Risk Factors of Febrile Urinary Tract Infection in Children

Abstract

Background: Urinary tract infection (UTI) implies invasion of the urinary tract by pathogens, which may involve the upper or lower urinary tract depending on the infection in the kidney or bladder and urethra. UTI are the third most common bacterial infection in children in developing countries. The risk factors that predispose to UTI are diverse and should be evaluated during diagnosis.

Objective: To estimate risk factors of febrile UTI patient in children.

Study design: In this case control study, a total of 50 cases and 50 controls were enrolled purposively and conveniently and was carried out in the Department of Pediatric Nephrology and Department of Pediatrics, of Comilla Medical College Hospital, Comilla, Bangladesh between February 2013 to December 2013.

Methods: Cases were selected following clinically suspected by fever, dysuria, urgency, frequency, incontinence, tender renal angle and on urine analysis having >10 pus cells/HPF. Then they were confirmed by urine culture whose urine growth were >105 colony forming unit per ml. Controls were selected from the patients admitted for other diseases in the Pediatrics ward. Cases and controls were searched for risk factors of urinary tract infection. Data were collected by a preformed structured questionnaire.

Results: Study showed that female sex (P<0.05), constipation (P<0.001), not taking anthelmintic (P<0.001), lack of toilet training (P<0.001) and inadequate water intake (P<0.01) were significant risk factors for urinary tract infection in children and there was no relationship between school going and UTI.

Conclusion: Female sex, constipation, not taking anthelmintic, lack of toilet training and inadequate water intake are significant risk factors for febrile urinary tract infection in children.

Introduction

Urinary tract infection (UTI) implies invasion of the urinary tract by pathogens, which may involve the upper or lower urinary tract depending on the infection in the kidney or bladder and urethra. UTI constitutes a common cause of morbidity in association with abnormalities of the urinary tract; resulted in long term complications, including hypertension and chronic renal failure. Prompt detection and treatment of UTI is of the utmost importance [1]. UTI may be defined by clinical presentation of UTI together with the presence of single pathogenic organism in urinary tract with growth of which is more than 10^5 cfu per ml [1]. It is the third most common bacterial infection in children in developing countries after those of the gastrointestinal and respiratory tract. Because of nonspecific signs and vague symptoms in very young children, they remain unrecognized and therefore precise data on incidence and prevalence of UTI are not available. The commonest age for the occurrence of the first symptomatic UTI is the first year of life and male affected more than female [1]. During infancy, the risk of developing UTI is equal in boys and girls and thereafter higher in girls [2].

About 90 percent of the first symptomatic UTI and 70 percent of recurrent infections are due to E. coli. Proteus species may be causative in about a third of boys with acute cystitis. Other organisms including Klebsiella, staphylococcus epidermidis and streptococcus faecalis may occasionally be responsible. Pathogens of low virulence and fungi may be causative agent in patients who are immuno-compromised. Candida albicans infections are particularly seen in preterm babies [2].

Most UTI’s are ascending infections. The bacteria arise from the fecal flora, colonize the perineum, and enter the bladder via the urethra. In uncircumcised boys the bacterial pathogens arise from the flora beneath the prepuce. In some cases, the bacteria causing cystitis ascend to the kidney to cause pyelonephritis. If bacteria ascends from the bladder to the kidney, acute pyelonephritis can occur. Normally the simple and compound papillae in the kidney have an anti-reflux mechanism that prevents urine flow from renal pelvis to collecting tubule. Infected urine then stimulate an immunologic and inflammatory response. Backflow of vesicoureteral reflux to developing kidney ultimately damage kidney that causes renal injury and scarring [1]. UTI may present with pyelonephritis and cystitis. Those who present with pyelonephritis has history of abdominal or flank pain, fever, chills and those who present with cystitis may have history of dysuria, frequency, urgency, suprapubic pain, incontinence and malodour urine [1].
UTI may be suspected based on symptoms and findings on urine analysis or both. A urine culture is necessary for confirmation and appropriate therapy [1]. There are several methods to obtain a urine sample. Some are more accurate than others. In toilet-trained children, a midstream urine sample usually is satisfactory; the introitus should be cleaned before obtaining the specimen. In uncircumcised boys, the prepubis must be retracted; if the prepuce is not retractable; avoided sample may be unreliable and contaminated with skin flora. In children who are not toilet-trained, a catheter sample is better but too invasive [1].

The risk factors of UTI are numerous; they may be age, gender, constipation, lack of circumcision, inadequate water intake, not taking antihelmintic and lack of toilet training. As male are more likely to be born with structural abnormalities of the urinary tract, UTI is more common in their first six months of life [3]. Few infections are encountered in boys in the age range of 1-5 years. In preschool years symptomatic infections occur 10 to 20 times more frequently in girls than in boys [4]. Constipation is a common problem of children worldwide. Estimates of the prevalence [5,6] rate of functional constipation in the pediatric population have varied from 4% to 37%. Enterobius vermicularis is one of the most prevalent worms found in children worldwide. Children, particularly when there are heavy worm burdens, neurological symptoms such as nervousness, restlessness, irritability and distraction may occur and these may influence on child growth[7] and may occur urinary tract infections [8]. Uncircumcised male infants appear to be at increased risk of UTI. The periurethral area was found to be more frequently and more heavily colonized with uropathogens, especially Escherichia coli, in uncircumcised infants than in circumcised infants [9, Winberg et al. [9] offered an explanation for the high incidence of UTIs in uncircumcised male infants in an intriguing article.

**Rationale**

Urinary tract infection is a great morbidity in children. It encompasses renal damage as well as mortality of children. Many risk factors are responsible for initial urinary tract infection and recurrent urinary tract infection. But there is scanty data in Bangladesh to estimate the risk factors of febrile urinary tract infection by case control study. The goal of our study was to shed some light in some risk factors which are common in urinary tract infection in children. So that it can create public awareness to prevent UTI and improve the well being of children.

**Patients and Methods**

The objectives of this study were to identify the risk factors of febrile urinary tract infection in children. This was a case control study carried out in the Department of Pediatric Nephrology and Department of Pediatrics, Comilla Medical College, Bangladesh from February 2013 to December 2013.

**Studied population**

All UTI patients of children were included in this study with the following inclusion and exclusion criteria.

**Inclusion criteria**

Children who were between 5 years to 12 years old, can give history of fever, abdominal pain, dysuria, urgency, frequency of micturition and able to give urine sample aseptically by mid stream urine and cleaning the external urethral orifice were included in this study.

**Exclusion criteria**

Children whose history were difficult to take from the parents, unable to give urine sample and having other cause of fever like viral respiratory tract infection, other cause of abdominal pain like acute appendicitis or gastro enteritis and who had congenital anomalies of urinary tract detected by ultra sonogram were excluded from the study.

**Sampling method and sample size**

Purposive sampling method was applied and conveniently 50 samples were taken as cases.

**Controls**

50 Controls were selected from the Department of Pediatrics who had no symptoms of urinary tract infection and excluded by laboratory investigations.

**Procedure**

Immediately after admission, detailed history and physical examination of the patient was done. Cases were selected from Pediatric Nephrology ward when clinically suspected as febrile UTI by fever, dysuria, urgency, frequency, abdominal pain, malodour urine, incontinence, tender suprapubic region, tender renal angle and on urine analysis having >10 pus cells/HPF. Then they were confirmed by urine culture whose urine growth were >10^7 colony forming unit per ml. Negative culture patients were excluded from the study. Controls were selected from the patients admitted for other diseases in the pediatrics ward. Cases and controls were searched for risk factors of urinary tract infection. Data were collected by a preformed structured questionnaire and were analyzed using computer software and SPSS. Significance of difference was calculated using, chi-square(χ²) test. Multiple regression analysis was also calculated out.

**Flow chart**

Initially 500 patients were included in this study based on clinical features → Among them 200 were found positive urine analysis → from those patient 50 patients were found positive urine culture → this 50 patients were searched for risk factors of UTI and were compared with 50 controls by statistical analysis.

**Ethical issues**

Ethical issues were addressed duly by taking permission of parents and approval from ethical committee of Comilla Medical College, Bangladesh. Total 50 cases and 50 controls were included in this study. The mean age of cases and control were 7.4 years and 7.8 years respectively. Regarding sex 22 were male and 28 were female among cases and 34 were male and 16 were female among controls. Among the cases 100% were having clinical features of UTI, positive urine analysis and positive urine culture. None of the cases and controls had congenital anomalies of urinary tract detected by ultra sonogram (Table 1-3).
Table 1: Baseline characteristics of Cases & Controls.

<table>
<thead>
<tr>
<th></th>
<th>Cases</th>
<th>Controls</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>Total number</td>
<td>50</td>
<td>50</td>
</tr>
<tr>
<td>2</td>
<td>Age (years)</td>
<td>7.5</td>
<td>7.7</td>
</tr>
<tr>
<td>4</td>
<td>Sex</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>Male</td>
<td>22</td>
<td>34</td>
</tr>
<tr>
<td></td>
<td>Female</td>
<td>28</td>
<td>16</td>
</tr>
<tr>
<td>5</td>
<td>Clinical features present (%)</td>
<td>100</td>
<td>-</td>
</tr>
<tr>
<td>6</td>
<td>Urine R/M/E (%)</td>
<td>100</td>
<td>-</td>
</tr>
<tr>
<td>7</td>
<td>Urine culture (%)</td>
<td>100</td>
<td>-</td>
</tr>
<tr>
<td>8</td>
<td>USG findings of urinary tract</td>
<td>nil</td>
<td>nil</td>
</tr>
</tbody>
</table>

Table 2: Effect of different risk factors independently associated with UTI.

<table>
<thead>
<tr>
<th>Factors</th>
<th>Case</th>
<th>Controls</th>
<th>x²</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Female Sex</td>
<td>28</td>
<td>16</td>
<td>5.8</td>
<td>&lt;0.05</td>
</tr>
<tr>
<td>Constipation</td>
<td>32</td>
<td>15</td>
<td>11.6</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Lack of Circumcision</td>
<td>7</td>
<td>12</td>
<td>4.9</td>
<td>&lt;0.05</td>
</tr>
<tr>
<td>Not Taking Anthelmintic</td>
<td>15</td>
<td>28</td>
<td>6.9</td>
<td>&lt;0.01</td>
</tr>
<tr>
<td>Lack of Toilet Training</td>
<td>10</td>
<td>14</td>
<td>23.3</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Inadequate Water Intake</td>
<td>5</td>
<td>17</td>
<td>8.3</td>
<td>&lt;0.01</td>
</tr>
<tr>
<td>Effect of School Going</td>
<td>37</td>
<td>37</td>
<td>0.00</td>
<td>&gt;.5</td>
</tr>
</tbody>
</table>

Interpretation: Female sex, constipation, lack of circumcision, not taking anthelmintic, lack of toilet training, inadequate water intake had significant effect independently on UTI but school going had no effect on UTI.

Table 3: Multiple Logistic Regression analysis results showing factors associated with UTI infection in children.

<table>
<thead>
<tr>
<th>Factors</th>
<th>OR (95% CI)</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Female Sex</td>
<td>1.5(1.33-1.58)</td>
<td>&lt;0.05</td>
</tr>
<tr>
<td>Constipation</td>
<td>5(2.89 - 9.57)</td>
<td>&lt;0.05</td>
</tr>
<tr>
<td>Lack of Circumcision</td>
<td>5(2.89 - 9.57)</td>
<td>&lt;0.05</td>
</tr>
<tr>
<td>Lack of Circumcision</td>
<td>1.2(1.10 - 3.65)</td>
<td>&lt;0.05</td>
</tr>
<tr>
<td>Inadequate Water Intake</td>
<td>1.3(1.23 - 4.41)</td>
<td>&lt;0.05</td>
</tr>
<tr>
<td>School Going</td>
<td>1.0(0.40 - 2.44)</td>
<td>1.00</td>
</tr>
<tr>
<td>Lack of Toilet Training</td>
<td>1.6(1.15 - 2.93)</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>

Interpretation: Multiple Logistic Regression analysis results showed multiple independent variables like female sex, constipation, lack of circumcision, inadequate water intake, not taking anthelmintic and lack of toilet training were significant risk factors for dependent variable UTI but school going had no effect on UTI.

Interpretation

Female sex, constipation, lack of circumcision, not taking anthelmintic, lack of toilet training, inadequate water intake had significant effect independently on UTI but school going had no effect on UTI.

Multiple Logistic Regression analysis results showed multiple independent variables like female sex, constipation, lack of circumcision, inadequate water intake, not taking anthelmintic and lack of toilet training were significant risk factors for
dependent variable UTI but school going had no effect on UTI.

Discussion
Urinary tract infections (UTIs) are more common among girls than boys. The predominant UTI risk factors in young girls are anatomic and physiologic factors. The short urethra of female permits easy entry of bacteria into the bladder and causes UTI. In our study we have found female baby are more prone to developed UTI than male baby (p<.05). Similar results are found by Foxman B [10]. Harrington and Hooton also supported our results [11]. This finding also supported by Nader et al. [12] and Steven L [13].

It has been shown that constipation with a dilated rectum causes the same pattern of voiding dysfunction as that encountered in children with persistence of an unstable bladder. Effective treatment of the constipation results in normalization of bladder function and cessation of UTIs [14]. In our study we have found constipation is a risk factor to developed UTI (p<0.001). A relationship between constipation and UTIs is well described by O’Regan et al. [15]. Farid Imanzadeh and Ali-Akbar found similar result [16].

In one study, appropriate hygiene decreased significantly the incidence of pinworm, adhesions, and inflammation [17]. We have found UTI are more common in male children who did not get circumcision (p<0.05). Herzog suggested in a case control study that there is a correlation between urinary tract infections and circumcision [18]. This also supported by many other studies [19,20].

Enterobiasis is most common in children between 5 years and 10 years of age. Serious complications like urinary tract infection may occur due to pin worm. Regular anthelmintic taking may reduce the prevalence of urinary tract infection. We have found taking anthelmintic reduce the infection of UTI (p<0.01). This is supported by study conducted by Okuz et al. [20] and Gokalp et al. [21]. Effect of lack of toilet training is a significant risk factor of UTI (P<0.001). We can conclude that lack of toilet training is extremely high relationship with UTI. This was supported by study conducted by Chen J et al. [22].

Bacterial eradication from the urinary tract is partially dependent on urine flow and voiding frequency. Therefore, it seems logical to postulate a connection between fluid intake and the risk of urinary tract infections (UTIs). In our study we found inadequate water intake predisposes to developed urinary tract infection (p<.01). Denman SJ & Burton JR suggested that mild dehydration is a risk factor of urinary tract infection [23].

Limitation of this study
Small sample size is one of the limitations of the study. Due to lack of kits we did not perform nitrate Patient did not co operate to do MCU to exclude VUR.

Conclusion
Female sex, constipation, not taking anthelmintic, lack of toilet training and inadequate water intake are significant risk factors for febrile urinary tract infection in children.

References


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enuresis nocturia and bacteruria in primary school girls. Indian Pediatr 28(8): 948-950.
