 (2.4)	79 (97.5)	79 (97.5)	00 (00)	00 (00)
<b>Amoxicillin</b>	5 (6.1)	5 (6.1)	73 (90.1)	73 (90.1)	3 (3.7)	3 (3.7)
<b>Piperacillin/tazobactam</b>	9 (11.1)	10 (12.3)	71 (87.6)	70 (86.4)	00 (00)	00 (00)
<b>Cefoperazone/sulbactam</b>	10 (12.3)	11 (13.5)	64 (79)	64 (79)	7 (8.6)	6 (7.4)
<b>Imipenem</b>	16 (19.7)	15 (18.5)	55 (67.9)	57 (70.3)	10 (12.3)	9 (11.1)
<b>Meropenem</b>	14 (17.2)	16 (19.7)	64 (79)	61 (75.3)	3 (3.7)	3 (3.7)
<b>Amikacin</b>	37 (45.6)	37 (45.6)	40 (49.3)	39 (48.1)	4 (4.9)	4 (4.9)
<b>Gentamicin</b>	30 (37)	30 (37)	46 (56.7)	46 (56.7)	5 (6.1)	5 (6.1)
<b>Ciprofloxacin</b>	6 (7.4)		73 (90.1)		2 (2.4)	
<b>Tigecycline</b>	56 (69.1)	55 (67.9)	23 (28.3)	21 (25.9)	2 (2.4)	2 (2.4)
<b>Nitrofurantoin</b>	31 (38.2)	31 (38.2)	47 (58)	47 (58)	3 (3.7)	3 (3.7)

It was observed 2, 5, 37, 30 and 31 among 81 gram negative bacterial isolates were sensitive to ampicillin, amoxicillin, amikacin, gentamicin and nitrofurantoin respectively in both conventional and automated vitek-2 method.

Sensitivity was shown by 9 and 10 isolates to piperacillin/ tazobactam and cefoperazone/ sulbactam respectively by conventional method however 1 more

in each case shown to be sensitive by automated vitek-2 method.

While 16 and 56 isolates were sensitive to imipenem and tigecycline respectively by conventional method, one less was shown to be sensitive by automated vitek-2 for both mentioned antibiotics. On the contrary sensitivity to meropenem was shown by 14 isolates by conventional method and 16 by automated

vitek-2 method.

It was observed that all isolates of *Acinetobacter baumannii*, *Enterobacter cloacae*, *Klebsiella pneumoniae* and *Pseudomonas aeruginosa* were resistant to ampicillin, amoxicillin and piperacillin/tazobactam.

We found that 34 (81%), 33 (80.5%), 28(66.7%), 27 (64.3%), 29(69%), 14(33.3%), 22(52.4%), 36(85.7%), 3(7.1%) and 11(26.2%), of 42 *E.coli* strains were resistant to Ampicillin and Amoxicillin,

Piperacillin/Tazobactam, Cefoperazone/Sulbactam, Imipenem, Meropenem, Amikacin, Gentamicin, Ciprofloxacin, Tigecycline, Nitrofurantoin respectively. As many as 29 (96.7%), 19 (63.3%), 26 (86.7%), 17(56.7%), 15(50%), 16 (53.3%), 28(93.3%) of 30 *Klebsiella pneumoniae* were resistant to Piperacillin/Tazobactam, Cefoperazone/Sulbactam and Ciprofloxacin, Imipenem, Meropenem, Amikacin, Gentamicin, Tigecycline and Nitrofurantoin respectively.

**Table 6: Incidence of CAUTI**

Month	Incidence of CAUTI	% positivity of ICU catheter samples
January 2020	25.8	80
February 2020	28.5	40
March 2020	28.2	87.5
April 2020	20	60
May 2020	29	90
June 2020	28.5	85.7
July 2020	32.2	100
August 2020	32.2	100
September 2020	33.33	100
October 2020	21.5	66.66
November 2020	25	75
December 2020	32.25	100
January 2021	28.22	77.7
February 2021	35.70	100
March 2021	32.25	100
April 2021	33.33	100
May 2021	32.25	100
June 2021	33.33	100

**Table 7: Association between day of sampling and type of bacterial isolate**

Day of sampling VS Bacterial isolate	Day 3		Day 5		Day 7		Total		P value
	N	%	N	%	N	%	N	%	
<i>Acinetobacter baumannii</i>	1	7.1	1	4	2	3.3	4	4	.045
<i>E.coli</i>	3	21.4	10	40	29	47.5	42	42	
<i>Enterobacter cloacae</i>	0	0	0	0	1	1.6	1	1	
<i>Enterococcus faecium</i>	1	7.1	1	4	2	3.3	4	4	
<i>Klebsiella oxytoca</i>	0	0	0	0	2	3.3	2	2	
<i>Klebsiella pneumoniae</i>	2	14.3	8	32	20	32.8	30	30	
<i>Pseudomonas aeruginosa</i>	0	0	2	8	0	0	2	2	
<i>Staph.hemolyticus</i>	1	7.1	1	4	0	0	2	2	
Yeast like organisms	6	42.9	2	8	5	8.2	13	13	
Total % within Day of Growth	14	100	25	100	61	100	100	100	

## DISCUSSION

CAUTI continues to be the most common health care associated infection worldwide and result from widespread misuse of urinary catheterization. The overall incidence rate observed in the present study was 30.94%, similar to the study conducted by Dund et al.<sup>5</sup> in 2013 which reported an the incidence rate of 32%. It was also in concordance to the studies conducted by Bagchi et al.<sup>6</sup> and Patil et al.<sup>7</sup> that reported the incidence rate to be 29.09% and 27.70% respectively. However results in present study were in sharp contrast to the studies conducted by Chih-Cheng Lu et al.<sup>8</sup> Taiwan in 2000 and Danchaivijitr S et al.<sup>48</sup>

who found overall incidence of CAUTI 57% and 73.3% respectively. In the study by Kulkarni et al.<sup>9</sup> in MGM MCH, Aurangabad the incidence was found to be lower at 21.47%. However in a study by Kaji et al.<sup>10</sup> the overall incidence of 4.59 per 1000 catheter days was much lower than the incidence observed in the current study. In present study the incidence rate was more as ours is a tertiary care hospital and patients are mostly admitted in critical conditions thereby increasing the incidence of debilitating diseases.

In present study, maximum patients were present in the age group 41 to 60 years (53.9 %) followed by 61-

80 years (24.3%),  $\leq 40$  (16.5%) and  $>80$  years (5.2%) making the average age of the patient  $55 \pm 15$  years. Similar results were shown by Patil et al.<sup>7</sup>, Kulkarni et al.<sup>9</sup> and Leelakrishna et al.<sup>11</sup> reported 54.5 years,  $46.48 \pm 17.82$  and 51.61 years respectively as the mean age of patients with CAUTI. Conversely, Chih-Cheng Lu, et al.<sup>8</sup> conducted a study in a tertiary care hospital in Taiwan and observed the mean age of the patients to be 75.9 years which was noticeably greater than present study.

The present study found no significant association between age and culture results implying that age was not a risk factor for development of CAUTI. Similarly, study conducted by Verma et al.<sup>12</sup> showed no significant correlation between age and growth of bacteria. However, Anggietal.<sup>13</sup> in 2019 concluded a significant relationship between CAUTI infection with age, with a value of  $p = 0.028$  ( $p < 0.05$ ), with the most research subjects being in the age group of 21 to  $\geq 60$  years. The subjects who aged  $\leq 50$  years had two times more risk than a subjects who aged  $>50$  years. Similarly, Leelakrishna and Karthik.<sup>11</sup> also observed that patients  $>50$  years were at more risk of developing CAUTI than those below 50 years of age ( $p = .001$ ).

In present study 68 (59.1%) of 115 patients were males and 47 (40.9%) patients were females almost same as in the study done by Bhayani et al.<sup>14</sup> in 2019 comprising more male patients (63%) compared to female patients (37%). Leelakrishna and Karthik.<sup>11</sup> also observed more no. of males (67.1%) in their study.

This study also noted that CAUTI developed more frequently (95.7%) in female patients compared to in male patients (80.9%). A significant association was seen in gender and culture result ( $p = .012$ ). In a study by Vincitorio et al.<sup>15</sup> 483 patients, the number of females were more being 297 (61.5%) and also had more prevalence of CAUTI. Conversely in present study the number of male patients was more although it was females who had more prevalence of CAUTI. Similarly study conducted by Mishra et al.<sup>16</sup> had more females 25 (67.6%). Verma et al.<sup>12</sup> in their study had more number of females, 38.04% and 21% of them

## CONCLUSION

Due to pooling of large number of referrals from other centers to our hospital, the development of CAUTI was common in our critically ill patients making incidence of CAUTI high further, most of the isolates were MDRO's. It can be managed through extra emphasis placed on good catheter care management, reducing the duration of catheterization and avoiding indiscriminate use of antibiotics, that may contribute to emergence of drug resistant strains in the environment. Active infection control team and efficient institutional infection control surveillance policy for non-compliance, strict adherence to preventive bundle practices and constant monitoring can go a long way in bringing down the incidence rate of CAUTI. Vitek-2 could be very useful as reference

developed CAUTI. Leelakrishna and Karthik.<sup>11</sup> reported that 64.4% males and 35.6% females to be affected with CAUTI. The difference was statistically non significant in the latter two studies in contrast to current study. Our finding is in strong agreement with the findings of other researchers that females have a stronger predilection for CAUTI compared to males.<sup>12</sup>

Sensitivity of gram negative organisms in present study to tigecycline, amikacin and nitrofurantoin is 69.1%, 45.6% and 38.2% respectively making them the most effective antibiotics.

It was observed that 3 and 1 of *Acinetobacter baumani*, were sensitive to tigecycline and nitrofurantoin respectively. In contrast, Michael Osthoff et al.<sup>17</sup> in his study showed 84 % sensitivity to nitrofurantoin. Sujatha et al.<sup>18</sup> in her study found that most of the Gram negative bacilli were sensitive to nitrofurantoin (97%) and amikacin (85%). The above mentioned results are opposite to current study where very less isolates were sensitive to these drugs.

Whereas, all isolates of *Acinetobacter baumani*, *Enterobacter cloacae*, *Klebsiella pneumoniae* and *Pseudomonas aeruginosa* were resistant to ampicillin, amoxicillin and piperacillin/tazobactam.

None of the isolates of *Enterobacter cloacae* and *Klebsiella oxytoca* was sensitive to any of the drugs used making the last two mentioned organisms pan resistant strains.

Similar results were obtained by Verma et al.<sup>12</sup> who showed that most of the Gram negative bacilli were resistant to commonly used antibiotics. Earlier studies have suggested that selective pressure from the use of antimicrobial agents is a major determinant for the emergence of resistant strains.

Peng et al.<sup>19</sup> in their study showed that some common pathogen groups of gram-negative bacteria, including *E.coli*, *Klebsiella spp*, *Acinetobacter spp*, and *Pseudomonas spp*, were generally resistant to ampicillin, ciprofloxacin, cefazolin, and cotrimoxazole. The present study also showed similar results where majority of the bacteria were resistant to most of antibiotics.

method for bacterial identification and antibiotic susceptibility. By decreasing the time required for accurate identification of common uropathogens and determination of their antibiotic susceptibility resulting in better clinical outcome as well as financial saving for the hospital.

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