

## ANTIBIOTIC RESISTANCE PATTERNS OF ESCHERICHIA COLI IN URINARY TRACT INFECTIONS AT A TERTIARY CARE HOSPITAL, LAHORE

Amina Asghar,<sup>1</sup> Saadia Ijaz,<sup>2</sup> Mariam Danish Iqbal,<sup>3</sup> Muhammad Masood Afzal,<sup>4</sup> Asma Ejaz<sup>5</sup>

### ABSTRACT

**Background and objective:** As a chronic nosocomial and community-acquired infections, Escherichia coli (E. coli) has become a major global health concern. Finding the pattern of antibiotic susceptibility in Escherichia coli isolated from urine specimens was the aim of the current study.

**Methods:** It was descriptive Cross-Sectional study conducted at Shalamar Teaching Hospital in Lahore in the Micro-biology section of the Pathology Department from June 1<sup>st</sup>, 2020 to November 30<sup>th</sup>, 2020. Patients exhibiting symptoms and indications of urinary tract infections (UTIs) had urine samples taken. The samples were processed on the Cystine Lysine Electrolyte-Deficient (CLED) Agar and identified as E. coli by biochemical profile. Antibiogram of E. coli was determined by Kirby-Bauer method.

**Results:** A total of 165 E. coli isolates were available for analysis. Among the tested antibiotics, Colistin exhibited the highest sensitivity of 164(99.4%), followed by Fosfomycin 156(94.5%), Imipenem 153 (92.7%), and Meropenem 150(90.9%). Nitrofurantoin demonstrated a sensitivity of 141(85.5%), while Gentamicin had a sensitivity of 110(66.7%). Piperacillin/Tazobactam, Cotrimoxazole, Ceftriaxone and Cefotaxime, Ciprofloxacin, Cefuroxime, Amoxicillin-Clavulanic Acid, and Ampicillin exhibited sensitivities of 60(36.4%), 31(18.8%), 28(17.0%), 28(17.0%), 24(14.5%), 13 (7.9%), 11( 6.7%), and 3(1.8%), respectively. Notably, E. coli was isolated more frequently from female patients (72.7%) than from male patients (27.3%).

**Conclusion:** Current study revealed that antibiotic resistance is emerging in E coli. However, Colistin, Fosfomycin, Imipenem, Meropenem, and Nitrofurantoin are the most effective drugs and, depending on the clinical situation, can be chosen empirically for the treatment of UTI caused by E. coli.

**Key words:** Urinary tract infections, E. Coli, Sensitivity pattern.

**How to cite:** Asghar A, Ijaz S, Iqbal MD, Afzal MM, Ejaz A. Antibiotic Resistance Patterns of Escherichia coli in Urinary Tract Infections at a Tertiary Care Hospital, Lahore. JAIMC. 2024;22(2): 40-44

In humans, pathogenic bacteria cause several infections. The Urinary Tract is a sterile system except for the lower urethra which may have some normal flora. The ascending invasion of the system by bacteria is the cause of urinary tract infections (UTIs). The UTI

caused at different levels of the tract by microbial colonization or inflammation is called Urethritis (Urethra), Cystitis (Urinary Bladder), and Pyelonephritis (Kidneys).<sup>1,2</sup>

The bacteria causing UTIs are many and varied. Of them, the most prevalent are the Gram-Negative Bacilli of the Enterobacteriales family, and the most widespread cause of UTIs is Escherichia coli.<sup>3</sup> Community and hospital acquired UTIs are brought about by E. Coli, which can also occasionally result in major secondary health problems.<sup>4,5</sup> It is thought to be the cause of about 85% of UTIs that occur in the community and 50% of UTIs that occur in hospitals.<sup>6</sup>

Understanding the E. Coli antibiogram is crucial

1. Shalimar Medical and Dental College/Shalamar Institute of Health Sciences
2. Services Institute of Medical Sciences (SIMS), Lahore
3. Rashid Latif Khan University Medical College, Lahore
4. Chughtai Institute of Pathology
5. Ph.D Scholar, UHS, Lahore

### Correspondence:

Mariam Danish Iqbal, Associate professor, Rashid Latif Khan University Medical College, Lahore.  
Email : dr.mariam.danish@gmail.com

Submission Date: 19-04-2024  
1st Revision Date: 02-05-2024  
Acceptance Date: 29-05-2024

for starting an efficacious empirical therapy.<sup>7,8</sup> In developing countries, treatment costs, morbidity, and mortality are increased by multidrug-resistant (MDR) E. coli UTIs.<sup>9,10</sup> Microorganisms use a variety of techniques, such as genetic material modification, recombination of foreign DNA in bacterial chromosomes, and horizontal gene transfer, to develop drug resistance.<sup>11</sup> National differences exist in the resistance patterns of microorganisms, ranging from community-acquired to hospital-acquired, and from smaller to larger hospitals.<sup>12</sup>

Antibiotic resistance is becoming an increasingly pressing issue in Pakistan as a result of improper and excessive antibiotic use.<sup>9,13</sup> Antibiotic resistance is not systematically monitored nationally, and there is not enough data available to accurately estimate the issue.<sup>14</sup> The current study set out to identify the sensitivity and resistance pattern of E. coli that causes UTI in our hospitalized patients. In order to avoid or lessen the likelihood of problems, the results will assist the doctors in initiating empirical therapy in these situations. The outcomes might also add to the national database.

**METHODS**

This descriptive cross-sectional study was conducted at Microbiology section, Pathology Department, Shalamar Teaching Hospital Lahore, Pakistan. The duration of study was 6 months from June 01<sup>st</sup>, 2020 to November 30<sup>th</sup>, 2020. Urine sample of the patients with symptoms of urinary tract infection received in Tertiary care Hospital Laboratory were analyzed for the presence of infection with E.Coli. All age groups from both genders were included. Duplicate samples from the same patient during the period of illness were excluded.

Samples of urine which were positive for E. Coli were analyzed further for sensitivity. Following a centrifugation of the samples, the sediments were grown mostly on a media that is specifically for urine ie CLED (cystine lysine electrolyte deficient) Agar and incubated for a full day at 37°C. Gram staining, the Oxidase reaction, and basic biochemical tests such inoculation on Triple Sugar Iron Agar slant, citrate

utilization, urease generation, and the Indole test were used to identify the isolates as E. coli.<sup>15</sup>

E. Coli's antibiotic susceptibility was evaluated on Mueller Hinton Agar using the Kirby Bauer Disc diffusion method.<sup>16</sup> On the Mueller Hinton Agar plate, 0.5 McFarland standard units of bacterial inoculum were added in order to produce a confluent growing lawn. Amoxicillin, Ampicillin, Cefotaxime, Ceftriaxone, Cefuroxime, Ciprofloxacin, Colistin, Co-Trimoxazole, Fosfomycin, Gentamicin, Imipenem, Meropenem, Nitrofurantoin, and Piperacillin-Tazobactam discs containing antibiotics were arranged on the plates and incubated for a whole day at 37°C. Based on their zones of inhibition, the isolates were categorized as sensitive or resistant in accordance with the Clinical Laboratory guidelines Institute (CLSI) guidelines. The statistical analysis was carried out utilizing the SPSS 20.0 program (SPSS Inc). Frequency distribution was calculated for qualitative data i.e. Gender and Antibiotic Sensitivity and resistance pattern.

**RESULTS**

Between June 1<sup>st</sup>, 2020 and November 30<sup>th</sup>, 2020, 165 E. Coli isolates were reported for the current investigation. According to the data below, of the 165 E. Coli isolates, the frequency was higher in females 120/165 (72.7%) than in males 45 (27.3%).

The results of current study revealed that E. coli had a high level of resistance against Ampicillin 162

*Table 1: Gender wise distribution of UTI infection caused by E.coli.*

Gender	Frequency
Male	45 (27.3%)
Female	120 (72.7%)

(98.2%), Amoxicillin 154 (93.3%), Cefuroxime 152 (92.1%), Ciprofloxacin 141 (85.5%), Ceftriaxone and Cefotaxime 137 (83.0%) and Co-trimoxazole 134 (81.2%). Whereas high rate of sensitivity was found against Colistin 164 (99.4%), Fosfomycin 156 (94.5%), Imipenem 153 (92.7%) and Meropenem 150 (90.9%) as shown in the table-2:

**Table 2:** Pattern of Antibiotic Susceptibility and Resistance in *E.coli*.

Antibiotics	Sensitive (%)	Resistant (%)
Amoxicillin	11 (6.7%)	154 (93.3%)
Ampicillin	3 (1.8%)	162 (98.2%)
Cefotaxime	28 (17%)	137 (83%)
Ceftriaxone	28 (17%)	137 (83.0%)
Cefuroxime	13 (7.9%)	152 (92.1%)
Ciprofloxacin	24 (14.5%)	141 (85.5%)
Colistin	164 (99.4%)	1 (0.6%)
Co-trimoxazole	31 (18.8%)	134 (81.2%)
Fosfomycin	156 (94.5%)	9 (5.5%)
Gentamicin	110 (66.7%)	55 (33.3%)
Imipenem	153 (92.7%)	12 (7.3%)
Meropenem	150 (90.9%)	15 (9.1%)
Nitrofurantoin	141 (85.5%)	24 (14.5%)
Piperacillin tazobactam	60 (36.4%)	105 (63.6%)

## DISCUSSION

Microbial invasion of the urinary system is the source of infections in the urinary tract (UTI's) and the problems that usually accompany them.<sup>17</sup> UTIs can strike anyone of any gender, however they seem to strike women more frequently. The prevalence of UTIs in women is thought to be between 50 and 70 percent.<sup>18</sup> Since women's urethras are shorter and their perineum are more anatomically shaped like that, they are more likely to suffer urinary tract infections.<sup>19</sup> There is a global increase in reports of *E. coli* and antimicrobial resistance. Growing concerns are being expressed about the rising rate of resistance in both wealthy and developing nations.<sup>20</sup> Microorganisms that are exposed to two or more antibiotic groups can develop multi-drug resistance (MDR) through creation of enzymes that are resistant to the antibiotics, modification of the drugs' target sites, efflux pump mechanism, and genetic mutation.<sup>21</sup> The treatment of UTI is becoming increasingly complex due to the growth in antibiotic resistance in bacteria, which is a global concern.<sup>22</sup>

The 165 *E. Coli* isolates included in this study were resistant to Ampicillin 162 (98.2%), Amoxicillin 154 (93.3%), Cefuroxime 152 (92.1%), Ciprofloxacin 141 (85.5%), Ceftriaxone and Cefotaxime 137 (83.0%), and Co-Trimoxazole 134 (81.2%) following suit. On the other hand, strong rates of sensitivity to Meropenem 150 (90.9%), Fosfomycin 156 (94.5%), Imipenem

153(92.7%), and Colistin 164 (99.4%) were discovered. While some investigations revealed 100% resistance to Penicillin, resistance to the Penicillin group of antibiotics is growing daily in many parts of the world.<sup>23</sup> Resistance to additional  $\beta$ -Lactam antibiotics, such as cefuroxime (92.1%), ceftriaxone (83%) and cefotaxime (83%), was also quite high, deeming these medications unsuitable for empirical prescription for the management of UTIs.

Penicillin and Cephalosporin are not effective against UTI infections in underdeveloped nations like Pakistan, and our findings imply that these antibiotics shouldn't be used empirically to treat UTIs. The fact that these antibiotics are ineffective does not imply that they are not utilized elsewhere in the globe to treat UTIs brought on by *E.coli*. Penicillin and cephalosporins were found to be effective against a greater percentage of *E. coli* in a few reports.<sup>24</sup> These medicines were effective against *E. coli*-caused UTIs in Pakistan ten years ago.<sup>25</sup>

A high level of antibiotic resistance to both cephalosporin and penicillin has been found in *E. coli*, according to numerous investigations carried out in Pakistan.<sup>26</sup> In the current study, gentamicin shown a low level of resistance of 33 percent. *E. Coli* has previously been treated with fluoroquinolones, most notably ciprofloxacin. Nonetheless, the current study's discovery that *E. Coli* exhibited 85% resistance to ciprofloxacin is in line with the findings of the previous publication.<sup>27</sup> The current study's findings demonstrate the efficacy of imipenem, colistin, meropenem, and fosfomycin as medications for treating UTIs caused by *E. coli*.

## CONCLUSION

According to the current study, *E. coli*-caused UTI infections had a higher rate of drug resistance. When treating a UTI brought on by multi drug resistant *E. coli*, Fosfomycin, Imipenem, Meropenem, advised as first-line antibiotics. Moreover, the presence of *E. Coli* that was isolated from UTIs exhibiting drug resistance suggests that antibiotics should be prescribed and closely monitored following sensitivity and culture testing.

**Ethical Approval:**

The ethical Approval was obtained vide letter no. SNDC-IRB/AL/90/2021

**Conflict of Interest:**

*None*

**Funding Source:**

*None*

**REFERENCES**

1. Flannery DD, Akinboyo IC, Mukhopadhyay S, Tribble AC, Song L, Chen F et al. Antibiotic susceptibility of Escherichia coli among infants admitted to neonatal intensive care units across the US from 2009 to 2017. *JAMA pediatrics*. 2021 Feb 1;175(2):168-75.
2. Nas FS, Ali M, Abdallah MS, Zage AU. Prevalence and antibiotic susceptibility pattern of Escherichia coli isolated from urine samples of urinary tract infection patients. *ARC Journal of Urology*, (2019); 4(1): 14-20
3. Naqid IA, Balatay AA, Hussein NR, Saeed KA, Ahmed HA, Yousif SH. Antibiotic susceptibility pattern of Escherichia coli isolated from various clinical samples in Duhok City, Kurdistan Region of Iraq. *International Journal of Infection*. 2020 Apr 30;7(3).
4. Tambat R, Mahey N, Chandal N, Verma DK, Jangra M, Thakur KG, Nandanwar H. A microbe-derived efflux pump inhibitor of the resistance-nodulation-cell division protein restores antibiotic susceptibility in Escherichia coli and Pseudomonas aeruginosa. *ACS infectious diseases*. 2022 Jan 19;8(2):255-70.
5. Muhammad A, Khan SN, Ali N, Rehman MU, Ali I. Prevalence and antibiotic susceptibility pattern of uropathogens in outpatients at a tertiary care hospital. *New Microbes and new infections*. 2020 Jul 1;36:100716.
6. Malik N, Sultan A, Nizamuddin S, Shameem F. Oral treatment options for patients with urinary tract infections caused by carbapenem-resistant Escherichia coli. *Infectious Diseases Journal of Pakistan*. 2024 Sep 27;33(3):102-6.
7. Hussaini IM, Suleiman AB, Olonitola OS, Oyi RA. Antibiotic Susceptibility Pattern of Carbapenem Resistant Escherichia coli and Klebsiella pneumoniae, 2019. *The Microbe*. 2024 Sep 24:100174.
8. Xiao R, Li Y, Liu X, Ding Y, Lai J, Li Y, Kang W, Zou P, Wang J, Du Y, Zhang J. Antibiotic susceptibility of Escherichia coli isolated from neonates admitted to neonatal intensive care units across China from 2015 to 2020. *Frontiers in cellular and infection microbiology*. 2023 May 22;13:1183736.
9. Parsons JB, Sidders AE, Velez AZ, Hanson BM, Angeles-Solano M, Ruffin F, Rowe SE, Arias CA, Fowler Jr VG, Thaden JT, Conlon BP. In-patient evolution of a high-persister Escherichia coli strain with reduced in vivo antibiotic susceptibility. *Proceedings of the National Academy of Sciences*. 2024 Jan 16;121(3):e23-14514121.
10. Jeje O, Ewunkem AJ, Jeffers-Francis LK, Graves Jr JL. Serving two masters: effect of Escherichia coli dual resistance on antibiotic susceptibility. *Antibiotics*. 2023 Mar 17;12(3):603.
11. Hasan TH, Aljanaby IA, Al-Labban HM, Aljanaby AA. Antibiotic susceptibility pattern of E. coli causing urinary tract infection in pregnant women in Al-Najaf Province, Iraq. In *AIP Conference Proceedings 2023 Dec 22 (Vol. 2977, No. 1)*. AIP Publishing.
12. Ait-Mimoune N, Hassaine H, Boulanoir M. Bacteriological profile of urinary tract infections and antibiotic susceptibility of Escherichia coli in Algeria. *Iranian journal of microbiology*. 2022 Apr;14(2):156.
13. Salamzade R, McElheny CL, Manson AL, Earl AM, Shaikh N, Doi Y. Genomic epidemiology and antibiotic susceptibility profiling of uropathogenic Escherichia coli among children in the United States. *Mosphere*. 2023 Sep 24;8(5):e00184-23.
14. Hasanli L, Dagi HT, Arslan U. Investigation of antibiotic susceptibility and virulence genes in Escherichia coli strains isolated from blood and urine samples. *Journal of Pediatric Infectious Diseases*. 2022 Mar;17(02):098-105.
15. Bhargava K, Nath G, Bhargava A, Kumari R, Aseri GK, Jain N. Bacterial profile and antibiotic susceptibility pattern of uropathogens causing urinary tract infection in the eastern part of Northern India. *Frontiers in Microbiology*. 2022 Aug 9;13:965053.
16. Hasan TH, Aljanaby IA, Al-Labban HM, Aljanaby AA. Antibiotic susceptibility pattern of E. coli causing urinary tract infection in pregnant women in Al-Najaf Province, Iraq. In *AIP Conference Proceedings 2023*. 2977:1. AIP publishing.
17. Miftode IL, Văță A, Miftode RȘ, Parângă T, Luca MC, Manciu C, Țimpău AS, Radu V, Roșu MF, Stămăteanu LO, Leca D. The Impact of Urinary Catheterization on the Antibiotic Susceptibility of ESBL-Producing Enterobacterales: A Challenging Duo. *Antibiotics*. 2024 May 17;13(5):462.
18. Fatima T, Rafiq S, Iqbal A, Husnain S. Prevalence and antibiogram of MDR E. coli strains isolated from UTI patients—1-Year retrospective study at Nishtar medical hospital, Multan. *SN Comprehensive Clinical Medicine*; 1-9.
19. Saleem Z, Haseeb A, Abuhussain SS, Moore CE, Kamran SH, Qamar MU, Azmat A, Pichierri G, Raees F, Asghar S, Saeed A. Antibiotic susceptibility surveillance in the Punjab Province of Pakistan: findings and implications. *Medicina*. 2023 Jun 28;59(7):1215.

20. Hefetz I, Bardenstein R, Rotem S, Zaide G, Bilinsky G, Shifman O, Zimhony O, Aloni-Grinstein R. Rapid Phenotypic Antibiotic Susceptibility Profiling of Clinical *Escherichia coli* and *Klebsiella pneumoniae* Blood Cultures. *Antibiotics*. 2024 Feb 29;13(3):231.
21. Chan A, Spradley M, Williams T, Morales G, Van Horn G, Schmitz JE. The Microbial Antibiogram as a Function of Testing Indication: Susceptibility Analysis of *Escherichia coli* from Symptomatic and Asymptomatic Bacteriuria Patients, 2020-2021. *Journal of Clinical and Translational Science*. 2024 Apr;8(s1):6-10
22. Neuenschwander FR, Groß B, Schubert S. Rapid antibiotic susceptibility testing of gram-negative bacteria directly from urine samples of UTI patients using MALDI-TOF MS. *Antibiotics*. 2023 Jun 12;12(6):1042.
23. Simon M, Fougnot S, De Monchy P, Duda A, Thilly N, Pulcini C, Bocquier A, Charmillon A, Chopard V, Delpuech M, Fagot-Campagna A. Impact of selective reporting of antibiotic susceptibility testing results for urinary tract infections in the outpatient setting: a prospective controlled before-after intervention study. *Clinical Microbiology and Infection*. 2023 Jul 1;29(7):897-903.
24. Yu CH, Chang CN, Wang CC. Causative microbes and antibiotic susceptibility of acute appendicitis in adults and children. *Pediatrics & Neonatology*. 2024 Mar 1;65(2):159-64.
25. Nosheen S, Bukhari NI, Ejaz H, Abbas N. Antibiogram and recent incidence of multi-drug resistant carbapenemase producing *Escherichia coli* isolated from paediatric patients. *Pakistan journal of medical sciences*, (2020); 36(2): 246.
26. Khatoon I, Khanam S, Azam A, Qadeer S, Naz S, Hassan NU. Incidence pattern, antibiotic susceptibility pattern and associated risk factors of bacterial Uropathogens among general population of Pakistan. *Infection and Drug Resistance*. 2023 Dec 31:4995-5005.
27. Sundaramurthy R, Tiruvanmalai R, Sivaraman ST, Arunagiri R, Charles J. Study on clinico microbiological profile and antibiotic susceptibility pattern of urinary tract pathogens with Special reference to susceptibility of *Escherichia coli* to fosfomycin. *Indian Journal of Microbiology Research*. 2023 Jan 24;5(2):258-65.