

Prevalence and Antimicrobial Sensitivity Pattern of Urinary Tract Infection in Febrile Children Aged 1 Month to 5 Years

Saravanan.S

[Assistant professor, Department of Pediatrics, Meenakshi medical college & research institute, MAHER University, Tamilnadu, India]

I. Introduction

Urinary tract infection (UTI) is a common infection in infants and children. During infancy, boys are more commonly affected than girls and thereafter, female preponderance is found. Presentation varies among different age groups. Clinical features in neonates and young infants are non-specific, manifest as septicemia where a high index of suspicion is needed. Older children typically present as simple or complicated UTI. Rapid diagnosis, institution of early treatment and further evaluation by imaging modalities are of utmost importance. This was a prospective study in febrile children from 1 month to 5 years at Meenakshi medical college hospital and research institute from Jan 2011 – Dec 2012.

II. Material And Methods

Children who presented with fever were included and those with known congenital genitourinary anomalies and those who received antibiotics 48 hours prior were excluded. Routine blood counts, urinary analysis was done. The urine specimens were centrifuged in a standard manner and in those showing (pus cells > 5 HPF in centrifuged urine), a clean catch mid stream urine sample was sent for culture which was incubated on Blood and MacConkey agar plates with a 0.01ml calibrated loop. All plates were incubated at 35-37°C for 24hrs under aerobic condition to obtain accurate colony count. On culture of mid stream sample of urine, a colony count of more than 10^5 /ml organisms of a single species was considered significant. Sample showing insignificant growth, mixed growth of two or more pathogens or growth of non-pathogens were not considered as culture positive. Antibiotic sensitivity was put up by the Kirby Bauer method following the clinical laboratory standard institute (CLSI)¹⁹ guidelines.

All Enterobacteriaceae and Acinetobacterspp. were tested against first line agents: gentamycin(10µg), amikacin(30µg), nitrofurantoin(300µg), trimethoprim sulphamethoxazole(1.25-23.75 µg), norfloxacin(10 µg), ciprofloxacin(5 µg), amoxycylav(20/10 µg) and tobramycin(10 µg), Pseudomonas aeruginosa against amikacin (30 µg), gentamicin (10 µg), ceftazidime (30 µg) and ciprofloxacin (5 µg). Staph spp was tested against vancomycin (30 µg). Second line antibiotics to all 1st line antimicrobials or specifically requested for by the attending physicians. These included: Imipenem (10 µg) and piperacillin-tazobactam (100/10 µg) for all Enterobacteriaceae, Acinetobacter spp. and Pseudomonas isolates.

III. Results And Analysis

A total of 630 Children (1 month to 5 yrs) were evaluated in the study. Of 630 cases 305(48.4%) were males, 325(51.5%) were females, 193 cases were <1 year (30.6%) [90(14.2%) males, 103(16.34%) females], 195(30.9%) [95(15%) males, 100(15.87%) females] cases were between 1-2 years and 242(38.4%) [120(19%) males, 122(19.36%) females] cases were more than 2 years. Minimum age in the study group was 1 month and maximum age in the study group was 60 months.

3.1 AGE AND SEX DISTRIBUTION OF SUBJECTS WITH URINE SHOWING > 5PUS CELLS/ HPF

TABLE -1

Age	Sex		Total
	Male	Female	
< 1 year	20(15.6%)	20(15.6%)	40(31.2%)
1 – 2 years	15(11.7%)	20(15.6%)	35(27.3%)
2 – 5 years	21(16.4%)	32(25%)	53(41.4%)
Total	56	72	128(100)

Significant pyuria was seen in 128 children, of whom children between 2-5 yrs of age were predominant group and further analysis showed patients with 8-10 pus cells /HPF being the highest among the group (table-2).

3.2 DISTRIBUTION OF PUS CELLS IN URINE

TABLE - 2

No of Pus cells in Urine	Male	Female	Total
6 - 8	22(17.1%)	26(20.3%)	48(37%)
8 - 10	25(19.5%)	30(23.4%)	55(43%)
Numerous	11(8.5%)	14(10.9%)	25(20%)
Total	58(45.3%)	70(54.6%)	128(100%)

3.3 DISTRIBUTION OF UTI IN ACCORDANCE WITH AGE

TABLE- 3

age	Total no of pts	Culture positivity
<1	193	22(11.3%)
1-2	195	18(9.2%)
2-5	242	29(11.9%)
	630	69(10.9%)

3.4 CORRELATION BETWEEN PYURIA AND UTI

TABLE-4

Age	Males with pyuria	Females with pyuria	Total children	Male UTI	Female UTI	Overall UTI
< 1 year	20(15.6%)	20(15.6%)	40(31.2%)	13(65%)	9(45%)	22(55%)
1 - 2 years	15(11.7%)	20(15.6%)	35(27.3%)	7(46%)	11(55%)	18(51.4%)
2 - 5 years	21(16.4%)	32(25%)	53(41.4%)	10(47%)	19(59%)	29(54.7%)
Total	56(43.7%)	72(56.2%)	128(100)			69(53.9%)

3.5 MICROBIOLOGICAL PROFILE AND PERCENTAGE DISTRIBUTION OF ISOLATES

TABLE-5

Culture report	Sex		Total
	Male	Female	
E coli	17(24.6%)	23(33.3%)	40(57.97%)
Klebsiella	3(4.3%)	5(7.2)	8(11.59%)
Pseudomonas	3(4.3%)	4(5.7%)	7(10.14%)
Proteus	2(2.8%)	2(2.8%)	4(5.79%)
CONS	2(2.8%)	2(2.8%)	4(5.79%)
Staph aureus	1(1.4%)	2(2.8%)	3(4.34%)
Acinetobacterspp	-	1(1.4%)	1(1.4%)
Citrobacter	1(1.4%)	-	1(1.4%)
Candida Spp.	-	1(1.4%)	1(1.4%)
	29(42.02%)	40(57.9%)	69(100%)

With reference to tables 3-5 UTI was overall more common in females and in age 2-5 years, when comparing UTI in children with pyuria, age 1month to one year was more common being 55% than age group 2-5 years(54.7%) .Culture positivity in patients with significant pyuria was high 53.9%. Most common organism isolated was E.coli 40(57.97%) followed by Klebsiella 8(11.59%), Pseudomonas7 (10.14%), Proteus 4(5.79%), CONS 4(5.79%), Staph aureus 3(4.34%), Acinetobacterspp 1 (1.4%), Citrobacter 1 (1.4%), Candida spp 1 (1.4%).

3.6 ANTIBIOTIC SUSCEPTIBILITY PATTERNS OF BACTERIA IN CHILDREN WITH UTI IN PERCENTAGE

TABLE-6

Isolate	IMP	CIP	CEF	AMK	NIT	AMC	GEM	TOB	VAN	COT	NOR	PTT
E.coli	100	100	-	97	94.5	87	73.3	93	-	28	61.1	100
Klebsiella	100	100	-	95	92	81	44	97	-	21	21	100
P. aeruginosa	100	89	100	89	76		28	100	-	0	32	100
P. mirabilis	100	86	-	82	51	88	34	98	-	16	83	100
CONS	-	97	-	79	36	87	53	62	100	0	52	-
S. aureus	40	91	-	88	27	78	70	67	100	0	47	-
Acinetobacterspp	100	100	-	100	0	0	0	100	-	0	100	100
Citrobacter	100	100	-	100	100	100	100	100	-	100	100	100

Imipenem (IMP), ceftazidime (CEF), ciprofloxacin (CIP), amikacin (AMK), nitrofurantoin (NIT), coamoxiclav (AMC), gentamycin (GEM), tobramycin (TOB), cotrimoxazole (COT), norfloxacin (NOR), vancomycin (VAN), piperacillin-tazobactam (PTT), (– not tested)

The antibiotic susceptibility of *E. coli*, which was the most common causative pathogen of UTI in children during the period of this study, was highest to Imipenem (100%), ciprofloxacin (100%) followed by amikacin (97%), nitrofurantoin (94.5%), and coamoxiclav (87%). *E. coli* showed low susceptibility to norfloxacin (61.1%) and gentamycin (73.3%). *Klebsiella* sensitivity pattern was almost similar to *E. coli*. *Proteus* was more resistant than *E. coli* to ciprofloxacin and amikacin. Other Enterobacteriaceae (*Acinetobacter*, *Citrobacter*) were universally sensitive to Imipenem, ciprofloxacin, amikacin and norfloxacin. Among CONS all were susceptible to vancomycin and was resistant to cotrimoxazole. *Staph aureus* was sensitive to vancomycin and ciprofloxacin.

IV. Discussion

Urinary tract infections are common, potentially serious infection of childhood. Community acquired urinary tract infections (UTI) cause significant illness in the first 2 years of life and are considered as common disease in school and pre-school children⁽¹⁻³⁾. Etiologic agents of UTI are variable and usually depend on time, geographical location and age of patients. However, Enterobacteriaceae species including *Escherichia coli*, *Proteus mirabilis*, *Enterobacter agglomerans*, *Citrobacter freundii* and *Klebsiella pneumoniae* account for over 70% cases⁽²⁻⁵⁾. Based on the microbial sensitivity test results, drugs that are usually administered against uropathogens include cotrimoxazole, amoxicillin, ampicillin, amino glycosides, cephalosporin's, nalidixic acid and nitrofurantoin. However, many reports have indicated the presence of multi-drug resistance in organisms causing UTI⁽⁶⁻⁹⁾. UTI cause acute morbidity as well as long term sequelae including hypertension and impaired renal function. A cause of occult febrile illness in up to 5% of young children they often remain undiagnosed.¹⁰

A total of 630 Children (1 month to 5 yrs) with fever were evaluated in the study of whom 69 (10.9%) children [29 male (42%) and 40 female (57.9%)] had culture proven UTI [68 bacterial and 1 due to yeast], of them 22 (31.8%) were 1 month to one year, 18 (26.08%) were 1-2 years, 29 (42%) were 2-5 years age. There was an overall female preponderance in cases of UTI (57.9%). Significant pyuria was seen in 128 (20.3%) children, of whom 72 (56.25%) were females, 56 (43.75%) were males. 40 (31.2%) of them were 1 month – 1 yrs, 35 (27.3%) were between 1-2 yrs and the rest 53 (41.4%) were 2-5 yrs of age. 48 (37%) of children with pyuria showed 6-8 pus cells / HPF and 55 (43%) showed 8-10 pus cells / HPF and 20% showed numerous pus cells. Most common organism isolated was *E. coli* 40 (57.97%) followed by *Klebsiella* 8 (11.59%), *Pseudomonas* 7 (10.14%), *Proteus* 4 (5.79%), CONS 4 (5.79%), *Staph aureus* 3 (4.34%), *Acinetobacter* spp (1.4%), *Citrobacter* 1 (1.4%), *Candida* spp (1.4%).

The antibiotic susceptibility of *E. coli*, which was the most common causative pathogen of UTI in children during the period of this study, was highest to Imipenem (100%), ciprofloxacin (100%) followed by amikacin (97%), nitrofurantoin (94.5%), and coamoxiclav (87%). *E. coli* showed low susceptibility to norfloxacin (61.1%) and gentamycin (73.3%). *Klebsiella* sensitivity pattern was similar to *E. coli*. *Proteus* was more resistant than *E. coli* to ciprofloxacin and amikacin. Other Enterobacteriaceae (*Acinetobacter*, *Citrobacter*) were universally sensitive to Imipenem, ciprofloxacin, amikacin and norfloxacin. Among CONS all were susceptible to vancomycin and was resistant to cotrimoxazole. *Staph aureus* was sensitive to vancomycin and ciprofloxacin.

Prevalence of febrile UTI in infants in our study was higher 11.3% than the study by Dharni Dharaka et al¹² (1993) who reported a prevalence of 5.4% in febrile infants and Hoberman et al¹¹ (1993) who reported prevalence of 5.3% in infants. Overall prevalence of UTI in febrile children in our study was 10.9% and 11.3% in children <5 years and infants respectively in contrast to study conducted by R.K. Kaushal et al¹³ (2003) who reported higher prevalence of 8.4% and 12.3% in children <5 years and infants respectively. Prevalence of febrile UTI in infants in our study (11.3%) was higher compared to report by Shaw K.N et al¹⁴ (1998) from USA who reported prevalence of 3.3% in febrile infants. In our study prevalence of UTI in <2 years age group was 10.25% which was higher than the study by Roberts K. et al¹⁵ (1983) who quoted prevalence of 4.1%. P.R. Srivastha et al¹⁶ (1996) reported a prevalence of 2.48% in children <2 years which was lowest reported from a developing country.

Bryan C.S et al¹⁷ reported *E. coli* as the common urinary pathogen in 85% of cases. According to Aravind Bagga et al¹⁸ (2000) 90% of first symptomatic urinary tract infection and 70% reoccurrence infection were due to *E. coli*. Hoberman et al¹¹ (1993) reported as *E. coli* as the most common bacterium isolated in his study. All these studies correlate with our study with *E. coli* being commonest isolate.

In our study 70% of children who showed numerous pus cells were culture positive and 54% were culture positive who showed 8 to 10 pus cells and 43% of children showing 6-8 pus cells were culture positive. Hence the presence of pyuria of >5 leukocytes/HPF in a centrifuged sample is a significant indicator of UTI. The study would have been more conclusive if urine culture were done in all febrile children screened, but economical constraints limited us to do urine culture only in those children showing significant pyuria of >5 pus cells/HPF of centrifuged urine sample and we found that 53% of febrile pyuric children were culture positive.

V. Conclusions

Our tertiary centre caters to a group of children at high risk of UTI as can be estimated from the 10.9 per cent culture positivity. Similar situations exist in other parts of India, albeit the load has not been studied in children^{20,21}. Important facts emanating from the present study include (i) infants (31.8%) represent a significant group vulnerable to UTI (ii) Male gender is clearly a risk factor towards acquiring UTI in infancy similar to taneja et al²¹ after which females predominant. E coli (47.1%) was the leading etiology of pediatric UTI at our center, multidrug resistant microbes (K. pneumonia, p. aeruginosa) were responsible for a substantial proportion of infections, however, staphylococci were not found to play a major role in UTI at our center (<2%) unlike reports from else were^{21,22}. Co-trimoxazole and Norfloxacin once the mainstay in treatment of UTIs, were no longer useful at our center. Our study would have been more conclusive if urine culture was done in all febrile children and those children with known genitourinary anomalies.

References

- [1]. Schlager T. Urinary tract infections in infants and children. *Infect Dis Clin North Am* 2003; 17: 353-365.
- [2]. Wald ER. Cystitis and pyelonephritis. In: Feigin RD, Chery JD, Demmier GJ, Kapian SL, eds. *Textbook of Pediatric Infectious Diseases*, 5th edn, Philadelphia: Saunders 2004; p 541-53.
- [3]. Fallahzadeh MH, Alamdarlu HM. Prevalence of urinary tract infection in pre-school febrile children. *Iranian J of Med Sci* 1999; 24: 35-39.
- [4]. Ma JF, Shortliffe LM. Urinary tract infection in children: etiology and epidemiology. *UrolClin North Am* 2004; 3: 517-526.
- [5]. Adjei O, Opoku C Urinary tract infections in African infants. *Int J Antimicrob Agents* 2004; 24 Suppl 1: S32-34.
- [6]. Yüksel S, Öztürk B, Kavaz A, Özçakar ZB, Acar B, Güriz H, et al. Antibiotic resistance of urinary tract pathogens and evaluation of empirical treatment in Turkish children with urinary tract infections. *Int J Antimicrob Agents* 2006; 28: 413-416.
- [7]. Yıldız B, Kural N, Durmaz G, Yazar C, Ak I, Akcar N. Antibiotic resistance in children with complicated urinary tract infection. *Saudi Med J* 2007; 28: 1850-1854.
- [8]. Mathai D, Jones RN, Pfaller MA. SENTRY Participant Group North America. Epidemiology and frequency of resistance among pathogens causing urinary tract infections in 1,510 hospitalized patients: a report from the SENTRY Antimicrobial Surveillance Program (North America). *DiagnMicrobiol Infect Dis* 2001; 40: 129-136.
- [9]. Kahlmeter G. Prevalence and antimicrobial susceptibility of pathogens in uncomplicated cystitis. The ECO SENS study. *Int J Antimicrob Agents* 2003; 22: 49-52.
- [10]. Zorc JJ, Kiddoo DA, Shaw KN. Diagnosis and management of Pediatric urinary tract infections. *clin microbiol rev* 2005; 18 : 417-422
- [11]. Hoberman A, Chao HP, Keller DM, Hickey R, Davis HW, Ellis D. Prevalence of urinary tract infection in febrile infants. *J pediatri* 1993; 123 : 17-23.
- [12]. Dharmidharka VR. Prevalence of bacteriuria in febrile infants. *Indian Pediatr* 1993; 30: 981-986
- [13]. Kaushal R.K, Bansal S, Sharma V.K, Sood A, Goyal A Urinary tract infection among children presenting with fever. *Indian Pediatr* 2003; 40:269-270.
- [14]. Shaw KN, Gorelick MH. Urinary tract infection in the pediatric patient. *Pediatric clinics of North America* 1999; 46: 6.
- [15]. Roberts KB, Charney E, Sweren RJ, Ahonkhai VI, Bergman DA, Coulter MP et al: Urinary tract infection in infants with unexplained fever: a collaborative study. *J Pediatr* 1983; 103: 864-867.
- [16]. Srivaths PR, Rath B, Krishanprakash S, Talukdar B et al: Usefulness of screening febrile infants for urinary tract infection. *Indian Pediatr* 1990; 33: 218-220
- [17]. Bryan CS, Reynolds KL. Community acquired Bacteremic urinary tract infection: Epidemiology and Outcome. *J Urol* 1984; 132: 490, 934.
- [18]. Bagga A, Sharma J. Urinary tract infections clinical features, evaluation and treatment. *Pediatr today* 2000; 3: 395-401
- [19]. Clinical and Laboratory Standards Institute. Performance standards for antimicrobial susceptibility testing; 17th informational supplement, CLSI M100-S17. Vol. 27 no.1. Wayne, PA: Clinical and Laboratory Standards Institute: 2007.
- [20]. Navaneeth BV, Belwadu S, Suganthi N. Urinary pathogens' resistance to common antibiotics: a retrospective analysis. *Trop Doct* 2002; 32 : 20-2.
- [21]. Taneja N, Chatterjee SS, Singh M, Singh S, Sharma M. Pediatric urinary tract infections in a tertiary care center from north India *Indian J Med Res*. 2010 Jan;131:101-105.
- [22]. Mohammed A, Mohammed S, Khan AU. Etiology and antibiotic resistance patterns of community acquired urinary tract infection in JNMC Hospital Aligah, India. *Ann Clin Microbiol Antimicrob* 2007; 6 : 4-11.