

## Prevalence of urinary tract infection in acutely unwell children in general practice:

a prospective study with systematic urine sampling

### Abstract

#### Background

Urinary tract infection (UTI) in children may be associated with long-term complications that could be prevented by prompt treatment.

#### Aim

To determine the prevalence of UTI in acutely ill children  $\leq 5$  years presenting in general practice and to explore patterns of presenting symptoms and urine sampling strategies.

#### Design and setting

Prospective observational study with systematic urine sampling, in general practices in Wales, UK.

#### Method

In total, 1003 children were recruited from 13 general practices between March 2008 and July 2010. The prevalence of UTI was determined and multivariable analysis performed to determine the probability of UTI.

#### Results

Out of 597 (60.0%) children who provided urine samples within 2 days, the prevalence of UTI was 5.9% [95% confidence interval (CI) = 4.3% to 8.0%] overall, 7.3% in those  $< 3$  years and 3.2% in 3–5 year olds. Neither a history of fever nor the absence of an alternative source of infection was associated with UTI ( $P = 0.64$ ;  $P = 0.69$ , respectively). The probability of UTI in children aged  $\geq 3$  years without increased urinary frequency or dysuria was 2%. The probability of UTI was  $\geq 5\%$  in all other groups. Urine sampling based purely on GP suspicion would have missed 80% of UTIs, while a sampling strategy based on current guidelines would have missed 50%.

#### Conclusion

Approximately 6% of acutely unwell children presenting to UK general practice met the criteria for a laboratory diagnosis of UTI. This higher than previously recognised prior probability of UTI warrants raised awareness of the condition and suggests clinicians should lower their threshold for urine sampling in young children. The absence of fever or presence of an alternative source of infection, as emphasised in current guidelines, may not rule out UTI in young children with adequate certainty.

#### Keywords

children; general practice; prevalence; symptoms; urinary tract infections.

### INTRODUCTION

The prevalence of urinary tract infection (UTI) in acutely ill children in general practice is unknown. UTI in young children is difficult to diagnose and many cases are probably missed.<sup>1–3</sup> The challenge is that young children with UTI often present with non-specific symptoms that are also present in non-specific illness and in many other common conditions. Clinicians may therefore not consider the diagnosis or obtain a urine sample.

Childhood UTI has been associated with renal scarring and serious long-term complications, including hypertension, pre-eclampsia, and renal failure.<sup>4–7</sup> A systematic review found renal scarring was present in approximately 15% of children following a first UTI.<sup>8</sup> It remains unclear exactly what causes renal scarring to develop in some children, or which groups of children are most at risk.<sup>8</sup> There is some evidence that even children without fever or those with a self-limiting UTI may nevertheless be at risk of renal scarring.<sup>9</sup>

Guidelines highlight the importance of prompt diagnosis and early treatment of UTI in children, to prevent renal scarring.<sup>2</sup> However, urine is infrequently sampled from young children in primary care.<sup>10</sup> GPs indicated that they would normally sample urine from only a small proportion of children presenting with acute illness, even when awareness of UTI and the non-specific presenting symptoms had been raised.<sup>11</sup> The UK National Institute for Health and Clinical Excellence (NICE)

guideline promotes increased urine sampling.<sup>2</sup> However, clinicians may not respond to this recommendation unless there is evidence that the prevalence of UTI in children warrants increased testing.

Studies reporting the incidence and prevalence of UTI in children have varied by population, sampling method, and diagnostic criteria. Rates therefore vary widely, from 0.25% in a small UK GP study<sup>12</sup> to 13.5% in a hospital-based study of febrile infants.<sup>13</sup> A meta-analysis of the prevalence of UTI included 18 studies and a total of 22 919 children.<sup>14</sup> It found a pooled prevalence of 7% in febrile children  $< 24$  months old, and 7.8% in children  $< 19$  years old with urinary symptoms.<sup>14</sup> The applicability of these findings to general practice is limited because most studies were based in emergency departments with narrow inclusion criteria and age range, many excluded children without fever, and many sampled urine using invasive methods unsuitable for general practice. For children over 12 months old, all but one of the studies relied on urine samples obtained only if the treating clinician suspected a UTI. One study systematically obtained urine samples, but excluded boys older than 12 months and girls without a fever of  $\geq 38.5^{\circ}\text{C}$ .<sup>15</sup>

Recent UK guidelines state that UTI is unlikely if there are symptoms or signs suggestive of an alternative diagnosis, but there is evidence that UTI cannot be excluded on this basis.<sup>11,15,16</sup> It is also unclear how important fever is as a diagnostic sign,

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## How this fits in

Urinary tract infection (UTI) in children may be associated with long-term complications that could be prevented by prompt treatment. The current strategy of suspicion-led urine sampling is likely to miss the majority of cases of UTI in young children. The absence of fever or presence of an alternative source of infection does not necessarily rule out the possibility of UTI. The probability of UTI in children under the age of 3 years is reasonably high, irrespective of the presenting symptoms and signs. Larger studies are needed but a lower threshold for urine sampling in young children appears to be indicated.

as most prevalence studies exclude afebrile children.<sup>14</sup> Data on the pretest probability of UTI in an acutely unwell child presenting in primary care would help clinicians manage these children. Systematically sampling urine from sequential, unselected children presenting in primary care with a wide range of non-specific symptoms is required to determine the prevalence of UTI. Potential risk factors and subsets of children can then be analysed to determine whether urine sampling can be more effectively targeted to those children most likely to have UTI.

The aims of this study were to identify the prevalence of UTI in acutely ill children under the age of 5 years presenting in UK primary care, and to explore patterns of presentations in terms of symptoms, signs, and previous history, in relation to clinical suspicion and clinical guideline-based sampling practice.

## METHOD

### General practices and patients

Following a pilot study of feasibility, general practices in South Wales were invited to participate in this study.<sup>11</sup> Thirteen practices and five NHS microbiology laboratories across Wales participated between March 2008 and July 2010. Not all practices joined the study at the same time, so the duration of patient recruitment varied by practice, and not all clinicians in participating practices recruited. Children were eligible if they were aged <5 years and had an acute illness of <28 days' duration. Children were excluded if they were on immunosuppressant treatment (chemotherapy or oral/intramuscular steroids for  $\geq 2$  weeks) or long-term antibiotic treatment (>28 days), or had previously taken part in the study.

### Data collection

Carers of acutely unwell children presenting at participating general practices were provided with study information, and given an opportunity to discuss the study. Nurses or clinical studies officers recorded the details of presenting symptoms, medical and family history, temperature, pulse, and respiratory rate, and these were recorded on a case report form. Examination findings and a working diagnosis were recorded on the case report form by the responsible clinician. A urine sample was sought from all children. Clean catch was the preferred method but if this was not feasible, a nappy pad was used as recommended by current guidelines.<sup>2</sup> If a urine sample was not obtained at the general practice, carers were given urine-sampling equipment to take home and asked to return the sample as soon as possible. The urine was sent to the routine NHS receiving laboratory and results sent to the responsible GP practice, following routine processes. Clinicians managed the child's illness according to their standards of usual care. Laboratories also sent a copy of the urine result to the research team.

All urine samples were sent to NHS laboratories using the standard sample containers recommended by that laboratory. Three of the five laboratories required urine samples to be collected in containers containing boric acid (one of these laboratories provided special paediatric bottles containing boric acid). Practices stored urine samples in their fridges overnight if they received them after the daily specimen collection from the practice (usually midday). Laboratories processed and cultured the samples according to their standard operating procedures. There were some differences in standard operating procedures between laboratories. For example, one laboratory did not routinely culture urine that was negative on microscopy.

### Definition of urinary tract infection

A positive culture was defined as pure or predominant bacterial growth of  $>10^5$  colony-forming units (cfu)/ml on culture.<sup>2</sup> All other results were considered negative for the main analyses. Additional sensitivity analyses defined a borderline culture as  $10^4$ – $10^5$  cfu/ml of a single organism (in laboratories that recorded growth at this level), or  $>10^5$  cfu/ml of two organisms. Urines with heavy mixed growths ( $>10^5$  cfu/ml of  $>2$  organisms) were considered contaminated. Only those urine samples received within 2 days of the initial consultation were included in the analysis.

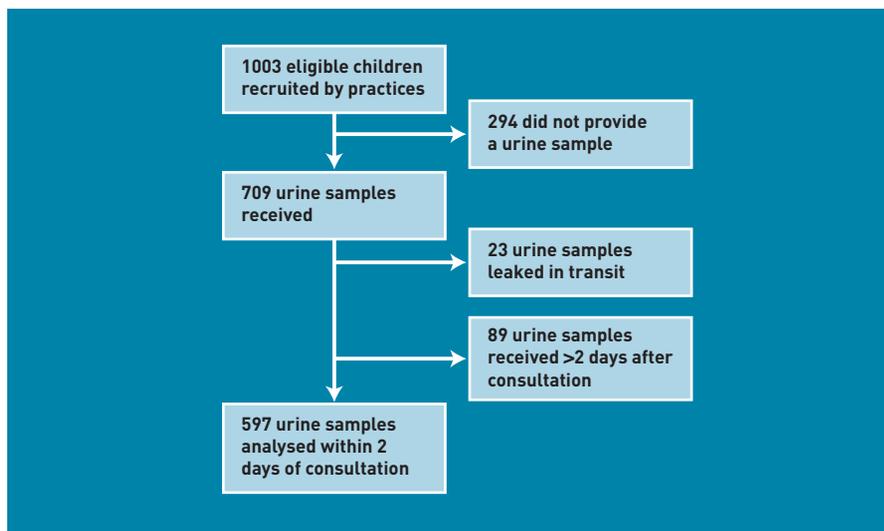


Figure 1. Numbers of participants recruited and urine samples collected.

#### Data entry

Data entry was double checked for 10% of all urine results and for 100% of those with positive and borderline cultures. Case report forms were collected from practices by research officers, and scanned using Cardiff Teleform (version 10.4.1). Data were combined, checked, anonymised, and analysed using SPSS (version 16).

#### Sample size requirement

To estimate the point prevalence of UTI in children under the age of 5 years presenting in general practice with an acute illness, with a 95% confidence interval (CI) of  $\pm 1\%$  around a predicted prevalence of 3%, a sample size of 1100 was needed.<sup>17</sup>

#### Analysis

The prevalence and 95% CIs were calculated using methods appropriate for proportions close to zero.<sup>17</sup>

$\chi^2$  and Fisher's exact tests were used to screen associations of symptoms, signs, and potential risk factors (for example, family history, previous medical history, age [categorised according to NICE <3 months,  $\geq 3$  months to <3 years,  $\geq 3$  years], sex, month of presentation, with the presence of UTI [Appendix 1]).<sup>2</sup>

All symptoms and signs with a *P*-value of <0.1 on univariate analysis were entered into a multivariable analysis, using forward stepwise logistic regression. All figures are presented to one decimal place except for *P*-values, which are presented to two decimal places.

#### Recruitment bias assessment

Clinicians were asked to keep 'recruitment logs' for eligible children who consulted, and to record the numbers of children invited and those consenting to participate. A check was made as to whether age and sex profiles were consistent with the practice population.

#### RESULTS

The recruitment rate varied between practices, from 1.5 per month to 11.7 per month. Four practices were not able to complete recruitment logs. The nine practices that did complete recruitment logs listed 63 eligible children who were approached but not recruited. The median age of non-recruited children was 1.6 years (interquartile range [IQR] = 1.0 to 3.1), and 34 (53.9%) were male.

Out of 1003 eligible children recruited (Figure 1), urine samples were obtained in 709 (70.7%). However, samples leaked in 23 (3.2%) cases and these were not analysed. Two hundred and ninety-four (41.5%) urine samples were received by laboratories on the day of recruitment. Only those urine samples received within 2 days of recruitment were used in this analysis (*n* = 597).

Five hundred and four 504 (50.2%) of the recruited children were male (Table 1) and their median age was 2.3 years (IQR = 1.00 to 3.49 years). Table 1 shows the

Table 1. Participant characteristics for the 597 children included in the analysis and the 406 not included

Characteristic	Those included in full analysis ( <i>n</i> = 597)	Those not included in full analysis ( <i>n</i> = 406)	<i>P</i> -value
<b>Age</b>			
Median age, years (IQR)	2.3 (1.0–3.5)	1.6 (0.8–3.3)	<0.01
<3 months, <i>n</i> (%)	32 (5.4)	28 (6.9)	
$\geq 3$ months and <3 years, <i>n</i> (%)	349 (58.5)	289 (71.2)	
$\geq 3$ years, <i>n</i> (%)	216 (36.2)	89 (21.9)	
<b>Sex, <i>n</i> (%)</b>			
Male	313 (52.4)	191 (47.0)	0.94
Female	284 (47.6)	215 (53.0)	
<b>GP working diagnosis, <i>n</i> (%)</b>			
Upper respiratory tract infection	177 (29.6)	121 (29.8)	0.35
Viral illness	90 (15.1)	55 (13.5)	
Lower respiratory tract infection	48 (8.0)	43 (10.6)	
UTI	41 (6.9)	13 (3.2)	
Tonsillitis	32 (5.4)	26 (6.4)	
Otitis	32 (5.4)	19 (4.7)	
Gastroenteritis	26 (4.4)	18 (4.4)	
Conjunctivitis	15 (2.5)	12 (3.0)	
Other	100 (16.8)	67 (16.5)	
No diagnosis	36 (6.0)	32 (7.9)	
<b>Common presenting symptoms, <i>n</i> (%)</b>			
Runny nose	423 (70.9)	301 (74.1)	0.20
Cough	413 (69.2)	280 (69.0)	0.94
Clingy	401 (67.2)	272 (67.0)	0.95
Irritable	383 (64.2)	275 (67.7)	0.24
Hot/feverish	355 (59.5)	232 (57.1)	0.46
Poor feeding	329 (55.1)	223 (54.9)	0.95

IQR = interquartile range.

**Table 2. Culture results for 597 children aged <5 years presenting to primary care with an acute illness**

Culture result	n (%)	% (95% CI)
<b>Positive</b>		
>10 <sup>5</sup> cfu/ml single organism	35 (5.9)	5.9 (4.3 to 8.0)
<b>Borderline</b>		
10 <sup>4</sup> –10 <sup>5</sup> cfu/ml single organism	11 (1.8)	2.8 (1.8 to 4.5)
>10 <sup>5</sup> cfu/ml two organisms	6 (1.0)	
<b>Negative</b>		
Heavy mixed growth >10 <sup>5</sup> cfu/ml	208 (34.8)	91.3 (88.8 to 93.3)
Mixed growth 10 <sup>4</sup> –10 <sup>5</sup> cfu/ml	81 (13.6)	
No growth or growth <10 <sup>4</sup> cfu/ml	216 (36.2)	
Not cultured as microscopy negative	40 (6.7)	

**Table 3. Prevalence of urinary tract infection by age**

Age range (NICE)	Proportion with UTI	% with UTI	95% CI
<3 months	4/32	12.5	5.0 to 28.1
≥3 months to <3 years	24/349	6.9	4.7 to 10.0
≥3 years	7/216	3.2	1.6 to 6.5

NICE = National Institute of Health and Clinical Excellence.

**Table 4. Presenting symptoms and signs in children entered into the logistic regression model**

Symptom	Proportion (%) of those with UTI with symptom	Proportion (%) of those without UTI with symptom	Odds ratio	P-value univariate analysis
Increased urinary frequency	11/35 (31.4)	75/562 (13.3)	3.0	<0.01
Wetting when previously dry	5/35 (14.3)	32/562 (5.7)	2.8	0.06
Pain/crying when passing urine	5/35 (14.3)	26/562 (4.6)	3.4	0.03
Irritable/grouchy	28/35 (80)	355/562 (63.2)	2.3	0.04
Temperature measured in surgery ≥38°C	15/35 (42.9)	163/562 (29.0)	1.8	0.08
Muscle aches or pains	0/35 (0)	55/562 (9.8)	0.1	0.03
Poor feeding/off food	24/35 (68.6)	305/562 (54.3)	1.8	0.10

**Table 5. Multivariable analysis: variables included in the model**

Symptom/characteristic	Odds ratio	95% CI	P-value
Urinary frequency	2.6	1.2 to 5.7	0.02
Pain on passing urine	3.3	1.1 to 9.8	0.03
NICE age range ≥3 years	1.0		Reference range
NICE age range ≥3 months to <3 years	2.4	1.0 to 5.8	0.06
NICE age range <3 months	5.5	1.5 to 21.0	0.01

P-value for model over the null model <0.01. NICE = National Institute of Health and Clinical Excellence.

characteristics of 597 children included in the analysis.

There was no significant difference in sex ( $P=0.94$ ), but there was a difference in age between children included in the main

analysis (those providing a urine sample within 2 days;  $n=597$ ) and those who were not included. Children who did not provide a urine sample within 2 days were approximately 8 months younger (median age 1.6 years) than those who did (median age 2.3 years). There was no statistical difference in GP diagnosis between those included and those not included in the main analysis. However, there was a greater proportion of those suspected of having a UTI among those included in the analysis, compared to those who were not.

In 431 (72.2%) cases the method of urine sampling used was indicated. Nappy pads were used in the majority of children less than 3 years old (100% aged <3 months; 74.3% aged ≥3 months to <3 years). Clean-catch collection was used in all children who were ≥3 years old. No difference was found in the prevalence of UTI according to urine-collection method when considering all children ( $P=0.15$ ), and there was no difference when considering the age range ≥3 months to <3 years alone (UTI prevalence 6.6% in clean catch and 5.1% in nappy pad;  $P=0.44$ ).

Urine samples were obtained before the child left the surgery for 318 (53.2%) children included in the analysis. It was much less likely that the urine sample was received within 2 days of the consultation if the sample was not obtained before leaving the surgery ( $P<0.01$ ). Antibiotics were prescribed in 31.1% (99/318) of children who provided urine samples prior to leaving the surgery and in 25.1% (70/279) of those who did not.

#### Prevalence of urinary tract infection

The prevalence of UTI, defined as the growth of one organism >10<sup>5</sup> cfu/ml was 5.9% overall (Table 2). A further 2.9% had a borderline culture result. Almost half of the samples (48.4%) had mixed growths, presumed to be contaminants, and regarded as negative. Heavy mixed growths were more common in nappy-pad samples (61.7%) compared with clean-catch samples (13.2%;  $P>0.01$ ). Forty (6.7%) were not cultured, owing to negative microscopy results, and so were classified as negative.

#### Risk factors and presenting symptoms and signs

Nineteen (6.7%) females and 16 (5.1%) males had UTI; this was not significantly different ( $P=0.41$ ). There was a trend towards a higher prevalence of UTI in the younger children ( $P=0.05$ ; Table 3).

Age category, symptoms, or signs that were associated with UTI with a  $P$ -value of

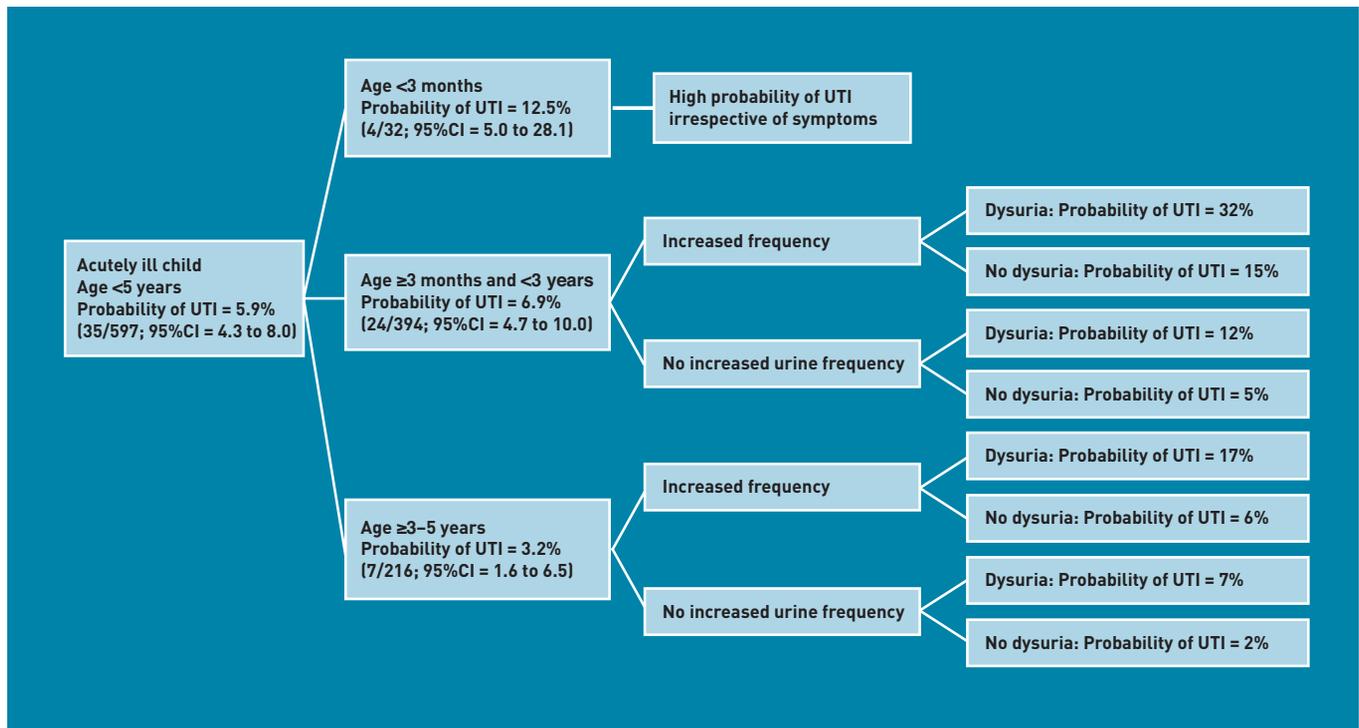


Figure 2. Probabilities of urinary tract infection based on the study model.

<0.1 on univariate analysis were entered into a logistic regression model (Table 4). Features not significantly associated with UTI included: history of being hot or feverish ( $P=0.69$ ), abdominal pain or vomiting, or a family history of UTI or kidney problems. An alternative site of infection [upper respiratory tract infection [URTI], lower respiratory tract infection [LRTI], tonsillitis, gastroenteritis, conjunctivitis, otitis] diagnosed by clinicians was not significantly associated with ruling out UTI ( $P=0.64$ ). Those with a UTI were less likely to have

a history of asthma ( $P=0.06$ ), and more likely to have a history of kidney or bladder disease ( $P=0.06$ ) than those without.

Multivariable analysis identified age range, pain or crying on passing urine, and increased urinary frequency or frequency of wet nappies as being associated with UTI (Table 5).

Based on the probability of UTI using the study model (Figure 2), an effective sampling strategy may therefore be to sample urine on all acutely ill children under the age of 5 years, except for those who are 3 years or older and have neither increased urinary frequency nor pain/crying on passing urine (Figure 2). Table 6 compares outcomes using the proposed sampling strategy (model) with either sampling based on GP suspicion of UTI or if current NICE guidelines had been followed, for the sample population. GP suspicion of UTI was based on the 'working diagnosis' question on the case report form, which was completed by GPs at the initial consultation and before urine culture results were available.

## DISCUSSION

### Summary

A 5.9% (95% CI = 4.3% to 8.0%) prevalence of UTI was found among systematically sampled acutely ill children under the age of 5 years presenting in primary care. This was higher in those under the age of 3 years (7.3%). A multivariable logistic

Table 6. Predicted outcomes of sampling based on GP suspicion, NICE guidance, and the proposed sampling strategy

Age group	Urine sample	GP suspicion, n	If NICE guidelines had been applied, n	Proposed sampling strategy, n
<3 months (n=32)	Urine samples	0	9	32
	UTIs diagnosed	0	1	4
	UTIs missed	4	3	0
≥3 months to <3 years (n=349)	Urine samples	19	150	349
	UTIs diagnosed	3	10	24
	UTIs missed	21	14	0
≥3 years (n=216)	Urine samples	22	77	33
	UTIs diagnosed	4	6	6
	UTIs missed	3	1	1
Total	Urine samples	41 (7%)	236 (40%)	414 (69%)
	UTIs diagnosed	7 (20%)	17 (49%)	34 (97%)
	UTIs missed	28 (80%)	18 (51%)	1 (3%)

NICE = National Institute of Health and Clinical Excellence.

regression model that included age band, increased urinary frequency, and pain or crying when passing urine found that the probability of UTI in children aged 3 years or older without increased urinary frequency or pain when passing urine was 2%. The probability of UTI was higher in all other groups of children ( $\geq 5\%$ ).

### Strengths and limitations

This was a prospective primary care-based prevalence study of UTI, in which acutely unwell children were well described clinically and had their urine systematically sampled. Urine samples were all analysed by the NHS laboratory local to the participating GP practice, keeping transport of samples and reporting of results consistent with normal practice.

The study did not recruit the target sample of 1100. Confidence intervals for the main estimate were wider ( $\pm 2\%$ ) than had been hoped (1% around a prevalence of 3%). However, the prevalence of UTI was higher in the study sample than the 3% used for the sample size calculation, which mitigated loss of precision. The study was not powered to accurately determine the predictive value of symptoms and signs, and this resulted in large confidence intervals for the odds ratios and probabilities in the multivariable model. Many potentially predictive variables were tested, which increased the likelihood that some would be statistically significant purely by chance.

Not all eligible children presenting in practices were recruited, and not all recruited children provided a urine sample. There was a difference in age between those who were included in the main analysis and those who were not, with younger children less likely to provide a urine sample within 2 days ( $P < 0.01$ ).

Nappy pads were commonly used for urine sampling. These were associated with an increased frequency of heavy mixed growth, implying contamination, compared to clean-catch specimens ( $P < 0.01$ ). Some urine results categorised as 'positive' may have been caused by contaminating bacteria (false positives) and some of those with heavy mixed growth and categorised as 'negative' may have been a true UTI.<sup>18</sup> It was found that UTI prevalence was higher in the clean-catch specimens, suggesting that false negatives may be more of a problem among nappy-pad specimens than false positives, but this difference was not statistically significant.

Antibiotic prescription in those children who did not provide a urine sample prior to leaving the surgery could potentially

have interfered with subsequent urine results. This most likely would have resulted in additional false negatives and an underestimation of UTI prevalence.

The screening logs showed that recruited children were older than non-recruited children. This probably reflects the increased difficulties of obtaining urine samples in younger children.<sup>19</sup> As there is a higher prevalence of UTI in younger children (in both this study and others), this may also indicate that the prevalence found in this study is an underestimation for the group overall.<sup>10</sup> Clinicians suspected UTI more often in children included in the main analysis compared with those who were not. This may also be due to the non-specific nature of symptoms in younger children, or it may reflect greater encouragement given to obtain urine samples if the clinician suspected a UTI.

### Comparison with existing literature

The 95% CI around the overall prevalence in this study (4.3% to 8.0%) excludes the prior estimate of 3% based on earlier studies.<sup>15,16</sup> The prevalence is slightly lower than the 7% pooled prevalence found in a systematic review and meta-analysis of studies that were largely based in secondary care with narrow inclusion criteria (for example, aged  $< 2$  years and fever  $> 38^{\circ}\text{C}$ ).<sup>14</sup> Both false-positive and false-negative urine results may have occurred. A large number of samples had heavy mixed growth on culture (classified as negative) and some of these may have masked a true UTI. Some of the positive cultures may represent 'asymptomatic bacteriuria'.<sup>20</sup> However, children were only eligible for the present study if they were acutely ill, and all met the laboratory diagnostic criteria for UTI as defined in UK NICE guidelines.<sup>2</sup> Some authors have suggested two consecutive urine samples should be obtained to determine UTI.<sup>21</sup> This would reduce the false-positive rate but could also increase the false-negative rate. Obtaining a single sample is already challenging in busy general practice. A requirement to obtain two consecutive samples for all children may reduce sampling in general practice. Both higher and lower cut-off values for bacterial counts have been suggested for achieving the ideal trade off between sensitivity and specificity, but for now  $> 10^5$  cfu/ml remains the standard for diagnosing UTI.<sup>2,18,21</sup>

The finding that fever was not associated with UTI within an acutely ill group of children is important, as most previous studies excluded children without fever. Coulthard *et al* also found that fever was

not present in all children with UTI and that fever or other clinical symptoms or signs could not be used to predict renal scarring.<sup>9</sup>

NICE guidelines recommend that a urine sample is not necessary at first presentation of an ill child if there is evidence of an alternative source of infection.<sup>2</sup> However, the presence or absence of an alternative source of infection was not associated with UTI in the present study. Other studies have also found that the presence of an alternative source of infection cannot reliably rule out UTI.<sup>11,15,16</sup>

#### Implications for research and practice

The findings of this study suggest that the probability of UTI in children under the age of 3 years is reasonably high, irrespective of the presenting symptoms and signs. There were no symptoms or signs that ruled in or ruled out UTI with adequate precision in this age group and setting. With a pre-test (pre-urine sample) probability of >5%, a urine sample appears justified in far more acutely unwell children under the age of 3 years presenting in primary care than previously thought or currently practised. Implementing such a recommendation would represent a considerable increase in urine sampling, with associated costs and inconvenience. It would also increase the possibility of false-positive results with associated further unnecessary, additional investigations. However, the current strategy of suspicion-led urine sampling is likely to miss the majority of cases of UTI, along with the opportunity to treat promptly, and hence minimise morbidity and possibly reduce the risk of future complications.

It was found that, in children older than 3 years, without urinary frequency or pain on passing urine, there was a low probability (2%) of UTI. It may be that routine urine sampling is not indicated in this group.

Using the proposed model would mean sampling urine in 10 times more children compared to sampling based on current practice/GP suspicion, and would mean sampling twice as many children as is currently recommended by NICE. However, only 3% of UTIs would be missed, compared with 51% when implementing NICE guidelines and 80% when sampling is based on GP suspicion alone.

The management of children has not been considered in this study, beyond whether or not a urine sample would need to be obtained to identify those at risk of UTI. A further study, powered to test the predictive value of symptoms, signs, and clinical history features, is needed to determine whether dipsticks or combinations

of symptoms and signs can be used to guide treatment without relying on urinary culture. It is not clear whether antibiotics can safely be delayed until the culture result is available. Increasing urine sampling may lead to increases in unnecessary antibiotic prescribing while waiting for culture results. Current guidelines advise that dipsticks are unreliable in children under 3 years of age, but may be useful in older children.<sup>2</sup> If clinicians wish to increase urine sampling in their own practices, it is advisable to obtain the appropriate equipment (large collection pots for clean catch, which can be put inside a potty, and nappy pads), and emphasise the importance of sampling irrespective of urinary symptoms, preferably by clean-catch specimens, and, wherever possible, obtaining the sample before the child leaves the surgery.

The proposed model should be validated in another large sample of children. The potential benefits of diagnosing more UTIs, but possibly not until culture results become available, need to be considered alongside the additional associated costs of increasing urine sampling. Currently, all children with UTI (acutely ill with bacteriuria on culture) are considered at risk of renal complications, but only a small proportion will develop them.<sup>2</sup> It remains unclear whether serious long-term sequelae will be prevented by increasing the diagnosis and treatment of UTI in children, and whether the required substantial increase in urine sampling would be cost effective.

A larger study that systematically samples acutely unwell children and follows them up over the longer term is needed to identify which children are most likely to have a UTI and which are at greatest risk for complications.<sup>2</sup>

It was found that almost 6% of acutely unwell children presenting to UK general practice met the criteria for a laboratory diagnosis of UTI. This prior probability is higher than previously recognised and should increase GPs' awareness of this condition. Current guidelines promote prompt diagnosis of UTI in children.<sup>2</sup> Based on the findings of this study, clinicians should have a lower threshold for urine sampling in acutely ill children, and consider sampling urine in all acutely unwell children less than 3 years old, irrespective of symptoms, and in those aged 3–5 years with either dysuria or urinary frequency. In contrast to the NICE guideline, the present study suggests that neither the absence of fever nor the presence of an alternative site of infection satisfactorily rules out UTI in young children.

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#### Ethical approval

Ethical approval for the study was obtained from the South East Wales Local Research Ethics committee (ref no. 08/WSE03/11). Informed, written consent was obtained from all participants.

#### Provenance

Freely submitted; externally peer reviewed.

#### Competing interests

The authors have declared no competing interests.

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## Appendix 1. Table of the symptoms, signs, and risk factors screened for association with urinary tract infection

Symptoms	<ul style="list-style-type: none"> <li>Runny/blocked nose</li> <li>Sore throat</li> <li>Earache or holding ear</li> <li>Cough</li> <li>Grunting or difficulty breathing</li> <li>Hot/feverish</li> <li>Rash</li> <li>Irritable/grouchy</li> <li>Clinginess/needing extra care</li> <li>Low energy/tired/lost interest in playing</li> <li>Poor sleep</li> <li>Muscle aches or pains</li> <li>Poor feeding/poor appetite</li> <li>Diarrhoea</li> <li>Constipation</li> <li>Vomiting</li> <li>Nausea</li> <li>Abdominal pain/tummy ache</li> <li>Colic/grimacing/pulling up legs</li> <li>Bed wetting/day wetting when previously dry</li> <li>Smelly urine</li> <li>Dark or cloudy urine</li> <li>Pain or crying on passing urine</li> <li>Increased urinary frequency or number of wet nappies</li> <li>Poor weight gain/weight loss</li> <li>Highest recorded temperature by parents</li> <li>How ill (0–4) do parents feel their child is</li> </ul>
Signs	<ul style="list-style-type: none"> <li>Pulse rate</li> <li>Respiratory rate</li> <li>Temperature measured in surgery</li> <li>Ear examination</li> <li>Throat examination</li> <li>Chest examination</li> <li>Abdomen examination</li> <li>General examination: dehydration</li> <li>General examination: rash</li> <li>Fontanelles</li> <li>Overall impression of how ill child is (0–4)</li> <li>Presence of alternative source of infection (from stated working diagnosis)</li> </ul>
Other potential risk factors	<ul style="list-style-type: none"> <li>Age</li> <li>Sex</li> <li>Month of presentation</li> <li>Townsend score</li> <li>Past history of urinary tract infection</li> <li>Past history of kidney/bladder disease</li> <li>Past history of asthma</li> <li>Past history of diabetes</li> <li>Past history of eczema</li> <li>Has child been circumcised</li> <li>Abnormalities of child's urinary tract on antenatal ultrasound scan</li> <li>Family history of urinary tract infection</li> <li>Family history of kidney or bladder disease</li> </ul>