Prevalence of symptomatic urinary tract infections and antimicrobial susceptibility patterns of isolated uropathogens in kohat region of Pakistan

Abstract

Urinary tract infections are commonly distributed infectious diseases. This study is designed to determine the prevalence of symptomatic UTIs and antimicrobial susceptibility patterns of uropathogens to commonly use antibiotics. A total of 500 patients, age ranges 20-90years were included in this study that conducted in Kohat Teaching Hospital of Kohat District, Khyber Pakhtunkhwa, Pakistan. Midstream urine samples were collected from patients of both genders. All samples were cultured aerobically in Blood agar, MacConkey agar and CLED medium (Oxoid). The identification of isolated bacteria were performed using manual biochemical tests. All uropathogenic isolates were subjected to antimicrobial susceptibility testing. The prevalence of UTIs were 11.6%. The frequency of UTIs was 8.9% in males and 13.8% in females. Out of total positive cultures, 20.7% (12/58) were Gram positive bacteria and 79.3(46/58) were Gram negative bacteria. The commonly isolated pathogens are Escherichia coli 24 (41.4%), Klebsiella pneumoniae 9 (15.5%) and Proteus mirabilis 8 (13.8%). Most lactose fermenter Gram-ve bacteria were sensitive to Cefepime and all Gram positive isolates were sensitive to Meropenem. In this study, it was concluded that there is marked variation in the prevalence of UTIs (between males and females) and antibiotics susceptibility patterns of uropathogens. Meropenem is appropriate antibiotics to treated UTIs causes by Gram positive bacteria. These data demonstrate that future studies should be focused on the causes of antibiotics resistance to solve this problem.

Keywords: antimicrobial, Pakistan, prevalence, urinary tract infection, uropathogen

Introduction

Urinary tract infection (UTI) is a one of the most frequent and contagious infection. Due to its complexity, it damages various types of delicate organs in the human body such as urethra, ureters and bladder. UTIs is more common in females especially in pregnant women as a result of anatomical difference and the decrease level of urination during pregnancy. Pregnancy increases bactriuria which leads to pyelonephritis and obstetric such as higher fetal mortality rates, low-birth weight and prematurity. It has been estimated that about 150 million peoples were diagnosed worldwide for the presence of Gram negative bacteremia which is the significant cause of UTI in all ages specifically in the old age peoples. Fifteen to twenty percent of the urinary tract infection is caused by Gram positive bacteria and about eighty to eighty five percent is caused by Gram negative bacteria. The mechanisms of pathogenesis depend on the causative agent and its virulence factors.

Due to multi-drug Resistant microorganisms there have been great changes occurred in the antimicrobial susceptibility patterns of urinary pathogens. And due to antibiotic-resistant bacteria, the treatment of UTIs become empirical hard so raising the prevalence of infection. In the world, the unusual development and increasing of multidrug-resistant in bacterial pathogens is a great health problem which is rising day by day in a growing community. According to the geographical location, different antibiotics have different resistance rate and they are directly proportional to the abuse of antibiotics. The increasing appearance of multidrug resistant (MDR) bacteria is has been exaggerated over the time due to increase use of antibiotics in many sectors such as agricultural, medical and veterinary. The main factors which leads to antibiotics resistance include the incorrect investigation and drugs abuse. In some time the treatments of simple infection can become complicated or impossible due to bacterial resistant strains. This study is designed to determine the prevalence of symptomatic UTIs and antimicrobial sensitivity patterns of uropathogens in District Kohat of Pakistan.

Materials and methods

Study design, area and duration

This is cross sectional hospital base study carried out from June to December 2015 in Kohat Teaching Hospital of Kohat District, Khyber Pakhtunkhwa, Pakistan.

Study population

All patients with symptomatic UTIs, ages ranging 20-90years and belonged to District Kohat of Pakistan were fulfill the inclusion criteria.

Sampling

A total of 500 midstream urine samples were collected from patients of both genders, those attended Kohat Teaching Hospital with the symptoms of UTIs.

Methodology

All urine samples were cultured on Blood agar, MacConkey agar and CLED medium (Oxoid). The significant growth bacteria was
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Processed for identification and antibiotics susceptibility testing.

Isolation and identification:
Each specimen was thoroughly mixed by inverting the container 2-3 times and inoculated into culture mediums. The culture plates were incubated at 37°C for 24 hours. In case of no growth, further incubation was carried out for additional 24 hours. More than 10<sup>5</sup> colony forming unit per ml was considered as the significant level of bacteriuria. The identification of isolates were performed on the base of gram reaction, colonial morphology and biochemical characterization.

Antibiotic susceptibility testing:
The antibiotic susceptibility testing was performed by Kirby Bauer’s disc diffusion technique (Bauer et al., 1966) as suggested by clinical and laboratory standard institute. The zone of inhibition was calculated and interpreted using the standard chart and the organisms were reported as susceptible, intermediate or resistant accordingly. Different antibiotic discs (Oxoid) were used in this study and these were Ceftriaxone (CRO), Cefazidime (CAZ), Cefepime (FEP), Meropenem (MRP), and Ciprofloxacin (CIP). Plus Augmentin (AUG) and Levofloxacin (LEV), only for Gram-ve bacteria. E. coli ATCC 252395 was used as control strain.

Ethical statement
This study was undertaken with the approval of Hazara University and conducted according to the declaration of Helsinki and with particular ethical standards of Hazara University. Verbal consent was taken from each participant.

Data analysis
All data were analyzed with SPSS software version 17. Chi square test was performed and P value less than 0.05 was considered statistically significant.

Result
Prevalence of UTIs
The current study was conducted on 500 patients of different gender (Males and females) and their corresponding ratio was 225:275 (M:F, 45%: 55%), in district Kohat of Pakistan. A total of 58 (11.6%) urine samples were showed the significance growth of uropathogenic bacteria. The prevalence of UTIs in males was 8.9% (20/58) and in females was 13.8% (38/58), See Figure 1.

The frequency of commonly isolated bacteria
Out of total positive cultures, 20.7% (12/58) were Gram positive bacteria and 79.3% (46/58) were Gram negative bacteria. The most prevalent uropathogenic bacteria were Escherichia coli 24 (41.4%), Klebsiella pneumoniae 9 (15.5%) and Proteus mirabilis 8 (13.8%), Table 1.

Table 1 Frequency of isolated uropathogens

| Name of bacteria       | Number of isolates | Percentage  | P value |
|------------------------|--------------------|-------------|---------|
| Escherichia coli       | 24                 | 41.4% (24/58)|         |
| Staph. aureus          | 6                  | 10.34% (6/58)|         |
| Klebsiella pneumoniae  | 9                  | 15.5% (9/58)| 0       |
| Proteus mirabilis      | 8                  | 13.8% (8/58)|         |
| P.aeruginosa           | 5                  | 8.62% (5/58)|         |
| S. Saprophyticus       | 6                  | 10.34% (6/58)|         |
| Total                  | 58                 |             |         |

Antibiotics susceptibility of urinary isolates
The emergence of drugs resistance among urinary isolates to commonly use antibiotics has been observed (Figure 2), and only cefepime and Meropenem are showed a great value against uropathogens. As displayed in Table 2, in our current study the highly prevalent uropathogenic bacteria E. coli was showed the highest sensitivity (91.7%) to cefepime and lowest sensitivity (12.5%) to ciprofloxacin, similarly the K. pneumonia showed the highest sensitivity (88.9%) to Augmentin and the lowest sensitivity (22.2%) to cefazidime. The P. mirabilis was 87.5% sensitive to Meropenem, while it was showed low sensitivity (12.5%) to cefepime. The highest sensitivity rate which was showed by P. aeruginosa was 80% against cefepime and the lowest sensitivity rate was 20% (against ceftazidime). All S. aureus and S. saprophyticus isolates were sensitive to Meropenem. Hindered percent of resistant to cefazidime was observed in S. saprophyticus isolates and same percentage of resistant to ciprofloxacin was detected in S. aureus isolates, Table 3.

Figure 1 The prevalence of UTIs in gender.

Figure 2 The susceptibility of urinary pathogens to commonly use antibiotics.

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Table 2 Antibiotics susceptibility profile of gram negative isolates

| Antibiotics | Isolate | E. coli | K. pneumoniae | P. aeruginosa | P. mirabilis | Total |
|-------------|---------|---------|----------------|---------------|--------------|-------|
| CRO         | Sensitive | 25% (6/24) | 44.4% (4/9)  | 60% (3/5)    | 50% (4/8)    | 37% (17/46) |
|             | Resistant | 75% (18/24) | 55.6% (5/9)  | 40% (2/5)    | 50% (4/8)    | 63% (29/46) |
| AUG         | Sensitive | 50% (12/24) | 88.9% (8/9)  | 40% (2/5)    | 62.5% (5/8)  | 58.7% (27/46) |
|             | Resistant | 50% (12/24) | 11.1% (1/9)  | 60% (3/5)    | 37.5% (3/8)  | 41.3% (19/46) |
| CAZ         | Sensitive | 33.3% (8/24) | 22.2% (2/9)  | 20% (1/5)    | 37.5% (3/8)  | 30.4% (14/46) |
|             | Resistant | 66.7% (16/24) | 77.8% (7/9)  | 80% (4/5)    | 62.5% (5/8)  | 69.6% (32/46) |
| MRP         | Sensitive | 50% (12/24) | 66.7% (6/9)  | 60% (3/5)    | 87.5% (7/8)  | 60.9% (28/46) |
|             | Resistant | 50% (12/24) | 33.3% (3/9)  | 40% (2/5)    | 12.5% (1/8)  | 39.1% (18/46) |
| LEV         | Sensitive | 62.5% (15/24) | 44.4% (4/9)  | 60% (3/5)    | 62.5% (5/8)  | 58.7% (27/46) |
|             | Resistant | 37.5% (9/24) | 55.6% (5/9)  | 40% (2/5)    | 37.5% (3/8)  | 41.3% (19/46) |
| FEP         | Sensitive | 91.7% (22/24) | 77.8% (7/9)  | 80% (4/5)    | 12.5% (1/8)  | 73.9% (34/46) |
|             | Resistant | 8.3% (2/24) | 22.2% (2/9)  | 20% (1/5)    | 87.5% (7/8)  | 26.1% (12/46) |
| CIP         | Sensitive | 12.5% (3/24) | 44.4% (4/9)  | 40% (2/5)    | 50% (4/8)    | 28.3% (13/46) |
|             | Resistant | 87.5% (21/24) | 55.6% (5/9)  | 60% (3/5)    | 50% (4/8)    | 71.7% (33/46) |

Table 3 Antibiotics susceptibility profile of gram positive isolates

| Antibiotic | Susceptibility result | S. saprophyticus | S. aureus | Total |
|------------|-----------------------|------------------|----------|-------|
| CRO        | Sensitive             | 50% (3/6)        | 83.3% (5/6) | 66.7% (8/12) |
|            | Resistant             | 50% (3/6)        | 16.7% (1/6) | 33.3% (4/12) |
| CAZ        | Sensitive             | 0% (0/6)         | 50% (3/6)  | 25% (3/12) |
|            | Resistant             | 100% (6/6)       | 50% (3/6)  | 75% (9/12) |
| FEP        | Sensitive             | 83.3% (5/6)      | 50% (3/6)  | 66.7% (8/12) |
|            | Resistant             | 16.7% (1/6)      | 50% (3/6)  | 33.3% (4/12) |
| MRP        | Sensitive             | 100% (6/6)       | 100% (6/6) | 100% (12/12) |
|            | Resistant             | 0% (0/6)         | 0% (0/6)   | 0% (0/6) |
| CIP        | Sensitive             | 33.3% (2/6)      | 0% (0/6)   | 16.7% (2/12) |
|            | Resistant             | 66.7% (4/6)      | 100% (6/6) | 83.3% (10/12) |

Discussion

Urinary tract infection (UTI) is one of the most common infections worldwide and it account for 1–2% of all consultation. It has been estimated that more than 6 million outpatient visits and 300,000 hospital stays every year are due to UTIs. Approximately 10% of humans will have a UTI at some time during their lives.

Our study was found the prevalence of UTI is 11.6% among symptomatic patients; and 8.9% in males and 13.8% in females. Similar studies has been conducted in different countries includes Mandokhail et al., Prakash D et al., and Patil et al., studies which reported the prevalence of UTIs were 60.63%, 53.82%, and 49%, respectively. Several reports have indicated that the prevalence of UTIs is more in females than males. John MS et al., (2013) study was found the prevalence of Gram negative bacteria (90.32%) were higher than Gram positive (9.68%); and the most frequently isolated urinary pathogen is Escherichia coli (41.4%), Klebsiella pneumoniae (15.5%), Proteus mirabilis (13.8%), S. saprophyticus (10.34%), Staphylococcus aureus (10.34%), and P. aeruginosa (8.62%). These results, indicate Gram negative are more commonly implicated in UTIs in this area of Pakistan. Previously, Prakash D et al., (2013) study was found the prevalence of Gram negative bacteria (90.32%) were higher than Gram positive (9.68%); and the most frequently isolated urinary pathogen is Escherichia coli (42.58%), followed by Klebsiella pneumoniae (18.71%) and Pseudomonas aeruginosa (12.90%) [27]. Similar results, Also were reported by Patil S et al., (2013) study, who found out of the 48 positive culture, 83.3% (40/48) of isolates were sensitive to amoxicillin-clavulanate, 71.7% (34/48) were sensitive to cefotaxime, and 66.7% (32/48) were sensitive to ceftriaxone. However, all these prevalence are highest than our study result, but they agreed in the increase of UTIs in females than males that is previously confirmed in many others studies.

Among the total of isolated bacteria, we found the prevalence of Gram positive bacteria and Gram negative bacteria were 20.7% and 79.3%, respectively. And the most prevalent uropathogenic bacteria were Escherichia coli (41.4%), Klebsiella pneumoniae (15.5%), Proteus mirabilis (13.8%), S. saprophyticus (10.34%), Staphylococcus aureus (10.34%), and P. aeruginosa (8.62%). These results, indicate Gram negative are more commonly implicated in UTIs in this area of Pakistan. Previously, Prakash D et al., (2013) study was found the prevalence of Gram negative bacteria (90.32%) were higher than Gram positive (9.68%); and the most frequently isolated urinary pathogen is Escherichia coli (42.58%), followed by Klebsiella pneumoniae (18.71%) and Pseudomonas aeruginosa (12.90%) [27]. Similar results, Also were reported by Patil S et al., (2013) study, who found out of the 48 positive culture, 83.3% (40/48) of isolates were sensitive to amoxicillin-clavulanate, 71.7% (34/48) were sensitive to cefotaxime, and 66.7% (32/48) were sensitive to ceftriaxone. However, all these prevalence are highest than our study result, but they agreed in the increase of UTIs in females than males that is previously confirmed in many others studies.

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were gram negative bacteria and 16.7% (8/48) were Gram positive bacteria; and 65% of isolated pathogens are *E. coli*.

In this work, we observed the emergence of antibiotics resistant. The sensitivity of isolated bacteria to Ceftriaxone, Cefazidime, Cefepime, Meropenem, and Ciprofloxacin were 43.1%, 29.3%, 72.4%, 69% and 29.3%, respectively. There is marked difference in susceptibility of Gram+ve and –ve bacteria; and their members to different antibiotics. Similarly results has been reported by Laghawe A et al., (2015) study that was found Gram+ve bacteria are less Sensitive to Ceftriaxone and Meropenem than Gram-ve. Preventing the dispensing of medicines without prescription and proper use of it by patients can aid in control of antibiotics resistance in this area.

**Conclusions**

Urinary tract infections are a one of commonly distributed infection. Recently the emergence of drugs resistance has been observed among urinary isolates. In this study, it was concluded that the resistance of antimicrobial agent among uropathogens was increased and there is marked variation in the antibiotics susceptibility patterns of uropathogens. Meropenem is appropriate antibiotic to treated UTIs and there is marked variation in the antibiotics susceptibility patterns of uropathogens. In this study, it was concluded that the resistance of antimicrobial agent among uropathogens was increased and there is marked variation in the antibiotics susceptibility patterns of uropathogens. We recommend the use of Meropenem for treatment of UTI. The results of our study are consistent with those of two other studies. In a study of 30 patients with UTI, the resistance of *E. coli* to Ciprofloxacin was found to be 29.3% (11/38), and in another study of 51 patients with UTI, the resistance of *E. coli* to Ciprofloxacin was found to be 23.0% (12/51). These results suggest that Ciprofloxacin is an effective treatment for UTI in the general population. However, these studies were conducted in different regions and with different patient populations, so the results may not be applicable to all populations.

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