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Prevalence and antibiotic resistance pattern of urinary tract bacterial infections in Dessie area, North-East Ethiopia

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Abstract

Background: Different studies have indicated that urinary tract infections frequently occur in both community and hospital environments and are of the most common bacterial infections in humans. the outcomes of urinary tract infections are increased hospitalization, increased direct patient costs and mortality. In Dessie, the prevalence of the commmon pathogens and antibiotic susceptibility pattern is not well studied sofar. Thus, the aim of this study is to address these gaps in the study area.

Methods: Retrospective study was conducted in Dessie regional health research laboratory from January 1-March 31, 2012. All culture and antibiotic susceptibility test results of patients' diagnosed with UTI from September 2002 to September 2011 G.C were included in the study. Data were abstracted using structured questionnaires and finally, entered into SPSS Windows version 16.0, and descriptive statistics was generated to meet the study objective.

Results: During the last ten years 680 (27.35%) bacteria were isolated in the regional laboratory. The most commonly isolated were *E. coli* 410 (60.29%), Pseudomonas species 59 (8.68%), Proteus species 53 (7.79%), S. *aurous* 50 (7.35%) and Klebsiella species 40 (5.88%). The *E.coli* were susceptible to Nitrofurantoin 43 (89.6%), furantoin 124 (87.3%), Nalidixic acid 91 (86.7%), kanamycin 116 (80%) & ciprofloxacin 66 (71.7%) but were almost resistant to Ampicillin, tetracycline, & trimethoprim-sulfamethoxazole. Similarly Pseudomonas and proteus species were resistant to almost all antibiotics except Gentamycin.

Conclusion: The E.coli, pseudomonas and proteus species were the commonly isolated bacteria in the regional health research laboratory. A majority of isolated bacterial microbes were resistant to antibiotics commonly used in clinical practices and generally available in the local economy without prescription. Culture results are necessary before initiating antibiotics.

Keywords: UTI, Prevalence, Antibiotic, Susceptibility, Resistance

Background

Urinary tract infections (UTI) are one of the most common bacterial infections in the human urinary system. Most of these infections involve the lower urinary tract, namely the bladder and the urethra. UTIs are much more common in elderly than younger individuals for a variety of reasons, and frequently occur both in community and hospital environments [1,2]. Urethritis, cystitis, acute pylonephritis, prostatitis, and intra -renal and peri-nephric

abscesses are the common types of UTIs [1-3]. The symptoms of UTI are non-specific and may lead to vague diagnosis [3,4].

Women are at three times greater risk for UTI than men because of short, straight anatomy of the urethra, and termination of female urethra beneath the labia resulting in colonization by colonic gram-negative bacilli [1,3,5]. There are different pathogens that cause UTI [1,3]. The *E.coli*, Klebsiella species, *P.aeruginosa*, and Enterococcus species were the most common bacterial pathogens isolated from the urinary tracts of infected patients [6-8]. UTIs are usually listed among common health care associated infections that occur mostly in health care

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facilities of low income countries due to cultural and environmental reasons [9-12].

The antibiotic sensitivity patterns of bacteria isolated from urinary tracts differ for different bacteria and antibiotics [6,7,13] with sensitivity to the Quinolones being usual while resistance is often seen to nitrofurantoin; Ampicillin and Cotrimoxazole. So far, *E.coli* has been reported as the most resistant microorganism almost for all antibiotics including Quinolones [6,14] leading to recurrent infections [15,16]. Irrational drug use such as long term use and low-dose antibiotic use due to lack of protocol for antibiotic use and empiric therapy due to lack of laboratory facility to determine sensitivity are the possible reasons for resistance and thus, recurrent infections and complicated UTIs [14-16].

Hence reporting common etiologic agents and respective antibiotic sensitivity is crucial for stakeholders to search for preventive and control measures against antibiotic resistance. There are few data regarding etiologic agents and respective antibiotic sensitivity in our country except limited studies on antimicrobial sensitivity. According to this study the overall prevalence of antibiotic resistance ranges from 65% to 72% [17,18]. However, to our knowledge there was no published data on etiologic agents and antibiotic sensitivity in South Wollo. Thus, the aim of this study is to determine common etiologic agents and respective antibiotic sensitivity in an undeveloped country with some effective health care, as is common to many areas of the world.

Methods

Study area

This retrospective study was conducted in Dessie regional health research laboratory from January 1 to March 31, 2012 G.C. Dessie regional health research laboratory is located in Dessie town, Amhara region, North-East Ethiopia, 401 km from capital city, Addis Ababa. It is the only regional health research laboratory in North-East Amhara region serving laboratory requests from different areas of the region both governmental and nongovernmental health organizations and from community too. According to Dessie town health administrative office, in the town there are 16 governmental health institutions (1 referral hospital, 1 primary hospital, 8 health centers and 6 health posts), and private health institutions (3 general hospitals, 6 higher clinics, 23 medium clinics, 15 pharmacies, and 24 drug stores).

Study subjects

All patient data recorded on patient recording books from September 2002 to September 2011 G.C in the laboratory were included in the study. Similarly, the antibiotic sensitivity for the specific pathogen isolated was included in the study. Data were collected using a

structured data collection format developed by the principal investigators. All the information including culture results and antibiotic sensitivity for isolated pathogens were collected.

Laboratory diagnostic methods

The records data showed mid-stream ("clean catch") urine specimen collected from each patient. Bacterial isolation and identification was performed according to standard operating procedures of Feingold and Martin [19]. Urine samples were inoculated on cystine lysine electrolyte deficient [CLED] agar and blood agar plates [Oxoid Basingstoke, UK] using standard wire loop [0.001 ml]. The inoculated agar plates were incubated at 37°C for 24 -48 hours. Then, the growths were inspected to identify the bacteria. Biochemical tests were performed on colonies from primary cultures for final identification of the isolates. In brief, Gram-negative bacteria were identified by performing a series of biochemical tests (Oxoid, LTD), namely triple sugar iron agar, indole, Simon's citrate agar, lysine iron agar, urea, mannitol and motility. Gram-positive bacteria were identified based on their gram reaction, noviobiocin, catalase and coagulase test results.

The antimicrobial susceptibility tests of the isolates were performed according to the national committee for clinical laboratory standards (NCCLS) method using Kibry-Bauer disk diffusion test on Mueller-Hinton agar. Results were interpreted after measuring the zone of inhibition and comparing with the standards. *Eshercia coli* ATCC 25922 and S. aureus ATCC 25923 were employed as quality controls for the antimicrobial susceptibility test [20]. Finally data were edited, coded, and entered into SPSS Windows version 16.0, and descriptive statistics was computed to meet the stated objective.

Ethical considerations

Ethical clearance was obtained from ethical review committee of Wollo University. The head of laboratory was requested for cooperation/permission by a formal letter from Wollo University. Patient privacy was protected by de-identification of records. Names of patients were replaced by initials. All data obtained in the course of the study were kept confidential, and used solely for the purpose of the study.

Results

Demographic characteristics of patients and tests performed

About 2486 culture results from urinary tract discharges and urine during the ten years period were included in the study. A majority of samples 1694 (68.1%) were from females, and 1373 (55.2%) were under age group of

16-35 years. Sex and age were unavailable for 34 (1.4%) and 218 (8.8%) of samples respectively (Table 1).

Urine 2007 (80.7%) and vaginal discharge 395(15.9%) were the most commonly used samples for culture, while, urethral discharge being used in 84 (3.37%) cases.

The number of samples cultured in each year tended to decrease over time. Thus, majority of samples were analyzed in between 2002-2004 G.C (12-14%), whereas, least number was during 2009-2011 G.C (5-8%). Majority of bacteria 71 (35.86%) were isolated in 2011 G.C but only 41 (17.52%) were isolated in 2006 G.C (Table 2, Figure 1).

Prevalence and common types of bacterial uropathogens isolated

About 680 (27.35%) bacteria were isolated from 2486 cultures. The most commonly isolated bacteria were *E. coli* 410 (60.29%) followed by Pseudomonas species 59 (8.68%), Proteus species 53 (7.79%), *S. aurous* 50 (7.35%) and *Klebsiella species* 40 (5.88%), respectively. Nevertheless, from the total positive samples of the patients about 21 (3.09%) of results were reported as 'pathogenic microorganisms detected', means the lab personnel were unable to identify the species of the microorganisms detected (Figure 2). About 528 (77.65%) and 372 (54.71%) of bacteria were isolated from females and in age group of 16-35 years respectively.

Approximately a quarter of the E.coli isolates were resistant to Amoxicillin, Ampicillin, tetracycline, and Trimetoprim-sulphamethoxazole (TMP-SMX) but were susceptible to Chloramphenicol, Ceftriaxone, Ciprofloxacin, Gentamicin, Kanamycin and Nalidixic acid. Antimicrobial susceptibility results are summarized in (Table 3).

Table 1 Sex distribution of patients in the Dessie regional health research laboratory, May, 2012

S. No	Patient characteristics	Frequency (%)				
	Sex of patients					
	Not mentioned	34 (1.4)				
	Male	758 (30.5)				
	Female	1694 (68.1)				
	Total	2486 (100)				
	Age of patients					
	Not mentioned	218 (8.8)				
	<5 years	62 (2.5)				
	5-15	151 (6.1)				
	16-35	1373 (55.2)				
	36-50	487 (19.6)				
	> = 51	195 (7.8)				
	Total	2486 (100)				

Table 2 Distribution of samples in years in Dessie regional health research laboratory, May, 2012

S. No	Year	Samples (%)					
1	2002	317 (12.8)					
2	2003	323 (13.0)					
3	2004	357 (14.4)					
4	2005	266 (10.7)					
5	2006	234 (9.4)					
6	2007	285 (11.5)					
7	2008	201 (8.1)					
8	2009	177 (7.1)					
9	2010	128 (5.1)					
10	2011	198 (8.0)					
11	Total	2486 (100)					

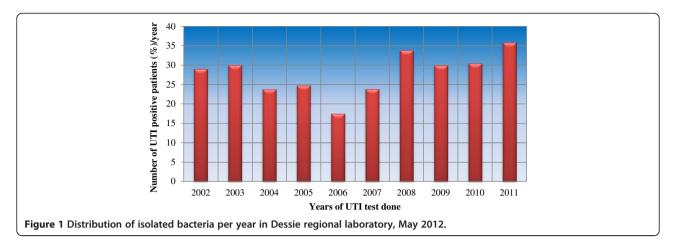
Discussion

According to this study out-off 680 [27.35%] cultures at least one bacterial species was isolated. The majority 528 (77.65%) were from females, and 372 (54.71%) from patients of 16-35 years old. This supports the idea women are more prone UTIs [1,3,5,21]. Likewise, UTIs were more common among women of reproductive age groups (16-35 years) which agrees with earlier studies in this country and abroad in Nigeria, India and Kuwait [21-26] identified sexually active and/or probably pregnant females in this age group are at high risk for UTI.

The prevalence of isolated bacteria 680 [27.35%] were higher in this study compared with similar studies in the country [23 -25,36]. This might be due to data in this study were collected from culture results of patients referred for diagnostic purpose which will select for infected cases with gram-negatives 598 [87.94%] bacteria, which were most commonly isolated in this study. This prevalence differs with studies in other parts of the country [23,24,27] and other countries like Kuwait, Nigeria and Tanzania [6,28,29]. This might be due to geographical and/or methodological differences as well large site coverage, difference in source of data, laboratory verses hospitals, from other studies done in Ethiopia or other countries.

In this study like many other similar studies, *E. coli* remain the most prominent uropathogens which was isolated in 60.29% of the cultures. This is consistent with other studies in Ethiopia and abroad though the figure is slightly higher in this study [30-32]. Among the other frequently isolated bacterial uropathogens the most common were Pseudomonas species [8.68%], Proteus species [7.79%], *S.aurous* [7.35%] and *Klebsiella species* [5.88%]. This is similar when compared with other studies in Ethiopia and outside Ethiopia [23,24,27-29].

This study has showed that approximately a quarter of *E. coli* isolates were resistant to Amoxicillin, Ampicillin,



Tetracycline, and Trimetoprim-sulphamethoxazole [TMP-SMX]. But were relatively sensitive to chloramphenicol, Ceftriaxone, Ciprofloxacin, Gentamycin, Kanamycin and Nalidixic Acid. This was partially in agreement with other studies in other parts of Ethiopia [23,24,27,33] but was comparable with a report from Nigeria [6,25]. The difference might be attributed to methodology, and pattern of management [34,35]. In the study area there are many private pharmacies, hospitals and clinics which might have contributed to overuse of antibiotics and self-medication resulting in drug resistance as reported in earlier studies [14-16].

In comparison with the developed world the resistance rate of *E.coli* was elevated for all antibiotic treatments that have been used in the study area for the past ten years to treat UTIs [36-38]. For example, in Europe Amoxicillin resistance ranges 48% in Netherland to 60% in Belgium and from 21% to 36% for the TMP-SMX, whereas in our case it was 84.6% and 75.8%, respectively [39]. Hence, as it has been indicated in various researches the reason for difference might be factors related to poverty, although,

difference in antibiotic use, patient population and prescribing rate can also play a role [36,40-42].

In this study most isolates of Gram-negative bacteria were resistant to two or more drugs [multi-drug resistance] similarly a reported for studies in other parts of the country [24,27]. This indicates that multi-drug resistance is increasingly becoming a major problem in the management of uropathogens in Ethiopia. This raises alarms to implement a nationwide antimicrobial surveillance and in-vitro susceptibility testing with strict adherence to antibiotic policy to inhibit the spread of drug resistant microbes in the country. Moreover, considering antibiotics with a resistance level of 10% or less are suitable for empiric therapy and the resistance level of antibiotics reported in this study were more than 10% for most common causative agents for UTI. Thus, these antibiotics are no longer appropriate for emparic management of UTIs. Hence, calls for nationwide study to know the exact level of antibiotics resistance among pathogenic bacteria is vital to make the right recommendation of alternative antibiotics and to support expert opinion [43].

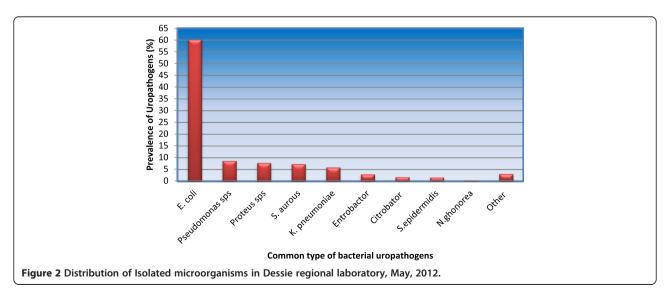


Table 3 Antibiotic Susceptibility pattern of isolated microorganisms in Dessie regional health research laboratory, May, 2012

Antibiotic	E. coli		Pseudomonas spps		Proteus spps		S. Aurous		Klebsiela spps		Entrobactor		S. epidermidis		N. gonorrhea		Citrobactor	
	T _t	%S	T _t	%S	T _t	%S	T _t	%S	T _t	%S	T _t	%S	T _t	%S	T _t	%S	T _t	%S
Amoxicillin	182	28(15.4)	8	1/8	14	0/14	3	1/3	20	6(30.0)	5	0/5	5	1/5	1	1/1	1	0/1
Tetracycline	314	56(17.8)	16	6(37.5)	23	6(26.1)	10	3/10	26	9(34.6)	19	8(42.1)	4	0/4	2	1/2	7	0/7
TMP-SMX	231	56(24.2)	27	9(33.3)	17	6(35.3)	15	4/15	23	8(34.8)	15	12/15	8	2/8	1	0/1	4	3/4
Cephalothin	71	41(57.7)	8	0/8	10	3/10	2	1/3	5	5/5	7	2/7	5	2/5	1	0/1	2	1/2
Ampicillin	45	9(20.0)	3	1/3	4	0/4	1	0/1	8	1/8	1	0/1	1	0/1	2	2/2	1	0/1
Chloramphenicol	201	104(51.0	15	12/15	16	5(31.3)	7	4/7	21	13(61.9	19	13(68.4)	4	2/4	2	2/2	5	3/5
Ceftriaxone	196	104(53.1	15	10/15	14	614	14	8/14	23	11(47.8	15	11/15	2	2/2	2	2/2	3	2/3
Ciprofloxacin	92	66(71.7)	43	41(95.3)	10	9/1	18	11(61.1)	10	6/4	17	15(88.2)	4	3/4	3	1/3	5	4/5
Doxycyclin	62	22(35.5)	10	2/10	7	2/7	3	2/3	7	4/7	1	1/1	-	-	-	-	1	0/1
Erythromycin	125	7(5.6)	39	7(17.9)	17	4(23.5)	27	10(37.0)	11	1/11	14	3/14	4	2/4	2	2/2	6	0/6
Gentamicin	362	239(66.0)	18	13(72.2)	25	16(64.0)	7	4/7	35	20(57.1)	19	16(84.2)	5	4/5	3	2/3	6	0/6
Kanamycin	145	116(80.0)	-	-	8	5/8	1	0/1	4	2/4	1	0/1	-	-	-	-	2	0/2
Naldixic acid	105	91(86.7)	-	-	8	5/8	-	-	9	9/9								
Nitrofurantoin	48	43(89.6)	4	4/4	4	4/4	3	3/3	6	4/6	6	5/6	-	-	1	1/1	1	1/1
Penicillin G	2	1/2	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Vancomycin	31	3(9.7)	9	1/9	3	0/3	3	0/3	1	0/1	-	-	-	-	-	-	1	0/1
Furantoin	142	124(87.3)	1	0/1	13	5/13	1	1/1	24	16(66.7)			1	1/1			4	4/4
Carbencillin	127	42(33.1)	5	0/5	14	5/14	2	0/2	3	0/3	2	2/2	3	1/3			2	0/2
Clindamycin	26	16(61.5)	5	1/5	3	1/3	-	-	1	0/1	3	2/3	-	-	-	-	2	1/2
Bacitracin	45	13(28.9)	-	-	7	2/7	-	-	4	1/4	-	-	1	0/1		-	-	-
Streptomycin	10	1/10	-	-	-	-	-	-	5	2/5	-	-	-	-	-	-	-	-
Oxacyclin	12	12/12	3	2/3	1	0/1	1	0/1	-	=	4	4/4	1	0/1	-	-	1	0/1

*Cut-off levels to establish sensitivity was done based on Oxoid the FDA susceptivity test interprative criteria available at: www.oxoid.com.

As a limitation, this study may not represent general population that live in South Wollo area since only those who visited the health research laboratory either by referral or by patients own interest were included in the study. Only those antibiotics tested in the regional health research laboratory were included in the study. Thus, it may not include all antibiotics used in clinical practice. Being a retrospective study incomplete records and illegible hand writing eliminated some culture results and antibiotics from the study.

Conclusion

In conclusion, the *E. coli*, pseudomonas and proteus species were the three commonly isolated microorganisms in the regional health research laboratory. Moreover, most isolated bacterial microbes were resistant to antibiotics used in clinical practices in the country. This calls for attention of health professionals and policy makers to consider the resistance pattern in their clinical practice, and policy making process respectively. Most importantly, these data may be used to control

trends of antibiotic susceptibilities, to develop local antibiotic policies and to assist clinicians in the rational choice of antibiotic therapy and thus, to prevent indiscriminate use of antibiotics.

Competing interests

The authors declare that they have no competing interests.

Authors' contributions

AAA, AAD: design the study, collect the data, analysis and interpreted the data, write and reviewed the initial and final drafts of the manuscript. AGM: participated in the design of the study, data collection, review the manuscript. All authors read and approved the final manuscript.

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^{**}Abbreviations: T_t Total tests, %S percent of sensitive results; for those samples \leq 15 total sensitive/total tests (T^s/T_t) was used, TMP-SMX- trimetoprim-sulphamethoxazole, spps species.

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