Bacterial associated urinary tract infection, risk factors, and drug susceptibility profile among adult people living with HIV at Haswassa University Comprehensive Specialized Hospital, Hawassa, Southern Ethiopia

Netsanet Nigusse Tessema¹, Musa Mohammed Ali²* & Mengistu Hyilemeriam Zenebe²

People living with human immunodeficiency virus (HIV) are more likely to develop urinary tract infections (UTI) due to the suppression of their immunity. The aim of this study was to determine the prevalence, risk factors of UTI, and drug susceptibility pattern of bacteria isolated among peoples infected with HIV. A hospital-based cross-sectional study was conducted among 224 HIV positive individuals attending Hawassa University Comprehensive Specialized Hospital (HUCSH) from September 17 to November 16, 2018. Midstream urine was collected from all study participants and inoculated on to Blood and MacConkey agar. Bacterial isolates were characterized by Gram stain and standard biochemical tests. Kirby-Bauer method was used for antimicrobial susceptibility testing. Sociodemographic and clinical data were collected by a semi-structured questionnaire. Data were analyzed using SPSS version 20. A bivariate and a multivariable regression model were employed to determine the association between dependent and independent variables. From the total 224 study participants, 23 (10.3%) (95% CI 6.7–14.7) had culture-confirmed UTIs. The distributions of the bacteria were as follows: *Escherichia coli* 16 (69.6%), *Staphylococcus aureus* 2 (8.7%), *Klebsiella pneumoniae* 2 (8.7%), *Enterobacter aerogenes* 2 (8.7%) and *Pseudomonas species* 1 (4.3%). UTI prevalence was also high among study participants with a previous history of UTI and CD4+ count < 200/mm³. Female study participants were about five times more likely to have UTI (AOR 5.3, 95% CI 1.5–19.2). Ninety-three percent of bacteria isolated were susceptible to nitrofurantoin, ceftriaxone, and gentamycin; 87.5% were susceptible to meropenem and norfloxacin; whereas 93.8%, 68.8%, and 62.5% of isolates were resistant to ampicillin, tetracycline, and cotrimoxazole respectively. Multidrug resistance (MDR) was seen in 18 (78.3%) of bacterial isolates.

**Abbreviations**
PDR Proliferative diabetic retinopathy  
DME Diabetic macular edema  
ADED Advanced Diabetic Eye Disease  
LMIC Low and middle-income countries  
PRP Panretinal photocoagulation  
NVE Retinal neovascularization

¹Hawassa University Comprehensive Specialized Hospital, Awassa, Ethiopia. ²School of Medical Laboratory Science, Hawassa University College of Medicine and Health Sciences, Awassa, Ethiopia. *email: ysnmss@yahoo.com
Urinary tract infection (UTI) is an important health problem that can affect HIV positive individuals. UTI is defined as the presence of significant number of bacteria (greater than or equal to $10^5$) in the urine. Individuals who are infected with Human Immunodeficiency Virus (HIV) are at increased risk of acquiring UTI because of several reasons, mainly due to suppression of immune system. UTI can be asymptomatic and asymptomatic. Among HIV infected individuals, asymptomatic UTI can progress to symptomatic UTI characterized by mild irritation during voiding to bacteremia, sepsis, and death.

Urinary tract infections are mostly caused by bacteria and it may progress to blood infection and pyelonephritis in individual with some underlying risks. UTI infection may lead to hospitalization of HIV infected patients. HIV infected population are disproportionately affected by UTI as result of immune suppression. Several bacteria and some fungus cause UTI among HIV-infected persons. However, most infections are caused by enteric pathogens such as *Escherichia coli*. There are few studies from Ethiopia: Jimma, Gondar, Harar, and two from Addis Ababa that reported UTI among HIV patients.

Urinary tract infections are a serious public healthcare problem that causes a high economic burden to the country and decreases the quality of life. Worldwide there are about 150 million individuals who are diagnosed with UTI. UTI affects HIV infected population considerably. HIV positive individuals, whose CD4 cell is less than 300 cells/mm$^3$ are commonly affected by UTI. According to study from Tanzania, a CD4$^+$ cell count less than 200 per microliter was significantly associated with UTI among HIV positive individuals. A report from Europe also indicated HIV infected individuals with a CD4$^+$ cell count less than 200/mm$^3$ are more likely to experience UTI. UTI commonly affect females than males among general population and also among HIV infected population. Management of other non-HIV associated diseases including UTI in people living with HIV has become increasingly important. Additionally, it is complicated by emergence of antibiotic resistant bacteria making treatment of UTI challenging. In these regard, UTI co-infection with HIV is becoming a major challenge and leads to additional costs.

There are scarce data regarding the causative agents of UTI among HIV positive individuals from the study area; therefore, this study was conducted to determine the prevalence of bacterial UTI, risk factors and drug susceptibility pattern of isolates among adult people living with HIV attending HUCSH, Hawassa Ethiopia.

**Methods**

**Study area.** Hawassa is the capital city of the south region and located 275 km from Addis Ababa, the capital city of Ethiopia. Hawassa University Comprehensive Specialized Hospital was established in November 2005 and it serves about 12 million peoples. Patients seeking medical care receive services at different outpatient and inpatient units (surgery, gynecology and obstetrics, internal medicine, pediatrics, ophthalmology, psychiatry, radiology, pathology). The hospital also provides different services for individuals living with HIV in the ART clinic. The total number of HIV infected individuals attending HUCSH was 2,856. In government hospital of Ethiopia including the study area, HIV infection is diagnosed by using repaid serological tests. The drugs used to the treatment of HIV infected individuals are: Zidovudine, Lamivudine, Abacavir, Efavirenz, Nevirapine, Lopinavir/ritonavir, TDF (Tenofovir Disoproxil fumarate), Dolutegravir, Atazanavir. Previously the treatment was initiated when CD4$^+$ cell count is less than 350/mm$^3$ but currently it the treatment is given based on viral load (> 1,000 copies/µl).

**Study design and period.** A Hospital-based cross-sectional study was conducted from September 17 to November 16, 2018.

**Source population.** All adult people living with HIV and those who attended HUCSH, ART clinic during the study period for follow up.

**Study population.** HIV positive adults (both symptomatic and asymptomatic for UTI) were selected by systematic random sampling method were the study population.

**Eligibility.** Individual aged greater or equal to 18 years and HIV positive were included in the study. Individuals on antibiotic therapy for the last 2 weeks prior to data collection were excluded.

**Variables of the study.** Dependent variables include the prevalence of UTI, drug susceptibility pattern; Independent variables include age, Sex, marital status, occupational status, educational level, clinical signs and symptoms of UTI, previous history of UTI, previous history of catheterization, cotrimoxazole usage, CD4$^+$ cell count.

**Sample size determination.** The required sample size was determined by using single population formula considering the following assumptions: A prevalence of 15.8% from the previous study conducted in Ethiopia.
and margin of error (d) 5%, 95 confidence interval, and 10% non-response rate. Based on the above assumption the total sample size was 224.

Operational definition. Urinary tract infection: is the presence of pathogenic microorganisms within the urinary tract in a significant quantity (≥ 10^5 cfu/ml)\(^2\). Multi-drug resistance bacteria: are bacteria resistant for greater than two different classes of drug categories\(^3\).

Sampling technique. To recruit study participants we used a systematic random sampling method. We followed patient flow at ART clinic for one week. The average patient flow per day was 22 and the data collection period was 2 months. By dividing the sample size (N = 224) for the data collection period (48 days) we arrived at the sample size that could be collected per day, which is equal to 5. K value was calculated by dividing the average number of participants per day (n = 22) to participants recruited by day (n = 5), K was 4. By using a lottery method one participant was selected from the 1st 4 attendants then systematically every 4 participants were selected until the required sample size was obtained.

Data collection. Sociodemographic, associated factors and clinical data were collected by attending nurses using semi-structured questionnaire. The participants' current CD4^+ cell value was taken from their medical records (It was performed by using BD FACSPresto).

Sample collection. A urine sample was collected after adequate explanation/information was provided by attending laboratory professionals. Participants were instructed to collect about 30 ml of midstream urine (MSU) for microbiological examination by giving a sterile, dry, wide-necked, leak-proof container. Urine samples were processed immediately at HUCSH microbiology laboratory. If there were a delay the samples were stored in the refrigerator at 2–8 °C.

Urine culture and biochemical test. Using a calibrated loop 0.001 ml of well-mixed un-centrifuged urine was inoculated on blood and MacConkey agar. The inoculated media were incubated at 35–37 °C for 24 h and examined for the growth of bacteria\(^2\). For identification of Gram negative bacteria, bacterial colony was sub-cultured onto nutrient broth. Then, the nutrient broth was inoculated on biochemical test culture medias such as Triple sugar iron agar, Simon's citrate agar, Lysine iron agar, Urea, Motility tests, and Indol. Identification of species was done by their characteristics in the respective culture media as per the standard. Species identification for Gram positive bacteria was carried out using catalase and coagulase test\(^2\).

Antimicrobial susceptibility testing. Drug susceptibility testing was done by following the Kirby-Bauer disk diffusion method on Mueller–Hinton agar (Oxoid Ltd, Hampshire, UK)\(^2\). The tested drugs includes ampicillin (10 µg), ciprofloxacin (5 µg), cotrimoxazole (23.75 µg), gentamicin (10 µg), meropenem (10 µg), nitrofurantoin (300 µg), augmentin (20 µg), ceftriaxone (30 µg), norfloxacin (10 µg), cefazidime (30 µg), tetracycline (30 µg), clindamycin (2 µg), penicillin (10 µg), erythromycin (15 µg), and cefoxitin (30 µg). Briefly, using a sterile wire loop, 3–5 pure similar colonies were mixed in 5 ml of normal saline until the turbidity of the suspension matches 0.5 McFarland standards. By using a dry, sterile cotton swab, a portion of suspension was inoculated on the surface of the Mueller Hinton agar plate (MHA). Selected antibiotics disks were placed on MHA by using forceps and incubated at 37 °C for 18–24 h. The zone of inhibition (diameter) around the antibiotic disk was measured; bacteria were classified as susceptible, intermediate and resistant according to Clinical Laboratory Standard Institute (CLSI) guidelines 2019\(^2\).

Data quality assurance. To ensure the quality of sociodemographic and clinical data, the semi-structured questionnaire was pretested and data collectors were trained. Sterility of culture media was checked by incubating 5% of culture media overnight at 35–37 °C without specimen inoculation. The performance of culture media was checked by using control strains. Any physical changes like cracks, excess moisture, color, hemolysis, dehydration, and contamination was assessed and expiration date was also checked. Standard strains of E. coli (ATCC 25922) and S. aureus (ATCC 25923) were used as quality control throughout the study for culture and antimicrobial susceptibility tests.

Data processing and analysis. The data was analyzed using SPSS version 20. The bivariate regression model was employed to examine the associations between dependent and independent variables. Based on the bivariate analysis a variables with P value ≤ 0.25 were selected for further analysis using a multivariable regression model. A P value < 0.05 was considered as statistically significant and results were presented by using odds ratio and 95% level of confidence.

Results Sociodemographic characteristics. In the current study, a total of 224 people with HIV participated with zero no response rates. The age of the participant ranged from 18 to 59 years with a mean and median age of 39 years. The majority of the study participants were females 131 (58.5%). Most of the participants belong to the age category 28–37 years. The majority of study participants completed elementary school (Table 1).

Prevalence of Urinary tract infection among HIV positive individuals. Out of the 224 study participants, bacteria were isolated from 23 giving an overall prevalence of 10.3% (95% CI 6.7–14.7) UTI. Symp-
tomatic and asymptomatic UTI was 11 (4.9%) and 12 (5.4%) respectively. Twenty-one (91.3%) of isolates were Gram-negative bacteria. *E. coli* was the most predominant isolates. The proportion of bacteria isolated is as follows: 16 (69.6%) *E. coli*, 2 (8.7%) *K. pneumoniae*, 2 (8.7%) *E. aerogenes*, 2 (8.7%) *S. aureus* and 1 (4.3%) *Pseudomonas* species.

**Factors associated with the prevalence of UTI among HIV positive individuals.** Based on bivariate regression analysis variables with *P* value ≤ 0.25 (sex, age and marital status) were selected for further multivariable regression analysis and female study participants showed at least five times more likely to have significant bacteriuria (AOR 5.3; 95% CI 1.5, 19.2) when compared to male study participants (*P* = 0.012). However, there was no statistically significant association with age and marital status (*P* > 0.05) (Table 2).

Based on bivariate logistic analysis variables with *P* value ≤ 0.25 such as fever, diabetics, previous history of UTI, previous history of catheterization and CD4 count were further selected for multivariable logistic analysis. Previous history of UTI (AOR 4.4; 95% CI: 1.6, 11.7) and CD4 count less than 200 (AOR 4.9; 95% CI 1.2, 18.5) were significantly associated with UTI. However, there was no statistically significant association between UTI and fever, diabetes and previous history of catheterization (*P* > 0.05) (Table 3).

**Antimicrobial susceptibility profile.** From the total of *E. coli* (n = 16) isolated in this study, 15 (93.8%), 11 (68.8%), 10 (62.5%) were resistant to ampicillin, tetracycline and to cotrimoxazole respectively. 16 (100%) of *E. coli* were susceptible to nitrofurantoin and ceftriaxone (Table 4).

**Multi-drug resistance pattern of isolated bacteria.** None of the isolates was susceptible or resistant to all the drugs in the testing panel. Among the total isolates (n = 23), 18 (78.3%) were MDR. From 21 g negative bacterial isolates 16 (76.2%) showed MDR of these, 12 (57.1%) *E. coli* were MDR (Table 5).

| Characteristics | Frequency | Percent (%) |
|-----------------|-----------|-------------|
| **Sex**         |           |             |
| Male            | 93        | 41.5        |
| Female          | 131       | 58.5        |
| **Age**         |           |             |
| 18–27           | 15        | 6.7         |
| 28–37           | 85        | 37.9        |
| 38–47           | 77        | 34.4        |
| 48–57           | 39        | 17.4        |
| ≥ 58            | 8         | 3.6         |
| **Residence**   |           |             |
| Urban           | 218       | 97.3        |
| Rural           | 6         | 2.7         |
| **Marital status** |          |             |
| Married         | 103       | 46          |
| Single          | 43        | 19.2        |
| Divorced        | 45        | 20.1        |
| Widowed         | 33        | 14.7        |
| **Occupational status** |     |             |
| Employee        | 52        | 23.2        |
| Daily laborer   | 42        | 18.8        |
| Merchant        | 66        | 29.5        |
| Housewives      | 53        | 23.7        |
| Others*         | 11        | 4.9         |
| **Educational status** |   |             |
| University graduate | 31    | 13.8        |
| High school     | 60        | 26.8        |
| Elementary      | 96        | 42.9        |
| Illiterate      | 37        | 16.5        |

Table 1. Sociodemographic characteristics of people living with HIV who were attending Hawassa University Comprehensive Specialized Hospital ART clinic, Hawassa, Ethiopia, from September 17 to November 16, 2018 (n = 224). *Others: driver, student and farmer.*
Discussion

The overall prevalence of UTI among HIV positive individuals in the current study was 10.3%. The result of this finding is consistent with other studies carried out in Gondar, Ethiopia (10.7%)\(^{12}\) and Jimma, Ethiopia (12%)\(^{11}\). While a high prevalence of UTI was recorded from India (77.5%)\(^{27}\), South Africa (48.7%)\(^{19}\), Warsaw (23.2%)\(^{5}\), and Nigeria (21.1%)\(^{6}\). On the other hand, a low prevalence of 5.8% was recorded from Jos metropolis, Nigeria\(^{10}\). This difference may be due to the difference in sample size, the degree of the immune status of the study participants, ART use and geographical variation. Even if CD4 cell count rise as a result of initiation of HIV treatment other factors such as old age, other chronic disease can increase the risk of UTI.

In this study, HIV infected females had about 5 times the chance of developing UTIs compared to HIV infected males (\(P = 0.012\)). The finding of this study is in line with reports from other parts of Ethiopia. A study from Jimma, Ethiopia reported a high prevalence of UTI among females than males HIV positive individuals\(^{11}\). According to a study from Addis Ababa, Ethiopia HIV infected female study participants were three times more likely to have significant bacteriuria\(^{14}\). Additionally, a study from Gondar found a high prevalence of UTI among females\(^{12}\). Evidence from various epidemiological studies showed that UTIs were more common in females than in males\(^{2-4}\). High prevalence of UTI among female participants may be due to females have shorter and wider urethra, lack of prostatic fluid, and having moist urethra. Additionally, mechanical introduction of pathogens into the bladder and trauma increase the risk of UTI among females irrespective of their HIV serostatus\(^{3,29}\). But the finding of the current study is not comparable to the study conducted in Nigeria that reported a high prevalence of UTI in males than females\(^{30}\).

In this study, HIV infected females had about 5 times the chance of developing UTIs compared to HIV infected males (\(P = 0.012\)). The finding of this study is in line with reports from other parts of Ethiopia. A study from Jimma, Ethiopia reported a high prevalence of UTI among females than males HIV positive individuals\(^{11}\). According to a study from Addis Ababa, Ethiopia HIV infected female study participants were three times more likely to have significant bacteriuria\(^{14}\). Additionally, a study from Gondar found a high prevalence of UTI among females\(^{12}\). Evidence from various epidemiological studies showed that UTIs were more common in females than in males\(^{2-4}\). High prevalence of UTI among female participants may be due to females have shorter and wider urethra, lack of prostatic fluid, and having moist urethra. Additionally, mechanical introduction of pathogens into the bladder and trauma increase the risk of UTI among females irrespective of their HIV serostatus\(^{3,29}\). But the finding of the current study is not comparable to the study conducted in Nigeria that reported a high prevalence of UTI in males than females\(^{30}\).

In this study, participants with the previous history of UTI were about 4 times more likely to develop UTI (\(P = 0.004\)). This finding agrees with the studies conducted in Gondar, Ethiopia\(^{12}\) and Addis Ababa, Ethiopia\(^{14}\). This might be due to the presence of resistant strains from those who had the previous history of UTI.

Urinary tract infections appear to be multifactorial in patients with HIV infections as CD4+ level declines\(^{13}\). In the current study, the distribution of UTI according to CD4+ count showed that study participants with CD4+ count < 200/mm\(^3\) had a chance of 4.9 times to develop UTI (\(P = 0.017\)). This finding was supported by studies

| Variables          | UTI |  |  |  |  |  |
|--------------------|-----|---|---|---|---|---|
| Sex                |     | Yes (n (%)) | No, n (%) | COR (95% CI) | P value | AOR (95% CI) | P value |
| Male               | 3 (3.2) | 90 (96.8) | 1 | 1 | 0.024 | 1 | 5.3 (1.5, 19.2) | 0.012 |
| Female             | 20 (15.3) | 111 (84.7) | 5.5 (1.3, 21.1) | 0.123 |
| Age                |     |  |  |  |  |  |
| 18–27              | 3 (20) | 12 (80) | 1 | 0.605 |
| 28–37              | 6 (7.1) | 79 (92.9) | 0.3 (0.1, 1.4) | 0.123 |
| 38–47              | 6 (7.8) | 71 (92.2) | 0.3 (0.1, 1.5) | 0.161 |
| 48–57              | 7 (17.9) | 32 (82.1) | 0.9 (0.2, 3.4) | 0.862 |
| ≥ 58               | 1 (12.5) | 7 (87.5) | 0.6 (0.1, 6.6) | 0.654 |
| Residence          |     |  |  |  |  |  |
| Urban              | 22 (10.1) | 196 (89.9) | 1 | 0.605 |
| Rural              | 1 (16.7) | 5 (83.3) | 1.8 (0.2, 15.9) | 0.123 |
| Educational status |     |  |  |  |  |  |
| University graduate | 2 (6.5) | 29 (93.5) | 1 | 0.750 |
| High school        | 5 (8.3) | 55 (91.7) | 1.3 (0.2, 7.2) | 0.299 |
| Elementary         | 13 (13.5) | 83 (86.5) | 2.3 (0.5, 10.7) | 0.795 |
| No formal education | 3 (8.1) | 34 (91.9) | 1.3 (0.2, 8.2) | 0.195 |
| Marital status     |     |  |  |  |  |  |
| Married            | 10 (9.7) | 93 (90.3) | 1 | 0.956 |
| Single             | 2 (4.7) | 41 (95.3) | 0.5 (0.2, 2.2) | 0.956 |
| Divorced           | 5 (11.1) | 40 (88.9) | 1.2 (0.4, 3.6) | 0.663 |
| Widowed            | 6 (18.2) | 27 (81.8) | 2.1 (0.7, 6.2) | 0.972 |
| Occupation         |     |  |  |  |  |  |
| Government employee | 6 (11.5) | 46 (88.5) | 1 | 0.999 |
| Daily laborer      | 5 (11.9) | 37 (88.1) | 1.0 (0.3, 3.7) | 0.663 |
| Merchant           | 6 (9.1) | 60 (90.9) | 0.8 (0.2, 2.5) | 0.972 |
| House wife         | 6 (11.3) | 47 (88.7) | 0.9 (0.3, 3.3) | 0.972 |
| Others\(^{a}\)      | 0 | 11 (100) | 0.0 (0.0, 0.0) | 0.999 |

Table 2. Bivariate and multivariate analysis of sociodemographic characteristics and UTI among people living with HIV who were attending Hawassa University Comprehensive Specialized Hospital ART clinic, Hawassa, Ethiopia, from September 17 to November 16, 2018 (n = 224). UTI urinary tract infection, COR crude odd ratio, AOR adjusted odd ratio, CI confidence interval, n number. \(^{a}\) Others: driver, student and farmer.
conducted in Ethiopia\textsuperscript{13,15}, Nigeria\textsuperscript{16} and India\textsuperscript{30}. These results imply that as CD4\textsuperscript{+} value declines the risk of UTI increases. There was no significant association between age, residence, educational status, occupation, and marital status with UTI in this study (\(P > 0.05\)).

In the current study, 91.3% of UTI was caused by Gram negative bacteria. We have noted that non-typical bacteria among UTI in our study. The predominant bacterium isolated in the current study was \textit{E. coli} (69.6%). A similar \textit{E. coli} predominance was reported from Jimma, Ethiopia (54.3%)\textsuperscript{11} and Gondar, Ethiopia (56.1%)\textsuperscript{12}. \textit{E. coli} predominance may be due to \textit{E. coli} is the most common microorganism in the vaginal and rectal area\textsuperscript{31}. In contrast, this study was inconsistent with the finding reported in Ebony State, Nigeria in which the predominant isolates were \textit{S. aureus} (45.33%)\textsuperscript{4}, Cape Coast, Ghana the predominant isolates were \textit{S. aureus} (40%) and \textit{S. saprophyticus} (21.8%)\textsuperscript{2}. Most of the isolates reported from Tamil Nadu, India were \textit{P. aeruginosa} (41.9%)\textsuperscript{27}. Our finding of \textit{K. pneumoniae} (8.7%), \textit{S. aureus} (8.7%) and \textit{Pseudomonas} species is greater than report from Warsaw\textsuperscript{5}. The variation in the type of bacterial isolate may be due to sample collection technique and personal and environmental hygiene, and underlying conditions\textsuperscript{13}.

In the present study, 80% of Gram negative bacteria were susceptible to ciprofloxacin, gentamycin, nitrofurantoin, and norfloxacin. The finding of this study is similar to findings reported from other areas\textsuperscript{4,12,14}. Whereas 95.2% Gram negative bacteria in this study showed resistance to ampicillin and 57.1% of them were resistant to cotrimoxazole and 69.9% were resistant to tetracycline. Among Gram-negatives isolates, 93.8% of \textit{E. coli} demonstrated resistance to ampicillin followed by tetracycline (68.8%) and co-trimoxazole (62.5%). Whereas, all isolates of \textit{E. coli} were susceptible to ceftriaxone and nitrofurantoin followed by gentamicin (93.8%), ciprofloxacin and norfloxacin (87.5%) each and augumentin (68.8%). All \textit{K. pneumoniae} were resistant to ampicillin.

| Variables                        | UTI      | Fever  | No, n (%) | Yes, n (%) | COR (95% CI) | \(P\) value | AOR (95% CI) | \(P\) value |
|----------------------------------|----------|--------|-----------|------------|--------------|-------------|-------------|-------------|
|                                  |          | No     | 18 (8.6)  | 191 (91.4) | 1            | 0.005       |             |             |
|                                 |          | Yes    | 5 (33.3)  | 10 (66.7)  | 5.3 (1.6, 17.2) |             |             |             |
| Dysuria                          |          | No     | 22 (10.7) | 183 (89.3) | 1            | 0.463       |             |             |
|                                 |          | Yes    | 1 (5.3)   | 18 (94.7)  | 0.5 (0.1, 3.6) |             |             |             |
| Diabetics                        |          | No     | 20 (9.5)  | 191 (90.5) | 1            | 0.132       |             |             |
|                                 |          | Yes    | 3 (23.1)  | 10 (76.9)  | 2.9 (0.7, 11.3) |             |             |             |
| Cytomyclose usage as prophylaxis |          | Yes    | 5 (12.5)  | 35 (87.5)  | 1            | 0.609       |             |             |
|                                 |          | No     | 18 (9.8)  | 166 (90.2) | 0.8 (0.3, 2.2) |             |             |             |
| Frequency of urination           |          | No     | 21 (10.8) | 174 (89.2) | 1            |             |             |             |
|                                 |          | Yes    | 2 (6.9)   | 27 (93.1)  | 0.6 (1.2, 2.8) | 0.525       |             |             |
| Urgency of urination             |          | No     | 22 (10.8) | 181 (89.2) | 1            |             |             |             |
|                                 |          | Yes    | 1 (4.8)   | 20 (95.2)  | 0.4 (0.5, 3.2) | 0.397       |             |             |
| Flank pain                       |          | No     | 21 (1.2)  | 166 (88.8) | 1            |             |             |             |
|                                 |          | Yes    | 2 (5.4)   | 35 (94.6)  | 0.5 (0.1, 2.0) | 0.298       |             |             |
| Renal stone                      |          | No     | 22 (10.0) | 197 (90)   | 1            |             |             |             |
|                                 |          | Yes    | 1 (20)    | 4 (80)     | 2.2 (0.2, 20.9) | 0.480       |             |             |
| Previous history of UTI          |          | No     | 14 (7.7)  | 169 (92.3) | 1            |             |             | 0.004       |
|                                 |          | Yes    | 9 (22)    | 32 (78)    | 0.3 (0.1, 0.7) | 0.009       | 4.4 (1.6, 11.7) |             |
| History of catheterization       |          | No     | 20 (9.20) | 198 (90.8) | 1            |             |             |             |
|                                 |          | Yes    | 3 (50)    | 3 (50)     | 9.9 (1.9, 52.3) | 0.007       |             |             |
| CD4 count                        |          | \(\geq 200\) | 17 (8.30) | 188 (91.7) | 1            |             |             | 0.017       |
|                                 |          | \(< 200\) | 6 (31.6)  | 13 (68.4)  | 5.1 (1.7, 15.1) | 0.003       | 4.9 (1.2, 18.5) |             |

Table 3. Bivariate and multivariate analysis of clinical characteristics and UTI among people living with HIV who were attending Hawassa University Comprehensive Specialized Hospital ART clinic, Hawassa, Ethiopia, from September 17 to November 16, 2018 (\(n = 224\)). AOR adjusted odds ratio, COR crude odd ration, AOR adjusted odd ratio, CI confidence interval, n number.
and augmentin and all of them were susceptible to co-trimoxazole, gentamycin, ceftazidime, nitrofurantoin, and norfloxacin. All S. aureus isolated in the current study were susceptible to ciprofloxacin, norfloxacin, gentamycin, penicillin, whereas, all of them were resistant to co-trimoxazole and tetracycline.

Generally, ciprofloxacin, ceftriaxone, gentamicin, nitrofurantoin, and norfloxacin were most effective for bacteria isolated in the current study while ampicillin, cotrimoxazole and tetracycline were less effective. This agrees with the finding from Ethiopia\textsuperscript{14} and South Africa\textsuperscript{19}.

In this study, MDR was observed among 78.3% of the isolated bacteria. This was higher compared to the finding reported in Mysore, India (58.3%)\textsuperscript{7} and Bangalre, India (48.55%)\textsuperscript{16}. But it was lower than a report from Gondar, Ethiopia (95%)\textsuperscript{12} and Portharcourt, Nigeria (92.8%)\textsuperscript{18}. The high prevalence of MDR seen to commonly prescribed antibiotics in this study might be due to the easy availability of drugs in the community, and inappropriate use of antimicrobial agents.

**Conclusion**

In the current study, the overall prevalence of UTI among people living with HIV was 10.3%. Factors such as sex, CD4\textsuperscript{+} count < 200/mm\textsuperscript{3}, and previous history of UTI were significantly associated with the prevalence of UTI. The isolated bacteria were E. coli, K. pneumoniae, S. aureus, E. aurogenes, and Pseudomonas spp. E. coli was the

---

**Table 4.** Antimicrobial susceptibility pattern of Gram-negative and Gram positive bacteria isolated from urine culture of people living with HIV attending Hawassa University Comprehensive Specialized Hospital ART clinic, Hawassa, Ethiopia, from September 17 to November 16, 2018 (N = 23).

| Bacterial isolates (n) | Pattern | AMP, n (% n) | CIP, n (%) | COT, n (%) | GN, n (%) | MER, n (%) | NIT, n (%) | AUG, n (%) | CTR, n (%) | NOR, n (%) | CAZ, n (%) | TET, n (%) | CLN, n (%) | PEN, n (%) |
|------------------------|---------|-------------|-----------|-----------|---------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|
| E. coli (16)           | S       | –           | 14 (87.5) | 6 (37.5)  | 15 (93.8)| 14 (87.5) | 16 (100)  | 11 (68.8) | 16 (100)  | 14 (87.5) | 8 (50)    | 11 (68.8) | NA        | NA        |
|                        | I       | 1 (6.3)     | 1 (6.3)   | –         | –        | –         | –         | –         | –         | –         | 1 (6.3)   | NA        | NA        | NA        |
|                        | R       | 15 (93.8)   | 1 (6.3)   | 10 (62.5) | 1 (6.3)  | 2 (12.5)  | –         | 5 (31.3)  | –         | 1 (6.3)   | 8 (50)    | 11 (68.8) | NA        | NA        | NA        |
| K. pneumoniae (2)      | S       | –           | 2 (100)   | 2 (100)   | 2 (100)  | 2 (100)   | –         | 1 (50)    | 2 (100)   | 2 (100)   | 1 (50)    | NA        | NA        | NA        |
|                        | I       | –           | –         | –         | –        | –         | –         | –         | –         | –         | –         | 1 (50)    | NA        | NA        | NA        |
|                        | R       | 2 (100)     | –         | –         | –        | –         | –         | –         | –         | –         | –         | –         | NA        | NA        | NA        |
| E. aurogenes (2)       | S       | –           | 2 (100)   | 1 (50)    | 2 (100)  | 2 (100)   | 1 (50)    | 1 (50)    | 2 (100)   | 1 (50)    | 1 (50)    | 1 (50)    | 1 (50)    | 1 (50)    |
|                        | I       | –           | –         | –         | –        | –         | 1 (50)    | 1 (50)    | –         | 1 (50)    | –         | 1 (50)    | NA        | NA        | NA        |
|                        | R       | 2 (100)     | –         | 1 (50)    | –        | –         | –         | –         | –         | 1 (50)    | –         | NA        | NA        | NA        | NA        |
| Pseudomonas spp. (1)   | S       | –           | 1 (100)   | –         | 1 (100)  | –         | –         | 1 (100)   | 1 (100)   | –         | 1 (100)   | NA        | NA        | NA        |
|                        | I       | –           | –         | –         | –        | –         | –         | –         | –         | –         | –         | NA        | NA        | NA        |
|                        | R       | 1 (100)     | –         | 1 (100)   | –        | –         | –         | –         | 1 (100)   | –         | 1 (100)   | NA        | NA        | NA        |
| S. aureus (2)          | S       | –           | 2 (100)   | –         | 2 (100)  | –         | –         | 2 (100)   | NA        | 2 (100)   | NA        | 2 (100)   | 2 (100)   | 2 (100)   |
|                        | I       | NA          | –         | –         | –        | –         | –         | –         | –         | –         | –         | –         | –         | –         |
|                        | R       | 2 (100)     | –         | –         | NA       | NA        | –         | NA        | NA        | –         | –         | –         | –         | –         |

**Table 5.** Multi-drug resistance pattern of bacterial isolates from urine culture of adult people living with HIV attending Hawassa University Comprehensive Specialized Hospital ART clinic, Hawassa, Ethiopia, from September 17–November 16, 2018 (N = 23).

| Bacterial isolate | Total n (%) | R1 | R2 | R3 | R4 | R5 | R6 | MDR n (%) |
|-------------------|-------------|----|----|----|----|----|----|-----------|
| Gram negative     | 21 (91.3)   | 1  | 4  | 6  | 2  | 6  | 2  | 16 (76.2) |
| E. coli           | 16 (76.2)   | 1  | 3  | 5  | 1  | 5  | 1  | 12 (57.1) |
| K. pneumoniae     | 2 (9.5)     | –  | 1  | 1  | –  | –  | –  | 2 (9.5)   |
| E. aurogenes      | 2 (9.5)     | –  | 1  | –  | –  | –  | 1  | 1 (4.8)   |
| Pseudomonas spp.  | 1 (4.8)     | –  | –  | –  | 1  | –  | 1  | 1 (4.8)   |
| Gram positive S. aureus | 2 (8.7) | –  | –  | –  | 2 | – | –  | 2 (100) |
|                   | 2 (100)     | –  | 2  | 2  | 2  | 2  | 2  | 2 (100) |
| Total             | 23 (100)    | 1  | 4  | 8  | 2  | 6  | 2  | 18 (78.3) |
predominant bacteria. Most of the bacterial isolates were susceptible to ciprofloxacin, ceftriaxone, gentamicin, nitrofurantoin, and norfloxacin. Most of the isolates were resistant to ampicillin, tetracycline and co-trimoxazole. Multi-drug resistant bacteria were common in the current study. As susceptibility of bacteria to various antibiotics vary, management of UTI among HIV infected individuals should be guided by antimicrobial susceptibility testing.

Ethics approval and consent to participate. Ethical clearance was obtained from the institutional review board (IRB) of Hawassa University College of medicine and health science. Then support letter were obtained from the hospital administration. Written informed consents were obtained from each study participants. All methods were carried out in accordance with relevant guidelines and regulation as mentioned by Declaration of Helsinki.

Data availability
All relevant data are available within the paper.

Received: 21 March 2020; Accepted: 10 June 2020
Published online: 01 July 2020

References
1. Ranjan, A., Sridhar, S. T. K., Matta, N., Chokkakkula, S. & Ansari, R. K. Prevalence of UTI among pregnant women and its complications in newborns. Indian J. Pharm. Pract. 10, 45 (2017).
2. Barnie, P., Akwetey, S., Swallah, M., Acheampong, D. & Kwakye-Nuako, G. Occurrence and distribution of bacterial uropathogens among antiretroviral therapy users and non-users, Cape Coast Teaching Hospital (2019).
3. Agersew, A. & Chandrashekhar, U. Prevalence and antimicrobial susceptibility pattern of urinary tract infection causing human pathogenic bacteria among symptomatic outpatients visiting Gondar University hospital Gondar, Northwest Ethiopia. Novus Int. J. Med. Sci. 2, 1–14 (2013).
4. Ifeanyi, I. et al. Frequency and antiobiotic of uropathogens isolated from urine samples of HIV infected patients on antiretroviral therapy. Afr. J. Biomed. Sci. 50, 50–53 (2013).
5. Skrzat-Klapaczynska, A. et al. Factors associated with urinary tract infections among HIV-1 infected patients. PLoS ONE 13(1), e0190564 (2018).
6. Olowe, O. A., Ojo-Johnson, B. B., Makanjuola, O. B., Olowe, R. A. & Mabayoje, V. O. Detection of bacteriuria among human immunodeficiency virus seropositive individuals in Osogbo, south-western Nigeria. Eur. J. Microbiol. Immunol. 3, 126–130 (2015).
7. Muriyegh, K., Deepa, S., Ravindranath, C. & Venkatesha, D. Multi drug resistant uropathogens in HIV: Are they a threat to community?. Int. J. Sci. Study 2(3), 38–42 (2014).
8. Michael, I. O., Abel, O. & Ukoh, G. Urinary tract infection in adolescent/young adult nigerians with acquired human immunodeficiency disease in Benin City. J. Med. Biomed. Res. 5(2), 55–60 (2006).
9. Emmanuel, B. & Wakjissa, D. Prevalence of urinary tract infections among HIV patients attending a non-governmental health facility in Jos, Plateau State, Nigeria. Int. J. Biomed. Adv. Res. 4, 528 (2013).
10. Sheyn, Z. et al. Prevalence of urinary tract infection in HIV patients on antiretroviral drugs in Jos Metropolis, Nigeria. World J. Public Health 3(2), 57 (2018).
11. Debalkie, S., Cheneke, W., Tassew, H. & Awole, M. (2014) Urinary tract infection among antiretroviral therapy users and nonusers in Jimma University Specialized Hospital, Jimma, Ethiopia. Int. J. Microbiol. 2014, 968716 (2014).
12. Agersew, A., Mulat, D., Meseret, A. & Mucheye, G. Uropathogenic bacterial isolates and their antimicrobial susceptibility patterns among HIV/AIDS patients attending Gondar University Specialized Hospital Gondar, Northwest Ethiopia. J. Microbiol. Res. Rev. 1(4), 42–51 (2013).
13. Marami, D., Balakrishnan, S. & Seyoum, B. Prevalence, antimicrobial susceptibility pattern of bacterial isolates, and associated factors of urinary tract infections among HIV-positive patients at Hiwot Fana Specialized Hospital, Eastern Ethiopia. Can. J. Infect. Dis. Med. Microbiol. 2019, 6780354 (2019).
14. Getu, Y., Ali, I., Lema, T., Belay, H. & Yesheleta, B. Bacteriuria and antimicrobial susceptibility pattern among HIV patients attending ALERT Center, Addis Ababa Ethiopia. Am. J. Health Res. 5(3), 76–82 (2017).
15. Fenta, G. M., Legese, M. H. & Weldearegay, G. M. Bacteriuria and their antibiotic susceptibility patterns among people living with HIV attending Tikur Anbessa Specialized and Zewditu Memorial Hospital ART Clinics, Addis Ababa, Ethiopia. J. Bacteriol. Parasitol. 7(5), 1–7 (2016).
16. Yadav, K. & Samreen, S. Asymptomatic urinary tract infection with multidrug resistant pathogens in retro positive patients on ART. Int. J. Med. Res. Rev. 5, 610–615 (2017).
17. Kucheria, R., Dasgupta, P., Sacks, S. H., Khan, M. S. & Sheerin, N. S. Urinary tract infections: New insights into a common problem. Postgrad. Med. J. 81, 83–86 (2005).
18. Frank-Peterside, N., Okerentugba, P. O., Nwodo, C. R. & Okonko, I. O. Prevalence of bacterial uropathogens in a cohort of HIV/AIDS patients attending Gondar University Specialized Hospital Gondar, Northwest Ethiopia. Can. J. Infect. Dis. Med. Microbiol. 2017, 4042666 (2017).
19. Klasinc, R., Rieger, A., Presterl, E., Wbra, T. & Diab-Elsalahawi, M. Epidemiology of urinary tract infections in HIV-positive patients at a tertiary care hospital in central Europe (2011–2016). In The Congress of ECCMID (2017).
20. Akinbami, A. et al. Prevalence of asymptomatic bacteriuria in HIV infected patients in a tertiary hospital in Lagos, Nigeria. World J. AIDS 03, 105–110 (2013).
21. Nerurkar, A., Solanki, P. & Naik, S. S. Bacterial pathogens in urinary tract infection and antibiotic susceptibility pattern. J Pharm Biomed Sci. 21 (2012).
22. Basak, S., Singh, P. & Rajurkar, M. Multidrug resistant and extensively drug resistant bacteria. J. Pathog. 216, 4065035 (2015).
23. Cheshbrough, M. District Laboratory practice in Tropical Countries (Cambridge University Press, Cambridge, 2006).
24. CLSI. Performance Standards for Antimicrobial Susceptibility Testing 28th edn. CLSI supplement M100. (Clinical and Laboratory Institute, Wayne, PA, 2018).
25. Xavier, T. F., Auxilia, A. & Kannan, M. Isolation and characterization of UTI pathogens from HIV positive patients of Karur Medical Institute, Wayne, PA, 2018).
26. Nwadioha, S. I., Nwokezi, E., Ekei, I., Egie, J. & Nwadioha, S. I. Prevalence of asymptomatic bacteriuria in HIV infected patients in a tertiary hospital. J. Med. Med. Sci. 11(11), 530–534 (2010).
29. Wasihun, A., Araya, T., Legese, H., Geberemariam, G. & Weldu, Y. Bacteriological profile, risk factors and antimicrobial susceptibility patterns of symptomatic urinary tract infection among students of Mekelle University, northern Ethiopia. *BMC Infect. Dis.* **19**, 11 (2019).

30. Inyang-Etoh, P. C., Udofia, G. C., Anyanwu, A. & Ndifreke, U. Asymptomatic bacteriuria in patients on antiretroviral drug therapy in calabar. *J. Med. Sci.* **9**, 270–275 (2009).

31. Ali, J. & Gholamreza, I. Asymptomatic urinary tract infection in pregnant women. *Iran. J. Pathol.* **4**, 105–108 (2009).

**Acknowledgements**

We would like to acknowledge Staff of Microbiology Laboratory of Hawassa Comprehensive Specialized Hospital for their cooperation during laboratory work. We acknowledge staff of ART clinic for facilitating during sociodemographic and clinical data collection. We also acknowledge all study participants for their willingness to participate in the study.

**Author contributions**

N.T. Conceived designed the experiments, laboratory work, data analysis M.M.A. Conceived the experiment, supervision, review, analysis and manuscript preparation M.H.Z. Review, supervision. All authors have read and approved the manuscript.

**Competing interests**

The authors declare competing interest.

**Additional information**

**Correspondence** and requests for materials should be addressed to M.M.A.

**Reprints and permissions information** is available at [www.nature.com/reprints](http://www.nature.com/reprints).

**Publisher’s note** Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons license, and indicate if changes were made. The images or other third party material in this article are included in the article’s Creative Commons license, unless indicated otherwise in a credit line to the material. If material is not included in the article’s Creative Commons license and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this license, visit [http://creativecommons.org/licenses/by/4.0/](http://creativecommons.org/licenses/by/4.0/).

© The Author(s) 2020