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## Microbiological profile and antimicrobial susceptibility pattern among patients with urinary tract infection in northern Tanzania

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### Abstract

**Background:** Urinary tract infection (UTI) is the most common bacterial infection, it causes morbidity as well as financial burden. There is a role of empirical treatment of UTI while awaiting urine culture and sensitivity result. The type of empirical antibiotics given varies from place to place depending on susceptibility pattern of the particular area. According to Tanzania standard treatment guidelines and essential medicine list of 2013, cystitis in adult is treated by ciprofloxacin while pregnant women and children are treated by amoxicillin/clavulanate. In case of pyelonephritis, women of reproductive age who does not require referral are treated by ciprofloxacin while all other cases that fulfill criteria for referral are given single dose of ceftriaxone before referral. However, most studies have shown increasing resistance to commonly used antibiotics as a result of misuse and overuse.

The study aimed at describing the microbiological profile and susceptibility pattern among patients with UTI at KCMC.

**Methods:** This was a seven-month hospital based descriptive, cross-sectional study conducted from June 2016 to December 2016 by using structured data extraction sheet. Data analysis was performed using SPSS after cleaning and checking for error(s).

**Results:** A total of 270 samples that tested positive for UTI were analyzed. Majority 233(86.3%) of the sample were obtained from male subjects, age of study subjects ranged from 1-94 years with the median of 64 years. Majority 235 (87%) of the sample came from urology department with E. coli being most common cause of UTI (39.3%), followed by Klebsiella spp, 20%. E. coli demonstrated high sensitivity against amikacin, ceftazidime and Nitrofurantoin

**Conclusion:** The commonest cause of UTI in our set up is E. coli that showed high resistance against most commonly used antibiotics in our setting.

**Keywords:** Urinary tract infection; Kilimanjaro Christian Medical Center; Microbiological Profile; Antimicrobial susceptibility pattern.

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## 1. Introduction

Urinary tract infections (UTIs) are the most prevalent bacterial infection rendering people in the community and in hospitals to morbidity and financial burden in healthcare seeking [1,2].

It is approximated about 150-250 million people per annum are identified to have UTI worldwide [1,3]. Furthermore, it is estimated that the global visits to outpatient clinics to be as many as 7 million visits, emergency department 1 million and 100,000 hospitalization annually [1,3,4]. The commonest causative organism of UTI is *Escherichia coli*. Others include *Staphylococcus aureus*, *Proteus mirabilis*, *Klebsiella spp.*, *Enterococcus faecalis*, *Proteus vulgaris*, *Pseudomonas aeruginosa*, *Enterobacter spp.*, *Staphylococcus epidermidis*, and *Providencia spp* [5]. The investigations for UTI include urinalysis and urine culture and sensitivity as the gold standard [6,7].

There is a role of empirical treatment while awaiting urine culture and sensitivity result. The type of empirical antibiotics given varies from place to place depending on microbiological profile and their sensitivity pattern. According to the European Association of Urology (EAU) guidelines of 2013, the recommended baseline drugs for UTI treatment are trimethoprim sulfamethoxazole, nitrofurantoin, trometamol, fosfomycin and pivmecillinam [5]. As per Tanzania standard treatment guidelines and essential medicine list of 2013, cystitis in adult is treated by ciprofloxacin, pregnant women and children are treated by amoxicillin/clavulanate. In case of pyelonephritis, women of reproductive age who does not require referral are treated by ciprofloxacin while all other cases that fulfill criteria for referral are given single dose of ceftriaxone before referral.

The World Health Organization (WHO) reported an increase in antimicrobial resistance across many different infectious agents, UTI being one of them [9]. The commonly used antibiotics are facing high rate of resistance. Globally, antimicrobial resistance is a life threatening condition, and it increases the treatment cost [3,10]. The cause of the resistance may be due to misuse and overuse of the antimicrobial. It can also occur naturally overtime due to genetic changes and the presence of drug resistant microbes in people, food, animals and hospital environments that spread either from person to person or between people and animals [9]. Microbiological profile and its susceptibility pattern should be known in every region to guide empirical treatment whenever needed [3,10]. The aim of the study is to describe the microbiological profile and antimicrobial sensitivity pattern among patients with UTI in our hospital.

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## 2. Patients and Methods

This hospital based descriptive, cross-sectional study was carried out from June 2016 to December 2016 at KCMC, Moshi Tanzania. This zonal referral hospital in Northern zone of Tanzania has more than 600 bed capacity in a catchment area of approximately 15.7 million people. It has a large clinical laboratory that runs a number of tests including urine culture and sensitivity. The laboratory participates in external quality assurance with public health agency of Canada (QASI-LI), is also internationally recognized and accredited with ISO number 15189.

The study evaluated all urine samples submitted to our lab during the study period and tested positive for UTI. Urine sample that grew multiple bacterial and the culture result form that were incomplete filled were excluded from the study. A structured questionnaire was used to collect data from clinical laboratory computer system.

Ethical clearance and permission was obtained from Kilimanjaro Christian Medical University College and clinical laboratory department at KCMC zonal referral Hospital. All patients' information was kept confidential; no patient's direct identifiers were used in the data extraction instrument. Data were entered into computer for analysis where by SPSS version 16 was used to analyze the information. Cross-tabulations were generated and comparisons were made.

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## 3. Results

A total of 270 urine culture positive results were reviewed. The age range of the patients was 1-94 years with the median age of 64 years. Majority of patients were in the age group of 51-75 years, 127(47.0%), followed by the age group of >75 years, 70(25.9%). The age group of 5-25 had the frequency of 13(4.8%). Under-fives (<5 years) were the fewest, accounted for (3.3%). Males accounted for (86.3%) with urology department having the largest number of sample (Table 1).

**Table 1** Socio-demographic characteristics (N=270)

Variable	Frequency	Percentage
	N	%
<b>Subjects</b>		
<b>Age (years)</b>		
<5	9	3.3
5-25	13	4.8
26-50	51	18.9
51-75	127	47
>75	70	25.9
<b>Median age=64</b>		
<b>Sex</b>		
Male	233	86.3
Female	37	13.7
<b>Departments</b>		
Urology	235	87
GOPD	3	1.1
Casualty	10	3.7
OG	1	0.4
Pediatrics	2	0.7
Medical	9	3.3
General surgery	8	3
RDTC	1	0.4
Orthopaedics	1	0.4

Eleven different types of bacterial pathogens were isolated with *E. coli* being the most prevalent (39.3%), followed by *Klebsiella spp* (20%). *Morganella spp* and *Serratia spp* were the least prevalent accounted for 1.9 and 1.1% respectively (Table 2).

**Table 2** Different uropathogens isolated (N=270)

Uropathogens isolated	n (%)
<i>E. Coli</i>	106 (39.3)
<i>Klebsiella pneumonia</i>	54 (20)
<i>Coliform</i>	27 (10)
<i>Pseudomonas aeruginosa</i>	25 (9.3)
<i>Citrobacter</i>	17 (6.3)
<i>Enterobacter</i>	13 (4.8)
<i>Proteus</i>	10 (3.7)
<i>Staphylococcus aureus</i>	10 (3.7)
<i>Morganella morganii</i>	5 (1.9)
<i>Serratia spp.</i>	3 (1.1)
Total	270 (100)

*E. coli* has high sensitivity against amikacin (97%), and ceftazidime while demonstrating high resistance against ampicillin/clavulanate (87%), trimethoprim-Sulfamethoxazole (75%), ciprofloxacin (69%). *Klebsiella spp* were only 38% sensitive to cefotaxime and 36% sensitive to amikacin while demonstrating resistance against trimethoprim-sulfamethoxazole (100%), ampicillin/clavulanate (96%) and ceftazidime (70%) (Table 3 and 4). Table 3 Below summarizes the resistance profile of these drugs

**Table 3** The pattern of drug resistance

Uropathogens	*Number of resistant sample n (%)								
	GEN	AMP	CET	SXT	CIP	AMK	CEZ	AMC	NIT
<i>E. Coli</i>	48(56)	41(87)	28(47)	18(75)	62(69)	2(3)	7(27)	37(55)	33(42)
<i>Klebsiella spp</i>	35(69)	22(96)	20(63)	13(100)	32(67)	1(3)	7(70)	16(64)	27(68)
<i>Coliform</i>	14(58)	10(77)	2(14)	7(100)	13(65)	2(9)	1(13)	12(80)	19(83)
<i>P. Aeruginosa</i>	16(70)	4(100)	6(43)	4(100)	12(52)	4(21)	1(11)	7(88)	11(91)
<i>Citrobacter</i>	12(80)	10(100)	6(67)	3(75)	5(39)	0(0)	1(25)	5(56)	10(77)
<i>Enterobacter spp.</i>	4(67)	6(100)	1(50)	2(67)	10(83)	0(0)	0(0)	2(100)	4(67)
<i>Proteus spp.</i>	3(30)	4(67)	0(0)	2(100)	4(50)	1(14)	0(0)	3(75)	5(83)
<i>S. Aureus</i>	1(20)	1(25)	0(0)	0(0)	1(33)	0(0)	0(0)	0(0)	2(67)
<i>Morganella spp.</i>	0(0)	1(50)	1(33)	0(0)	2(50)	1(25)	1(50)	1(50)	1(33)
<i>Serratia spp.</i>	2(67)	0(0)	1(50)	0(0)	2(100)	1(33)	0(0)	0(0)	0(0)
<b>TOTAL</b>	<b>135(59)</b>	<b>99(85)</b>	<b>66(46)</b>	<b>50(85)</b>	<b>143(64)</b>	<b>12(10)</b>	<b>18(31)</b>	<b>83(62)</b>	<b>112(60)</b>
<b>TOTAL</b>	<b>135(59)</b>	<b>99(85)</b>	<b>66(46)</b>	<b>50(85)</b>	<b>143(64)</b>	<b>12(10)</b>	<b>18(31)</b>	<b>83(62)</b>	<b>112(60)</b>

\*Resistance includes intermediate and resistance; Abbreviations: GEN, gentamicin; AMP, ampicillin; CET, cefotaxime; SXT, trimethoprim-sulfamethoxazole; CIP, ciprofloxacin; AMK, amikacin; CEZ, ceftazidime; AMC, amoxicillin/clavulanate and NIT, nitrofurantoin.

**Table 4** Sensitivity profile of the isolates

Uropathogens	No of sensitive sample n (%)								
	GEN	AMP	CET	SXT	CIP	AMK	CEZ	AMC	NIT
<i>E. Coli</i>	38(44)	3(13)	32(53)	6(25)	28(31)	69(97)	19(73)	30(44)	45(57)
<i>Klebsiella spp.</i>	16(31)	1(4)	12(38)	0(0)	16(33)	1(4)	3(30)	9(36)	13(33)
<i>Coliform</i>	10(42)	3(23)	12(86)	0(0)	7(35)	21(91)	7(88)	3(20)	4(17)
<i>P. Aeruginosa</i>	7(30)	0(0)	8(57)	0(00)	11(48)	15(79)	8(89)	1(13)	1(8)
<i>Citrobacter</i>	3(20)	0(0)	3(33)	1(25)	8(62)	13(100)	3(75)	4(44)	3(23)
<i>Enterobacter spp.</i>	2(33)	0(0)	1(50)	1(33)	2(17)	6(100)	0(0)	0(0)	2(33)
<i>Proteus spp.</i>	7(70)	2(33)	6(100)	0(0)	4(50)	6(86)	0(0)	1(25)	1(17)
<i>S. Aureus</i>	4(80)	3(75)	3(100)	0(0)	2(67)	0(0)	0(0)	2(100)	1(33)
<i>Morganella spp.</i>	5(100)	1(50)	2(67)	1(50)	2(50)	3(75)	0(0)	1(50)	2(67)
<i>Serratia spp.</i>	1(33)	2(100)	0(0)	0(0)	0(0)	2(67)	0(0)	1(100)	3(100)

Abbreviations: GEN, gentamicin; AMP, ampicillin; CET, cefotaxime; SXT, trimethoprim-sulfamethoxazole, CIP, ciprofloxacin; AMK, amikacin; CEZ, ceftazidime; AMC, amoxicillin/clavulanate and NIT, nitrofurantoin

#### 4. Discussion

Majority of urine sample analysed came from male subjects (86.3%) with the most prevalent (47%) age group being 51-75 years. This is different from other studies where by female are mostly affected [11,12]. The difference is due to the fact that majority of analysed sample in our study came from urology department (87%), in which more males are affected by urological condition than females. Moreover, males of this age group are more prone to UTI probably because they have prostate enlargement causing resistance to the flow of urine.

*E. coli* was the most prominent uropathogen isolated in 39.3% of the urine culture. This finding concurs with other studies in Tanzania and abroad though the figure is a bit lower [11,13,14]. Other frequently cultured uropathogens were *Klebsiella spp* (20%), *Pseudomonas spp.* (9.3%) and *Citrobacter Spp* (6.3%). This is different from other studies though the second most common uropathogens is the same [11,13]. This difference could be due to difference in geographical location, study population and sample size.

Most of isolated uropathogens showed high overall resistance against ampicillin, trimethoprim- sulfamethoxazole, ciprofloxacin and amoxicillin/clavulanate. *E. coli* were 87%, 75% and 69% resistant to ampicillin, trimethoprim-sulfamethoxazole and ciprofloxacin respectively. *Klebsiella Spp* were 96%, 100%, 67% and 64% resistant to ampicillin, trimethoprim- sulfamethoxazole, ciprofloxacin and amoxicillin/clavulanate respectively. Our finding is partly similar to other study that was done in another tertiary hospital in Tanzania that found high resistance against ampicillin, trimethoprim-sulfamethoxazole and amoxicillin/clavulanate [15]. In the index study, resistance against ciprofloxacin was higher compared to the previous study. The high resistance against these antibiotics is because they are the most common antibiotics used in the primary health facility but also they are accessed easily in the pharmacies even without doctor's prescription. In addition to that, high resistance to trimethoprim- sulfamethoxazole can be due to the fact that it has been in use for a long time for long-term prevention of Pneumocystis Jiroveci Pneumonia (PJP) in HIV positive patients. The difference observed between the index and previous study is due to the fact that in the latter study most of their participants were pediatric patient that usually are not treated by ciprofloxacin because of its side effects.

*E. coli* showed resistance of 42% against nitrofurantoin. This figure is much higher than other studies [16,17]. Our high resistance of 68% against nitrofurantoin amongst *Klebsiella Spp* isolates differs also from other studies in east Africa [18].

Resistance against gentamycin was high; for example, *E. coli* and *Klebsiella Spp* were 56% and 69% resistance against gentamicin respectively. This is much higher than what was reported in Dar es Salaam Tanzania by other investigator, where *E. coli* and *Klebsiella* were 38.4% and 28.8% resistant against gentamycin respectively [15]. The difference could be due to small sample size in our study and hence small number of isolates tested for gentamycin leading into overestimation.

Our finding of low resistance to amikacin is comparable to other study in the country and abroad [13,15,19]. The low resistance in our country is because this drug is not commonly used, hence less chance of microbes to develop resistance against it.

The antimicrobial sensitivity patterns vary from different geographical locations. In this index study, *E. coli* showed high rate of sensitivity to amikacin, ceftazidime and Nitrofurantoin; the sensitivity was 2 to 3 times lower to cefotaxime, amoxicillin/clavulanate, gentamicin, ciprofloxacin, trimethoprim-sulfamethoxazole and ampicillin. This is similar to the study done in Italy in which the sensitivity of *E. coli* to amikacin was high followed by third generation cephalosporin (ceftazidime and cefotaxime) and amoxicillin/clavulanate [20], however trimethoprim-sulfamethoxazole and ciprofloxacin showed some deviation in which it was among the least sensitive in our study [20].

In this study *Klebsiella spp.* had the highest sensitivity to amikacin that was 2 times lower compared to the study done in Italy but the sensitivity to the third generation cephalosporin and amoxicillin/clavulanate is almost similar [20].

#### Limitations of the study

The result of this study should be discussed in the light of the following limitation; this study was conducted in single centered zonal referral hospital and therefore the result cannot be generalized to all health facilities, but to the health facility with similar context.

Despite this limitation, our findings remained comparable to many other studies.

## 5. Conclusion

About two in five of bacterial isolates causing UTI is *E. Coli*. The resistance is high against most commonly used antibiotics; ampicillin, trimethoprim- sulfamethoxazole, ciprofloxacin and amoxicillin/clavulanate. *E. coli* had a high sensitivity against amikacin, ceftazidime and Nitrofurantoin, therefore we recommend use of these antibiotics whenever empirical treatment for UTI is required. In this study most urine sample came from urology department, therefore a prospective study is recommended for urological patients with UTI in order to find out the possible cause of UTI and their susceptibility profile. Since this was a single centered study, generalizability is limited, therefore a large multicenter study is recommended.

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## Compliance with ethical standards

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Authors would like to extend their sincere thanks to Kilimanjaro Christian Medical University College ethical clearance committee for granting ethical clearance for this study. Furthermore, they also want to express their deep gratitude to all workers of KCMC hospital for their tireless assistance.

### *Disclosure of conflict of interest*

All authors listed in this article declare that they have no conflict of interest for this publication.

### *Statement of ethical approval*

The ethical clearance was obtained for Kilimanjaro Christian Medical university ethical review committee.

### *Statement of informed consent*

This study involved already collected and analyzed urine samples, therefore, informed consent was not required and hence not obtained.

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