Clinical Characteristics and Microbiological Analysis of Urinary Tract Infection in Children of 2 Months to 12 Years: A Prospective Study

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ABSTRACT

Objectives: To study the clinical characteristics and etiological agents of urinary tract infection (UTI) in children of 2 months to 12 years, and to find the prevalence of vesicoureteric reflux (VUR) in children with UTI.

Materials and methods: This prospective study was conducted among 47 children with culture-positive UTI in children between 2 months to 12 years. All children attending the pediatric outpatient department (OPD) of Government Medical College Hospital, Kannur, Kerala, India, were enrolled. Clinical and bacteriological profile of all cases were evaluated and ultrasonogram (USG) abdomen/micturating cystourethrogram (MCU) were done for these patients to find the predisposing factors associated with recurrent infections like VUR and posterior urethral valve.

Results: A total of 47 children were enrolled, of which 25 (53.19%) children were below 1 year of age. Male to female ratio was 2.1:1. The most common symptom was fever, and Escherichia coli (65.9%) was the most common organism isolated. Within the cohort, 23.40% of children had recurrent UTI and 19.14% showed VUR.

Conclusion: Urinary tract infection is a very common infection in infants and children. It is equally important that we treat the child with antibiotics and investigate for any underlying causes such as VUR or obstructive uropathy.

Keywords: Micturating cystourethrogram, Urinary tract infection, Vesicoureteric reflux.

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INTRODUCTION

Urinary tract infection (UTI) is a common cause of febrile illness among the pediatric age-group (third most common), and it accounts for 0.7% of all outpatient’s clinic visits.¹ In the emergency department, up to 14% of children are admitted because of UTI either due to direct infection or indirectly due to its associated complications.² Knowledge of the exact incidence and prevalence of UTI among children is essential for pediatricians because the diagnosis of the diagnosis may be challenging on a clinical basis, and the incidence and prevalence will determine the cost–benefit effect of investigating the condition. Incidence of UTI is around 1% in boys and 1–3% in girls. During infancy the male to female ratio is 2.8–5.4:1. After which, there is a female preponderance, with a male to female ratio of 1:10. In boys, most UTIs occur within the first year of life; UTIs are much more common in uncircumcised boys, especially within 1 year of age. In girls, the first UTI usually occurs by age of 5 years, which peaks during infancy and toilet training period.³

Colonic bacteria are the most common etiological agent for UTI. Approximately 75–90% of UTI in girls are caused by E. coli, followed by Klebsiella and Proteus species. Although E. coli is the most common organism in males, some studies have reported that in boys older than 1 year, Proteus is as common a cause as E. coli; others report a preponderance of gram-positive organisms in boys. Staphylococcus saprophyticus and Enterococcus are uropathogens in both sexes. The risk factors associated with UTI are females, uncircumcised boys, vesicoureteral reflux (VUR), dysfunctional voiding, obstructive uropathies, urethral instrumentation and manipulation, wiping from back to front in girls, tight clothing, pinworm infestation, constipation, bacteria with P fimbriae, anatomic abnormality (labial adhesion), and neuropathic bladder.³

Acute pyelonephritis has been diagnosed in two third of febrile UTIs in early childhood.⁴ Published data which have used renal parenchyma nuclear scans like dimercaptosuccinic acid (DMSA) and dihydrenetramine pentaacetic acid (DTPA) conclude that more than 75% of children under 5 years of age with febrile UTI have pyelonephritis.⁵–⁷ Pyelonephritis leads to renal scarring in 27% to 64% of children with UTI, even in the absence of the underlying urinary tract abnormalities.⁸,⁹ Most UTIs that lead to scarring or diminished kidney growth occurred in children less than 4 years, especially within the first year of life.⁶,⁹ It is essential to diagnose UTIs in febrile children at the earliest and intervene to reduce the potential of lifelong morbidity. Optimum treatment results in symptomatic relief, prevents progressive renal damage, and urosepsis with immediate bacterial eradication.¹⁰ Progressive renal damage from unrecognized pyelonephritis may lead to

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hypertension and chronic renal failure later in life. A study from Sweden showed that focal renal scarring caused by pyelonephritis in children carried 23% risk of hypertension, 10% risk of end-stage renal disease, and 15% risk of toxemia during pregnancy as an adult. Approximately, 13–15% of end-stage renal disease is thought to be related to UTI in childhood, which was often unrecognized and therefore not adequately treated.

Vesicoureteral reflux is defined as the backward flow of urine from the bladder into the ureter. Vesicoureteric reflux is present in approximately 30% of females who had a UTI and in 5–15% of infants with antenatally detected hydronephrosis. Vesicoureteric reflux predisposes to pyelonephritis by facilitating the transport of bacteria from the bladder to the upper urinary tract. The inflammatory reaction caused by pyelonephritis can result in renal injury or scarring, which is also termed reflux-related renal injury or reflux nephropathy. Of the children with febrile UTI, those with VUR are three times more likely to develop renal injury compared to those without VUR, which can be identified by doing an micturating cystourethrogram (MCU).

The present study aimed at identifying the clinical and bacteriological profile of children with UTI in the age-group of 2 months to 12 years.

**Materials and Methods**

It was a prospective study. Inclusion criteria were to take all culture-positive cases of UTI between the age-group of 2 months and 12 years. We excluded children whose parents were not willing to take part in the study. Children with culture-positive UTI between the age-group of 2 months and 12 years attending the pediatric OPD of Government Medical College Hospital, Kannur during a 1-year period (from April 1, 2018, to March 31, 2019) were enrolled. The sample size was calculated using the formula

\[ p(1 - p)/d^2 \]

where \( p \) is the prevalence and \( d \) is the absolute precision. A detailed proforma for collecting data related to age, sex, birth weight, bowel habits, etc., were prepared. A detailed clinical history related to the onset of fever, duration, associated symptoms such as nausea, vomiting, diarrhea, urinary disturbances, and other system involvement was obtained. Routine physical examination was done in all enrolled children.

Complete blood counts, urine analysis, and urine culture and sensitivity were done for all patients. Midstream urine samples were used for routine and culture analysis. Urine culture was done by BACTEC method. In patients whose urine routine was positive for UTI (pus cells >5) were started on oral empirical antibiotics, such as cefixime, and after culture reports antibiotics were adjusted accordingly. An USG of the abdomen, renal function test, and MCU were done in all cases to identify abnormalities such as VUR and posterior urethral valve per the guidelines laid down by the Indian Society of Nephrology (Tables 1 to 3).

**Statistical Analysis**

The data were entered to Microsoft Excel and were analyzed using SPSS software. The results were described using frequencies and proportions. Inferential statistics such as Chi-square test and Fisher exact test were used to test the significance of association between various etiological factors and occurrence of UTI.

**Results**

A total of 47 children studied. Of them, 25 (53.19%) children were below 1 year and 22 (46.80%) above 1 year of age; 26 (55.31%) were males and 21 (44.6%) were females. Of the 26 male children, only 5 (19.2%) males had undergone circumcision.

Symptomatically 63.82% of children presented with fever, 40.42% with dysuria, 17.02% with vomiting, 10.6% with increased frequency of micturition and urgency, and 4.25% with irritability. During the antenatal period, 14.8% of them had abnormal findings. Presence of fever was noted in 72.97% children, 96.77% had a positive symptom, and 5.45% were below 6 months and 4.25% above 6 months, which was important as children unrecognised and therefore not adequately treated.

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| Table 1: Clinical profile of children with urinary tract infection |
|---------------------------------------------------------------|
| **Clinical features**          | **Number of patients (n = 47)** | **Percentage** |
|--------------------------------|---------------------------------|---------------|
| Fever                         | 30                              | 63.82         |
| Dysuria                       | 19                              | 40.42         |
| Vomiting                      | 8                               | 17.02         |
| Irritability                  | 2                               | 4.2           |
| Increased frequency of micturition | 5                            | 10.63         |
| Urgency                       | 5                               | 10.63         |
| Constipation                  | 6                               | 12.76         |

| Table 2: Demographic data |
|---------------------------|
| **Age**                   | **Percentage** |
| <1 year                   | 25 (53.19%)    |
| >1 year                   | 22 (46.80%)    |
| **Gender**                | **Percentage** |
| Male                       | 26 (55.31%)    |
| Female                     | 21 (44.6%)     |

| Table 3: Bacteriological profile of children with urinary tract infection |
|--------------------------------------------------------------------------|
| **Bacterial isolates** | **Number of isolates** | **Percentage** |
|------------------------|------------------------|---------------|
| *Escherichia coli*      | 31                     | 65.95         |
| *Klebsiella spp.*      | 8                      | 17.02         |
| *Proteus mirabilis*    | 4                      | 8.51          |
| *Coagulase-negative Staphylococcus*                                   | 2                      | 4.25          |
| *Pseudomonas*          | 2                      | 4.25          |

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*Escherichia coli* was the most common organism (65.95%) identified in all culture positive followed by *Klebsiella* (17.02%), *Proteus* species (8.51%), coagulase-negative *Staphylococci* (4.25%), and *Pseudomonas* in 4.25%. *Escherichia coli* was sensitive to ofloxacin, amikacin, nitrofurantoin, ceftriaxone, and piperacillin–tazobactam. *Klebsiella* was sensitive to nitrofurantoin, cefixime, ofloxacin, and amikacin. Both *E. coli* and *Klebsiella* were resistant to amoxicillin–clavulanic acid. Among the cases, 8.5% had to be treated with multiple drugs in order to get a sterile culture. Average duration of treatment was about 7–10 days.

Urine stream was good in 87.27% children but was poor in 12.76% (as observed by parents), which was important as children with poor urine stream have more chances of having lower urinary tract anomalies such as posterior urethral valve which in turn can increase the chances of UTI. Proportion of children with
constipation was 12.76%, 29.7% (14) children had pallor, 42.55% (20) of children had recurrent UTI out of which only 3 (6.3%) had antenatal hydronephrosis. Of all the children with UTI, the USG abdomen showed abnormal findings (hydronephrosis and hydroureteronephrosis) in 13 (27.65%), and 9 (19.14%) children had VUR.

Among children with VUR, 22% had grade VUR, 44% grade III, and 33% grade IV. For grades I and II, antibiotic prophylaxis was given with cephalixin and MCU was repeated; whereas in cases of grades III and IV, after starting on prophylaxis if the child was symptomatic, the treatment of choice was pyeloplasty.

**DISCUSSION**

Urinary tract infections are one of the most common, potentially serious infections of childhood. They can cause acute morbidity as well as long-term complications including hypertension and impaired renal function. Prevalence of culture-positive UTI in our study, <1 year—25 (53.19%) and >1 year—22 (48.80%), which is comparable to Gupta et al.13 and Ghagade et al.14 who found maximum number of patients (56.5 and 37.7%, respectively) in the age-group of less than 1 year. However, Sharma et al.,15 Malla et al.,16 Singh et al.,17 and Chand et al.18 have reported maximum number of patients between 1 year and 5 years (varying from 35 to 50%) in this age-group. Our study had shown male to female ratio of 1:2.1, which is actually not consistent with other studies which documented higher rates of infection in females, with a male to female ratio varying from 1.1:3 to 1:2. This is probably because our study had more affected children <1 year, and males are more susceptible to UTI in the first year of life; 80.76% were uncircumcised males compared to 19.24% circumcised males.

Fever was the most common symptom in our study (63.82%). Other studies have also found it to be the most common symptom, i.e., reported in 45–87% of their patients.13–15 The other common symptoms noted were 40.42% with dysuria, 17.02% with vomiting, 10.63% with increased frequency of micturition and urgency, and 4.25% with irritability that correlates with other studies. Infants presented with nonspecific symptoms such as fever, vomiting, poor feeding, diarrhea, and irritability, while older children had urinary symptoms and pain in abdomen with or without fever.

In our study, *E. coli* (65.95%) was the commonest organism isolated in both the genders and in all the age-groups, followed by *Klebsiella* (17.02%). This finding has also been observed in studies from other parts of India.15–17 Therefore, *E. coli* remains the most common uropathogen. As also reported in other studies, in our study *Proteus* was isolated only in males. This is because of the frequent colonization of the prepuce in males.

Twelve (25.5%) children had recurrent UTI and 9 children (19.14%) showed VUR. According to previous studies done in Indian children,18 primary VUR occurs in less than 1% of the general population and as many as 50% of children who present with recurrent UTI will have VUR. More than 75% of children under 5 years of age with febrile UTI have pyelonephritis, and higher risk of renal scarring is seen in children under 3 years of age with recurrent UTI.18

**CONCLUSION**

Urinary tract infection is a very common infection in infants and children. It is equally important to treat the child with UTI and also look for any underlying cause like VUR or obstructive uropathy which should not be missed. Prevalence of UTI is more common in <1 year males. The most common organism isolated was *E. coli*.

The most common symptom on presentation was fever. In all culture-positive UTI, it is always important to do an USG scan and MCU to look for pyelonephritic changes, renal scarring, and VUR, and treat them at the earliest.

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