Original Research Article

Microbial Profile and Antimicrobial Susceptibility Pattern of Uropathogens Isolated From Catheter Associated Urinary Tract Infection (CAUTI)

Deepa Bhani*, Rekha Bachhiwal, Rajni Sharma and Rakesh Kumar Maheshwari

Department of Microbiology, SMS Medical College, Jaipur, Rajasthan, India

*Corresponding author

ABSTRACT

Urinary tract infections are the most commonly acquired bacterial infections accounting for an estimated 25-40% of the nosocomial infections. Catheterization is a potential predisposing factor of CAUTI. Study was carried out over a period of one year included 100 urine samples obtained from catheterized patients. The urine samples obtained were immediately processed in microbiology laboratory by semi-quantitative method as per standard protocol. Following isolation and identification, all the isolates were subjected to antimicrobial susceptibility testing for commonly used antimicrobial drugs. Out of 230 urine samples from catheterised patients 43.5% of cases developed catheter associated urinary tract infection (CAUTI). The most common organism causing CAUTI was found to be E. coli in 27.2% of cases, followed by Pseudomonas (20%), Klebsiella (15.4%), Enterobacter aerogenes (9%), Enterobacter cloacae (0.09%), Proteus spp. (2.7%) and Citrobacter spp. (2.7%). Gram positive cocci accounted only in 10% of CAUTI cases and Candida spp. in 11.8% of cases. Antimicrobial susceptibility testing revealed Imipenem (66.7%-94.1%) and nitrofurantoin (73.3%-88.2%) to be the most sensitive antimicrobials for frequently isolated Gram negative bacteria like E. coli and Klebsiella spp. Pseudomonades isolated were sensitive to imipenem and gentamicin (31.8% each). Gram positive cocci were completely sensitive to linazolid and vancomycin. This study reported a high rate of CAUTI (43.5%) and resistance to commonly used antimicrobials among CAUTI isolates, hence it is necessary to reduce urinary infection in catheterized patients by limiting the use of catheter only for proper indications and remove catheters promptly when no longer required.

Keywords
Catheter associated urinary tract infection (CAUTI), Urinary Tract Infections (UTIs), Device Associated Infection (DAI).

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Introduction

Urinary Tract Infection constitutes a serious health problem that affect millions of people each year1.

Urinary Tract Infections (UTIs) are defined as diseases which are caused by a microbial invasion of the genitourinary tract that extends from the renal cortex of the kidney to the urethral meatus2. Among UTIs acquired in the hospital, approximately 75% are associated with a urinary catheter3. Catheter associated urinary tract infection (CAUTI) is one of the most important Device associated infection (DAI) and it is one of the leading causes of morbidity and mortality in hospitalized patients4. The risk of acquiring catheter associated urinary tract infection (UTI) depends on method and duration of catheterization, the quality of catheter care and host susceptibility5,6,7. Host factors including female sex, prolonged catheterization, severe underlying illness,
disconnection of the catheter and drainage tube, other types of faulty catheter care, and lack of systemic antimicrobial appear to increase the risk of catheter associated urinary tract infection (CAUTI)\(^8,9\). The incidence of CAUTI rises from 0.5 to 1% for a single “in-and-out” catheterization to 10 to 30% for catheters in place for up to 4 days and up to 95% for catheters in place for 30 days or more\(^10\).

CAUTI if left untreated, it can lead to complications such as cystitis, pyelonephritis, Gram negative bacteremia, prostatitis, epididymitis, endocarditis, vertebral osteomyelitis, septic arthritis, endophthalmitis and meningitis\(^11\). Additionally CAUTIs also result in prolonged hospital stay, increased cost and mortality\(^12\).

Present study was undertaken to find out rate of CAUTI, organisms causing it and their antimicrobial susceptibility pattern.

**Materials and Methods**

The present study was conducted in the Department of Microbiology, SMS Medical College and Attached Hospitals, Jaipur (Rajasthan).

A total of 230 urine samples collected from catheterised patients were included in the study. The diagnosis of CAUTI was done as per the CDC guidelines.\(^3\)

**Inclusion criteria**

Patients from urology and surgery ward/ICU, catheterized for \(\geq 48\) hr with baseline

Urine culture sterile.

**Exclusion criteria**

Patients with prior symptoms of UTI.

Patients catheterized for \(< 48\) hr.

Urine samples were collected aseptically using a sterile needle and syringe from the distal edge of catheter tube into a sterile universal container and transported to the microbiology laboratory for analysis with minimum delay. The samples were processed as per routine standard laboratory procedure. This included microscopy, culture, and antimicrobial susceptibility testing.

Urine samples were inoculated with an inoculating loop of internal diameter of 2mm delivering 0.001 ml of urine for semi quantitative analysis on blood agar and MacConkey agar. Culture plates were evaluated after 18-24 hours of aerobic incubation at 37˚C\(^14\).

Isolates were identified on the basis of Colony morphology, Gram staining and Biochemical reactions\(^15,16,17\). Urine samples with no more than two species of organisms, at least one of which is a bacteria of \(\geq 10^5\)CFU/ml were taken into consideration.\(^3\)

Antimicrobial susceptibility testing of isolated organism was performed by Kirby Bauer disk diffusion method\(^18\).

**Results and Discussion**

Out of 230 catheterised patients under study, 100 (43.5%) cases developed CAUTI.

A total of 110 uropathogens were isolated from the 100 urine samples from CAUTI cases. Of the 100 samples, single isolate was obtained in 90 samples and in the remaining 10 samples more than 1 isolate (i.e. 2 isolates) were obtained.

Among the 110 uropathogens Gram negative bacilli showed predominance.

\(E.\ coli\) (27.2%) were the most frequently isolated pathogens followed by \(Pseudomonas\)
Among the Gram positive pathogens isolated, **Enterococcus spp** (6.3%) was the commonest pathogen isolated followed by Coagulase positive **Staphylococci** and Coagulase negative **Staphylococci** (1.8% each). **Candida spp** accounted for 11.8% cases (Table 1).

Among the commonly used antimicrobials, imipenem and nitrofurantoin were the most sensitive antimicrobials to **E. coli** (66.7% and 73.3%) and **Klebsiella spp** (94.1%, and 88.2%). Sensitivity to other commonly used antimicrobials including Aminoglycosides, fluoroquinolones, 3rd and 4th generation cephalosporins were found to be lower than imipenem and nitrofurantoin. Ampicillin showed complete resistance to **Escherichia coli**, **Klebsiella spp**, **Enterobacter cloacae**, **Proteus spp** and **Citrobacter** (Table 2).

Table 3 shows antimicrobial sensitivity pattern of **Pseudomonas spp**. **Pseudomonas** isolated were most sensitive to imipenem and gentamicin (31.8% each) followed by Aztreonam and Pipercillin + Tazobactam (18.2% each) and were completely resistant to levofloxacin.

Antimicrobial sensitivity pattern of **Enterococcus** and **Staphylococci** spp revealed that both of these uropathogens were completely sensitive to vancomycin and linezolid. Sensitivity of **Enterococcus** spp for nitrofurantoin, penicillin and norfloxacin were found to be 57.6%, 42.8% and 28.6% respectively.

Sensitivity of **Staphylococci** for doxycycline, gentamicin was 75% each while levofloxacin and nitrofurantoin showed sensitivity of 50% each. **Staphylococci** were least sensitive to norfloxacin, erythromycin and cefoxitin (25% each) (Table 4).

Catheter associated urinary tract infections are the most common nosocomial infections, 75% cases of UTI in catheterised patients is due to use of urinary catheter\(^3\). A urinary catheter provides bacteria, a portal of entry into the urinary tract. Bacteria may ascend into the tract via the external or internal surface of the catheter\(^11\). Also urinary catheter use is often associated with catheter-related biofilms\(^19\).

In the present study rate of CAUTI in catheterized patients was found to be 43.5%. Chaudhari et al.,\(^21\) reported 44% and Poudel et al.,\(^22\) reported 55% rate of CAUTI in their study which is in agreement with the present study. However studies conducted by Chatterjee et al.,\(^23\) and Dimri et al.,\(^24\) reported a higher rate of CAUTI (86.6% and 86.1% respectively). A lower rate of CAUTI has been reported by Bhatia et al.,\(^25\) (28.8%), Prajapati et al.,\(^26\) (13.14%) and Roel et al.,\(^27\) (15.5%).

The source of microorganisms causing CAUTI can be endogenous typically via meatal, rectal, or vaginal colonization or exogenous, such as via equipment or contaminated hands of healthcare personnel\(^11\).

In the present study Gram negative bacteria (78.18%) were predominantly isolated from CAUTI cases. Gram positive bacteria accounted for only 10% of cases. **Candida** spp constituted 11.8% of uropathogens isolated.

Amongst the GNB, **E. coli** was found to be the most common uropathogen (27.2%), followed by **Pseudomonas** spp. (20%), **Klebsiella** spp. (15.4%), **Enterobacter aerogenes** (9%), **Enterobacter cloacae** (0.09%), **Proteus** spp. (2.7%) and **Citrobacter** spp. (2.7%).

(20%) and **Klebsiella** spp (15.4%). Other GNB accounted for 0.09%-9 % of cases (Table 1).
**Table 1.** Uropathogens isolated from CAUTI cases

| Organisms                | Number | Percentage% |
|--------------------------|--------|-------------|
| *Escherichia coli*       | 30     | 27.2%       |
| *Pseudomonas spp*        | 22     | 20%         |
| *Klebsiella spp*         | 17     | 15.4%       |
| *Enterobacter aerogenes* | 10     | 9%          |
| *Enterobacter cloacae*   | 1      | 0.09%       |
| *Proteus spp*            | 3      | 2.7%        |
| *Citrobacter spp*        | 3      | 2.7%        |
| *Enterococcus spp*       | 3      | 2.7%        |
| *CONS*                   | 2      | 1.8%        |
| *CPS*                    | 2      | 1.8%        |
| *Candida spp*            | 13     | 11.8%       |

**Table 2.** Antimicrobial sensitivity pattern of isolated Gram negative bacteria

| Organism                | Ampicillin | Pip-Tazo | Ceftazidime | Cefotaxime | Cefepine | Imipenem | Doxycycline | Choramphenicol | Amikacin | Gentamicin | Nitrofurantoin | Levofloxacin | Norfloxacin |
|-------------------------|------------|----------|-------------|------------|----------|----------|-------------|----------------|----------|------------|----------------|--------------|--------------|
| *Escherichia coli*      | 0          | 4 (13.3%)| 2 (6.7%)    | 2 (6.7%)   | 9 (30%)  | 20 (66.7%)| 3 (10%)     | 9 (30%)        | 6 (20%)  | 22 (73.3%)| 8 (26.7%)      | 2 (6.7%)     |              |
| *Klebsiella spp*        | 0          | 3 (17.6%)| 2 (11.8%)   | 6 (35.3%)  | 8 (47%)  | 16 (94.1%)| 1 (6%)      | 8 (47%)        | 9 (52.9%)| 15 (88.2%)| 5 (29.4%)      | 5 (29.4%)    |              |
| *Enterobacter aerogenes*| 1 (10%)    | 2 (20%)  | 1 (10%)     | 2 (20%)    | 3 (30%)  | 6 (60%)  | 4 (40%)     | 5 (50%)        | 3 (30%)  | 8 (80%)   | 1 (10%)        | 2 (20%)      |              |
| *Enterobacter cloacae*  | 0          | 0        | 0           | 0          | 0        | 1 (100%) | 0           | 1 (100%)       | 0        | 0         | 0              | 0            |              |
| *Proteus spp*           | 0          | 1 (33.3%)| 0           | 1 (33.3%)  | 2 (66.7%)| 3 (100%) | 0           | 2 (66.6%)      | 2 (66.7%)| 2 (66.7%)| 1 (33.3%)      | 2 (66.7%)    |              |
| *Citrobacter spp*       | 0          | 1 (33.3%)| 2 (66.7%)   | 1 (33.3%)  | 2 (66.7%)| 3 (100%) | 1 (33.3%)   | 3 (100%)       | 1 (33.3%)| 1 (33.3%)| 2 (66.7%)      | 1 (33.3%)    |              |

**Table 3.** Antimicrobial sensitivity pattern of *Pseudomonas spp* (N=22)

| Antibiotic                | No (22) | %   |
|---------------------------|---------|-----|
| Piperacillin + Tazobactam | 4       | 18.2|
| Ceftazidime               | 2       | 9   |
| Cefepime                  | 3       | 13.6|
| Aztreonam                 | 4       | 18.2|
| Imipenem                  | 7       | 31.8|
| Amikacin                  | 2       | 9   |
| Gentamicin                | 7       | 31.8|
| Levofoxacin               | 0       | 0   |
| Norfloxacin               | 5       | 22.7|
Table.4 Antimicrobial sensitivity pattern of *Staphylococcus* spp. (N=4, CONS=2 and CPS=2) and *Enterococcus* spp (N=7)

| Antimicrobial drug | Staphylococci N =4 | Enterococci N=7 |
|--------------------|-------------------|----------------|
| Cefoxitin          | 1 (25%)           | --             |
| Doxycycline        | 3 (75%)           | --             |
| Gentamicin         | 3 (75%)           | --             |
| Erythromycin       | 1 (25%)           | --             |
| Vancomycin         | 4 (100%)          | 7 (100%)       |
| Linezolid          | 4 (100%)          | 7 (100%)       |
| Nitrofurantoin     | 2 (50%)           | 4 (57.2%)      |
| Levofloxacín       | 2 (50%)           | --             |
| Norfloxacin        | 1 (25%)           | 2 (28.6%)      |
| Penicillin         | --                | 3 (42.8%)      |

Poudel *et al.*, 22 also reported *E. coli* as predominant isolate (40.7%), followed by *Pseudomonas* spp (11.1%) and *Klebsiella* spp (11.1%) in their study.

*E. coli* is the most common organism reported from CAUTI cases, the source of these strains of *E. coli* is frequently the patient’s own intestinal flora.22 Virulence factors of uropathogenic *E. coli* (UPEC) include the ability to adhere to uroepithelial cells and certain specific serotypes O and K antigens that are resistant to phagocytosis and bactericidal action of normal serum.29

In the present study *Escherichia coli* was isolated in 27.2 % of CAUTI cases. Similar results have been reported in studies performed by Chaudhari *et al.*, 21, and Roel *et al.*, 27, with rate of isolation to be 21% and 27.2% respectively. However Poudel *et al.*, 22 have reported a higher isolation rate of *E. coli* (40.7%).

Antimicrobial sensitivity pattern of isolates showed that imipenem (60-100%) and nitrofurantoin (73.2%-88.2%) were the most sensitive antimicrobial agents among GNB. Other antimicrobials including ceftazidime, cefotaxime, cefepime, gentamicin, amikacin levofloxacin and doxycycline showed moderate to low sensitivity. Ampicillin was found to be least sensitive (Table 2).

Sensitivity of imipenem to *Escherichia coli* was 66.7%, for *Klebsiella* spp 94.1% and 100% each for *Enterobacter cloacae*, *Proteus* spp and *Citrobacter* spp. Nitrofurantoin showed 73.3% sensitivity for *Escherichia coli*, 88.2% for *Klebsiella* spp and 80% for *Enterobacter aerogenes*.

Poudel *et al.*, 22, Dimri *et al.*, 24, Manish *et al.*, 30, Roel *et al.*, 27 also reported imipenem as the most effective antibiotic among GNB (91-100%).

*Pseudomonas* spp was found to be the second most frequently isolated microorganism in the present study after *E. coli* accounting for 20% (22/110) of all uropathogens isolated. These finding are broadly in agreement with the finding of Chaudhari *et al.*, 21 and Roel *et al.*, 27, who reported the isolation rate to be 21%, and 18.2% respectively. A little higher rate was reported by Chatterjee *et al.*, 23 (30.67%). While lower rate of isolation was reported by Manish *et al.*, 30, Dimri *et al.*, 24 (10.8%).

*Pseudomonas* causes catheter-associated urinary tract infections through biofilm formation on the surface of indwelling catheters.

*Pseudomonas* isolated were maximally sensitive to imipenem and gentamicin (31.8% each) followed by norfloxacin (22.7%),
aztreonam (18.2%) and piperacillin + tazobactam (18.2%). These demonstrated a low sensitivity to cefepime (10.6%), amikacin (9%), ceftazidime (9%) and levofloxacin (0%). Sensitivity to Gentamicin (31.8%) to Pseudomonas in the present study is broadly in agreement with that reported by Dimri et al.,24 (30%) and Chatterjee S. et al.,23 (30.44%), while sensitivity to imipenem was lower than the other studies. Sensitivity to Aztreonam (18.2%) correlates well with that reported by Dimri et al., (20%).24

Among the GPC, Enterococci accounted for 6.3% cases and Coagulase positive Staphylococcus and Coagulase negative Staphylococcus (CONS) each for 1.8% cases. These were completely sensitive to vancomycin and linezolid.

CA-UTI is an important device-associated health care acquired infection. In the present study the rate of CAUTI was found to be 43.5% and most of the isolates were GNB 78.18%, Escherichia coli being the predominant isolate. Most of organisms isolated were found to be resistant to commonly used antimicrobials.

Considering such a high rate of CAUTI and increasing resistance among CAUTI isolates, it is necessary to reduce urinary infection in catheterized patients by limiting the use of catheter only for proper indications and remove catheters promptly when no longer required. Infection control programs must develop, implement, and monitor policies and practices to minimize infections associated with use of these devices.

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