



UNVEILING THE DIFFERENCES: A CLINICO MICROBIOLOGICAL COMPARISON OF RECURRENT URINARY TRACT INFECTION IN PREMENOPAUSAL AND POSTMENOPAUSAL WOMEN

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ABSTRACT:

Recurrent urinary tract infection (rUTI) is defined as ≥ 2 episodes within 6 months or ≥ 3 episodes within 12 months. Urinary tract infections affect up to 50% of women in their lifetime, with almost half of these women experiencing a recurrence in 6–12 months. A subset of women experiences rUTI, with estimates ranging from 2 to 10%.

A cross-sectional study was performed to compare and assess the clinical characteristics and microbiological pattern of Recurrent Urinary Tract Infection in premenopausal women and postmenopausal women. A data collection was prepared to collect data from inpatient medical records regarding current presentation, susceptibility pattern of causative organisms and the prescription pattern of antimicrobials in Recurrent Urinary Tract Infection in a tertiary care hospital. Overall, 120 female patients from inpatient settings, diagnosed with recurrent UTI were included in the study. Of these, 50 patients were pre-menopausal and 70 were post-menopausal women.

Fever (51.6%), dysuria (41.6%) and urinary incontinence (20.8%) were the most common symptoms of the patients in UTI. Diabetes mellitus (61.66%) was the most common comorbidity found in the patients followed by hypertension (53.33%). The study data showed that the patients in both the groups were anaemic. Elevated RBS was observed in post-menopausal women (112.41 ± 45.03) when compared to pre-menopausal women (193.41 ± 132.40). The mean serum urea level (28.33 ± 23.58) and mean serum creatinine (0.55 ± 0.18) was higher in post-menopausal women. A significant difference in the proportions of patients with abnormal level of protein, RBC and epithelial cells in the urine between two groups was observed.

Escherichia coli [58 (48.33%)] was the most found causative agent of rUTI, followed by Enterococcus species [32 (26.67%)] and Klebsiella species [28 (23.33%)]. Multi-drug resistance was seen in 61 (50.83%) of the total 120 cases. The most resistant class of drugs found after assessing the culture reports are Penicillin antibiotics, Cephalosporins, Fluoroquinolones, Nitrofurantoin and Carbapenems. Ceftriaxone was the most prescribed antibiotic for rUTI followed by Piperacillin and tazobactam, Ciprofloxacin, Nitrofurantoin, Trimethoprim and Sulfamethoxazole, and Levofloxacin.

Keywords: Urinary tract infection, Antimicrobial susceptibility tests, Antimicrobial resistance, Multi drug resistance.

Introduction:

A urinary tract infection (UTI) is an infection in any part of the urinary system. It is defined as bacteriuria along with urinary symptoms. Urinary tract infection (UTI) is one of the most common infections in women. It is diagnosed in over 7 million women annually [1]. In acute cystitis, even 1,000 cfu/ml and, in acute pyelonephritis, 10,000 cfu/ml may be sufficient for diagnosis in a symptomatic patient. Recurrent urinary tract infection (rUTI) is defined as ≥ 2 episodes within 6 months or ≥ 3 episodes within 12 months [2]. Urinary tract infections affect up to 50% of women in their lifetime, with almost half of these women experiencing a recurrence in 6–12 months. A subset of women experiences rUTI, with estimates ranging from 2 to 10%. [3]

Cystitis (lower UTI) typically presents with local symptoms dysuria, frequency, urgency, pain, and/or bladder tenderness. Pyelonephritis (upper UTI) is associated with flank pain and signs of general infection such as fever, nausea and vomiting, or costovertebral angle tenderness, with or without cystitis symptoms. [4]

When bacterial virulence increases or host defense mechanisms decrease, bacterial inoculation, colonization, and infection of the urinary tract occurs. Careful diagnosis and treatment results in successful resolution of infections in most instances. Treatment of recurrent urinary tract infection requires

understanding of the pathogenesis of UTI and the role of host and bacterial factors.[5] UTIs can be caused by (Gram-positive or Gram-negative) bacteria, viruses or fungi, with more than 80% of UTIs being of bacterial origin. Most UTIs (75%) are caused by *Escherichia coli*, followed by *Klebsiella pneumoniae*, *Staphylococcus saprophyticus*, *Enterococcus faecalis* and *Proteus mirabilis*. [6]

According to several studies, risk factors associated with recurrent UTI in sexually active premenopausal women are sexual intercourse, use of spermicides, the age of first UTI (less than 15 years of age indicates a greater risk of rUTI). Menopause predisposes women to recurrent UTI (rUTI), as normally lower levels of estrogen lead to changes in the urogenital epithelium and subsequently the urogenital microbiome [7]. In postmenopausal women Utero-vaginal prolapse, urinary incontinence and post void residual urine are important risk factors [8].

Methodology:

Study design: A cross-sectional study was performed to compare and assess the clinical characteristics and microbiological pattern of Recurrent Urinary Tract Infection in premenopausal women and postmenopausal women. A data collection was prepared to collect data from inpatient medical records regarding current presentation, susceptibility pattern of causative of organisms and the prescription pattern of antimicrobials in Recurrent Urinary Tract Infection in a tertiary care hospital.

Study site: Srinivas Institute of Medical Science and Research Centre, Mukka, Mangalore.

Study design: A hospital-based observational study.

Study period: 6 months

Study sample size: 120

Ethics committee: Srinivas Institute of Medical Science and Research Centre, Mukka –574146

Study criteria:

Inclusion criteria:

- Female patients from inpatient settings diagnosed with rUTI
- Patients with positive urine culture and sensitivity report

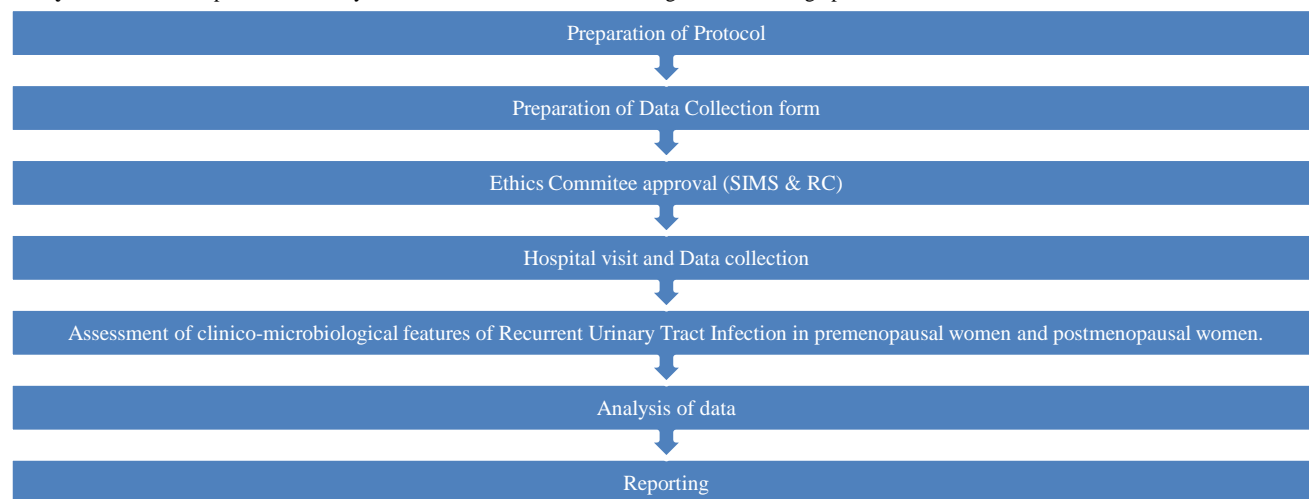
Exclusion criteria:

- Patients below 18 years of age
- Patients suffering from renal impairment

Data source: Medical records from inpatient and outpatient settings

Materials used: Data collection form

Study method: A hospital-based study which will be conducted according to the following operational modalities.



Statistical analysis:

Results were expressed as mean \pm standard error of mean. All the data were analysed using a one-way analysis of variance, followed by Dunnett's test (* $p < 0.05$)

Results

Patient Socio-demographics

Overall, 120 female patients diagnosed with recurrent UTI were included in the study. Of these, 50 patients were pre-menopausal and 70 were post-menopausal women (Figure 01). The mean age of pre-menopausal women was 24.91 ± 12.83 years and post-menopausal was 61.10 ± 10.54 years.

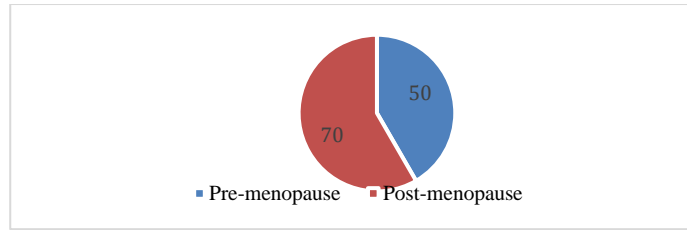


Figure 01: Distribution of subjects based on menopause status

Clinical Characteristics

Fever (51.6%), dysuria (41.6%) and urinary incontinence (20.8%) were the most common clinical presentation of the patients in UTI. Comparison of clinical characteristics between pre and post-menopausal patients are shown in Table 01 and Figure 02.

Table 01: Clinical characteristics between pre and post-menopausal patients

| Symptoms | Total (%) | Pre-menopausal women n (%) | Post- menopausal women n (%) | P value |
|----------------------|------------|----------------------------|------------------------------|---------|
| Fever | 62 (51.6) | 25 (50) | 37 (52.8) | >0.05 |
| Dysuria | 50 (41.6) | 23 (46) | 27 (38.5) | >0.05 |
| Urinary incontinence | 25 (20.83) | 08 (17) | 17 (24.2) | >0.05 |
| Abdominal pain | 18 (15) | 11 (22) | 7 (10) | >0.05 |
| Vaginal Thrush | 11 (09) | 8 (17) | 3 (04) | >0.05 |
| Vomiting | 7 (06) | 3 (06) | 4 (05) | >0.05 |
| Skin rashes | 3 (02) | 3 (06) | 0 (0) | >0.05 |

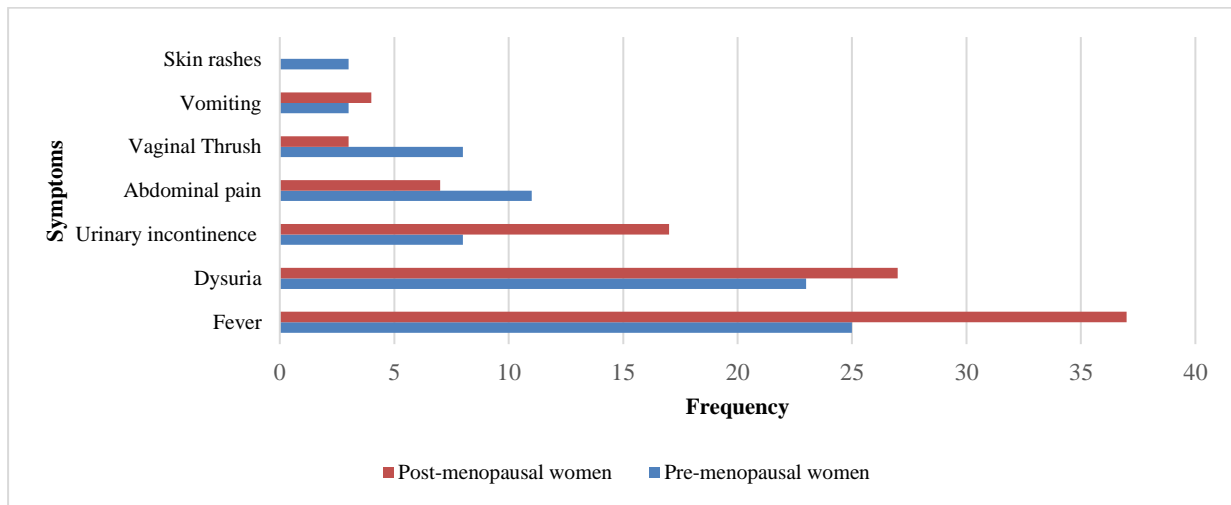


Figure 02: Clinical characteristics between pre and post-menopausal patients

Medical history of the patients was studied and it was found that out of 120 patients, 74 (61.66%) were diabetic. 64 (53.33%) were suffering from hypertension. Other comorbidities present in the patients were Hypothyroidism (17.5%), IHD (15%), COPD (3.33%), Atrial fibrillation (3.33%) and CAD (5.83%), as shown in Table 02 and Figure 03.

Table 02: Frequency of comorbidities

| Sl. No | Co-Morbidities | Frequency | Percentage |
|--------|---------------------|-----------|------------|
| 1 | Hypertension | 64 | 53.33 |
| 2 | Diabetes | 74 | 61.66 |
| 3 | IHD | 18 | 15 |
| 4 | COPD | 4 | 3.33 |
| 5 | Atrial fibrillation | 4 | 3.33 |
| 6 | CAD | 7 | 5.83 |
| 7 | Hypothyroidism | 21 | 17.5 |

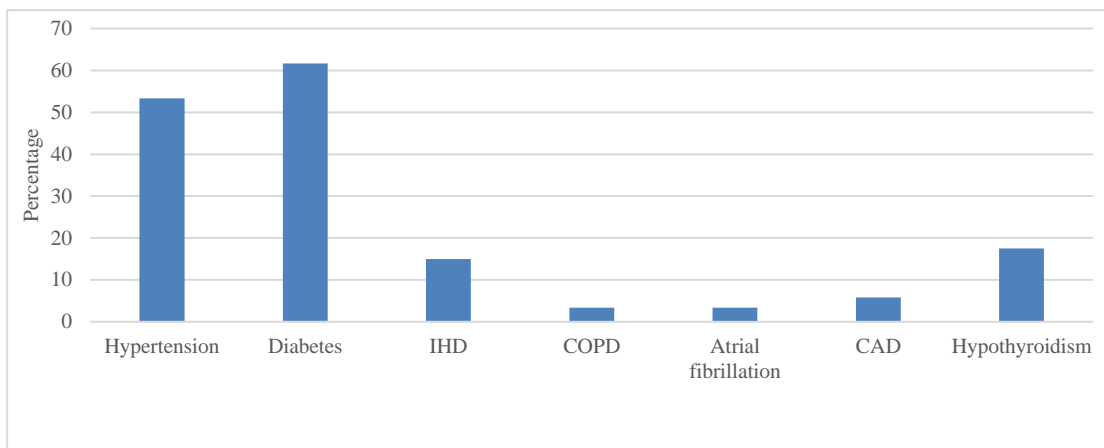


Figure 03: Frequency of comorbidities

The study data showed that the patients in both the groups were anaemic and there was no significant difference in the mean hemoglobin levels and RBC count between the groups. The mean Total Leukocyte Count was 11662.5 ± 6804.92 and 10126.21 ± 5342.05 in pre and post-menopausal women respectively which was higher than the normal range (3000-10000 cells/cumm of blood). However, the difference was found to be non-significant. According to the study data, patients of both the groups had elevated Erythrocyte Sedimentation Rate. In comparison, there was a significant increase of the Erythrocyte Sedimentation Rate in post- menopausal women (22.58 ± 19.84) than pre-menopausal women (55.96 ± 64.07).

Biochemical investigations showed significant difference of RBS, serum urea and serum creatinine between pre-menopausal women and post-menopausal women as shown in Table 03. Elevated RBS was observed in post-menopausal women (112.41 ± 45.03) when compared to pre-menopausal women (193.41 ± 132.40). The mean serum urea level was observed to be higher post-menopausal women (28.33 ± 23.58) in contrast to pre-menopausal women (44.4 ± 10.54). It was observed that the mean value of serum creatinine was higher in post-menopausal women (0.55 ± 0.18) compared to pre-menopausal women (1.41 ± 1.36).

Table 03: Comparison of laboratory parameters between pre and post-menopausal patients

| Parameters | Pre-menopause mean±SD | Post-menopause mean±SD | P value |
|--------------------------------|--------------------------|---------------------------|-----------|
| Hemoglobin | 10.72±1.95 | 10.87±2.49 | 0.71 |
| RBC | 3.95 ± 0.67 | 3.81±0.75 | 0.29 |
| Total Leukocyte Count | 11662.5 ± 6804.92 | 10126.21±5342.05 | 0.16 |
| Erythrocyte Sedimentation Rate | 22.58± 19.84 | 55.96±64.07 | <0.01** |
| RBS | 112.41 ± 45.03 | 193.41±132.40 | <0.001*** |
| Urea | 28.33 ± 23.58 | 44.4 ± 10.54 | <0.001*** |
| Creatinine | 0.55 ± 0.18 | 1.41 ± 1.36 | <0.001*** |

Values are expressed as mean ±SEM; n=6. One-way ANOVA followed by Dunnet’s t-test compared to control. *p<0.05, **p<0.01, ***p<0.001.

Out of the 120 patients, 6 (05%) patients had brownish coloured urine (1 in pre and 5 in post menopausal women). In total 80 (66.8), 29 (24.2) and 11 (09.2) patients had urine pH value of 5,6 and 6.5 respectively. Total of 92 (76.7%) patients had turbid urine. 31 (25.8%) of the patients had sugar present in urine. 31 (25.8%) out of 120 patients also had presence of protien in the urine.

Microscopical analysis of urine showed that 61 (50.8%) out of the total 120 patients had higher than normal levels of RBC in the urine. 80 (67%) had increased amount of pus cells and 54 (45%) had abnormal value of epithelial cells in the urine. The study data from urine analysis revealed significant difference in the proportions of patients with abnormal level of protien, RBC and epithelial cells in the urine between pre and post-menopausal women, as shown in Table 04.

Table 04: Comparison of Urine analysis between pre and post-menopausal patients

| Parameters | | Total n(%) | Pre-menopause n (%) | Post-menopause n (%) | P value |
|------------|-------------|---------------|------------------------|-------------------------|---------|
| Colour | Pale yellow | 114 (95) | 49 (98) | 65 (92.8) | 0.20 |
| | Brownish | 6 (05) | 1 (2) | 5 (7.14) | |
| pH | 5 | 80 (66.8) | 32 (63.63) | 48 (68) | 0.69 |

| | | | | | |
|--|---------|-----------|------------|------------|---------|
| | 6 | 29 (24.2) | 14 (27.27) | 15 (21.4) | |
| | 6.5 | 11 (09.2) | 4 (9.09) | 7 (10.71) | |
| Transparency | Clear | 28 (23.3) | 13 (26) | 15 (21.42) | 0.56 |
| | Turbid | 92 (76.7) | 37 (74) | 55 (78.6) | |
| Sugar | Present | 31 (25.8) | 9 (18) | 22 (32) | 0.098 |
| | Absent | 89 (74.2) | 41 (82) | 48 (68) | |
| Blood | Present | 34 (28.3) | 14 (27.27) | 20 (28.57) | 0.945 |
| | Absent | 86 (71.7) | 36 (72.73) | 50 (71.42) | |
| Protein | Present | 31 (25.8) | 4 (9.09) | 27 (39.2) | <0.01** |
| | Absent | 89 (74.2) | 46 (90.91) | 43 (60.71) | |
| RBC | 0-2/hpf | 59 (49.2) | 4 (9.09) | 53 (75) | <0.01** |
| | >2/hpf | 61 (50.8) | 46 (90.91) | 17 (25) | |
| Pus cells | 0-5/hpf | 40 (33) | 18 (36.36) | 22 (32.14) | 0.60 |
| | >5/hpf | 80 (67) | 32 (63.63) | 48 (67.86) | |
| Epithelial cells | 0-3/hpf | 66 (55) | 13 (27.27) | 53 (75) | <0.01** |
| | >3/hpf | 54 (45) | 37 (72.73) | 17 (25) | |
| Values are expressed as mean \pm SEM; n=6. One-way ANOVA followed by Dunnet's t-test compared to control. *p<0.05, **p<0.01, ***p<0.001. | | | | | |

MICROBIOLOGICAL PROFILE IN RECURRENT UTI

By assessing the culture reports of the patients, it was found that in both pre and post-menopausal women *Escherichia coli* [58 (48.33%)] was the most found causative agent of rUTI, followed by *Enterococcus* species [32 (26.67%)] and *Klebsiella* species [28 (23.33%)]. However, the incidence of *E. coli* caused rUTI was significantly higher in pre-menopausal women compared to post-menopausal women. List of organisms isolated by urine culture of the patients are given in Table 05 and Figure 04.

Table 05: Frequency and distribution pattern of pathogens

| Sl. No. | Organism | Total | | Pre-menopausal women | | Post-menopausal women | |
|---------|---|-----------|-------|----------------------|----|-----------------------|-------|
| | | Frequency | % | Frequency | % | Frequency | % |
| 1. | <i>Escherichia coli</i> | 58 | 48.33 | 19 | 38 | 39 | 55.71 |
| 2. | <i>Enterococcus faecium</i> | 28 | 23.33 | 8 | 08 | 20 | 28.57 |
| 3. | <i>Enterococcus faecalis</i> | 4 | 3.33 | 0 | 00 | 4 | 5.71 |
| 4. | <i>Klebsiella pneumonia</i> | 21 | 17.5 | 6 | 12 | 15 | 21.42 |
| 5. | <i>Klebsiella oxytoca</i> | 7 | 5.83 | 0 | 00 | 7 | 10 |
| 6. | <i>Pseudomonas aeruginosa</i> | 8 | 6.67 | 2 | 00 | 6 | 10 |
| 7. | Group b Beta Haemolytic <i>Streptococcus agalactiae</i> | 11 | 9.17 | 2 | 04 | 9 | 12.85 |
| 8. | <i>Proteus mirabilis</i> | 11 | 9.17 | 3 | 06 | 8 | 11.42 |
| 9. | Coagulase Negative <i>Staphylococcus</i> | 4 | 3.33 | 4 | 08 | 0 | 0 |
| 10. | Gram B Beta Haemolytic <i>Streptococci</i> | 4 | 3.33 | 4 | 08 | 0 | 0 |
| 11. | <i>Candida species</i> | 9 | 7.5 | 6 | 12 | 3 | 4.28 |

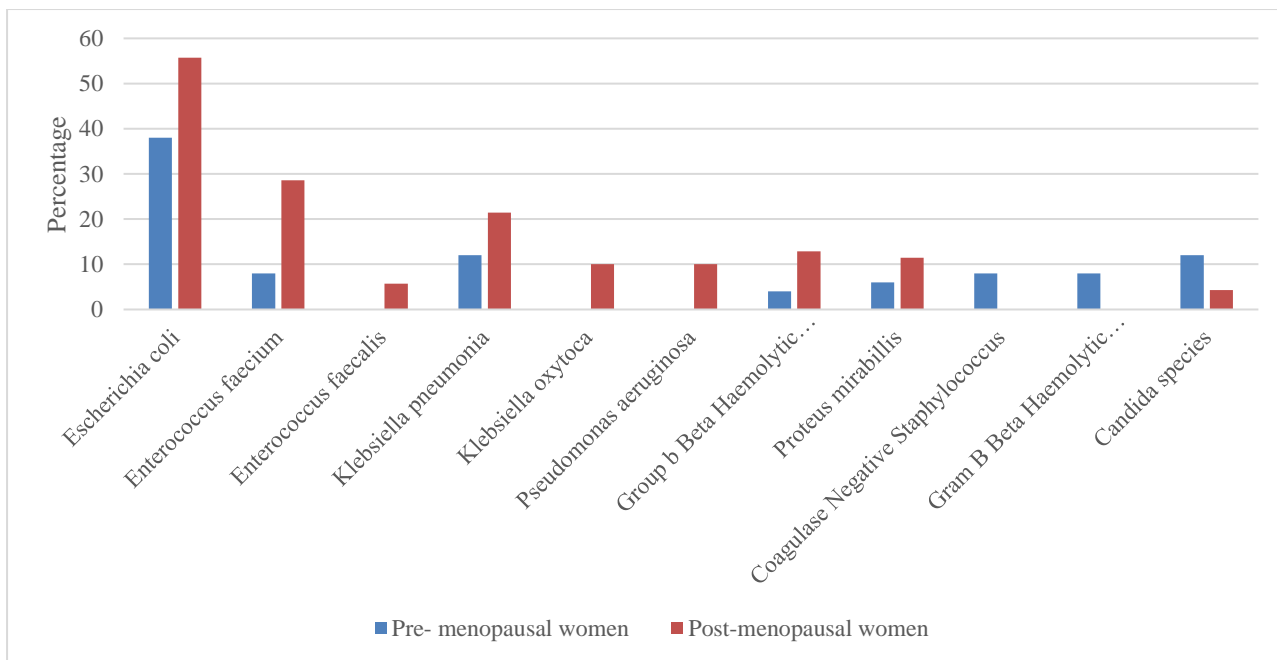


Figure 04: Frequency and distribution pattern of pathogens

Multi-drug resistance was seen in 61 (50.83%) of the total 120 cases. Prevalence of drug resistance pathogens are depicted in Table 06 and Figure 05. It was found that there was significant difference in the prevalence of Multi-drug Resistance (MDR) in pathogens ($p < 0.05$).

Table 06: Prevalence pattern of Multi-drug Resistance (MDR) in pathogens

| Sl. No. | Organism | Total | | Pre-menopausal women | | Post-menopausal women | | P value |
|---------|--|-----------|-------|----------------------|----|-----------------------|-------|---------|
| | | Frequency | % | Frequency | % | Frequency | % | |
| 1. | Escherichia coli | 25 | 20.83 | 10 | 20 | 15 | 21.42 | <0.05* |
| 2. | Enterococcus faecium | 21 | 17.5 | 5 | 10 | 16 | 22.85 | |
| 3. | Enterococcus faecalis | 6 | 5 | 1 | 2 | 5 | 7.14 | |
| 4. | Klebsiella pneumonia | 4 | 3.33 | 2 | 4 | 2 | 2.85 | |
| 5. | Klebsiella oxytoca | 7 | 5.83 | 1 | 2 | 6 | 8.57 | |
| 6. | Pseudomonas aeruginosa | 21 | 17.5 | 13 | 26 | 8 | 11.42 | |
| 7. | Group b Beta Haemolytic Streptococcus agalactiae | 32 | 26.67 | 13 | 26 | 19 | 27.14 | |
| 8. | Proteus mirabilis | 8 | 6.67 | 2 | 4 | 6 | 8.57 | |
| 9. | Coagulase Negative Staphylococcus | 1 | 0.83 | 0 | 0 | 1 | 1.43 | |
| 10. | Gram B Beta Haemolytic Streptococci | 1 | 0.83 | 0 | 0 | 1 | 1.43 | |

Values are expressed as mean \pm SEM; n=6. One-way ANOVA followed by Dunnet's t-test compared to control. * $p < 0.05$, ** $p < 0.01$, *** $p < 0.001$.

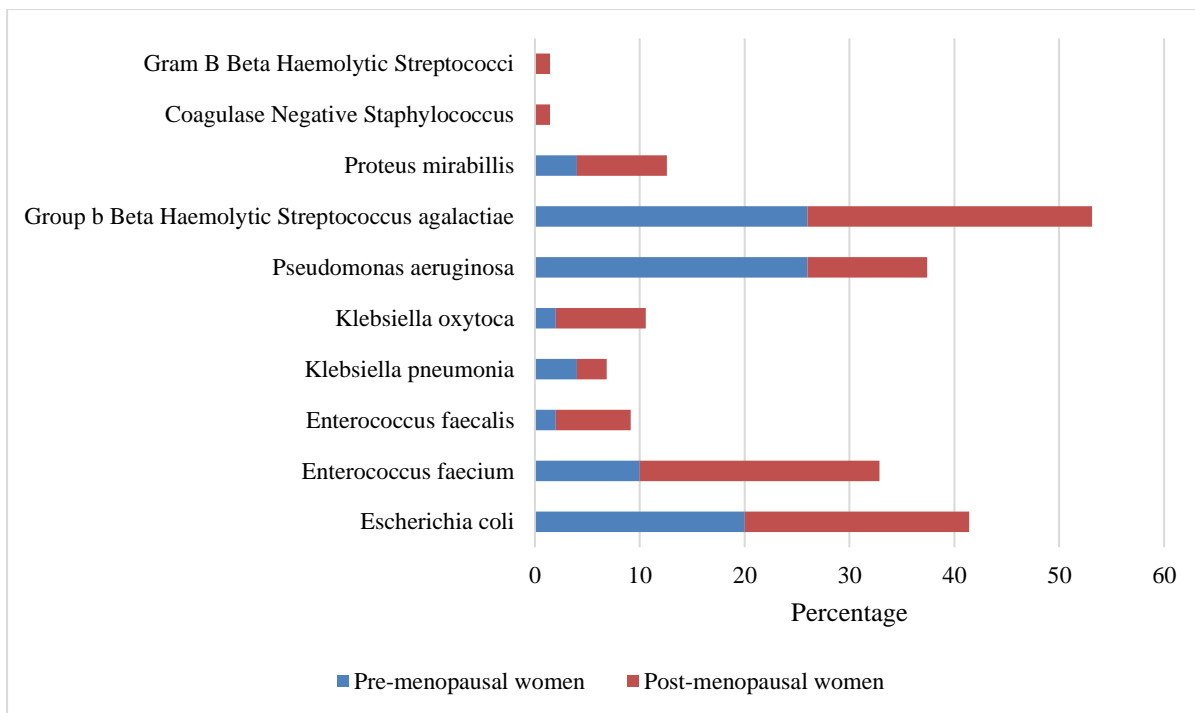


Figure 05: Prevalence pattern of Multi-drug Resistance (MDR) in pathogens

The most resistant class of drugs found after assessing the culture reports are Penicillin antibiotics, Cephalosporins, Fluoroquinolones, Nitrofurantoin and Carbapenems. Resistance pattern and its distribution in pre and post-menopausal women is depicted in Table 07 and Figure 06. Resistance pattern of individual drugs in pre and post-menopausal women is shown in Table 08.

Table 07: Resistance pattern and distribution of the pathogens to various antibiotic class

| Sl. No. | Antimicrobial class | Total | | Pre-menopausal women | | Post-menopausal women | |
|---------|-----------------------------------|-----------|-------|----------------------|----|-----------------------|-------|
| | | Frequency | % | Frequency | % | Frequency | % |
| 1. | Penicillin antibiotics | 67 | 55.83 | 35 | 70 | 32 | 45.71 |
| 2. | Cephalosporins | 55 | 45.83 | 25 | 50 | 30 | 42.85 |
| 3. | Fluoroquinolones | 53 | 44.17 | 15 | 30 | 38 | 54.28 |
| 4. | Nitrofurantoin | 43 | 35.83 | 20 | 40 | 23 | 32.85 |
| 6. | Carbapenems | 40 | 33.33 | 20 | 40 | 20 | 28.57 |
| 7. | Aminoglycosides | 29 | 24.16 | 20 | 40 | 9 | 12.85 |
| 8. | Trimethoprim and sulfamethoxazole | 15 | 12.5 | 3 | 6 | 12 | 17.14 |
| 9. | Linezolid | 8 | 6.66 | 4 | 8 | 4 | 5.71 |
| 10. | Clindamycin | 6 | 5 | 0 | 0 | 6 | 8.57 |
| 11. | Glycopeptides | 6 | 5 | 0 | 0 | 6 | 8.57 |

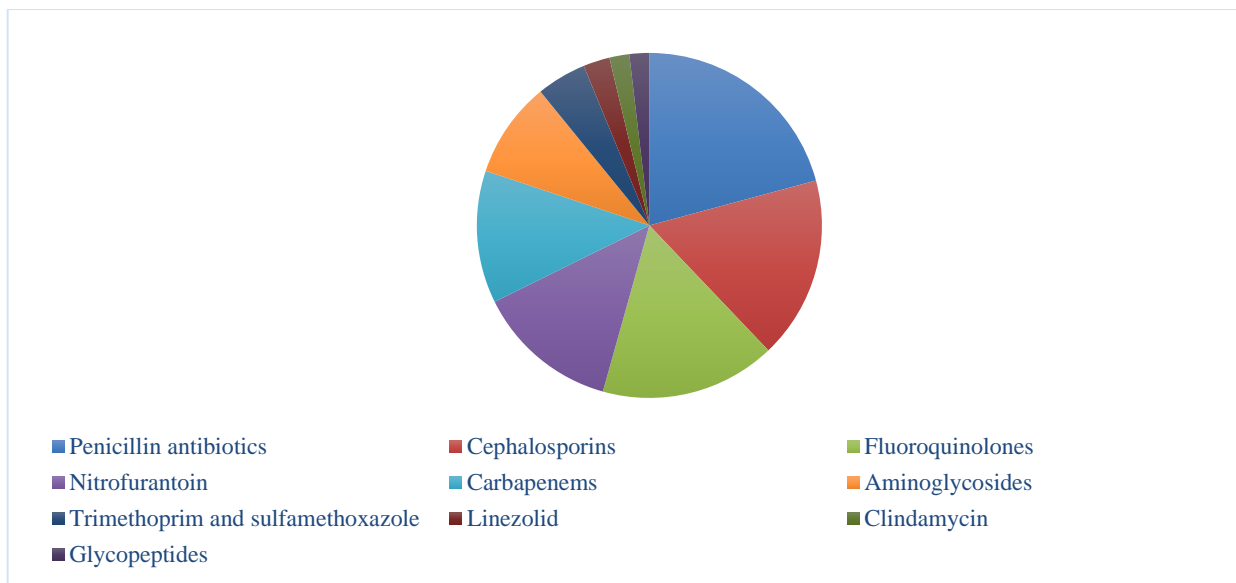


Figure 06: Resistance pattern and distribution of pathogens to various antibiotic class

Table 08: Resistance pattern and distribution of the pathogens to various antibiotics

| Sl. No. | Antimicrobial | Total | | Pre-menopausal women | | Post-menopausal women | |
|---------|---------------------------------------|-----------|-------|----------------------|----|-----------------------|-------|
| | | Frequency | % | Frequency | % | Frequency | % |
| 1. | Ampicillin | 67 | 55.83 | 35 | 70 | 32 | 45.71 |
| 2. | Ciprofloxacin | 55 | 45.83 | 25 | 50 | 30 | 42.85 |
| 3. | Levofloxacin | 53 | 44.17 | 15 | 30 | 38 | 54.28 |
| 4. | Cefazolin | 45 | 37.5 | 15 | 30 | 30 | 42.85 |
| 5. | Nitrofurantoin | 43 | 35.83 | 20 | 40 | 23 | 32.85 |
| 6. | Imipenem | 40 | 33.33 | 20 | 40 | 20 | 28.57 |
| 7. | Amoxicillin and Potassium Clavulanate | 36 | 30 | 10 | 20 | 26 | 37.14 |
| 8. | Gentamicin | 35 | 29.16 | 15 | 30 | 20 | 28.57 |
| 9. | Meropenem | 30 | 25 | 10 | 20 | 20 | 28.57 |
| 10. | Cefuroxime | 30 | 25 | 10 | 20 | 20 | 28.57 |
| 11. | Amikacin | 29 | 24.16 | 20 | 40 | 9 | 12.85 |
| 12. | Cefoperazone and Sulbactam | 25 | 20.83 | 5 | 10 | 20 | 28.57 |
| 13. | Piperacillin | 22 | 18.33 | 5 | 10 | 17 | 24.28 |
| 14. | Cefotaxime | 20 | 16.66 | 5 | 10 | 15 | 21.43 |
| 15. | Trimethoprim and sulfamethoxazole | 15 | 12.5 | 3 | 6 | 12 | 17.14 |
| 16. | Ceftriaxone | 10 | 8.33 | 4 | 8 | 6 | 8.57 |
| 17. | Linezolid | 8 | 6.66 | 4 | 8 | 4 | 5.71 |
| 18. | Clindamycin | 6 | 5 | 2 | 4 | 4 | 5.71 |
| 19. | Vancomycin | 6 | 5 | 0 | 0 | 6 | 8.57 |
| 20. | Teicoplanin | 6 | 5 | 0 | 0 | 6 | 8.57 |
| 21. | Ampicillin + gentamycin synergy | 6 | 5 | 0 | 0 | 6 | 8.57 |

| | | | | | | | |
|-----|-------------|---|------|---|---|---|------|
| 22. | Cefixime | 3 | 2.5 | 1 | 2 | 2 | 2.85 |
| 23. | Ceftazidime | 2 | 1.66 | 1 | 2 | 1 | 1.42 |

Antibiotic prescription pattern in recurrent uti

Assessment of the prescriptions of the patients showed that Ceftriaxone (40.83%) was the most prescribed antibiotic for rUTI followed by Piperacillin and tazobactam (16.67%), Ciprofloxacin (10.83%), Nitrofurantoin (7.5%), Trimethoprim and Sulfamethoxazole (5.83%), Levofloxacin (5.83%), Meropenem (3.33%), Amikacin (2.5%), Amoxicillin and Potassium Clavulanate (2.5%), Clindamycin (0.83%), Linezolid (0.83%), Cefixime (0.83%), Ceftazidime (0.83%) and Cefuroxime (0.83%), as shown in Table 09 and Figure 07.

Table 09: Prescription pattern of antimicrobials in pre and post-menopausal women

| Sl. No. | Antimicrobial | Total | | Pre-menopausal women | | Post-menopausal women | |
|---------|---------------------------------------|-----------|-------|----------------------|----|-----------------------|-------|
| | | Frequency | % | Frequency | % | Frequency | % |
| 1. | Ceftriaxone | 49 | 40.83 | 21 | 42 | 28 | 40 |
| 2. | Piperacillin | 20 | 16.67 | 9 | 18 | 11 | 15.71 |
| 3. | Ciprofloxacin | 13 | 10.83 | 10 | 20 | 3 | 4.28 |
| 4. | Nitrofurantoin | 9 | 7.5 | 8 | 16 | 1 | 1.43 |
| 5. | Trimethoprim and sulfamethoxazole | 7 | 5.83 | 2 | 04 | 5 | 7.14 |
| 6. | Levofloxacin | 7 | 5.83 | 7 | 14 | 0 | 0 |
| 7. | Meropenem | 4 | 3.33 | 1 | 02 | 3 | 4.28 |
| 8. | Amikacin | 3 | 2.5 | 0 | 0 | 3 | 4.28 |
| 9. | Amoxicillin and potassium clavulanate | 3 | 2.5 | 3 | 06 | 0 | 0 |
| 10. | Clindamycin | 1 | 0.83 | 1 | 02 | 0 | 0 |
| 11. | Linezolid | 1 | 0.83 | 0 | 0 | 1 | 1.43 |
| 12. | Cefixime | 1 | 0.83 | 1 | 02 | 0 | 0 |
| 13. | Ceftazidime | 1 | 0.83 | 1 | 02 | 0 | 0 |
| 14. | Cefuroxime | 1 | 0.83 | 1 | 02 | 0 | 0 |

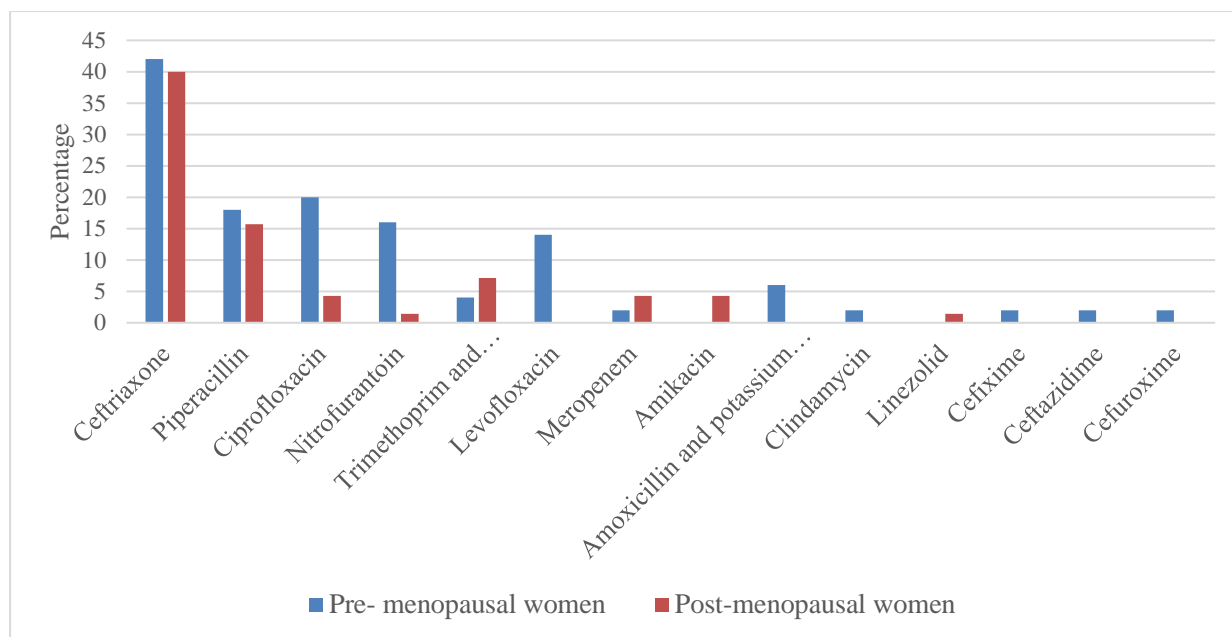


Figure 07: Prescription pattern of antimicrobials in pre and post-menopausal women

Discussion :

Recurrent UTI remains an important public health problem in women of all ages. The data reported here and in other recent studies of recurrent UTI suggest that the factors underlying rUTI differ by age and functional status. The study aimed at comparing the clinico-microbiological pattern between pre and post-menopausal women. A total of 120 patients suffering from rUTI were included in the study, out of which 50 were in pre-menstrual period and 70 were in post-menstrual period.

According to the study data, it was found that fever, dysuria and urinary incontinence were the most common clinical presentation of the patients in UTI. These results were consistent with study done by Eshwarappa M. *et al.*, which showed that most observed clinical symptoms of rUTI in women were fever and dysuria. [9] Assessment of medical history of the patients was study showed that most of the patients were diabetic. Diabetes may cumulate to the risk of developing rUTI, as indicated by a study done by Subramani J. *et al.*, which showed that besides the micro and macrovascular complications, diabetes can dysregulate the immune mechanisms and thereby increasing the risk of UTIs.[10]

It was observed that both the group had lower levels of hemoglobin. This can be explained by the study done by Shobha B. *et al.*, which showed that the hemoglobin levels in pre-menopausal women is affected by menstruation, whereas co-morbidities in old age might affect the hemoglobin levels of women after menopause. Interpretation of the study data showed that there was a significant increase of the Erythrocyte Sedimentation Rate in post- menopausal women compared to pre-menopausal women.[11]

Similarly, biochemical investigations such as RBS, serum urea and serum creatinine between pre-menopausal women and post- menopausal women showed significant difference. According to the study, most common abnormal results obtained by urine routine and microscopy were turbid urine, proteinuria and presence of elevated pus and epithelial cells. A possible explanation for these variations could be explained by the results of the study done by Oluboyo A.O *et al.*, which concluded that metabolic changes and the appearance of many diseases tend to occur following the onset of menopause that may cause further damage to kidney functions.[12]

The study data revealed that the most common pathogen of rUTI was E. coli, followed by Enterococcus species and Klebsiella species. These results were consistent with a study conducted by Bhuvanesh S. K *et al.* in a tertiary care hospital of South India.[13]

The proportion of causative organisms differed significantly pre-menopausal women in comparison with post-menopausal women. these results were consistent with the study conducted by Mazokopakis E. *et al.*, where there was noticeable difference in the percentages of causative organisms. Decrease in estrogenic levels after menopause leads to decrease in rate of replenishment of normal flora of urinary tract, allowing infections by various microorganisms.[14]

Assessment of urine cultures of the patients revealed that a significant proportion of isolated bacteria were multi-drug resistant. Prevalence of drug resistance pathogens were significantly higher in post-menopausal women compared to pre-menopausal women. these results might be the result of imbalanced natural flora of urinary system and prolonged exposure to antibiotics leading to increased antibiotic resistance in post-menopausal women. According to susceptibility reports studied, most resistant class of drugs to pathogens were Penicillin antibiotic followed by Cephalosporins, Fluoroquinolones, Nitrofurantoin and Carbapenems. Similar results were concluded in a study conducted by Amna K *et al.*[15]

Most prescribed antibiotics were Ceftriaxone followed by Piperacillin and tazobactam, Ciprofloxacin, Nitrofurantoin, Trimethoprim and Sulfamethoxazole, Levofloxacin, Meropenem and Amikacin. These results were consistent with the findings of a study carried out by Chowta M. *et al.*[16] The possible reason is that these agents are used as first-line drugs for the treatment of UTIs in the country.

Future Prospectives:

The results obtained from our study can help in the following aspects:

- Identification of the right drug for initiating empiric treatment of Recurrent UTI
- Aiding in early detection and diagnosis of Recurrent UTI.
- Creating awareness in healthcare professionals concerning antimicrobial resistance.

Conclusion :

The present study aimed at providing information of current scenario of clinical features and microbiological susceptibility data of pre- menopausal and post-menopausal women suffering from rUTI. Clinical symptoms of rUTI were similar between both the groups. However, Biochemical investigations showed that rUTI had higher physiological effects on the patients after menopause. Most of the pathogens isolated by urine culture were found to be multi-drug resistance indicating the need for proper diagnosis and rational use of antibiotics.

The cause for such increase in resistance to various antibiotics can be controlled if proper empiric treatment is being administered to the patients. This would also help to minimize further complications and better recovery from rUTI.

Periodic assessment of antibiotic susceptibility of pathogens causing UTI is necessary to revise the guidelines for administering initial treatment. The prevalence of pathogen and its resistance can vary between various ethnicities and geographical locations. Hence, it is recommended to regularly evaluate the susceptibility of infective microorganisms.

REFERENCES:

1. Vagios S, Hesham H, Mitchell C. Understanding the potential of lactobacilli in recurrent UTI prevention. *Microbial Pathogenesis*. 2020 Nov 1;148:104544.
2. Aydin A, Ahmed K, Zaman I, Khan MS, Dasgupta P. Recurrent urinary tract infections in women. *International urogynecology journal*. 2015 Jun;26:795-804.
3. Al-Badr A, Al-Shaikh G. Recurrent urinary tract infections management in women: a review. *Sultan Qaboos University Medical Journal*. 2013 Aug;13(3):359.
4. Grabe M, Bjerklund-Johansen TE, Botto H, Wullt B, Cek M, Naber KG, Pickard RS, Tenke P, Wagenlehner F. Guidelines on urological infections. *Eur Assoc Urol*.
5. Masson P, Matheson S, Webster AC, Craig JC. Meta-analyses in prevention and treatment of urinary tract infections. *Infectious disease clinics of North America*. 2009 Jun 1;23(2):355-85.
6. Kang CI, Kim J, Park DW, Kim BN, Ha US, Lee SJ, Yeo JK, Min SK, Lee H, Wie SH. Clinical practice guidelines for the antibiotic treatment of community-acquired urinary tract infections. *Infection & chemotherapy*. 2018 Mar;50(1):67.
7. Jung C, Brubaker L. The etiology and management of recurrent urinary tract infections in Postmenopausal women. *Climacteric*. 2019 May 4;22(3):242-9.
8. Scholes D, Hooton TM, Roberts PL, Stapleton AE, Gupta K, Stamm WE. Risk factors for recurrent urinary tract infection in young women. *The Journal of infectious diseases*. 2000 Oct 1;182(4):1177-82.
9. Eshwarappa M, Dosegowda R, Aprameya IV, Khan MW, Kumar PS, Kempegowda P. Clinico-microbiological profile of urinary tract infection in south India. *Indian journal of nephrology*. 2011 Jan;21(1):30.
10. Jagadeesan S, Tripathi BK, Patel P, Muthathal S. Urinary tract infection and Diabetes Mellitus—Etiological profile and antibiogram: A North Indian perspective. *Journal of Family Medicine and Primary Care*. 2022 May;11(5):1902.
11. Bembalagi S, Sojitra MP. Study of relationship between anemia and urinary tract infection in pregnant women. *Al Ameen J Med Sci*. 2018;11(1):71-5.
12. Oluboyo AO, Ucheagwu DC, Meludu SC, Oluboyo BO, Olayanju AO, Odewusi OO. EVALUATION OF SOME INFLAMMATORY AND MUSCLE MARKERS IN PREMENOPAUSAL AND POSTMENOPAUSAL WOMEN. *World Journal of Pharmaceutical and Life Sciences*. 2017 Jul 2;3(1):1-8.
13. Kalal BS, Nagaraj S. Urinary tract infections: a retrospective, descriptive study of causative organisms and antimicrobial pattern of samples received for culture, from a tertiary care setting. *Germes*. 2016 Dec;6(4):132.
14. Mazokopakis E, Potolidis E. Recurrent Urinary Tract Infections (Rutis) In Pre-Menopausal And Post-Menopausal Women. A Retrospective Study. *The Internet Journal of Urology*. 2012;9(3).
15. Kausar A, Akram M, Shoaib M, Mehmood RT, Abbasi MN, Adnan M, Aziz H, Asad MJ. Isolation and identification of UTI causing agents and frequency of ESBL (extended spectrum beta lactamase) in Pakistan. *Amer JPhytomed Clin Ther*. 2014;2:963-75.
16. Chowta DM. Drug Utilization Pattern in Urinary tract infections: A Retrospective study. *Journal of Medical Science & Clinical Research*. 2011;1:13-9.