Bacterial Profile And Antibiotic Susceptibility Pattern Of Urinary Tract Infection Among Children Attending Felege Hiwot Referral Hospital, Bahir Dar, Northwest Ethiopia

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Background: Urinary tract infection (UTI) is a common and important clinical problem in pediatrics. Recurrent UTIs may lead to renal scarring, hypertension, and end-stage renal dysfunction later in life. The objective of the study was to determine bacterial profile and antimicrobial susceptibility pattern of urinary tract infections (UTIs) among children attending Felege Hiwot Referral Hospital (FHRH).

Methods: A cross-sectional study was conducted from February 2013 to May 2013 among children 5–15 years of age with symptoms of UTI. Samples were processed for culture and identification. Antimicrobial susceptibility was done for positive urine cultures by the Kirby-Bauer’s disk diffusion method based on standards of the Clinical Laboratory Standard Institute (CLSI). Data were entered into Epi-data version 3.2.1 and exported to the Statistical Package for the Social Science (SPSS) version 20 statistical software. Fisher’s exact test and binary logistic regression test results were used.

Results: A total of 259 urine samples were collected from children with UTI. The result revealed 41 (15.8%) samples had significant bacteriuria, among which the most prevalent pathogen was E. coli 14 (34.1%) followed by Pseudomonas species. Gram-negative bacteria showed high level of sensitivity to ciprofloxacin (70), norfloxacin (63.4%) and ceftriaxone (60%), whereas the level of resistance was high to ampicillin (80%) and nitrofurantoin (70%). Gram-positive isolates showed high sensitivity to ciprofloxacin (77.8%), penicillin (72.8%) and erythromycin (72.7%). Multiple drug resistance (MDR) for Gram-positive and Gram-negative bacteria was 100% and 83.1%, respectively.

Conclusion: E. coli is the predominant bacteria isolated in the present study. The results showed that the prevalence of resistance to at least one antibiotic to commonly prescribed antimicrobials was high. Hence, the guidelines for empiric treatment of UTI should be re-evaluated periodically based on local studies.

Keywords: urinary tract infection, significant bacteriuria, antimicrobial susceptibility, children
Urinary tract infection is a common and important clinical problem in pediatrics, as recurrent UTIs may lead to renal scarring, hypertension and end-stage renal dysfunction later in life. UTIs account for 0.7% of physician office visits and 5–14% of emergency departments visits by children annually. Diagnosis in younger children may be difficult, however, older children often complain of urinary symptoms that are classically associated with UTI, such as dysuria, frequency, urgency and may present with suprapubic tenderness on examination. Since childhood UTI is the most important factor affecting adult health, children should be treated immediately with appropriate antibiotics.

Antimicrobial therapy is often initiated empirically before the urine culture results are available. Furthermore, UTIs are often treated with broad-spectrum antibiotics when one with a narrow spectrum of activity may be appropriate because of concerns about infection with resistant organisms. The etiology of UTI and the antibiotic resistance of uropathogens have been changing over the past years, both in community and nosocomial infection.

Given there are both geographical and historical variability of antimicrobial resistance patterns, the knowledge of the sensitivity pattern of common uropathogens according to local epidemiological studies is necessary for selection of an appropriate antibiotic for empirical treatment. Furthermore, studies recommended that the policies for treatment of UTI in children should be re-evaluated every 5 years according to local resistance rates.

There are studies on bacterial profile and antibiotic susceptibility pattern of UTI in different parts of Ethiopia, but these have been conducted mostly on adults and pregnant women. To our knowledge, there has been no study done around our targeted age group (age 5–15) focused on understanding the antimicrobial resistance pattern of pathogens are more frequently isolated in UTI locally. Consequently, studies on the assessment of bacterial profile and antibiotic susceptibility pattern of UTI in children in our geographical area (Bahir Dar) were limited. Hence, this study aimed to determine the bacterial profile and antibiotic susceptibility pattern of urinary tract infection among children attending at Felege Hiwot Referral Hospital, Bahir Dar, Northwest Ethiopia.

Methods And Materials

Study Setting And Period

The study was conducted from February 2013 to May 2013 at Felege Hiwot Referral Hospital, Bahir Dar, Amhara, Northwest Ethiopia. Bahir Dar is the capital city of Amhara region with total population of 155,428 according to 2007 census. The hospital includes inpatient, outpatient, and emergency wards. There are Surgical, Gynecological, Medical, Pediatric, Orthopedic and other departments in the hospital. All the procedures for isolation and antimicrobial susceptibility tests were performed at Amhara Regional Health Laboratory and Research Center. It is a well established and accredited laboratory center by Ethiopian drug administration and control authority.

Study Design And Participants

A cross-sectional study was conducted among randomly selected symptomatic UTI patients aged 5–15 years who presented to Felege Hiwot Referral Hospital. The sample size was determined using a single population proportion formula with the following assumption: proportion of symptomatic UTI in the population to be 21.5% margin of sampling error tolerated 5% (0.05), and critical value at 95% confidence interval of certainty (1.96), and 10% for non-response making the final sample size of 259 children.

Participants were selected consecutively among children’s visiting pediatric departments until required samples were obtained.

The study was cleared from the ethical review board of Addis Ababa University. Official recognition and support was secured from the relevant organizations and departments. The interviewees have also been informed about the aim of the study. Additionally, written and verbal consent were secured from the study participants and/or caregivers. Finally, confidentiality was ascertained by justifying that no information was disclosed individually without the full willingness of the respondent.

Data Collection Procedure

Clinical Examination

After obtaining an informed consent from children’s parent, socio-demographic data and clinical data were collected from patient chart using pre-structured questioners by nurses.

Specimen Collection

A freshly voided midstream urine sample (10–20 mL) was collected in a wide mouthed sterile container after instructing the parents or guardians of enrolled children to clean their genitals with soap and water.
Culturing And Identification Procedure
Urine specimens obtained from children were inoculated on blood, MacConkey and CLED agar (Oxoid, Ltd., Basingstoke, Hampshire, England) by using a calibrated loop (0.001mL). Cultures were incubated in aerobic atmosphere at 37°C for 24 hrs. A positive urine culture was defined as colony count ≥10^5 CFU/mL for midstream urine. A Stuart scientific colony counter was used for counting. All positive cultures were further identified by their colony characteristics, Gram staining was done to identify Gram positives from Gram negatives. Final confirmation was done by their pattern of biochemical reactions and rapid identification methods using the standard microbiological procedures.

Antimicrobial Susceptibility Testing
The antimicrobial susceptibility testing of all isolates was done by the standard Kirby-Bauer disk diffusion method,11 using commercial disks (Oxoid) according to Clinical and Laboratory Standards Institute (CLSI). Briefly, when pure culture was obtained, a loopful of bacteria was taken from a colony and transferred to a tube containing 5mL of normal saline and mixed gently until it forms a homogeneous suspension. Then, the turbidity of the suspension was adjusted to the density of a McFarland 0.5 (Mary-l’Etoil, France) in order to standardize the inoculum size. A sterile cotton swab was dipped into the suspension and the excess was removed by gentle rotation of the swab against the surface of the tube. The swab was then used to distribute the bacteria evenly over the entire surface of Mueller-Hinton agar (Oxoid). The inoculated plates were left at room temperature to dry for 3–5 mins. With the aid of sterile forceps, the following concentration of antibiotic discs were put on the surface of Mueller-Hinton agar (Oxoid): ampicillin (10µg), amoxicillin clavulanic acid (30µg), ciprofloxacin (5µg), gentamicin (10µg), ceftriaxone (30µg), nalidixic acid (30µg), nitrofurantoin (300µg), cefoxitin (1µg), trimethoprim-sulfamethoxazole (25µg) and ceftazidime (30µg). These antimicrobial agents were selected because of the frequent empirical treatment used by the clinicians. The plates were incubated at 37°C for 24 hrs. The result was interpreted as resistant, intermediate and sensitive based on CLSI guidelines. Multidrug resistance was defined as resistance to two or more of the antimicrobials tested. Positive results from urine culture and antimicrobial sensitivity test results were reported to the attending physician for subsequent treatment and follow up.

Data Processing And Analysis
Quantitative data were cleaned, edited and entered onto Epi-data version 3.2.1 and exported to the Statistical Package for the Social Sciences (SPSS) version 20 statistical software for further analysis. Fisher’s exact test and binary logistic regression test results were used. A P-value less or equal to 0.05 was employed to declare the statistical significance.

Results
Socio-Demographic Characteristics
Of the 259 children investigated, 83 (32%) were males and 176 (68%) were females. We observed a mean age of 12 with 179 (69.1%) of the study subjects between 11 and 15 years and the remaining were between 5 and 10 years. Majority of the study subjects were from rural areas 161 (62.2%) (Table 1).

Culture Results
Significant bacteriuria was observed in 41 of 259 (15.8%) urine samples cultured. Of these, 11 (26.8%) were Gram-positive and 30 (73.2%) were Gram-negative bacteria. Out of total positives, 9 (22%) were from males and 32 (78%) were from females (P=0.18). Twenty-eight (68.3%) of children with significant bacteriuria were between 11 and 15 years while 13 (31.7%) of them were between 5 and 10 years (P=0.9). Nine (21.9%) of children with significant bacteriuria have parents with previous history of UTI and the rest 32 (79.1%) did not have a parent with a history of UTI. Interestingly, there was statistical significant association between the rate of UTI and previous history of UTI in children’s parents (p<0.05). (Table 2).

Table 1 Sociodemographic Characteristics Of The Study Subjects Investigated For UTI In Felege Hiwot Referral Hospital (February 2013 To May 2013)

| Variables | Frequency | Percentage |
|-----------|-----------|------------|
| Sex       |           |            |
| Male      | 83        | 32         |
| Female    | 176       | 68         |
| Age       |           |            |
| 5–10      | 80        | 30.9       |
| 11–15     | 179       | 69.1       |
| Residence |           |            |
| Urban     | 98        | 37.8       |
| Rural     | 161       | 62.2       |
Table 2 The Isolation Rate Of UTI In Children Attending Felege Hiwot Referral Hospital In Relation With Their Associated Risk Factors And Gram Stain (February 2013 To May 2013)

| Category                                   | Significant Bacteriuria |   |   |   |   |
|--------------------------------------------|-------------------------|---|---|---|---|
|                                            | Yes (N (%))             | No (N (%)) | Total | P-value |
| Sex                                        |                         |             |       |         |
| Male                                       | 9 (22)                  | 74 (33.9)   | 83    | 0.18    |
| Female                                     | 32 (78)                 | 144 (66.1)  | 176   |         |
| Age (years)                                |                         |             |       |         |
| 5–10                                       | 13 (31.7)               | 67 (30.7)   | 80    | 0.9     |
| 11–15                                      | 28 (68.3)               | 151 (69.3)  | 179   |         |
| Urinary tract abnormality in children      |                         |             |       |         |
| Yes                                        | 6 (14.6)                | 41 (18.8)   | 47    | 0.6     |
| No                                         | 35 (85.4)               | 177 (81.2)  | 212   | 35 (85.4) |
| History of UTI in parents                  |                         |             |       |         |
| Yes                                        | 9 (21.9)                | 23 (10.6)   | 32    | 0.034   |
| No                                         | 32 (79.1)               | 195 (89.4)  | 227   |         |
| Previous history of catheterization        |                         |             |       |         |
| Catheterized                               | 0 (0)                   | 3 (1.4)     | 3     | 0.6     |
| Non-catheterized                           | 41 (100)                | 215 (98.6)  | 256   |         |
| Gram stain                                 |                         |             |       |         |
| Gram positive bacteria                      | 11 (26.8)               | -           | 11    |         |
| Gram negative bacteria                      | 30 (73.2)               | -           | 30    |         |
| Total                                      | 41 (100)                | 218 (100)   | 259 (100) |         |

Bacterial Etiologies

A total of 41 uropathogens were isolated. Out of this, 11 (26.8%) were Gram-positive bacteria and 30 (73.2%) were Gram-negative bacteria. *E. coli* 14 (34.1%) was the most frequently isolated uropathogen followed by *Pseudomonas* spp. 9 (22.0%) and *S. saprophyticus* 6 (14.6%). Among Gram-positive bacteria, *S. saprophyticus* 6 (54.6%) was the most frequently isolated bacteria followed by *S. aureus* 3 (27.3%) as shown in Figure 1.

![Figure 1](image1.png)

**Figure 1** Distribution of uropathogens isolated from children attending Bahir Dar Felege Hiwot Referral Hospital, February 2013 to May 2013.

Antimicrobial Susceptibility Testing

Antimicrobial susceptibility test was performed for all urine culture positive samples. Gram-negative bacteria isolated were tested against 10 antibiotics while Gram-positive isolates were tested against 7 antibiotics via disk diffusion (Tables 3 and 4, respectively).

Overall, the percentage of sensitivity for Gram-negative were high for ciprofloxacin (70%), norfloxacin (63.4%) and for ceftriaxone (60%) among selected antibiotics, whereas a high level of resistance was observed for ampicillin (80%) and nitrofurantoin (70%). *E. coli*, the most frequently isolated bacteria, was highly resistant for ampicillin (100%). Furthermore, more than 80% of Gram-negative isolates showed multiple resistances to the antibiotics tested (Table 3).

Overall, the percentage of sensitivity for Gram-positive bacterial isolates were relatively high for ciprofloxacin, penicillin and erythromycin, each accounted 7/9 (77.8%), 8/11 (72.7%) and 8/11 (72.7%), respectively. Overall resistance was high for ampicillin 9/11 (81.8%). All isolates showed multiple resistances to the antibiotics tested (100%) (Table 4).
| Isolates          | Antimicrobial Agents N (%) | Total (n=30) | Amp       | CIP       | CRO       | AMC       | CN        | F         | SXT       | NOR       | NA        | CAZ       |
|------------------|---------------------------|-------------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|
| E. coli (n=14)   |                           |             | S 0 (0)   | 10 (71.4)| 8 (57.1)  | 6 (42.9)  | 6(42.9)   | 3 (21.4) | 6(42.9)   | 10(71.4)  | 6(42.9)  | ND        |
|                  |                           |             | I 0 (0)   | 1 (7.2)   | 1 (7.2)   | 0 (0)     | 1(7.1)    | 0(0)     | 0(0)     | 0(0)     | 0(0)     | ND        |
|                  |                           |             | R 14 (100)| 3 (21.4)  | 5 (35.7)  | 8 (57.1)  | 7(50)     | 11(78.6) | 8(57.1)   | 4(28.6)   | 7(50)    | ND        |
| Pseudomonas spp. N=9 |                         |             | S 6 (66.7)| 7 (77.8)  | 5 (55.6)  | 5 (55.6)  | 7(77.8)   | 0(0)     | 0(0)     | 2(14.2)   | 6(66.7)  | ND        |
|                  |                           |             | I 0 (0)   | 2 (14.2)  | 2 (14.2)  | 2 (14.2)  | 0(0)     | 5(55.6)  | 1(7.1)    | 1(7.1)    | ND        | 0(0)     |
|                  |                           |             | R 3 (33.3)| 0 (0)     | 0 (0)     | 0 (0)     | 1(7.1)   | 4(44.4)  | 2(14.2)   | 6(66.7)   | ND        | 6(66.7)  |
| Klebsiella spp. n=5 |                         |             | S 0 (0)   | 3 (60)    | 4 (80)    | 0 (0)     | 0(0)     | 1(20)    | 4(80)     | 1(20)     | 3(60)    | ND        |
|                  |                           |             | I 0 (0)   | 1 (20)    | 1 (20)    | 5 (100)   | 4(80)    | 4(80)    | 0(0)      | 0(0)      | ND        | ND       |
|                  |                           |             | R 5 (100) | 1 (20)    | 1 (20)    | 5 (100)   | 4(80)    | 4(80)    | 0(0)      | 0(0)      | ND        | ND       |
| Enterobacter spp. n=1 |                       |             | S 0 (0)   | 0 (0)     | 0 (0)     | 0 (0)     | 1(100)   | 1(100)   | 1(100)    | 1(100)    | 0(0)     | ND        |
|                  |                           |             | I 0 (0)   | 0 (0)     | 0 (0)     | 0 (0)     | 1(100)   | 1(100)   | 1(100)    | 1(100)    | 0(0)     | ND        |
|                  |                           |             | R 1 (100) | 1 (100)   | 1 (100)   | 1 (100)   | 1(100)   | 1(100)   | 1(100)    | 1(100)    | 0(0)     | ND        |
| Citrobacter spp. n=1 |                     |             | S 0 (0)   | 0 (0)     | 0 (0)     | 0 (0)     | 1(100)   | 1(100)   | 1(100)    | 1(100)    | 0(0)     | ND        |
|                  |                           |             | I 0 (0)   | 0 (0)     | 0 (0)     | 0 (0)     | 0(0)     | 0(0)     | 0(0)      | 0(0)      | 0(0)     | ND        |
|                  |                           |             | R 1 (100) | 1 (100)   | 1 (100)   | 1 (100)   | 1(100)   | 1(100)   | 1(100)    | 1(100)    | 0(0)     | ND        |
| Total, n=30      |                           |             | S 6 (20)  | 21 (70)   | 18 (60)   | 12 (40)   | 15(50)   | 3 (10)   | 10(33.3)  | 19(63.4)  | 11(52.3) | 3(33.3)  |
|                  |                           |             | I 0 (0)   | 4 (13.3)  | 3 (10)    | 2 (6.7)   | 2(6.7)  | 6(20)    | 1(3.3)    | 1(3.3)    | 1(4.8)   | 0(0)     |
|                  |                           |             | R 24 (80) | 5 (16.6)  | 9 (30)    | 16 (53.3) | 13(43.3) | 21(70)   | 19(63.4)  | 10(33.3)  | 9(42.9)  | 6(66.7)  |

**Abbreviations:** Amp, ampicillin; CIP, ciprofloxacin; CRO, ceftriaxone; AMC, amoxicillin-clavulanate; CN, gentamicin; F, nitrofurantoin; NOR, norfloxacin; SXT, cotrimoxazole; NA, nalidixic acid; CAZ, ceftazidime; ND, not done.
Multidrug resistance (MDR) to two or more drugs was observed for 25/30 (83.3%) of Gram-negative bacteria and 11/11 (100%) Gram-positive bacteria. Table 5 shows the overall prevalence of MDR in both groups was 37/41 (90.3%).

Discussion
In the present study, the overall prevalence of UTI in children was 15.8%. It is comparable with the study conducted in Tanzania, Nigeria and Gondar, 20.3%\(^\text{12}\), 20.5%\(^\text{13}\) and 17.5%\(^\text{14}\), respectively. A lower prevalence was reported from Turkey, 7.1%\(^\text{4}\) India, 10.9%\(^\text{15}\) Iran, 7.8%\(^\text{16}\) Bahir Dar, 9.4%\(^\text{17}\) and Addis Ababa, 11.6%\(^\text{18}\). The variation might be due to difference in sample size and habit of personal hygiene.

Whereas a higher prevalence was reported in Italy, 22.6%\(^\text{19}\) People’s Republic of China, 36.5%\(^\text{20}\) Nepal, 45.5%\(^\text{21}\) Gondar, 28.1%\(^\text{22}\) and another study in Cameron also showed 58.3%\(^\text{23}\). This difference might be due to difference in age of the study subjects, sample size and presence of complication, for example, in the study conducted in People’s Republic of China with nephritic syndrome.

In the present study, Gram-negative bacteria were predominantly isolated with a rate of 73.2% which is in agreement with Madagascar, 65.5%\(^\text{24}\) Kenya, 78.9%\(^\text{25}\) and Bahir Dar, 61.9%,\(^\text{17}\) but lower than in a study reported from Mekelle, 83%\(^\text{26}\) India, 92%\(^\text{15}\) and South Africa which is 87.5%\(^\text{27}\). In most studies conducted elsewhere in the world, the commonest uropathogens were Gram-negative bacteria. The frequency of isolation of uropathogens varies with different geographical locations.

Among the overall bacterial isolates, \textit{E. coli} 34.1% was the most frequently isolated uropathogens in our study. Our study agrees with other studies conducted in Iran,\(^\text{28}\) India,\(^\text{15}\) Germany,\(^\text{29}\) Tanzania,\(^\text{12}\) Gondar\(^\text{22}\) and Tikur Anbesa Specialized Hospital,\(^\text{18}\) but in a study conducted in People’s

Table 4 Antimicrobial Susceptibility Pattern Of Gram-Positive Bacteria Isolated From Children With UTI Attending Felege Hiwot Referral Hospital (February 2013 To May 2013)

| Isolates       | Antimicrobial Agents N (%) |   |   |   |   |   |   |
|----------------|---------------------------|---|---|---|---|---|---|
|                | AMP | Ox | F | P | SXT | CIP | E |
|----------------|-----|----|---|---|-----|-----|---|
| \textit{S. aureus} N=3 |     |     |     | 2 (66.7) | 2 (66.7) | 0 (0) | 0 (0) | 3 (100) | 2 (66.7) |
| S               | 0 (0) | 2 (66.7) | 0 (0) | 0 (0) | 0 (0) | 3 (100) | 2 (66.7) |
| I               | 0 (0) | 2 (66.7) | 0 (0) | 0 (0) | 0 (0) | 3 (100) | 2 (66.7) |
| R               | 3 (100) | 1 (33.3) | 1 (33.3) | 1 (33.3) | 1 (33.3) | 1 (33.3) | 1 (33.3) |
| \textit{S. saprophyticus} N=6 |     |     |     | 2 (33.3) | 2 (33.3) | 6 (100) | 6 (100) | 4 (66.7) | 5 (83.3) |
| S               | 1 (16.7) | 4 (66.7) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| I               | 0 (0) | 0 (0) | 2 (33.3) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| R               | 5 (83.3) | 2 (33.3) | 2 (33.3) | 2 (33.3) | 2 (33.3) | 2 (33.3) | 2 (33.3) |
| \textit{Enterococcus} spp. N=2 |     |     |     | 2 (100) | 2 (100) | 2 (100) | 2 (100) | 2 (100) | 2 (100) |
| S               | 1 (50) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| I               | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| R               | 1 (50) | ND | ND | ND | ND | ND | ND |
| Total, N=11     |     |     |     | 8 (72.8) | 8 (72.8) | 7 (77.8) | 8 (72.8) | 8 (72.8) | 8 (72.8) |

Table 5 Multidrug Resistance Pattern Of Gram-Positive And Gram-Negative Bacteria Isolated From Urine Culture Of Children Attending Felege Hiwot Referral Hospital (February 2013 To May 2013)

| Number Of Isolated Gram-Negative Bacteria | Number Of MDR, n (%) |   |   |   |   |   |   |
|------------------------------------------|----------------------|---|---|---|---|---|---|
| \textit{E. coli} (n=14)                  | 13 (92.9)            |   |   |   |   |   |   |
| \textit{Pseudomonas} spp. (n=9)          | 5 (55.6)             |   |   |   |   |   |   |
| \textit{Klebsiella} spp. (n=5)           | 5 (100)              |   |   |   |   |   |   |
| \textit{Enterobacter} spp. (n=1)         | 1 (100)              |   |   |   |   |   |   |
| \textit{Citrobacter} spp. (n=1)          | 1 (100)              |   |   |   |   |   |   |
| Sub-total (n=30)                          | 25 (83.3)            |   |   |   |   |   |   |
| Number Of Isolated Gram-Positive Bacteria| Number Of MDR, n (%) |   |   |   |   |   |   |
| \textit{S. aureus} (n=3)                 | 3 (100)              |   |   |   |   |   |   |
| \textit{S. saprophyticus} (n=6)          | 6 (100)              |   |   |   |   |   |   |
| \textit{Enterococcus} spp. (n=2)         | 2 (100)              |   |   |   |   |   |   |
| Sub-total (n=11)                          | 11 (100)             |   |   |   |   |   |   |
| Total (n=41)                              | 37 (90.3)            |   |   |   |   |   |   |

Abbreviations: AMP, ampicillin; ox, cefoxitin; F, nitrofurantoin; p, penicillin; SXT, trimethoprim sulfamethoxazole; CIP, ciprofloxacin; E, erythromycin; CL, clindamycin; ND, not done.
Republic of China,\textsuperscript{20} Nairobi\textsuperscript{30} and Jimma,\textsuperscript{8} Enterococcus spp., S. aureus and Klebsiella spp. were the dominant pathogens isolated, respectively. Generally in the present study and in most other studies conducted elsewhere, E. coli was the most frequently isolated uropathogens. It is the most common cause of UTI and accounts for 75–90\% of UTIs in children. However, its relative frequency varies in different areas.\textsuperscript{31} E. coli is the commonest flora of the gastrointestinal tract and bowel from which it ascends to urinary tract and it has well characterized virulence factors to colonize the urinary tracts. The second leading bacterial isolate in our study was Pseudomonas spp. (22\%) followed by S. saprophyticus (14.6\%). But in other studies reported Klebsiella spp., S. aureus and Enterococcus faecalis were the second leading uropathogens in Iran,\textsuperscript{28} Germany\textsuperscript{29} and TikurAnbessa specialized hospital,\textsuperscript{8} respectively.

The first leading bacterium among Gram positives was S. Saprophyticus (54.6\%) followed by S. aureus (27.3\%). Our study disagrees with Germany\textsuperscript{29} Enterococcus spp.; at Gondar\textsuperscript{32} and Bahir Dar,\textsuperscript{17} S. aureus were the first leading pathogens. Bacterial etiologies vary with different geographical regions even vary over time within a population.

In the present study from children having significant bacteriuria, 21.9\% of them have parents with previous history of UTI (p<0.05). Similarly in a study conducted in Turkey showed that children with a family history of UTI had a 3.8-fold risk of developing UTI (P<0.05) compared to those without.\textsuperscript{4} Family history of UTI is the risk factor for UTI in children. Because some abnormalities and renal defects in parents may be acquired by children congenitally. Example children acquired vulvo – urethral reflux before birth.

In the present study, sensitivity of Gram-negative bacteria were relatively high for ciprofloxacin (70\%), norfloxacin (63.4\%) and ceftriaxone (60\%). A study in Jimma\textsuperscript{8} is in agreement with our study which showed that ciprofloxacin controlled 75\% of Gram-negative bacteria and in Gondar\textsuperscript{14} ciprofloxacin was relatively best drug. In other study, ciprofloxacin is a widely used fluoroquinolone with high bacterial activity against uropathogens and well-established clinical efficacy in the treatment of UTIs.\textsuperscript{33}

High level of resistance of Gram-negative bacteria was also observed for ampicillin (80\%) and nitrofurantoin (70\%) in the present study. This is in agreement with the study in India\textsuperscript{15} which showed high level of resistance to ampicillin; Mekelle (>80\%\textsuperscript{26} and Jimma\textsuperscript{8} (88\%) of the isolates were resistant to ampicillin. In contrast with our study, Gram-negative isolates were susceptible to nitrofurantoin in studies conducted in Jimma 75\%\textsuperscript{26} and Addis Ababa >85\%.\textsuperscript{18} Availability and use of these drugs in the community may vary in different areas.

E. coli, the first frequently isolated bacterium, was (100\%) resistant for ampicillin and (78.6\%) for nitrofurantoin. Our study is comparable with India\textsuperscript{15} that resistance for nitrofurantoin was (80\%). On the contrary in a study conducted in Iran,\textsuperscript{7} nitrofurantoin had the highest activity against E.coli. On the other hand, ampicillin resistance against E. coli in Tanzania\textsuperscript{12} revealed similar finding with the present study. The high resistance of Gram-negative bacteria to ampicillin might be due to ease availability, frequently prescribed drug and its low cost.

In the present study the overall percentage of sensitivity for Gram-positive isolates were high for penicillin, ciprofloxacin and erythromycin, 72.8\%, 77.8\% and 72.8\%, respectively. Our study is almost comparable with Gondar\textsuperscript{14} and Addis Ababa.\textsuperscript{18} Whereas (>80\%) Gram-positive isolates were resistant for ampicillin and cotrimoxazole. The present study disagrees with that of Gondar where ampicillin and cotrimoxazole resistance were 17.1\% and 57.1\%, respectively.\textsuperscript{14}

All isolated S. aureus were resistant for ampicillin, penicillin and cotrimoxazole. A study in Gondar showed that among Gram positives, S.aureus showed high level of resistance for cotrimoxazole (53.3\%), ampicillin (43\%) and penicillin (36.7\%).\textsuperscript{22} The present study is almost comparable with this study.

Multidrug resistance (MDR) to two or more drugs was observed in 83.3\% and 100\% of Gram-negative and Gram-positive isolates, respectively. Overall, MDR in both groups was 90.3\%. This study is almost similar with the study in Bahir Dar which was 93.1\%\textsuperscript{34} and in Gondar, 85.7\%.\textsuperscript{22} This high MDR in the present study might be due to widespread misuse of antibiotics, inappropriate prescription of drugs and lack of knowledge about drug resistance in the study area.

Limitation Of The Study

Our study has some limitations like antimicrobial susceptibility test was not done because of getting antibiotics disks and few bacterial isolates were not identified at a species level. The other limitation was using only Kirby-Bauer method to perform antimicrobial susceptibility instead of micro-dilution method, which makes more sensitive.

Conclusion

The result showed that the most predominant bacterial isolate was E. coli. Overall, the percentage of sensitivity of Gram-
negative bacteria to all antibiotics tested was relatively low, except for ciprofloxacin, norfloxacin and ceftriaxone, which were relatively high; whereas Gram-positive bacteria showed relatively high percentage of sensitivity for ciprofloxacin, penicillin and erythromycin among selected antibiotics tested in the study. Therefore, these antibiotics may be used for empirical therapy of urinary tract infection (UTI) when culture and sensitivity are unavailable.

The present study also showed a high incidence of resistance to most of the antimicrobials agents tested. The prevalence of multiple drug resistance was also high. This calls for a continuous monitoring and review of antimicrobial policy in the hospital and the country at large.

### Abbreviations

AAU, Addis Ababa University; FHRH, Felege Hiwot Referral Hospital; ARHRLC, Amhara Regional Health Research Laboratory Center; CAUTIs, catheter-associated urinary tract infection; CFU, colony-forming unit; CLED, Cysteine Lactose Electrolyte Deficient media; CI, confidence interval; CLSI, Clinical & Laboratory Standards Institute; ESBL, extended-spectrum beta-lactamases; MDR, multiple drug resistance; OPD, outpatient department; UTI, urinary tract infection; TMP-SMX, trimethoprim sulfamethoxazole.

### Ethics Approval And Consent To Participate

The MSc research project proposal was ethically cleared by the Department Research and Ethical Review Committee and was approved by the Department of Microbiology, Immunology and Parasitology, School of Medicine, Addis Ababa University. Official permission from the study site was obtained. Written informed consent was obtained from each child’s parents/guardians. Urine culture and antimicrobial sensitivity test results were reported to the attending physician for subsequent treatment and follow up.

### Availability Of Data And Materials

The data that support the findings of this study will be available from the corresponding author upon reasonable request in the form of Statistical Package for the Social Sciences (SPSS).

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### Authors Contributions

YB: Principal investigator of the study, study design, data collection, laboratory work, data analysis and drafted the manuscript; DA and YWA: study design, supervision of data collection, data analysis, and drafted the manuscript; GY and AG: critically reviewed the data and drafted the manuscript. All authors gave final approval of the version to be published, and agree to be accountable for all aspects of the work.

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### Disclosure

The authors declare that they have no conflicts of interest in this work.

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