

A study of asymptomatic bacteriuria in Egyptian school-going children.

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Abstract

Background: Urinary tract infections (UTI) are a common and important clinical problem in childhood. Upper urinary tract infections (i.e., acute pyelonephritis) may lead to renal scarring, hypertension, and end-stage renal disease. Despite the presence of simple and reliable methods of preliminary screening of children's urine, urinary tract infection continues to be under diagnosed.

Objectives: The aim of this study was to establish prevalence rates of significant bacteriuria in asymptomatic school children by simple urine tests in comparison to standard urine culture techniques in Giza, Egypt.

Patients and methods: A total of 1000 apparently healthy school going children (6-12) years, 552 boys (55.2%) and 448 girls (44.8%), were enrolled in this cross-sectional prevalence survey.

Results: Overall prevalence of significant bacteriuria was 6%. Higher prevalence occurred in girls (11.4%) than boys (1.6%). *Escherichia coli* was isolated in 35 (58%) cases (3 boys and 32 girls), *Staph. aureus* in 13 (22%) cases (3 boys and 10 girls), *Enterobacter* in 6 girls (10%), *Kelbsiella pneumoniae* in 3 boys (5%) and *Proteus vulgaris* in 3 girls (5%)

Conclusion: Asymptomatic bacteriuria could be detected by urine screening program at school age. Overall prevalence of significant bacteriuria was 6%, with predominance in girls than boys.

Keywords: Bacteriuria, asymptomatic, prevalence, children, school, male, female, simple urine tests.

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Introduction

Asymptomatic bacteriuria is defined as a significant bacterial count (usually 10^5 organism /ml) present in the urine of a person without symptoms.¹ Prevalence of asymptomatic bacteriuria differs according to different socioeconomic status and geographical areas. The prevalence of asymptomatic bacteriuria in Egyptian school children was 7% in a previous study done by El Gamal and Saleh² another study done in India by Kondapaneni et al. revealed a prevalence rate of 16.5%.³

Infection with symptoms will alert a person to take treatment in time to prevent further complications. But in case of asymptomatic persons, they may come to hospital with complications. Early detection of infection in asymptomatic children and prompt therapeutic intervention are essential to prevent cases of asymptomatic

UTI from becoming symptomatic with resultant pyelonephritis and renal damage.³

A previous study reported that the number of children with bacteriuria was significantly reduced at follow-up six months after antibiotic treatment.⁴

Nitrofurantoin and pefloxacin are the most active agents against all bacterial isolates.⁵

There are several rapid tests for the detection of UTI in children. These include leukocyte esterase, nitrate test and catalase test.⁶

For any given test administered to a given population it is important to calculate the sensitivity, specificity, positive predictive value (PPV) and negative predictive value (NPV) in order to determine how useful the test is to detect a disease in a given population. Sensitivity is the probability that an individual with the disease will test positive.⁷ Specificity is the probability that an individual without the disease will test negative.⁷ Positive predictive value (PPV) refers to the probability that a positive test correctly identifies an individual who actually has the disease.⁷ Negative predictive value (NPV) refers to the probability that a negative test correctly identifies an individual who does not have the disease.⁷

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Urine dipstick testing alone may provide an adequate initial UTI screen⁸ however urine culture is the standard test for the diagnosis of UTI⁹.

This study aimed to establish prevalence rates of significant bacteriuria in asymptomatic school children by simple urine tests in comparison to standard urine culture techniques in Giza, Egypt.

Patients and methods

Study design and subjects: This Cross-sectional study was carried out in Giza governorate in Egypt from October 2009 to January 2011. A total of 1000 apparently healthy school going children (6-12) years including 552 males (55.2%) and 448 females (44.8%) were enrolled in this study. All males had undergone religious based circumcision before enrollment in the study.

They were randomly selected by simple random selection method from 10 primary public schools after obtaining an informed consent from child parents and school managers. The study was approved by the Ethics Committee of the Pediatrics Department, Faculty of Medicine, Ain Shams University.

Randomization steps: (simple random method)

- 1- Target population was determined as children attending 10 primary public schools in Giza, Egypt.
- 2- A frame including list of all children was done.
- 3- Sample size was determined to be 1000.
- 4- Random numbers were developed by specific software used for selection of cases through Spread sheet.

We excluded any child who was feverish or complained from any UTI symptom (dark urine, dysuria, frequency, renal pain, changes of urine volume and suprapubic pain), or received antibiotics during 48 hours prior to sample collection.

The children were clinically examined to exclude any child who had any of the following abnormalities (high temperature, high blood pressure, pallor, jaundice, skin rash, edema and evaluating its extent (puffiness of eye lids), lower limb edema, abdominal tenderness, palpable masses or ascites).

Methods

All children enrolled in the study were subjected to the following:

Mid stream urine specimens were collected after implementing standard precautions of clean catch urine specimen as follows: For girls, the labia was spread and

the perineum cleaned two to three times with antiseptic solution (A cotton sponge soaked with benzalkonium hydrochloride is useful and non-irritating for this purpose.)¹⁰ or liquid soap and gauze pads. For boys, the meatus was cleansed in a similar fashion. Cleaning the perineum with soap prior to urine collection may decrease the rate of contamination.¹¹ The child was instructed verbally to urinate into a toilet or urinal. Midway through urination, a specimen was collected in a sterile container. Midstream urine collection demonstration video was shown to children prior to sample collection.¹²

Each specimen was divided into three parts: the first part of urine was tested with urine dipstick tests for nitrite and leukocyturia, the second part of urine was tested by catalase test, and the third part for culture and sensitivity.

1. Detection of nitrite and leukocyte esterase: a dipstick test (Medi-Test Combi 10®SGL, Macherey Nagel, Germany) was performed on fresh urine specimen with a reagent strip designed to react and progressively produce color changes in 30 seconds. The results were obtained by visual comparison of the test strip with a color chart provided on the bottle label. The test result was considered positive if the dipstick turned pink or red for nitrites or purple for leukocytes within 2 minutes of contact with the urine.

2. Catalase test: a few drops of 10% hydrogen peroxide were added to urine tube. A positive test consists of appearance of foam on the surface of urine within two minutes.¹³

3. Urine culture: The remaining part of urine was subjected to microbiologic investigations at the microbiology department, faculty of medicine, Ain shams university (All the urine samples were transported to the laboratory within half an hour to one hour. The distance from laboratory to school is about 20 to 40 minutes. Hence not much time was lost for transportation of samples).

Standard quantitative culture was performed on CLED agar, incubated aerobically at 37°C for 24 hours and the developed colonies were counted to detect significant bacteriuria "more than 100,000 organisms/ml"¹⁴ and resulting colonies were identified according to standard methods of identification. (After counting of the bacteria colonies, bacteria were identified by taking advantage of the different biochemical properties of bacteria. Bacteria were incubated in different medias with color reactions indicating e.g. enzyme activity (lactase, fermentation, urease), motility or use of citrate as an

energy source. The results of the biochemical properties were assigned to bacteria species with the help of a table.)¹⁵

Statistical analysis

Data was collected; tabulated and Statistical analysis was done by using statistical package for social science (SPSS) version 17 and Microsoft Excel 2007.

Sensitivity is the number of patients with a positive test who have the disease [true positives (TP)] divided by all patients who have the disease [TP/ (TP+FN)]. A test with high sensitivity will not miss many patients who have the disease (i.e., low false negative rate). Whereas TP= true positives, FN= false negatives.^{7,16}

Sensitivity for Nitrite test: $28 / (28+32) = 28/60 = 46.67\%$

Sensitivity for Catalase test: $19 / (19+41) = 19/60 = 31.67\%$

Sensitivity for Leukocyte esterase: $15 / (15+45) = 15/60 = 25\%$

Sensitivity for Nitrite, Leukocyte esterase and/or Catalase $44 / (44+16) = 44/60 = 73.33\%$

Specificity is the number of patients who have a negative test and do not have the disease (true negatives) divided by the number of patients who do not have

the disease [TN/ (TN +FP)]. A test with high specificity will infrequently identify patients as having a disease when they do not (i.e., low false positive results). Whereas TN= true negatives, FP= false positives.^{7,16}

Specificity for Nitrite test: $872 / (872+68) = 872/940 = 92.77\%$

Specificity for Catalase test: $843 / (843+97) = 843/940 = 89.6\%$

Specificity for Leukocyte esterase: $865 / (865+75) = 865/940 = 92\%$

Specificity for Nitrite, Leukocyte esterase and/or Catalase: $781 / (781+159) = 781/940 = 83.08\%$

Positive predictive value (PPV) is computed from two-by-two tables: true positives / (true positives + false positives) [TP/ (TP+FP)].^{7,16}

Negative predictive value (NPV) is computed from two-by-two tables: true negatives / (false negatives + true negatives) [TN/ (TN+FN)].^{7,16}

Results

Nitrite test was positive in 96 (9.6 %) urine specimens, leukocyte esterase was positive in 90 (9%) specimens, catalase test was positive in 116 (11.6%) and 203 (20.3%) urine specimens were positive for nitrite and/or leukocyte esterase and / or catalase test. (Table 1).

(Table 1): Culture results of urine samples.

| | Positive culture (n= 60) | Negative culture (n= 940) |
|--------------|--------------------------|---------------------------|
| Boys (552) | 9 (1.6%) | 543 (98.4%) |
| Girls (448) | 51 (11.4%) | 397 (88.6%) |
| Total (1000) | 60 (6%) | 940 (94%) |

Sixty out of 1000 urine specimens (i.e., 6%) were positive for significant bacteruria by culture with predomi-

nance in girls (11.4%) compared to boys (1.6%) (Table 2).

(Table 2): Results of different screening tests used.

| Tests | Positive | Percentage |
|---|----------|------------|
| Nitrite test | 96 | 9.6% |
| Leukocyte esterase | 90 | 9% |
| Catalase test | 116 | 11.6% |
| Nitrite, leukocyte esterase and/or catalase tests | 203 | 20.3% |

E.coli was isolated in 35(58%) cases (3 boys and 32 girls), *bacter* in 6 girls (10%), *Kelbsiella pneumoniae* in 3 boys (5%) *S. aureus* in 13 (22%) cases (3 boys and 10 girls), *Entero-* and *Proteus vulgaris* in 3 girls (5%) (Table 3).

(Table 3): Types of bacterial isolates in cases of asymptomatic bacteruria.

| Types of bacterial isolates | Sex | | Total | % |
|----------------------------------|------|-------|-------|-----|
| | Boys | Girls | | |
| <i>Escherichia coli (E.coli)</i> | 3 | 32 | 35 | 58% |
| <i>Staphylococcus aureus</i> | 3 | 10 | 13 | 22% |
| <i>Enterobacter spp.</i> | - | 6 | 6 | 10% |
| <i>Klebsiella pneumoniae</i> | 3 | - | 3 | 5% |
| <i>Proteus vulgaris</i> | - | 3 | 3 | 5% |

Out of 96 positive samples by nitrite test, 28 were also positive by culture with sensitivity, specificity, positive predictive value (PPV) and negative predictive value (NPV), 46.67%, 92.77%, 29.1% and 96.5%, respectively. As regards the catalase test, out of 116 positive samples, 19 of them were also positive by culture with sensitivity, specificity, PPV and NPV, 31.67%, 89% 16.4% and 95.4%, respectively. Out of 90 leukocyte esterase

positive samples only 15 were positive by culture with sensitivity, specificity, PPV and NPV, 25%, 92%, 16.7% and 95% respectively. While for the combined results of nitrite, leukocyte esterase and/ or catalase out of 203 positive samples 44 samples were also positive by culture with sensitivity, specificity, PPV and NPV, 73.33%, 83.16%, 21.7%, and 98%, respectively (Table 4).

(Table 4): Results of screening tests compared to culture.

| Test | | Culture | | Sensitivity | Specificity | PPV | NPV |
|---|------------|-----------|----------|-------------|-------------|-------|-------|
| | | (-ve) 940 | (+ve) 60 | | | | |
| Nitrate test | (-ve) 904 | 872 | 32 | 46.67% | 92.77% | 29.1% | 96.5% |
| | (+ve) 96 | 68 | 28 | | | | |
| Catalase test | (-ve) 884 | 843 | 41 | 31.67% | 89% | 16.4% | 95.4% |
| | (+ve) 116 | 97 | 19 | | | | |
| Leukocyte esterase | (-ve) 910 | 865 | 45 | 25% | 92% | 16.7% | 95% |
| | (+ve) 90 | 75 | 15 | | | | |
| Nitrite, Leukocyte esterase and/or Catalase | (-ve) 797 | 781 | 16 | 73.33% | 83.16% | 21.7% | 98% |
| | (+ve) 203 | 159 | 44 | | | | |

Discussion

The prevalence of asymptomatic bacteriuria in school-going children was 6%. Consistent with that obtained by El Gamal and Saleh² Where the prevalence of asymptomatic bacteriuria in Egyptian school children was 7% and that reported as 7.3% by Jombo et al.¹⁷

This was much less than the prevalence detected by Kondapaneni et al.³ which was 16.5% and was also inconsistent with the studies carried by Jalali et al.¹⁸, Al-Momani,¹⁹ and Jha and Singh¹ who reported the prevalence of bacteriuria to be 3.36%, 1.39% and 1.2%, respectively. Such differences may be attributed to different socioeconomic status and geographical areas.

In this study, there is predominance of asymptomatic bacteriuria in girls 11.4% compared to 1.6% in boys. This comes in accordance to the studies carried by Kumar et al.²⁰ and Jombo et al.¹⁷ who reported predominance 7.6% in females to 2.9% in males and 7.9% compared to 6.6% in males, respectively. However, such results disagree with Bakr et al.²¹ who reported that there was no difference according to sex.

E.coli constituted 58% of the total bacterial isolates consistent with the previously obtained results by Jalali et al.^{18,15}, Al-Momani¹⁹ and Jha and Singh¹ who reported *E.coli* isolates to be 50%, 57.16% and 72% respectively. However, it is more than the results obtained by Kumar et al.²¹ and Kondapaneni, et al.³ who reported *E.coli* isolate to be 32.8% and 27.27%, respectively.

Noteworthy, *E.coli* is the most common pathogen causing UTI. *E.coli* has different virulence factors like K antigen and capsular polysaccharides which resist immune factors and antibiotics and predisposes to chronic urinary tract infection.²²

S. aureus constituted 22% of total bacterial isolates, this comes in accordance to the results obtained by Kondapaneni, et al.³ and El Gamal and Saleh² who reported *S. aureus* isolate to be 15% and 30% respectively.

The nitrite test was found to have sensitivity of 46.67% and specificity of 92.77%. The reliability of the nitrite test for urinary tract infections has been investigated widely and it was concluded that false positive results were rare and that the test had a higher specificity for urinary tract infections about 98% The sensitivity of the test has been reported to be about 50%.^{9,23,24}

The leukocyte esterase test has sensitivity 25% and specificity 92% which is different from previous studies that showed higher sensitivity about 84% and lower specificity of 78%.^{9,23,24}

In our study all the screening tests alone or combined showed high negative predictive value of more than 95% , low positive predictive value (16.4-29.1%), high specificity (83.16-92.77%) and lower sensitivity (25-73.33%).

In a study of 6394 febrile infants aged 1-90 days, Glissmeyer and colleagues found evidence that urine dipstick testing alone may provide an adequate initial UTI screen, comparing well in terms of positive predictive value and specificity with urine microscopy alone or both tests combined.^{8,25} Although the different screens each had a negative predictive value of more than 98%, the dipstick screen had a higher positive predictive value (66.8%) than did the combined test (51.2%) or microscopy alone (58.6%).^{8,25} The dipstick test by itself also had a higher specificity (93.8%) compared with the combined test (87.6%) or microscopy (91.3%), although it did have a lower sensitivity than the combined test (90.8% vs. 94.7%, respectively).^{8,25}

Limitations

The studied group were selected from public schools while private schools were not included in the study, we didn't investigate for risk factors of urinary tract infection (for example: anatomical anomalies and vesicoureteral reflux), another limitation for this study is that we did our best to elaborate to children how to take mid stream urine sample by verbal explanation and video demonstration but we didn't assure this by observing them while collecting the sample.

Conclusion

Asymptomatic bacteriuria could be detected by urine screening program at school age. Screening for asymptomatic bacteriuria in school-going children helps to know the prevalence. The overall prevalence of significant bacteriuria was 6%, with predominance in girls (11.4%) than boys (1.6%), *Escherichia coli* was the most common isolated organism in our study (58%) cases.

Competing interests

None.

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