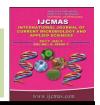


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A Study on Bacteriological Profile of Urinary Tract Infection in Diabetes Mellitus Patients in a Tertiary Care Hospital

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ABSTRACT

Keywords

UTI, Diabetic patients, Drug resistance, Imipenem, *E. coli*

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Urinary Tract Infection (UTI) is an inflammatory response of the urothelium to bacterial invasion that is usually associated with bacteriuria and pyuria. UTI is upto three times more common in diabetic than in non-diabetic women and increases with age. Urine from 60 diabetic patients with UTI was collected and subjected to Gram staining, culture & sensitivity by routine methods. Among Diabetic patients, the males constituted 51.66% and were predominant. The maximum number of isolates was present in 51-60yrs with 35% and next was 41-50yrs age group with 20%. In Diabetes mellitus, *E. coli* was the predominant isolate with 36.6% followed by *Klebsiella spp*, *Pseudomonas spp & Acinetobacter spp*. In Diabetes patients, Imipenem, Amikacin and Nitrofurantoin had greater sensitivity than other antibiotics. In Gram positive cocci isolated from Diabetic patients, Amikacin and cephalexin had a greater sensitivity. This alarmingly high resistance rate has to be approached cautiously, by formulating an antibiotic drug policy. This has to include giving effective drugs after the culture & sensitivity and ensuring that the patients receive full course of antibiotics.

Introduction

Urinary Tract Infection (UTI) is an inflammatory response of the urothelium to bacterial invasion that is usually associated with bacteruria and pyuria. Urinary tract is the second most common site for bacterial infection in human. Bacterial virulence factors play a subsidiary part in the genesis of disease.

UTI refer to those infections that occur in the presence of anatomic or functional abnormalities in the kidney, bladder or collecting systems; obstruction to normal urine flow (including renal, ureteric and bladder calculi and Benign Prostatic

Hypertrophy); cystic renal disease; specific diseases like Diabetes Mellitus and abnormalities in host defense mechanisms and those infections that follow surgery or procedures done in urinary tract.

UTI is upto three times more common in diabetic than in non- diabetic women and increases with age.

Many factors that predispose the diabetic patient to infections in the urinary tract have been described. Diabetes mellitus is associated with increased risk of these infections as a result of poorly controlled

plasma glucose concentrations, which in turn may impair granulocyte function and cellmediated immunity. Also, the neurologic dysfunction associated with diabetic neuropathy may result in a neurogenic bladder with incomplete bladder emptying, urinary stasis, and retention. The increased likelihood of urethral instrumentation may predispose these patients to infection, as may diabetic microangiopathy, which contribute to local ischemia and impaired host defences (Rayfield EJ et al., 1982, Wheat et al., 1980).

Asymptomatic bacteriuria has been described as occurring two to four times more frequently in diabetic woman (Wheat et al., 1980; Forland, 1977) and as being more prevalent in diabetic women than men (Kass et al., 1956). Other renal parenchymal complications of UTI in diabetic patients include Pyelonephritis, **Emphysematous** pyelonephritis, papillary necrosis Perinephric abscesses; these conditions should be considered in the evaluation of nonresponse to appropriate antibiotic therapy for urinary infection in diabetic patients.

Most of the bacteria responsible for urosepsis in diabetics are Gram-negative rods, with *E. coli and Klebsiella spp*, accounting for about 70%. *Klebsiella spp* are isolated twice as frequently in diabetic patients with bacteremic urinary infections, and a large proportion of these patients had indwelling urinary bladder catheters.

Madras Medical College and Rajiv Gandhi Government Hospital Chennai is a tertiary care hospital and a referral centre to the hospitals in the surrounding districts. Knowledge of the microbial organisms and antibiotic susceptibility pattern of Diabetic patients with UTI is essential for defining empirical treatment. Hence the present study focuses on the bacteriological profile and its

resistance patterns in cases of UTI in Diabetic patients for a better clinical approach.

The main objectives of this study includes: To determine the Bacteriological profile in Diabetic patients with Urinary Tract Infection. To determine the Antibiotic susceptibility pattern using Kirby Bauer Disc diffusion method. And to analyse the drug resistant pattern in uropathogens in Diabetic patients with UTI.

Materials and Methods

Sample collection

Midstream urine specimens

Urine samples were most commonly collected by sampling the midstream flow by the clean - catch technique. Urine collection from women by this technique requires personal supervision for best results.

Laboratory methods

Urine microscopic analysis

Five ml of the urine specimen was centrifuged at 2500 rpm. Supernatant was removed and sediment used for microscopy analysis. The sediment was transferred on the clean slide and viewed for the presence of epithelial cells, red cells and casts.

Gram staining

Gram staining of urine smear was done and the presence of Gram negative bacilli or Gram positive cocci was initially observed. The presence of pus cells were also noted.

Urine culturing and bacterial identification

One ml of diluted (1/1000) urine specimen was used for culturing on CLED, Blood, and

Mac Conkey agar. Specimen was inoculated using a calibrated loop. The plate was incubated at 35°C for 24 hrs-48 hrs.

Specimens with more than 10⁵ colony forming unit [CFU]/ ml were considered as positive samples. The isolates were identified based on colony morphology on culture plates, Gram's staining and by standard biochemical tests. Antibiotic susceptibility testing was done by standard methods according to CLSI guidelines.

Antimicrobial susceptibility testing

Antibiotic susceptibility testing was performed by the Kirbey Bauer disc diffusion method on MHA (Mueller Hinton Agar) according to CLSI protocols. The diameters of zone of inhibition were interpreted according to CLSI standards for each organism. Media and disks were tested for quality control with standard strains.

Results and Discussion

The study undertaken at the Institute of Microbiology, Madras Medical College, Chennai among samples received from Diabetic patients of Rajiv Gandhi Government General Hospital, Chennai with UTI showed the following results. Sixty patients were included in this study.

Among diabetic patients, the males constituted 51.66% and were predominant. The male to female prevalence ratio is 1.06: 1 (Table 1). This was in comparison with the study conducted by Sibi *et al.*, (2011) where the males were affected with UTI more than women in Diabetic patients.

In Gram positive cocci (GPC) isolated from Diabetic patients, Amikacin and cephalexin had a greater sensitivity. The majority of the patients presented with dysuria and was found in 37.5% patients. The next presentation was

with fever in 30.8%; followed by pain abdomen, frequency of micturition, hematuria, and oliguria in the decreasing order of presentation. These symptoms were comparable with the study by Mahesh *et al.*, (Mahesh *et al.*, 2010).

The *E. coli* was the commonest organism isolated with 36.6% (Table 3). The study conducted by Sibi *et al.*, (2011) also had *E. coli* accounting for 39.14% and another study reported 47% of *E. coli* in their isolate (Andy *et al.*, 2003). These results were in comparison with the present study.

The Klebsiella pneumoniae 20%, Klebsiella oxytoca 5%, Pseudomonas aeruginosa 11.6%, Acinetobacter spp 10%, Staphylococcus aureus 6% and Citrobacter freundii 5% and Enterococcus faecalis 5% were the other isolates in these diabetic patients (Table 3). The study by Sibi et al., (2011) had Klebsiella spp 15.7% which was comparable to this study.

The E. coli isolated in these patients had higher sensitivity 90.9% for Amikacin and 81.80% for Nitrofurantoin. But the resistance to cephalosporins and Fluoroquinolones were high, 75% and 80% respectively. The Imipenem was 100% sensitive (Table 4). Shill et al., (2010) in his study also reported 66% resistance to cepholosporins 79% resistance to Fluoroguinolones; 3% resistance to Amikacin and 11.9% resistance to Nitrofurantoin and 100% sensitive to Imipenem. This study results were similar to the present study. Yet the resistance pattern to Cephalosporins and Fluoroquinolones in these patients is highly alarming and Amikacin and Imipenem offer promise in treating these E. coli infected Diabetes mellitus patients.

The *Pseudomonas aeruginosa* was the third commonest isolate in these patients. They were 42.8% sensitive to Amikacin and

Cephalosporin, 14.2% sensitive to Fluroquinolones (Table 4).

In a similar study by Mahesh *et al.*, (2010) 74.5% resistance rate to Fluoroquinolones and 64.7% resistance to Amikacin and Gentamicin were noted for *Pseudomonas spp*.

This was in comparison to the present study. In the present study, *Pseudomonas spp* were 100% sensitive to Imipenem whereas 70% of the isolates were sensitive to Carbapenems in the study by Mahesh *et al.*, This is not coherent with the present study.

Table.1 Gender distribution in diabetes mellitus

Gender	Male	Female
No of patients	31	29
Percentage	51.66%	48.33%

Table.2 Distribution of patients with diabetes mellitus according to age group [n=60]

Age groups	No of patients	Percentage
12-20	6	10%
21-30	4	6%
31-40	7	11.6%
41-50	12	20%
51-60	21	35%
61-70	4	6%
71-80	6	10%
Total	60	

The maximum number of isolates were present in 51-60yrs with 35% and next was 41-50yrs age group with 20%

Table.3 Uropathogens isolated in diabetes mellitus

Name of the organisms	No of patients	Percentage
E.coli	22	36.6%
K pneumoniae	12	20%
K oxytoca	3	5%
Pseudomonas aeruginosa	7	11.6%
Acinetobacter spp	6	10%
Citrobacter freundii	3	5%
Staph.aureus	4	6%
Enterococcus faecalis	3	5%
Total	60	

In Diabetes mellitus, *E. coli* was the predominant isolate with 36.6% followed by *Klebsiella spp*, *Pseudomonas spp* and *Acinetobacter spp*.

Table.4 Gram negative bacilli - antibiotic sensitivity in diabetes mellitus patients (n = 53)

	Name of the Organisms											
Antibiotics	E. ce	oli	Kleb	siella	K.ox	ytoca	Pseua	lomonas	Acin	eto	Citro	
			pneu	moniae			aerug	inosa	bacte	er spp	bacte	r
	n=22	2	n=12 $n=3$		_		n =	6	freund	freundii n =3		
	No	%	No	%	No	%	No	%	No	%	No	%
												100%
Amikacin	20	90.9%	6	50%	3	100 %	3	42.8%	6	100%	3	
												100%
Nitrofurantoin	18	81.8%	6	50%	-	-	-		2	33.3%	3	
												100%
Norfloxacin	4	18.1%	3	25%	1	33.3	3	42.8%	3	50%	3	
Ceftazidime		31.8%	3	25%	3		3	42.8%	4	6.6%	3	100%
	7					00%						
		9.09%	1	8.3%				14.2%		33.3%	3	100%
Ciprofloxacin	2				-	-	1		2			
		18.1%	3	25%	-	-	-	-	2	33.3%	3	100%
Ofloxacin	4											
	9	40.9%	-	-	1	33.3	1	14.2%	1	16.6%	-	-
Cotrimoxazole						%						
	4	18.1%	3	25%	3	100	3	42.8%	6	100%	-	-
Gentamicin						%						
	22	100%	12	100	3	100	7	100%	6	100%	3	100%
Imipenem				%		%						

In Diabetes patients, Imipenem, Amikacin and Nitrofurantoin had greater sensitivity than other antibiotics.

Table.5 Gram positive cocci – antibiotic sensitivity in diabetes mellitus patients (n=7)

Antibiotics	Staphylococcus aureus [n= 4]	Enterococcus. faecalis [n=3]
Amikacin	3 [75%]	2 [66.6%]
Amoxy-clavulanic acid	2 [50%]	2 [66.6%]
Erythromycin	-	-
Cephalexin	2 [50%]	-
Amoxycillin	1 [25%]	-
Cotrimoxazole	1 [25%]	-
Ciprofloxacin	2 [50%]	-
Nitrofurantoin	-	3 [100%]

Out of the Gram positive cocci isolated, Staphylococcus aureus constituted the majority followed by *Enterococcus faecalis*. *Staphylococcus aureus* was found to be sensitive to Amikacin (75%) and Cephalexin (50%) and resistant to cotrimaxazole, amoxicillin and Ciprofloxacin. *Enterococcus faecalis* was found to be 66.6% sensitive to

Amikacin and amoxyclavulanic acid and 100% to Nitrofurantoin.

The poorly controlled glucose concentration may lead to decreased granulocyte function and cellular immunity. Diabetic neuropathy results in urinary stasis. Diabetic microangiopathy leads to impaired host

defenses. All these reasons contribute to the colonization of these resistant uropathogens.

To summarise, Among Diabetic patients, the males constituted 51.66% and were predominant.

The maximum number of isolates was present in 51-60yrs with 35% and next was 41-50yrs age group with 20% (Table 2). In Diabetes mellitus, *E. coli* was the predominant isolate with 36.6% followed by *Klebsiella spp*, *Pseudomonas spp* and *Acinetobacter spp*.

In Diabetes patients, Imipenem, Amikacin and Nitrofurantoin had greater sensitivity than other antibiotics. In GPC isolated from Diabetic patients, Amikacin and cephalexin had a greater sensitivity.

This alarmingly high resistance rate has to be approached cautiously, by formulating an antibiotic drug policy. This has to include giving effective drugs after the culture and sensitivity and ensuring that the patients receive full course of antibiotics.

An ongoing surveillance of UTI should be encouraged for an updated knowledge about the uropathogens causing infection and their antibiotic resistance pattern to keep the medical community informed about the emerging antibiotic resistance.

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