



Profile of Uropathogens and their Antibiogram in Pregnant Women with Suspected Urinary Tract Infections

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ABSTRACT

Urinary tract infections (UTIs) are common illnesses that develop when bacteria enter the urethra and infect the urinary tract. Factors such as age, sex, catheterization, pregnancy and hospitalization all have an impact on the relative incidence of the pathogens. Antimicrobial resistance has been reported among several uropathogens, inappropriate use of antimicrobials being one of the major reasons, thus making the treatment of UTI a serious public health concern, particularly in developing nations. Thus, the present research was undertaken to study the spectrum of uropathogens and their antibiogram. The present study included all consecutive urine samples from pregnant women with clinical suspicion of UTI received in the Bacteriology section of Microbiology laboratory for culture and sensitivity during January to June 2022. Identification of uropathogens and their antibiogram were determined by standard bacteriological techniques. A total of 283 urine samples from pregnant women with clinical suspicion of UTI were included in the study. Of the total samples, 28.3% (n=80) were culture positive while 71.7% (n=203) were culture negative. Among the uropathogens from pregnant women, *E. coli* was the most common isolate (60%, n=48), followed by *Klebsiella* spp. (25%, n=20), *Enterococcus* spp. (10%, n=8) and *Candida* spp. (5%, n=4). The isolates showed varying degrees of resistance to different antibiotics. Urinary tract infections (UTIs) are a common complication that can occur during pregnancy due to the many physiological changes that occur. As a result, it is critical to screen pregnant women on a regular basis in order to reduce the risk to both mother and child.

Keywords: Urinary Tract Infection, Pregnant Women, Uropathogens, *Escherichia coli*

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INTRODUCTION

Urinary tract infections (UTIs) are common illnesses that develop when bacteria enter the urethra and infect the urinary tract [1]. Based on site of infection, it can be classified into lower UTI (bladder and urethra are involved) and upper UTI (kidneys, pelvis, and ureter are involved). The majority of UTIs are caused by an ascending infection. Asymptomatic bacteriuria, acute cystitis, and acute pyelonephritis are the three most common clinical manifestations of UTIs in pregnancy [2]. Significant bacteriuria ($>10^5$ CFU/mL of urine) without clinical symptoms of UTIs (such as frequent urination, painful urination or fever) or any other findings is referred to as asymptomatic bacteriuria. Factors such as age, sex, catheterization, pregnancy and hospitalization all have an impact on the relative incidence of the pathogens. Globally, the prevalence of UTI has been estimated to be approximately 150 million people per year. In developing countries including India, its rate is reported to be more than 20%. Each year, India reported almost seven million hospital visits, one million of which are emergency visits; this leads to roughly 100,000 hospitalizations due to UTI. Between 2% and 10% of pregnant women are thought to experience asymptomatic UTI globally [5]. According to reports, women in India have a prevalence of UTI that varies between 3% and 24% overall while in the second half of pregnancy, 1 to 4% of pregnant women will experience acute pyelonephritis, and 1 to 2% will experience severe acute pyelonephritis [6]. Significant morbidity is linked to pregnancy-related UTI in both the mother and the foetus [7]. During pregnancy, women are more likely to get UTIs, starting at week six and peaking between weeks 22 and 26 [8]. UTIs are more likely to occur during pregnancy because the uterus, lying between the bladder and the rectum, expands and pushes the bladder forward, making it narrower. This increases the frequency of irritation and, as a result, the volume of urine that can be totally emptied. Pregnancy's combination of hormonal, mechanical, and physiological changes results in significant changes to the urinary tract [9].

Among the bacterial uropathogens, *Escherichia coli*, *Klebsiella pneumoniae*, *Proteus* spp., *Staphylococcus saprophyticus*, *Enterococcus faecalis* and *Staphylococcus aureus* are frequently isolated bacteria [10].

Antimicrobial resistance has been reported among several uropathogens, inappropriate use of antimicrobials being one of the major reasons, thus making the treatment of UTI a serious public health concern, particularly in developing nations [11]. Additionally, in substantial episodes of UTI, empirical antimicrobial therapy is initiated before the laboratory findings of urine culture are available; thereby contributing to the prevalence of antibiotic resistance among uropathogens [4]. Third-generation cephalosporins (cefotaxime, cephadrine, ceftazidime, and cefaclor), nitrofurantoin, fosfomycin, trimethoprim-sulfamethoxazole, and penicillin are some of the antimicrobial agents used to treat UTIs. Among these drugs, trimethoprim-sulfonamides and fluoroquinolones are not recommended during pregnancy [12]. Thus, the present study was undertaken in order to monitor the spectrum of uropathogens and their antibiotic sensitivity among the pregnant women with suspected UTIs.

MATERIAL AND METHODS

This study was a prospective observational study that took place from January to June 2022 in a tertiary care hospital in North India. The study included all consecutive urine samples received in the Microbiology laboratory for urine culture and sensitivity from pregnant women with clinical suspicion of UTI. The current study was approved by the institutional ethics committee of SGT Medical College, hospital, and research institute before it was carried out.

Urine samples were processed within two hours after collection. The samples were inoculated onto Cystine Lactose Electrolyte-Deficient (CLED) agar, and the plates were incubated for 24 hours at 37°C under aerobic conditions. Semi quantitative analysis of colonies grown on CLED was performed and interpreted as per Kass criteria [13]. The colony morphology of the significant bacterial growth was observed after incubation, and then subjected for identification by standard microbiological techniques [14]. The antibiotic susceptibility testing for the bacterial isolates was performed using the Kirby-Bauer disc diffusion method on Muller Hinton agar, and the results were interpreted according to Clinical Laboratory Standard Institute (CLSI) standards [15]. For Gram positive cocci, the following antibiotic discs were used: cefoxitin (10µg), ciprofloxacin (5µg), fosfomycin (200µg), nitrofurantoin (300µg), tetracycline (30µg), teicoplanin (30µg), gentamicin (10µg), minocycline (30µg), ampicillin (10µg), piperacillin-tazobactam (100/10µg), amoxy/clavulanic acid (20µg), erythromycin (10µg), co-trimoxazole (25µg), vancomycin (30µg) and linezolid (30µg). While for Gram negative bacterial isolates, the following antibiotic discs were used: ampicillin (10µg), gentamicin (10µg), amikacin (30µg), amoxyclav-clavulanic acid (20µg), piperacillin-tazobactam (100/10µg), cefepime (30µg), ciprofloxacin (5µg), ertapenem (10µg), nitrofurantoin (300µg), fosfomycin (200µg), cefepime-sulbactam (30/15µg), ceftioxaone (30µg), levofloxacin (30µg), aztreonam (30µg), tigecycline (15µg) and tetracycline (30µg).

Statistical analysis

Data were entered and recorded in Microsoft excel 2016. The prevalence of UTI and antimicrobial susceptibility were expressed in terms of (number) percentage.

RESULTS

The study included 283 urine samples from pregnant women with clinical suspicion of UTI. Of the total samples, 28.3% (n=80) were culture positive while 71.7% (n=203) were culture negative. Majority of the culture positivity (65%, n=52) was observed in the age group of 18-25 years, followed by 26-32 years (20%, n=16) and 33-40 years (15%, n=12). When the educational status of the study participants with culture positive was analyzed, majority (40%, n=32) were observed among the women who have not received any formal education, followed by up to 5th standard 30% (n=24), 6th to 10th standard 17.5% (n=14), up to 11th standard and above 12.5% (n=10). Highest prevalence of UTI was found among women with multigravida 38.7% (n=58) followed by primi gravida 16.92% (n=22). When the incidence of UTI was analyzed in relation to gestational age, highest prevalence was found in 2nd trimester 45% (n=36), followed by 3rd trimester 30% (n=24) and 1st trimester 25% (n=20). Among the uropathogens from pregnant women, *E.coli* was the most common isolate (60%, n=48), followed by *Klebsiella spp.* (25%, n=20), *Enterococcus spp.* (10%, n=8) and *Candida spp.* (5%, n=4) shown in fig.1.

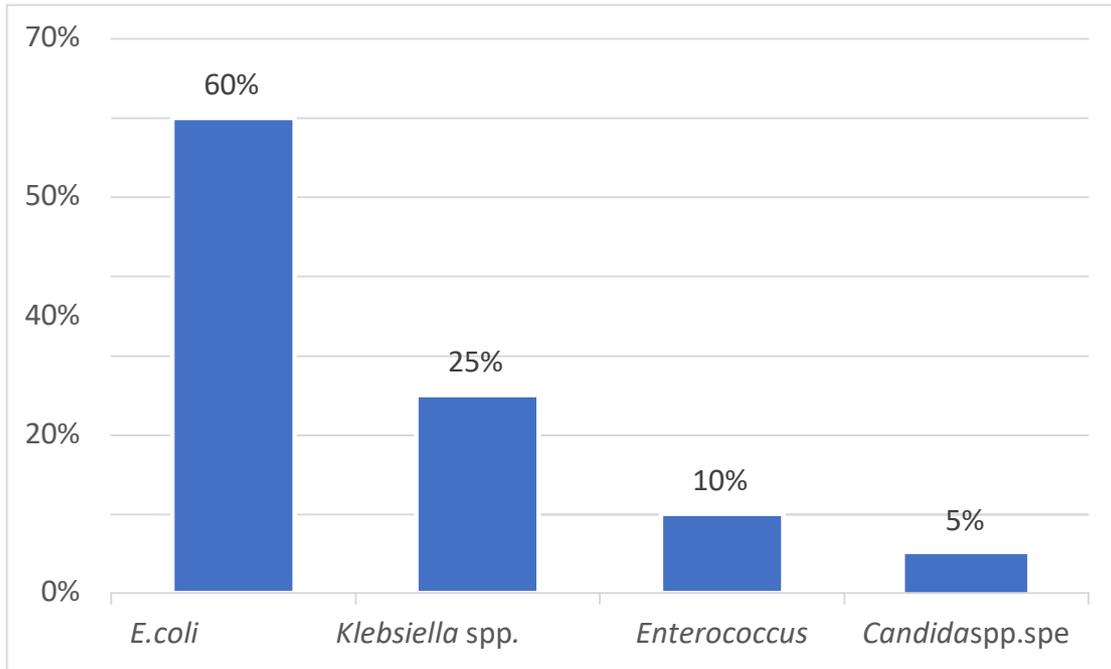
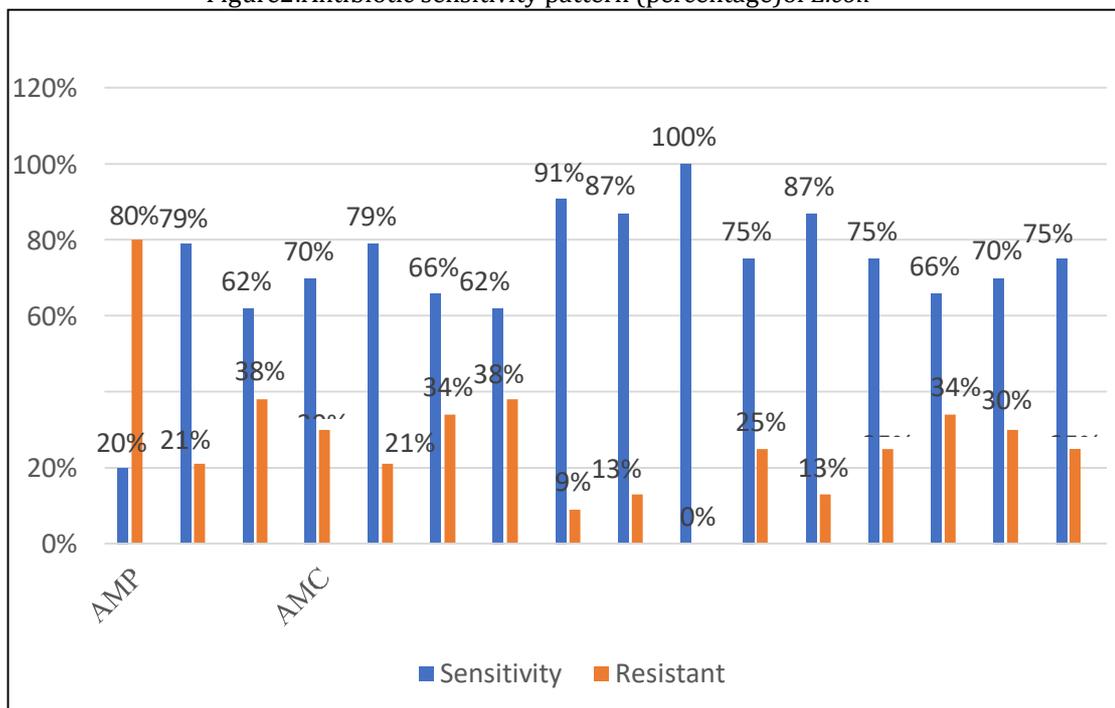


Figure1:Distribution of various isolates obtained from urine culture

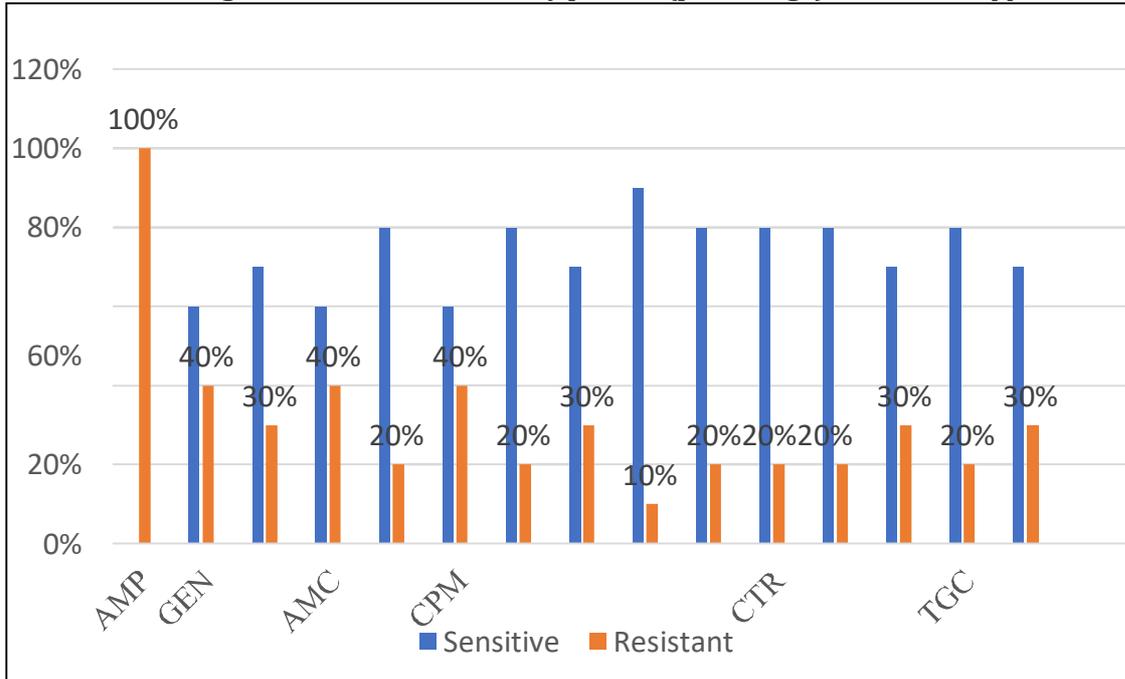
Antibiotic sensitivity rates for *E. coli* isolates to various antibiotics showed 100% sensitive to fosfomycin, 91% toertapenem, 87% tonitrofurantoin and ceftriaxone, 79% topiperacillin-tazobactam and gentamicin and 75% to levofloxacin, tetracycline and cefoperazone-sulbactam as depicted in fig. 2.

Figure2:Antibiotic sensitivity pattern (percentage)of *E.coli*



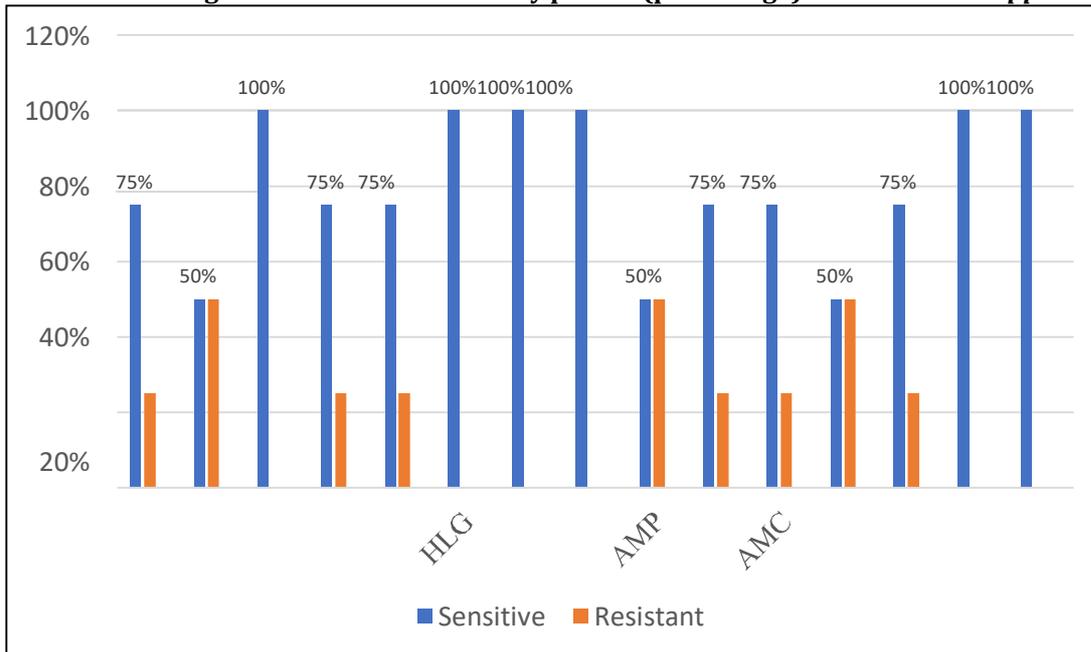
Antibiotic sensitivity pattern of *Klebsiella* isolates showed 90% sensitive to nitrofurantoin, 80% sensitive to piperacillin-tazobactam, cefepazone-sulbactam, ciprofloxacin, tetracycline and ceftriaxone as demonstrated in fig. 3.

Figure 3:Antibiotic sensitivity pattern (percentage) of *Klebsiellaspp.*



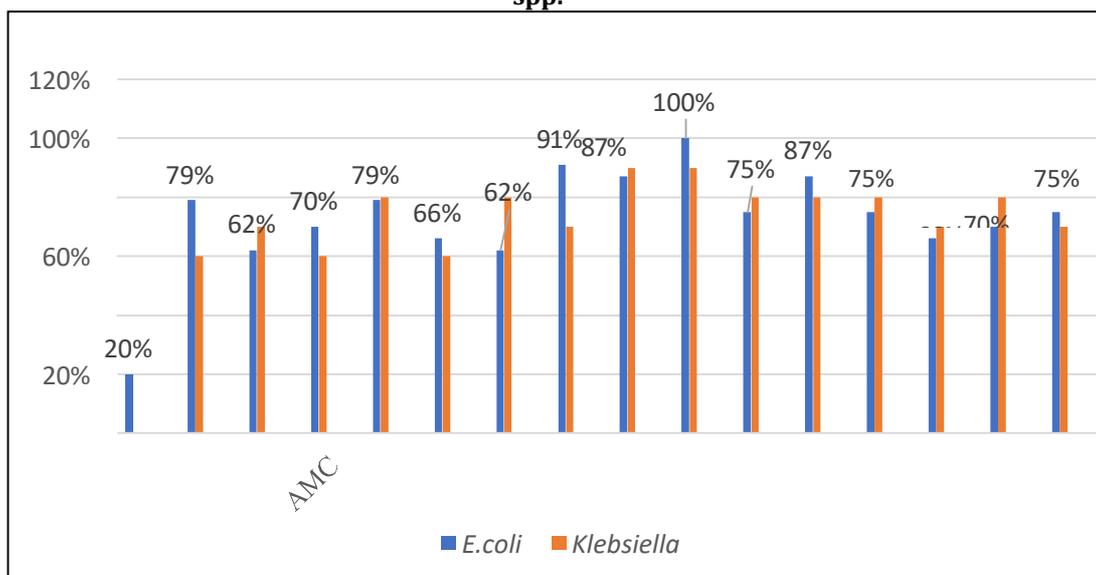
Antibiotic sensitivity pattern of *Enterococcus* isolates showed 100% sensitive to gentamycin, minocycline, teicoplanin, vancomycin and linezolid. 75% sensitive tonitrofurantoin, tetracycline, piperacillin-tazobactam and cotrimoxazole as demonstrated in fig. 4.

Figure4: Antibiotic sensitivity pattern(percentage) of *Enterococcuspp.*



Overall *E. coli* isolates showed higher susceptibility to most of the antibiotics tested when compared with *Klebsiella spp.*

Figure 5: Comparison of Antibiotic sensitivity pattern (percentage) between *E.coli* and *Klebsiella* spp.



DISCUSSION

UTIs are a significant burden on the health care system, owing to their prevalence in both community and hospital settings. Pregnant women are at an increased risk of urinary tract infections, which begins at week 6 and peaks between weeks 22–26.¹⁶ Due to a natural increase in plasma volume during pregnancy, a woman's urine concentration can drop by up to 70%, called glycosuria, and it is one of the main reasons bacteria grow in urine [17]. UTIs can result in severe obstetric complications, such as intrauterine growth retardation, preeclampsia, and preterm birth, which have negative maternal and perinatal consequences.

Moreover, asymptomatic bacteriuria has been associated with cystitis and pyelonephritis, which can induce severe respiratory distress, temporary renal failure, sepsis, and shock during pregnancy [18].

The current study enrolled 280 pregnant women attending an antenatal clinic, with a positive urine culture being detected in 28.3% (n=80) of pregnant women. Yasmin *et al.* reported a 28% prevalence of urinary tract infection (UTI) among symptomatic and asymptomatic pregnant women in Bihar in 2018 [18]. In rural Haryana, Kant *et al.* reported a prevalence of 33.3% among pregnant women in 2017 [5]. The preceding studies corroborate our findings. Arumaikannu *et al.* 2021 reported a low prevalence of 18% among pregnant women in Tamil Nadu [20], whereas Johnson *et al.* 2021 reported a prevalence of 35% among pregnant women with both symptomatic and asymptomatic UTI in the South-Western Uganda region [21].

A number of risk factors for UTI have been identified, with age, particularly in the female age group, being a major factor. In our study, the age group 18-25 years had the highest incidence of urinary tract infection (65%), followed by 22% in the 26-32 age group. In the Saudi Arabian region, Lele *et al.* (2015) found a 28.6% prevalence, which was particularly high in the age group 26-30 years [22]. Nwachukwu *et al.* (2018) found a prevalence of 48% in the age group 26-30 years in Nigeria [23]. The reason for this could be that many women in this age group are likely to have had several children prior to the current pregnancy, and it has been reported that multiparity is a risk factor for bacteriuria during pregnancy. Sexual activity and certain contraceptive methods have also been linked to an increased risk, and women of this age are the most sexually active. Arumaikannu *et al.* reported a high prevalence in the age group of 21-25 years in Tamil Nadu in recent studies [21]. Similar findings were found in a study conducted by Yashmin *et al.* in Bihar in 2018, which found a high incidence in the 21–25-year age group [19].

In terms of the level of education variable, the findings revealed that there was a significant link between women's UTI and their level of education. The prevalence of UTI was found to be higher among illiterate women in our study (40%). It could be due to a lack of concern for personal hygiene and personal care. Sheikh *et al.* found that education had no effect on the incidence rate of UTIs, whereas Gunes *et al.* observed that UTI was significantly higher among women with less than a secondary level of education. According to Dimetry *et al.*, those who were illiterate had the highest rate of UTIs among pregnant women, at 61.5% [24].

Multiparity has been linked to a twofold rise in UTI rates in pregnant women. This is because pregnancy causes major physiologic changes to the entire urinary system, which have a considerable impact on the natural history of UTI during gestation. These alterations differ from patient to patient and are more likely

to occur in women who have several pregnancies in a short period of time. According to our study, women with multiple pregnancy had the highest frequency (38.7%). Other studies, such as those conducted by Kant et al. (2017) on pregnant women in Haryana and Negussie et al. (2018) on pregnant women in the Ethiopia region, found an association between multiparity and UTI [5, 25]. In our study, there was a higher rate of infection in the second trimester (45%). Urinary tract infections are more common in the second trimester of pregnancy than in the first and third trimesters. This variation could be due to a shift in urine stasis and vesicoureteral reflux, or a drop in urinary progesterone and oestrogen levels throughout pregnancy. These findings are comparable to those reported by Onyango et al. (2018) in South Africa.²⁶

The most common uropathogens in our study were *E.coli* (60%), followed by *klebsiella species* (25%), *Enterococcus species* (10%) and *Candida species* (5%). Recent studies were conducted by Arumaikannu et al. (2021) among pregnant women of Tamil Nadu where the prevalence of *E. coli* was 50% and *Klebsiella* spp. was 20% [20]. Kaushal et al. (2021) reported a study in the region of Maharashtra among pregnant women where *E.coli* was the most commonly isolated bacteria at 62.5% and *Klebsiella* was 6.25% respectively [27]. Similar findings have been reported by Yashmin et al. (2018) in Bihar among pregnant women where prevalence of *E. coli* and *klebsiella species* was 57.14% and 8.16%, respectively [19, 27, 20].

Antibiotic resistance is a major public health concern around the world. Antibiotic resistance has emerged in the selection and spread of antibiotic resistant strains of bacterial pathogens, including uropathogens, due to the widespread use and misuse of antibiotics. Susceptibility testing of uropathogens, particularly *E. coli*, is advised, as is knowledge of local resistance and surveillance studies to monitor rising patterns of resistance [28]. In contrast, Kaushal et al. 2020 showed a susceptibility pattern of *E.coli* that was highly sensitive to meropenem (100%) and piperacillin-tazobactam (80%). In the study, *Klebsiella* was completely sensitive to meropenem and piperacillin tazobactam, while *Enterococcus* was completely sensitive to vancomycin [27]. A similar sensitivity pattern has been reported by Ali et al. 2020, where 96.7% of *E.coli* were highly sensitive to meropenem (96%), nitrofurantoin (80%), and gentamicin (78%). *K. pneumonia* showed sensitive to meropenem (100%), gentamycin (83%) and nitrofurantoin (66.7%) [29]. In Lucknow India Sonkar et al. 2021 reported that all Gram-negative isolates were mostly sensitive to most of the drugs like piperacillin-tazobactam, cefepime, nitrofurantoin, and meropenem [30]. The *E. coli* and *Klebsiella* species were highly sensitive to Nitrofurantoin (97.08%) reported by Shamim et al. 2018 among pregnant women of Bangalore India which gives a similar sensitivity pattern. Multidrug resistance was found in many of the isolates [31]. As a result, the current study provides insight into the common pattern of developing antibiotic resistance among uropathogens in this region, which could be the result of indiscriminate under dose antibiotic use. As a result, this information may aid physicians in the proper management of urinary tract infections and the avoidance of antibiotic resistance.

CONCLUSION

Due to enhancement of physiological changes during pregnancy UTI being the common complication. In our study isolated bacterial pathogens were shown to be resistant to routinely used antimicrobial drugs so it is important to do early screening of pregnant women for UTI infections and their antibiotic susceptibility pattern to prevent complications that could endanger both the mother and baby. The antenatal care physician should emphasize to all pregnant women about the importance of personal cleanliness and antibiotic resistance.

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