

Prevalence of urinary tract pathogens and antimicrobial susceptibility patterns in children at hospitals in Iran

Enayatollah Kalantar^{1*}, Mohammad esmaeel Motlagh², Hamid Lornejad², Naser Reshadmanesh³

¹Department of Microbiology, Faculty of Medicine, Kurdistan University of Medical Sciences, Sanandaj, Iran.

²Department of Family Health and Population, Ministry of health and Medical Education, Tehran, Iran

³Department of Statistics, Faculty of Health, Kurdistan University of Medical Sciences, Sanandaj, Iran.

ABSTRACT

Background: Urinary tract infections (UTIs) are one of the most frequent bacterial infections in children. Most commonly, members of Enterobacteriaceae, particularly uropathogenic strains of *E. coli* and *Enterobacter* spp. are the primary causative organisms of UTIs in different parts of the world.

Patients and methods: A total of 55 hospitals from 12 provinces in Iran participated in this study. 1696 children with UTI aged 0 to 5 years referred to these hospitals were included in this study. Urine cultures were carried out and the isolates were identified by gram staining and conventional biochemical methods. Antimicrobial susceptibility testing was performed by disk diffusion method according to the current National Committee for Clinical Laboratory Standards (NCCLS) guidelines.

Results: In general 438 urine isolates were obtained. *E. coli* was the most frequently occurring pathogen (54.80%), followed by *Klebsiella pneumoniae* (16.0%), coagulase negative *Staphylococci* (11.2%), *Enterobacter* spp. (9.6%), *Proteus* spp. 1.4% and *P. aeruginosa* (1.4%). Resistance rates of *E. coli* isolates were 85.9% to co-trimoxazole, 80.0% to penicillin, 77.0% to ampicillin, 68.0% to chloramphenicol, 12.9% to ciprofloxacin, 12.9% to ceftriaxone, 12.9% to cephalotin, and 14.0% to amikacin. *K. pneumoniae* isolates were 18.5%, 20.0%, 24.0% and 29.0% resistant to cephalothin, ceftriaxone, amikacin and gentamicin respectively; however, penicillin (88.5%), co-trimoxazole (74.2%), and ampicillin (68.5%) were the least effective drugs.

Conclusion: High prevalence of drug-resistant urinary tract pathogens, particularly to ampicillin and co-trimoxazole among Iranian children suggests cautious use of antibiotic therapy for the treatment. Finally, we suggest that empirical antibiotic selection should be based on knowledge of the local prevalence of bacterial organisms and antibiotic sensitivities rather than on universal guidelines.

Keywords: *Urinary tract infection, Antimicrobial susceptibility pattern*
(Iranian Journal of Clinical Infectious Diseases 2008;3(3):149-153).

INTRODUCTION

Urinary tract infections (UTIs) are one of the most frequent infections in children. At least 8% of

girls and 2% of boys will have a urinary tract infection (UTI) in childhood, and between 30% and 40% will have another episode within two years (1,2).

Several studies has demonstrated that the geographical variability of pathogen occurrence in

Received: 5 August 2007 Accepted: 27 April 2008

Reprint or Correspondence: Enayatollah Kalantar, MD.
Department of Microbiology, Faculty of Medicine, Kurdistan
University of Medical Sciences, Sanandaj, Iran.

E-mail: ekalantar@hotmail.com

cases of UTI among inpatients and outpatients populations is limited by the predominance of gram-negative species usually Enterobacteriaceae and particularly *E. coli* and *Enterobacter* spp. in various regions of the world (3,4).

However, *Pseudomonas aeruginosa* and Enterococcal spp. are usually ranked among the top five pathogens and the resistance patterns of these pathogens can vary significantly between hospitals, countries and continents (4,5).

Antimicrobial resistance among urinary tract isolates has recently been reported with an increased frequency all over the world (6-9). Several studies in Iran (10,11) also have reported that the incidence of UTI has been increasing recently and its treatment has become more complicated because of the emergence of pathogens with increasing resistance to antimicrobial agents.

It is necessary to identify the causative agent and spectrum of its antimicrobial susceptibilities in order to treat UTI. Since this spectrum may vary among geographical locations, hospitals and also in different age groups, each institution should carefully plan their antibiotic therapy.

Unfortunately, little has been published regarding Iranian scenario of the range and antimicrobial susceptibility patterns in urinary tract pathogens particularly among children.

This study was performed to find out the frequency of different urinary tract pathogens in children of age group 0 to 5 years and the antimicrobial susceptibility profiles.

PATIENTS and METHODS

A total of 1696 urine specimens from children aged 0 to 5 years with UTI were collected at 55 hospitals from 12 provinces representing different geographical areas in Iran between July and August 2006 (fig. 1). The following cases were excluded: previous history of urinary infection, known

urinary malformations (according to prenatal ultrasound and previous medical records), chronic illness, or current prophylactic treatment with antibiotics.

All urine samples were obtained by midstream clean-catch, catheterization or from urine bags. These samples were processed on blood agar and MacConkey media with a standard loop and were incubated at 37 °C overnight. Significant growth was determined as $\geq 10^5$ colony-forming units (CFU)/mL of midstream urine and bag urine samples, and $\geq 10^2$ CFU/mL of a catheter specimen (12).

Isolates were identified by Gram staining and conventional biochemical methods (12). Antimicrobial susceptibility testing by disc diffusion was carried out according to the Clinical and Laboratory Standard Institute guidelines (13) using the following antibiotics: oxacillin, penicillin, erythromycin, ciprofloxacin, ceftriaxone, co-trimoxazole, gentamicin, cefalotin, vancomycin, amikacin, cephradine, and ampicillin. Data management and statistical analysis were performed using SPSS software.

RESULTS

There were 867(51.9%) boys and 805 (48.1%) girls giving a total of 1669 children who enrolled in this study. Their age ranged from 0 to 5 years. The age distribution is showed in table 1.

Table 1. Prevalence of UTI in different age groups

Age Group (months)	UTI Positive	UTI Negative
Up to 1 (n=68)	14 (20.6%)	54 (79.4%)
>1- 12 (n=734)	300 (40.9%)	434 (59.1%)
> 12 - 36 (n=590)	101 (17.1%)	489 (82.9%)
> 36 - 60 (n=304)	23 (7.6%)	281 (92.4%)
Total (n=1696)	438	1258

The male to female ratio was 1.07:1. Occurrence of bacterial urinary tract infections was highest in the age group one month to 1 year (40.9%) and the lowest in the age > 36 to 60 months (7.6%).

Eight different types of pathogens were isolated from the urine samples. The frequency of pathogens and the ranking order among the most prevalent pathogens is shown in table 2.

Table 2. Prevalence of pathogens isolated on urine culture

Pathogen	No. of isolates
<i>E. coli</i>	240 (54.8%)
<i>K. pneumoniae</i>	70 (16%)
Coagulase negative Staphylococcus	50 (11.2%)
<i>Enterobacter spp</i>	42 (9.6%)
<i>Proteus spp</i>	20 (4.6%)
<i>S. aureus</i>	6.0 (1.4%)
<i>Pseudomonas aeruginosa</i>	6.0 (1.4%)
<i>Candida spp</i>	4.0 (1%)
Total	438 (100%)

Escherichia coli, which comprised 54.8% (240) of the isolates, showed high resistance to cotrimoxazole, (85.9%), penicillin (80%), ampicillin (77%). *Klebsiella pneumoniae* was resistant to penicillin in 88.5%, to ampicillin in 68.5%, and to co-trimoxazole in 74.2% of cases.

Among gram-positive isolates, Coagulase negative Staphylococcus comprised 11.2% of all

isolates followed by *Staphylococcus aureus* (1.4%) showed 80% and 66.7 % resistance to ampicillin respectively, but highly sensitive to Vancomycin (table 3).

DISCUSSION

Urinary tract infection in children is a significant source of morbidity. It is generally agreed that children with UTI require further investigation and continuing urinary surveillance to minimize future complications.

Escherichia coli is the most common cause of urinary tract infection (14). Other microorganisms include *Klebsiella spp.*, *Enterobacter spp.*, *Serratia sp.*, *Pseudomonas aeruginosa* and other *Pseudomonas spp.*, Enterococci, *Staphylococcus saprophyticus*, *Staphylococcus aureus*, *Staphylococcus epidermidis*, *Acinetobacter spp.*, and *Streptococci spp.* (15-17). Our findings are consistent with these reports.

Gram-positive organisms have received more attention recently as a cause for bacteriuria and urinary tract infection. Coagulase negative *Staphylococcus*, *Streptococci* and *Enterococci* are seen in small numbers but they are recognized as important causes of urinary tract infection (18, 19).

Table 3. Frequency of in vitro antimicrobial resistance spectrum of pathogens isolated from urine cultures at various hospitals in Iran*

	<i>P. aeruginosa</i>	<i>E. coli</i>	<i>K. pneumoniae</i>	CoNS	<i>Enterobacter spp</i>	<i>Proteus spp</i>	<i>S. aureus</i>
Penicillin	89	80	89	71	89	91	78
Ampicillin	93	77	69	80	69	79	70
Oxacillin	ND	ND	ND	58	ND	ND	34
Vancomycin	64	70	58	18	ND	ND	28
Gentamicin	52	37	29	68	12	71	75
Ciprofloxacin	25	12	ND	61	ND	ND	65
Amikacin	33	14	24	ND	ND	25	ND
Co-trimoxazole	70	85.9	74	ND	80	79	ND
Chloramphenicol	71	68	71	59	68	71	59
Ceftriaxone	56	13	20	45	46	ND	ND
Cephalotin	21	13	19	ND	ND	11	27

* % of isolates resistant to antimicrobial agents

We reported the occurrence rate of coagulase negative *Staphylococcus* to be 11.2%.

Increasing antibiotic resistance among urinary tract isolates (mostly against ampicillin) has been reported from many countries. The resistance rates to ampicillin were found to be 45%, 50% and 100% in children from Canada, Europe and Africa, respectively (20-23). In our study, the frequency of resistance to ampicillin of UTI isolates was 74.2%. This finding is similar to other reports (24,25).

The Results of the present study indicate a high incidence of microbial resistance to ampicillin and co-trimazole in urinary tract infections among Iranian children and suggest the physicians to be cautious about treatment with antibiotics.

Regular monitoring is required to establish reliable information about resistance pattern of urinary pathogens for optimal empirical therapy of patients with UTIs. Finally, we suggest that empirical antibiotic selection should be based on the knowledge of local prevalence of bacterial organisms and antibiotic sensitivities rather than on universal guidelines.

ACKNOWLEDGEMENTS

The authors would like to thank Mrs. Kakaie S for reviewing the data coding and entry to computer. We also thank Dr. S. Diwanay of the Department of Microbiology, Abasaheb Garware College, University of Pune, India, for critical review of the manuscript and suggestions.

REFERENCES

1. Larcombe J. Urinary tract infection in children. *BMJ* 1999;319:1173-75.
2. Shaw KN, Gorelick M, McGowan KL, Yakscoe NM, Schwartz JS. Prevalence of urinary tract infection in febrile young children in the emergency department. *Pediatrics* 1998;102:e16.
3. Bachur R, Harper MB. Reliability of the urinalysis for predicting urinary tract infections in young febrile children. *Arch Pediatr Adolesc Med* 2001;155:60-5.
4. Twaij M. Urinary tract infection in children: a review of its pathogenesis and risk factors. *J R Soc Health* 2000;120:220-26.
5. Azubike CN, Nwamadu OJ, Oji RU, Uzoiye N. Prevalence of urinary tract infection among school children in a Nigerian rural community. *West Afr J Med* 1994;13:48-52.
6. Zorc JJ, Kiddoo DA, Shaw KN. Diagnosis and management of pediatric urinary tract infections. *Clin Microbiol Rev* 2005;18:417-22.
7. Turnidge J, Bell J, Biedenbach DJ, Jones RN. Pathogen occurrence and antimicrobial resistance trends among urinary tract infection isolates in the Asia-Western Pacific Region: report from the SENTRY Antimicrobial Surveillance Program, 1998-1999. *Int J Antimicrob Agents* 2002;20:10-7.
8. Zhanel GG, Karlowsky JA, Schwartz B, Jensen SB, Hoban DJ. Mecillinam activity compared to ampicillin, trimethoprim/sulfamethoxazole ciprofloxacin and nitrofurantoin against urinary tract isolates of gram-negative bacilli. *Chemotherapy* 1998;44:391-96.
9. Bahram F, Farhad H, Mohammad E, Marzieh A, Farrok A, Bahram K. Detection of vancomycin resistant enterococci (vre) isolated from urinary tract infections (UTI) in Tehran, Iran. *Daru* 2006;14:141-45. [In Persian]
10. Sorkhi H. Causes of hydronephrosis in pyelonephritic children. *Indian J Pediatr.* 2005;72:1058-59.
11. Ladhani S, Gransden W. Increasing antibiotic resistance among urinary tract isolates. *Arch Dis Child* 2003;88:444-45.
12. Forbes BA. *Bailey and Scott's diagnostic microbiology*. 10th ed. St. Louis, Missouri: Mosby; 1998; p:283-304.
13. Clinical and Laboratory Standards Institute. 2006. Performance standards for antimicrobial susceptibility testing; 16th informational supplement. M100-S16. Clinical and Laboratory Standards Institute, Wayne, PA.
14. Esmaeili M. Antibiotics for causative microorganisms of urinary tract infections. *Iranian Journal of Pediatric infection.* 2005;15:163-83.
15. Fluit C, Mark J, Franz-Josef S, Jacques A, Renu G, Verhoef J. Antimicrobial resistance among urinary tract infection (UTI) isolates in Europe: results from the SENTRY Antimicrobial Surveillance Program 1997. *Antonie Van Leeuwenhoek* 2000;77:147-52.
16. Jalali M, Asteraki T, Emami-Moghadam E, Kalantar E. Epidemiological study of asymptomatic

bacteriuria among nursery school children in Ahwaz, Iran. *Afr J Clin Expt Microbiol* 2005;6:159-61.

17. Esmacili M. Antibiotics for causative microorganisms of urinary tract infections. *Journal of Iranian Pediatric Disease*. 2005;15:165-73.

18. Schlager TA. Urinary tract infections in infants and children. *Infect Dis Clin North Am* 2003;17:353-65.

19. Warren JW. Catheter-associated urinary tract infections. *Int J Antimicrob Agents* 2001;17:299-303.

20. Adjei O, Opoku C. Urinary tract infections in African infants. *Int J Antimicrob Agents* 2004;24:32-4.

21. Allen U, MacDonald N, Fuite L, Chan F, Stephens D. Risk factors for resistance to first-line antimicrobials among urinary tract isolates of *Escherichia coli* in children. *CMAJ* 1999;160:1436-40.

22. Prais D, Straussberg R, Avitzur Y, Nussinovitch M, Harel L, Amir J. Bacterial susceptibility to oral antibiotics in community acquired urinary tract infection. *Arch Dis Child* 2003;88:215-18.

23. Haller M, Brandis M, Berner R. Antibiotic resistance of urinary tract pathogens and rationale for empirical intravenous therapy. *Pediatr Nephrol* 2004;19: 982-86.

24. Haghi-Ashteiani M, Sadeghifard N, Abedini M, Soroush S, Taheri-Kalani M. Etiology and antibacterial resistance of bacterial urinary tract infections in children's medical center, Tehran, Iran. *Acta Medica Iranica* 2007;45:153-57.

25. Mostafa S, Abdulla K, Sedigheh R, Navid A. microbial sensitivity pattern in urinary tract infections in children: a single center experience of 1117 urine culture. *Jpn J Infect Dis* 2006;59:380-82.