

Appendices

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Clinical effectiveness and cost-effectiveness of tests for the diagnosis and investigation of urinary tract infection in children: a systematic review and economic model

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**Health Technology Assessment
NHS R&D HTA Programme**





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Appendix I

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Appendix 2

Detailed search strategies

This appendix presents the detailed searches carried out to inform the review.

Chosen strategy: strategy illustrated will run in the Ovid interfaces of MEDLINE

- 1 exp urinary tract infections/
- 2 bacterial infections/ or exp pseudomonas infections/ or exp klebsiella infections/ or gram negative infections/ or exp escherichia coli/ or exp proteus/ or exp enterococcus/
- 3 exp Staphylococcus/
- 4 exp leukocytes/
- 5 (microbial infection? or bacterial infection?).ti,ab.
- 6 (urinary or urine or urethra or bladder or ureter? or kidney or kidneys or renal).ti,ab.
- 7 exp urinary tract/
- 8 or/2-5
- 9 or/6-7
- 10 8 and 9
- 11 1 or 10
- 12 exp child, preschool/ or exp infant/
- 13 (infant? or baby or babies or toddler? or preschooler?).ti,ab.
- 14 or/12-13
- 15 11 and 14
- 16 (risk assessment? or exam or examination or feeding or slow weight gain or fever or vomiting or diarrh?).ti,ab.
- 17 (((sepsis or failure) adj2 thrive) or malaise or frequent urination or abdominal discomfort or abdominal pain).ti,ab.
- 18 (delayed bladder control or dysuria or (pain adj3 urination) or painful urination or difficult urination).ti,ab.
- 19 (urinalysis or urine analysis or urine sample? or urine specimen? or (urine adj3 collect?)).ti,ab.
- 20 (urine bags or dipstick? or dip stick? or urine microscopy).ti,ab.
- 21 (reagent strip? or colorimetric test? or gas analysis or impedance or luminescence).ti,ab.
- 22 (immunological test? or elisa or enzyme test? or bacterial oxygen consumption or turbidimetry or urine culture).ti,ab.
- 23 (bacterial culture or dipslide? or renal ultrasonography or planar imaging or radiography or urography or pyelography or kub or bladder imaging).ti,ab.
- 24 (cystography or cystourethrography or nuclear medicine or scintigraphy or cystogram?).ti,ab.
- 25 exp physical examination/ or exp fever/ or exp body weight changes/ or exp abdominal pain/ or exp urological manifestations/ or failure to thrive/
- 26 exp vomiting/ or diarrhea/ or exp sepsis/ or urinalysis/
- 27 exp microscopy/ or exp "indicators and reagents"/
- 28 colorimetry/ or electric impedance/ or exp immunoassay/ or exp fluorescent antibody technique/
- 29 exp diagnostic imaging/
- 30 exp nuclear medicine/ or exp cystoscopy/ or exp diagnostic techniques, urological/
- 31 or/16-30
- 32 15 and 31
- 33 vesico-ureteral reflux/ or pyelonephritis/ or bacteriuria/ or cystitis/
- 34 (failure adj2 thrive).ti,ab.
- 35 sepsis.tw.
- 36 ultrasonography.ti,ab.
- 37 exp succimer/ or exp organometallic compounds/ or technetium/ or exp sulfhydryl compounds/ or exp culture media/
- 38 urinary catheterization/ or ammonium chloride/ or c-reactive protein/ or urodynamics/ or urine/mi
- 39 (dmsa or urogram? or ultrasound? or (renal adj scan?)).ti,ab.
- 40 (spect or (planar adj image?) or (dip adj slide?) or cystoscopy).ti,ab.
- 41 ((bladder adj aspiration) or (acidification adj test?) or (cortical adj echogenicity)).ti,ab.
- 42 workup.ti,ab.
- 43 (radiographic or cystomanometry).ti,ab.
- 44 (bladder adj3 (investigat? or detect?)).ti,ab.
- 45 (kidney adj3 (investigat? or detect?)).ti,ab.
- 46 (urethra adj3 (investigat? or detect?)).ti,ab.
- 47 (renal adj3 (investigat? or detect?)).ti,ab.
- 48 (kidneys adj3 (investigat? or detect?)).ti,ab.
- 49 (urinary adj3 (investigat? or detect?)).ti,ab.
- 50 (infection? adj3 (urinary or urine or urethra or bladder or ureter? or kidney or kidneys or renal)).ti,ab.
- 51 (2 or 3 or 4 or 33) and 7
- 52 1 or 50 or 51

- 53 52 and 14
- 54 or/34-49
- 55 53 and 54
- 56 55 not 32

MEDLINE

First draft of search

(1966 to October week 5 2002)

- 1 exp urinary tract infections/ (27032)
- 2 bacterial infections/ or exp pseudomonas infections/ or exp klebsiella infections/ or gram negative infections/ or exp escherichia coli/ or exp proteus/ or exp enterococcus/ (217644)
- 3 exp Staphylococcus/ (41409)
- 4 exp leukocytes/ (398776)
- 5 (microbial infection? or bacterial infection?).ti,ab. (11874)
- 6 (urinary or urine or urethra or bladder or ureter? or kidney or kidneys or renal).ti,ab. (553654)
- 7 exp urinary tract/ (251201)
- 8 or/2-5 (645127)
- 9 or/6-7 (633796)
- 10 8 and 9 (27291)
- 11 1 or 10 (49809)
- 12 exp child, preschool/ or exp infant/ (827649)
- 13 (infant? or baby or babies or toddler? or preschooler?).ti,ab. (175142)
- 14 or/12-13 (857927)
- 15 11 and 14 (7594)
- 16 (risk assessment? or exam or examination or feeding or slow weight gain or fever or vomiting or diarrh?).ti,ab. (390002)
- 17 (((sepsis or failure) adj2 thrive) or malaise or frequent urination or abdominal discomfort or abdominal pain).ti,ab. (20335)
- 18 (delayed bladder control or dysuria or (pain adj3 urination) or painful urination or difficult urination).ti,ab. (1587)
- 19 (urinalysis or urine analysis or urine sample? or urine specimen? or (urine adj3 collect?).ti,ab. (17696)
- 20 (urine bags or dipstick? or dip stick? or urine microscopy).ti,ab. (1074)
- 21 (reagent strip? or colorimetric test? or gas analysis or impedance or luminescence).ti,ab. (16858)
- 22 (immunological test? or elisa or enzyme test? or bacterial oxygen consumption or turbidimetry or urine culture).ti,ab. (48330)
- 23 (bacterial culture or dipslide? or renal ultrasonography or planar imaging or radiography or urography or pyelography or kub or bladder imaging).ti,ab. (25490)

- 24 (cystography or cystourethrography or nuclear medicine or scintigraphy or cystogram?).ti,ab. (28553)
- 25 exp physical examination/ or exp fever/ or exp body weight changes/ or exp abdominal pain/ or exp urological manifestations/ or failure to thrive/ (369317)
- 26 exp vomiting/ or diarrhea/ or exp sepsis/ or urinalysis/ (88329)
- 27 exp microscopy/ or exp "indicators and reagents"/ (477710)
- 28 colorimetry/ or electric impedance/ or exp immunoassay/ or exp fluorescent antibody technique/ (320665)
- 29 exp diagnostic imaging/ (811099)
- 30 exp nuclear medicine/ or exp cystoscopy/ or exp diagnostic techniques, urological/ (68980)
- 31 or/16-30 (2116054)
- 32 15 and 31 (2893)

Second draft of search

(1966 to October week 5 2002). This draft excludes items already found. Results in fullmed4.txt and fullmed5.txt.

- 1 exp urinary tract infections/ (27032)
- 2 bacterial infections/ or exp pseudomonas infections/ or exp klebsiella infections/ or gram negative infections/ or exp escherichia coli/ or exp proteus/ or exp enterococcus/ (217644)
- 3 exp Staphylococcus/ (41409)
- 4 exp leukocytes/ (398776)
- 5 (microbial infection? or bacterial infection?).ti,ab. (11874)
- 6 (urinary or urine or urethra or bladder or ureter? or kidney or kidneys or renal).ti,ab. (553654)
- 7 exp urinary tract/ (251201)
- 8 or/2-5 (645127)
- 9 or/6-7 (633796)
- 10 8 and 9 (27291)
- 11 1 or 10 (49809)
- 12 exp child, preschool/ or exp infant/ (827649)
- 13 (infant? or baby or babies or toddler? or preschooler?).ti,ab. (175142)
- 14 or/12-13 (857927)
- 15 11 and 14 (7594)
- 16 (risk assessment? or exam or examination or feeding or slow weight gain or fever or vomiting or diarrh?).ti,ab. (390002)
- 17 (((sepsis or failure) adj2 thrive) or malaise or frequent urination or abdominal discomfort or abdominal pain).ti,ab. (20335)
- 18 (delayed bladder control or dysuria or (pain adj3 urination) or painful urination or difficult urination).ti,ab. (1587)

- 19 (urinalysis or urine analysis or urine sample? or urine specimen? or (urine adj3 collect?)).ti,ab. (17696)
- 20 (urine bags or dipstick? or dip stick? or urine microscopy).ti,ab. (1074)
- 21 (reagent strip? or colorimetric test? or gas analysis or impedance or luminescence).ti,ab. (16858)
- 22 (immunological test? or elisa or enzyme test? or bacterial oxygen consumption or turbidimetry or urine culture).ti,ab. (48330)
- 23 (bacterial culture or dipslide? or renal ultrasonography or planar imaging or radiography or urography or pyelography or kub or bladder imaging).ti,ab. (25490)
- 24 (cystography or cystourethrography or nuclear medicine or scintigraphy or cystogram?).ti,ab. (28553)
- 25 exp physical examination/ or exp fever/ or exp body weight changes/ or exp abdominal pain/ or exp urological manifestations/ or failure to thrive/ (369317)
- 26 exp vomiting/ or diarrhea/ or exp sepsis/ or urinalysis/ (88329)
- 27 exp microscopy/ or exp "indicators and reagents"/ (477710)
- 28 colorimetry/ or electric impedance/ or exp immunoassay/ or exp fluorescent antibody technique/ (320665)
- 29 exp diagnostic imaging/ (811099)
- 30 exp nuclear medicine/ or exp cystoscopy/ or exp diagnostic techniques, urological/ (68980)
- 31 or/16-30 (2116054)
- 32 15 and 31 (2893)
- 33 vesico-ureteral reflux/ or pyelonephritis/ or bacteriuria/ or cystitis/ (23125)
- 34 (failure adj2 thrive).ti,ab. (2130)
- 35 sepsis.tw. (28242)
- 36 ultrasonography.ti,ab. (30181)
- 37 exp succimer/ or exp organometallic compounds/ or technetium/ or exp sulfhydryl compounds/ or exp culture media/ (204118)
- 38 urinary catheterization/ or ammonium chloride/ or c-reactive protein/ or urodynamics/ or urine/mi (30758)
- 39 (dmsa or urogram? or ultrasound? or (renal adj scan?)).ti,ab. (63607)
- 40 (spect or (planar adj image?) or (dip adj slide?) or cystoscopy).ti,ab. (12053)
- 41 ((bladder adj aspiration) or (acidification adj test?) or (cortical adj echogenicity)).ti,ab. (149)
- 42 workup.ti,ab. (3809)
- 43 (radiographic or cystomanometry).ti,ab. (38227)
- 44 (bladder adj3 (investigat? or detect?)).ti,ab. (246)
- 45 (kidney adj3 (investigat? or detect?)).ti,ab. (242)
- 46 (urethra adj3 (investigat? or detect?)).ti,ab. (7)
- 47 (renal adj3 (investigat? or detect?)).ti,ab. (984)
- 48 (kidneys adj3 (investigat? or detect?)).ti,ab. (63)
- 49 (urinary adj3 (investigat? or detect?)).ti,ab. (479)
- 50 (infection? adj3 (urinary or urine or urethra or bladder or ureter? or kidney or kidneys or renal)).ti,ab. (22555)
- 51 (2 or 3 or 4 or 33) and 7 (14093)
- 52 1 or 50 or 51 (48512)
- 53 52 and 14 (9186)
- 54 or/34-49 (398248)
- 55 53 and 54 (1988)
- 56 55 not 32 (1121)

Third draft of search

(1966 to October week 5 2002). This draft excludes items already found. Results in fullmed4.txt and fullmed5.txt.

- 1 exp urinary tract infections/di or pyelonephritis/di or bacteriuria/di or cystitis/di (6612)
- 2 bacterial infections/ or exp pseudomonas infections/ or exp klebsiella infections/ or gram negative infections/ or exp escherichia coli/ or exp proteus/ or exp enterococcus/ or exp staphylococcus/ (250261)
- 3 exp urinary tract/ (251201)
- 4 2 and 3 (2911)
- 5 (infection? adj3 (urinary or urine or urethra or ureter? or bladder or kidney or kidneys or renal)).ti,ab. (22555)
- 6 urinary tract infections/ or pyelonephritis/ or bacteriuria/ or cystitis/ (36723)
- 7 or/4-6 (47969)
- 8 (risk assessment? or exam or examination or feeding or slow weight gain or fever or vomiting or diarrh\$).ti,ab. (421461)
- 9 (sepsis or (failure adj2 thrive) or malaise or frequent urination or abdominal discomfort or abdominal pain).ti,ab. (48312)
- 10 (delayed bladder control or dysuria or (pain adj2 urination) or painful urination) or difficult urination).ti,ab. (1578)
- 11 (urinalysis or urinalyses or urine analys\$ or urine sampl\$ or (urine adj2 specimen?) or (urine adj2 collect\$)).ti,ab. (23602)
- 12 (urine bag or urine bags or dip stick? or urine microscopy).ti,ab. (223)
- 13 (reagent strip? or colorimetric test\$ or gas analysis or impedance or luminescence).ti,ab. (16858)
- 14 (immunological test? or elisa or enzyme test? or bacterial oxygen consumption or turbidimetry or urine culture?).ti,ab. (49212)

- 15 (bacterial culture? or dipstick? or ultrasound or ultrasonography or dipstick? or planar imaging or radiography or urography or pyelography or kub or bladder imaging).ti,ab. (110841)
- 16 (cystography or cystourethrography or scintigraphy or cystogram?).ti,ab. (24072)
- 17 exp physical examination/ or exp fever/ or exp body weight changes/ or exp abdominal pain/ or exp urological manifestations/ or failure to thrive/ (369317)
- 18 exp vomiting/ or diarrhea/ or exp sepsis/ or urinalysis/ (88329)
- 19 exp microscopy/ or exp "indicators and reagents"/ (477710)
- 20 colorimetry/ or electric impedance/ or exp immunoassay/ or exp fluorescent antibody technique/ (320665)
- 21 exp diagnostic imaging/ or exp cystoscopy/ or exp diagnostic techniques, urological/ (857129)
- 22 exp succimer/ or exp organometallic compounds/ or technetium/ or exp sulfhydryl compounds/ or exp culture media/ (204118)
- 23 urinary catheterization/ or ammonium chloride/ or c-reactive protein/ or urodynamics/ or urine/mi (30758)
- 24 (dmsa or urogram? or renal scan\$ or spect or planar image? or dip slide? or cystoscopy).ti,ab. (14739)
- 25 (bladder aspiration or acidification test? or cortical echogenicity).ti,ab. (149)
- 26 (workup or radiographic or cystomanometry).ti,ab. (41941)
- 27 (bladder adj2 (investigat\$ or detect\$ or diagnos\$)).ti,ab. (2851)
- 28 (kidney adj2 (investigat\$ or detect\$ or diagnos\$)).ti,ab. (3829)
- 29 (urethra adj2 (investigat\$ or detect\$ or diagnos\$)).ti,ab. (186)
- 30 (renal adj2 (investigat\$ or detect\$ or diagnos\$)).ti,ab. (8746)
- 31 (kidneys adj2 (investigat\$ or detect\$ or diagnos\$)).ti,ab. (878)
- 32 (urinary adj2 (investigat\$ or detect\$ or diagnos\$)).ti,ab. (4439)
- 33 animal/ not (human/ and animal/) (2607560)
- 34 infant/ or child,preschool/ (630875)
- 35 adult/ not (adult/ and 34) (2177067)
- 36 exp neoplasms/ (1414496)
- 37 36 not (36 and 6) (1411481)
- 38 exp prostatic hyperplasia/ (10993)
- 39 exp pregnancy complications/ (202248)
- 40 exp pregnancy/ (482868)
- 41 exp diabetes/ (154314)
- 42 41 not (41 and 6) (153295)
- 43 exp multiple sclerosis/ (20918)
- 44 43 not (43 and 6) (20869)
- 45 exp spinal cord injuries/ (19647)
- 46 45 not (45 and 6) (19234)
- 47 exp prostate/ (15881)
- 48 exp aged/ (1257452)
- 49 48 not (48 and 34) (1180413)
- 50 middle age/ (1853948)
- 51 50 not (50 and 34) (1741644)
- 52 exp pneumonia,bacterial/ (7628)
- 53 52 not (52 and 6) (7534)
- 54 contraception/ or contraceptive devices/ or cesarean section/ (54227)
- 55 urinary tract infections/th (818)
- 56 bacterial infections/dt (13677)
- 57 antibiotics/tu (41019)
- 58 exp parasitic diseases/ (178865)
- 59 exp urologic surgical procedures,male/ (24061)
- 60 55 not (55 and 1) (513)
- 61 56 not (56 and 1) (13595)
- 62 57 not (57 and 1) (40585)
- 63 59 not (59 and 1) (24024)
- 64 or/8-32 (2350391)
- 65 7 and 64 (20524)
- 66 1 or 65 (23205)
- 67 66 not (33 or 35 or 37 or 38 or 39 or 40 or 42) (13699)
- 68 67 not (44 or 46 or 47 or 49 or 51 or 53) (11362)
- 69 68 not (54 or 60 or 61 or 62 or 58 or 63) (10052)
- 70 from 69 keep 1-1000 (1000)

This strategy seeks to find studies that are not explicitly (at least in terms of words in the title, abstract and subject indexing) about children. The yield is therefore large and has been reduced by careful limits such as removing records that focus on conditions and issues (such as contraception) that will not tend to be related to children. The search has low precision and after exploration the final strategy used with the other databases was based on MEDLINE draft 2.

PREMEDLINE

This was searched via Ovid on the web. The PREMEDLINE issue searched was dated 18 November 2002. Seven records were downloaded to PREMED.

The second MEDLINE strategy above was used: subject headings produced zero hits, but the search logic worked well. The final line was changed to read 55 or 32.

Biosis

Searched from 1985 to 11 December 2002, on the Edina service; 1225 hits downloaded in annual batches.

((((((((((((((al: (tomography or diagnosis or diagnosing or diagnostic+)) or (al: (spect or ct scan* or bladder aspiration or acidification or cortical echogenicity or workup* or cystomanometry or cat or computer assisted or tomograp))) or (al: (succimer or organometallic or technetium or sulfhydryl or catheteri* or ammonium chloride or c reactive protein or urodynamics or dmsa))) or (al: (cystogram+ or cystourethography or nuclear medicine or scintigraphy or cystoscopy))) or (al: (urography or urogram+ or pyelography or kub or bladder imag* or cystography))) or (al: (bacterial culture+ or ultrasonogra* or ultrasound or diagnostic imaging or echography or planar imag* or radiography))) or (al: (elisa or immunoassay+ or enzyme test+ or oxygen consumption or turbidimetry or urine culture+))) or (al: (colorimetric or colorimetry or gas analysis or impedence or luminescence or fluorescen* or immunological test*))) or (al: (dipstick+ or dip stick+ or dipslide+ or dip slide+ or urine microscopy or reagent+ or test strip+))) or (al: (dysuria or urinalysis or urinalyses or urine sample+ or urine specimen+ or urine collection or collecting urine or urine bag+))) or (al: (sepsis or thrive or malaise or frequent urination or polyuria or abdominal pain or abdominal discomfort))) or (al: (risk assessment+ or physical exam* or feeding or vomiting or diarrh*))) and (((al: (child or children or infant+ or baby or babies or toddler+ or pediatric or paediatric)) and (((al: (bladder or kidney+ or ureter or renal or urinary tract or cystitis or vesicoureteral reflux or pyelonephritis or bacteriuria)) and (al: (bacterial infection+ OR microbial infection+ or pseudomonas or klebsiella or gram negative infection+ or eschrichia coli or proteus or enterococcus or staphylococcus)))))) or (al: (urinary tract infection+))))))

Pascal

Pascal was searched using the Dialog search interface and for the period 1973 to January week 4 2003.

- 1 10003 (URINARY(W)TRACT(W) INFECTION? ?) OR PYELONEPHRITIS OR BACTERIURIA OR CYSTITIS
 2 15142 (URINARY OR URINE OR URETHRA OR URETER? ? OR BLADDER OR KIDNEY? ? OR RENAL)(4N)(BACTERIAL OR PSEUDOMONAS OR

- KLEBSIELLA OR GRAM(W)NEGATIVE OR ESCHERICHIA(W)COLI OR PROTEUS OR ENTEROCOCCUS OR STAPHYLOCOCCUS OR INFECTION OR INFECTIONS)
 3 19491 S1 OR S2
 4 260198 RISK(W)ASSESSMENT OR RISK(W)ASSESSMENTS OR EXAM OR EXAMINATION OR FEEDING OR SLOW(W)WEIGHT(W)GAIN OR FEVER OR VOMITING OR DIARRHEA OR DIARRHOEA
 5 17101 SEPSIS OR (FAILURE(W)THRIVE) OR MALAISE OR FREQUENT(W)URINATION OR ABDOMINAL(W)DISCOMFORT OR ABDOMINAL(W)PAIN
 6 534 DELAYED(W)BLADDER(W) CONTROL OR DYSURIA OR (PAIN(4N)URINATION)
 7 3 PAINFUL(W)URINATION OR DIFFICULT(W)URINATION
 8 944 URINALYSIS OR URINALYSES
 9 6839 URINE(W)(ANALYSIS OR ANALYSES OR SAMPLE? ? OR SAMPLING OR SPECIMEN? ? OR BAG? ? OR MICROSCOPY OR CULTURE? ?)
 10 653 URINE(4N)(COLLECTION OR COLLECTING)
 11 27 DIP(W)STICK? ?
 12 558 REAGENT(W)STRIP? ?
 13 66 COLORIMETRIC(W)(TEST OR TESTS OR TESTING)
 14 2826 GAS(W)ANALYSIS
 15 71394 IMPEDANCE OR LUMINESCENCE
 16 394 (IMMUNOLOGICAL OR ENZYME)(W)(TEST? ? OR TESTING)
 17 37532 ELISA OR BACTERIAL(W)OXYGEN(W)CONSUMPTION OR TURBIDIMETRY
 18 1228 BACTERIAL(W)CULTURE? ?
 19 138552 DIPSLIDE? ? OR ULTRASOUND OR ULTRASONOGRAPHY OR DIPSTICK? ? OR PLANAR(W)IMAGING OR RADIOGRAPHY OR UROGRAPHY OR PYELOGRAPHY OR KUB OR BLADDER(W)IMAGING
 20 31238 CYSTOGRAPHY OR CYSTOURETHROGRAPHY OR SCINTIGRAPHY OR CYSTOGRAM?
 21 964 DMSA OR UROGRAM? ?

- 22 295 (PLANAR OR RENAL)(W)(SCAN? ? OR SCANNING)
- 23 6955 SPECT OR DIP(W)SLIDE? ? OR CYSTOSCOPY
- 24 20 BLADDER(W)ASPIRATION
- 25 29 ACIDIFICATION(W)(TEST? ? OR TESTING) OR CORTICAL(W)ECHOGENICITY
- 26 14010 WORKUP OR RADIOGRAPHIC OR CYSTOMANOMETRY
- 27 2296 BLADDER(4N)(INVESTIGATION? ? OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 28 6350 KIDNEY? ?(4N)(INVESTIGATION? ? OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 29 565 URETHRA(4N)(INVESTIGATION? ? OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 30 6351 RENAL(4N)(INVESTIGATION? ? OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 31 9125 URINARY(4N)(INVESTIGATION? ? OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 32 449207 INFANT? ? OR PRESCHOOL OR PRESCHOOLER? ? OR CHILD OR CHILDREN OR BABY OR BABIES OR PAEDIATRIC? ? OR PEDIATRIC?
- 33 574159 S4:S31
- 34 1268 S3 AND S33 AND S32

LILACS

LILACS was searched on the BIREME website on 25 February 2003. The following search terms were used:

(Urinary tract infection\$ or uti or infeccion urinaria) AND (child or children or baby or babies or infant or infants or paediatric\$ or pediatric\$ or ninos or ninas)

In total, 279 records were identified and scanned; 100 records were added to the Endnote Library.

COCHRANE LIBRARY

The 2002/4 issue of the Cochrane Library was searched using the CD-ROM.

1. (URINARY next (TRACT next INFECTION*))
2. (((URINARY-TRACT-INFECTIONS*:ME or PYELONEPHRITIS:ME) or BACTERIURIA:ME) or CYSTITIS:ME)
3. ((BACTERIAL-INFECTIONS:ME or PSEUDOMONAS-INFECTIONS*:ME) or KLEBSIELLA-INFECTIONS*:ME)
4. (((GRAM-NEGATIVE-INFECTIONS:ME or ESCHERICHIA-COLI*:ME) or PROTEUS*:ME) or ENTEROCOCCUS*:ME) or STAPHYLOCOCCUS*:ME)
5. (URINARY and TRACT*:ME)
6. URINARY-TRACT*:ME
7. ((#3 or #4) and #6)
8. (INFECTION* near URINARY)
9. (INFECTION* near URINE)
10. (INFECTION* near URETHRA)(INFECTION* near URETER*)
11. (INFECTION* near BLADDER)
12. (INFECTION* near KIDNEY*)
13. (INFECTION* near RENAL)
14. (((((((#1 or #2) or #7) or #8) or #9) or #10) or #11) or #12) or #13) or #14)
15. "RISK ASSESSMENT* OR EXAM OR EXAMINATION* OR FEEDING OR "SLOW WEIGHT GAIN" OR FEVER OR VOMITING OR DIARRH*
16. (((((((RISK next ASSESSMENT*) or EXAM) OR EXAMINATION*) OR FEEDING) OR (SLOW NEXT (WEIGHT NEXT GAIN))) OR FEVER) OR VOMITING) OR DIARRH*)
17. (((((SEPSIS or (FAILURE near THRIVE)) OR MALAISE) OR (FREQUENT NEXT URINATION)) OR (ABDOMINAL NEXT DISCOMFORT)) OR (ABDOMINAL NEXT PAIN))
18. (DELAYED AND ((BLADDER OR ((CONTROL or DYSURIA) OR (PAIN near URINATION))) AND ((PAINFUL OR URINATION) AND (DIFFICULT AND URINATION))))
19. (((((URINALYSIS or URINALYSES) OR (URINE next ANALYS*)) OR (URINE NEXT SAMPL*)) OR (URINE NEAR SPECIMEN*)) OR (URINE NEAR COLLECT*))
20. ((URINE or BAG) AND ((URINE or BAGS) AND ((DIP or STICK*) AND (URINE and MICROSCOPY))))
21. (((((REAGENT next STRIP*) or (COLORIMETRIC next TEST*)) OR (GAS

- NEXT ANALYSIS)) OR IMPEDANCE) OR LUMINESCENCE)
22. ((IMMUNOLOGICAL or (TEST* or ELISA)) AND ((ENZYMES or TEST*) AND (BACTERIAL AND ((OXYGEN or (CONSUMPTION or TURBIDIMETRY)) AND (URINE and CULTURE*))))))
 23. ((BACTERIAL or (((CULTURE* or DIPSLIDE*) or ULTRASOUND) or ULTRASONOGRAPHY) or DIPSTICK*)) AND ((PLANAR or (((IMAGING or RADIOGRAPHY) or UROGRAPHY) or PYELOGRAPHY) or KUB)) AND (BLADDER and IMAGING))
 24. (((CYSTOGRAPHY or CYSTOURETHROGRAPHY) or SCINTIGRAPHY) or CYSTOGRAM*)
 25. (((((PHYSICAL-EXAMINATION*:ME or FEVER*:ME) or BODY-WEIGHT-CHANGES*:ME) or ABDOMINAL-PAIN*:ME) or UROLOGICAL-MANIFESTATIONS*:ME) or FAILURE-TO-THRIVE*:ME)
 26. (((VOMITING*:ME or DIARRHEA*:ME) or SEPSIS*:ME) or URINALYSIS*:ME)
 27. (MICROSCOPY*:ME or INDICATORS-AND-REAGENTS*:ME)
 28. (((COLORIMETRY*:ME or ELECTRIC-IMPEDANCE*:ME) or IMMUNOASSAY*:ME) or FLUORESCENT-ANTIBODY-TECHNIQUE*:ME)
 29. ((DIAGNOSTIC-IMAGING*:ME or CYSTOSCOPY*:ME) or DIAGNOSTIC-TECHNIQUES-UROLOGICAL*:ME)
 30. (((SUCCIMER*:ME or ORGANOMETALLIC-COMPOUNDS*:ME) or TECHNETIUM*:ME) or SULFHYDRYL-COMPOUNDS*:ME) or CULTURE-MEDIA*:ME)
 31. (((URINARY-CATHETERIZATION*:ME or AMMONIUM-CHLORIDE*:ME) or C-REACTIVE-PROTEIN*:ME) or URODYNAMICS*:ME)
 32. ((((((DMSA or UROGRAM*) OR (RENAL next SCAN*)) OR SPECT) OR (PLANAR next IMAGE*)) OR (DIP next SLIDE*)) OR CYSTOSCOPY)
 33. ((BLADDER or ASPIRATION) AND ((ACIDIFICATION or TEST*) AND (CORTICAL and ECHOGENICITY)))
 34. ((WORKUP or RADIOGRAPHIC) or CYSTOMANOMETRY)
 35. ((INVESTIGAT* or DETECT*) or DIAGNOS*)
 36. ((((((((((#16 or #17) or #18) or #19) or #20) or #21) or #22) or #23) or #24) or #25) or #26) or #27)

37. ((((((((((#28 or #29) or #30) or #31) or #32) or #33) or #34) or #35) or #36) or #37)
38. (#15 and #38)
39. ((((((CHILD or CHILDREN) or BABY) or BABIES) or INFANT) or INFANTS) or NEWBORN*)
40. (PEDIATRIC* or PAEDIATRIC*)
41. (CHILD-PRESCHOOL*:ME or INFANT*:ME)
42. ((#40 or #41) or #42)
43. [0]
44. (#39 and #43)

Science Citation Index

The database was searched using the Datastar search interface for the period 1980 to 17 January 2003.

- 1 SCZZ 22615 (URINARY ADJ TRACT ADJ (INFECTION OR INFECTIONS)) OR PYELONEPHRITIS OR BACTERIURIA OR CYSTITIS
- 2 SCZZ 25913 (URINARY OR URINE OR URETHRA OR URETER OR URETERS OR BLADDER OR KIDNEY OR KIDNEYS OR RENAL) WITH (BACTERIAL OR PSEUDOMONAS OR KLEBSIELLA OR GRAM ADJ NEGATIVE OR ESCHERICHIA ADJ COLI OR PROTEUS OR ENTEROCOCCUS OR STAPHYLOCOCCUS OR INFECTION OR INFECTIONS)
- 3 SCZZ 31650 1 OR 2
- 4 SCZZ 357504 RISK ADJ ASSESSMENT OR RISK ADJ ASSESSMENTS OR EXAM OR EXAMINATION OR FEEDING OR SLOW ADJ WEIGHT ADJ GAIN OR FEVER OR VOMITING OR DIARRHEA OR DIARRHOEA
- 5 SCZZ 46980 SEPSIS OR (FAILURE ADJ THRIVE) OR MALAISE OR FREQUENT ADJ URINATION OR ABDOMINAL ADJ DISCOMFORT OR ABDOMINAL ADJ PAIN
- 6 SCZZ 808 DELAYED ADJ BLADDER ADJ CONTROL OR DYSURIA OR (PAIN WITH URINATION)
- 7 SCZZ 1992 URINALYSIS OR URINALYSES
- 8 SCZZ 11525 URINE ADJ (ANALYSIS OR ANALYSES OR SAMPLE OR SAMPLES OR SAMPLING OR

- 9 SCZZ SPECIMEN OR SPECIMENS OR BAG OR BAGS OR MICROSCOPY OR CULTURE OR CULTURES) 1543 URINE WITH (COLLECTION OR COLLECTING)
- 10 SCZZ 64 DIP ADJ (STICK OR STICKS)
- 11 SCZZ 349 REAGENT ADJ (STRIP OR STRIPS)
- 12 SCZZ 121 COLORIMETRIC ADJ (TEST OR TESTS OR TESTING)
- 13 SCZZ 3049 GAS ADJ ANALYSIS
- 14 SCZZ 103131 IMPEDANCE OR LUMINESCENCE
- 15 SCZZ 671 (IMMUNOLOGICAL OR ENZYME) ADJ (TEST OR TESTS OR TESTING)
- 16 SCZZ 2587 BACTERIAL ADJ (CULTURE OR CULTURES)
- 17 SCZZ 137428 DIPSLIDE OR DIPSLIDES OR ULTRASOUND OR ULTRASONOGRAPHY OR DIPSTICK OR DIPSTICKS OR PLANAR ADJ IMAGING OR RADIOGRAPHY OR UROGRAPHY OR PYELOGRAPHY OR KUB OR BLADDER ADJ IMAGING
- 18 SCZZ 27154 CYSTOGRAPHY OR CYSTOURETHROGRAPHY OR SCINTIGRAPHY OR CYSTOGRAM OR CYSTOGRAMS
- 19 SCZZ 1600 DMSA OR UROGRAM OR UROGRAMS
- 20 SCZZ 470 (PLANAR OR RENAL) ADJ (SCAN OR SCANS OR SCANNING)
- 21 SCZZ 31940 SPECT OR DIP ADJ SLIDE OR DIP ADJ SLIDES OR CYSTOSCOPY
- 22 SCZZ 20 BLADDER ADJ ASPIRATION
- 23 SCZZ 54 ACIDIFICATION ADJ (TEST OR TESTS OR TESTING) OR CORTICAL ADJ ECHOGENICITY
- 24 SCZZ 29384 WORKUP OR RADIOGRAPHIC OR CYSTOMANOMETRY
- 25 SCZZ 3464 BLADDER WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 26 SCZZ 5429 KIDNEY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 27 SCZZ 205 URETHRA WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 28 SCZZ 12240 RENAL WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 29 SCZZ 1301 KIDNEYS WITH (INVESTIGATION OR INVESTIGATION OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 30 SCZZ 8298 URINARY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
- 31 SCZZ 694427 INFANT OR INFANTS OR PRESCHOOL OR PRESCHOOLER OR PRESCHOOLERS OR CHILD OR CHILDREN OR BABY OR BABIES OR PAEDIATRIC OR PAEDIATRICS OR PEDIATRIC OR PEDIATRICS
- 32 SCZZ 11 PAINFUL ADJ URINATION OR DIFFICULT ADJ URINATION
- 33 SCZZ 45960 ELISA OR BACTERIAL ADJ OXYGEN ADJ CONSUMPTION OR TURBIDIMETRY

In total, 2612 records were downloaded.

Conference proceedings Index on Dialog

This database is no longer available on Dialog. Conference papers were searched via Biosis and BL Inside Conferences.

BL Inside Conferences on Datastar

This database was searched for the period 1993 to 17 January 2003.

00001 (URINARY ADJ TRACT ADJ (INFECTION OR INFECTIONS)) OR

00002	PYELONEPHRITIS OR BACTERIURIA OR CYSTITIS (URINARY OR URINE OR URETHRA OR URETER OR URETERS OR BLADDER OR KIDNEY OR KIDNEYS OR RENAL) WITH (BACTERIAL OR PSEUDOMONAS OR KLEBSIELLA OR GRAM ADJ NEGATIVE OR ESCHERICHIA ADJ COLI OR PROTEUS OR ENTEROCOCCUS OR STAPHYLOCOCCUS OR INFECTION OR INFECTIONS)	00020	UROGRAPHY OR PYELOGRAPHY OR KUB OR BLADDER ADJ IMAGING CYSTOGRAPHY OR CYSTOURETHROGRAPHY OR SCINTIGRAPHY OR CYSTOGRAM OR CYSTOGRAMS
00003	1 OR 2	00021	DMSA OR UROGRAM OR UROGRAMS
00004	RISK ADJ ASSESSMENT OR RISK ADJ ASSESSMENTS OR EXAM OR EXAMINATION OR FEEDING OR SLOW ADJ WEIGHT ADJ GAIN OR FEVER OR VOMITING OR DIARRHEA OR DIARRHOEA	00022	(PLANAR OR RENAL) ADJ (SCAN OR SCANS OR SCANNING)
00005	SEPSIS OR (FAILURE ADJ THRIVE) OR MALAISE OR FREQUENT ADJ URINATION OR ABDOMINAL ADJ DISCOMFORT OR ABDOMINAL ADJ PAIN	00023	SPECT OR DIP ADJ SLIDE OR DIP ADJ SLIDES OR CYSTOSCOPY
00006	DELAYED ADJ BLADDER ADJ CONTROL OR DYSURIA OR (PAIN WITH URINATION)	00024	BLADDER ADJ ASPIRATION
00007	PAINFUL ADJ URINATION OR DIFFICULT ADJ URINATION	00025	ACIDIFICATION ADJ (TEST OR TESTS OR TESTING) OR CORTICAL ADJ ECHOGENICITY
00008	URINALYSIS OR URINALYSES	00026	WORKUP OR RADIOGRAPHIC OR CYS TOMANOMETRY
00009	URINE ADJ (ANALYSIS OR ANALYSES OR SAMPLE OR SAMPLES OR SAMPLING OR SPECIMEN OR SPECIMENS OR BAG OR BAGS OR MICROSCOPY OR CULTURE OR CULTURES)	00027	BLADDER WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
00010	URINE WITH (COLLECTION OR COLLECTING)	00028	KIDNEY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
00011	DIP ADJ (STICK OR STICKS)	00029	URETHRA WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
00012	REAGENT ADJ (STRIP OR STRIPS)	00030	RENAL WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
00013	COLORIMETRIC ADJ (TEST OR TESTS OR TESTING)	00031	KIDNEYS WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
00014	GAS ADJ ANALYSIS	00032	URINARY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)
00015	IMPEDANCE OR LUMINESCENCE	00033	INFANT OR INFANTS OR PRESCHOOL OR PRESCHOOLER OR PRESCHOOLERS OR CHILD OR
00016	(IMMUNOLOGICAL OR ENZYME) ADJ (TEST OR TESTS OR TESTING)		
00017	ELISA OR BACTERIAL ADJ OXYGEN ADJ CONSUMPTION OR TURBIDIMETRY		
00018	BACTERIAL ADJ (CULTURE OR CULTURES)		
00019	DIPSLIDE OR DIPSLIDES OR ULTRASOUND OR ULTRASONOGRAPHY OR DIPSTICK OR DIPSTICKS OR PLANAR ADJ IMAGING OR RADIOGRAPHY OR		

CHILDREN OR BABY OR BABIES OR
 PAEDIATRIC OR
 PAEDIATRICS OR PEDIATRIC OR
 PEDIATRICS
 00034 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19
 20 21 22 23 24 25 26 27 28 29 30 31
 00035 3 AND 33 AND (32 OR 34)

Six records were identified.

SIGLE

This was searched via the ARC WinSPIRS service for 1980 to 2002/06.

1 9 urinary tract with infection*
 2 17 (urine or ureter* or kidney* or renal or bladder) with infection*
 3 6 uti
 4 0 (urine or ureter* or kidney* or renal or bladder) with pseudomonas
 5 0 (urine or ureter* or kidney* or renal or bladder) with klebsiella
 6 0 (urine or ureter* or kidney* or renal or bladder) with gram negative
 7 5 (urine or ureter* or kidney* or renal or bladder) with coli
 8 2 (urine or ureter* or kidney* or renal or bladder) with escherichia
 9 0(urine or ureter* or kidney* or renal or bladder) with proteus
 10 0 (urine or ureter* or kidney* or renal or bladder) with enterococcus
 11 1 (urine or ureter* or kidney* or renal or bladder) with staphylococcus
 12 0 vesicoureteral reflux
 13 13 pyelonephritis
 14 3 bacteriuria
 15 3 cystitis
 *16 53 #1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15

Dissertation Abstracts

This database was searched using the Datastar interface for the period 1861 to date.

00001 (URINARY ADJ TRACT ADJ (INFECTION OR INFECTIONS)) OR PYELONEPHRITIS OR BACTERIURIA OR CYSTITIS
 00002 (URINARY OR URINE OR URETHRA OR URETER OR URETERS OR BLADDER OR KIDNEY OR KIDNEYS OR RENAL) WITH (BACTERIAL OR

PSEUDOMONAS OR KLEBSIELLA OR GRAM ADJ NEGATIVE OR ESCHERICHIA ADJ COLI OR PROTEUS OR ENTEROCOCCUS OR STAPHYLOCOCCUS OR INFECTION OR INFECTIONS)
 00003 1 OR 2
 00004 RISK ADJ ASSESSMENT OR RISK ADJ ASSESSMENTS OR EXAM OR EXAMINATION OR FEEDING OR SLOW ADJ WEIGHT ADJ GAIN OR FEVER OR VOMITING OR DIARRHEA OR DIARRHOEA
 00005 SEPSIS OR (FAILURE ADJ THRIVE) OR MALAISE OR FREQUENT ADJ URINATION OR ABDOMINAL ADJ DISCOMFORT OR ABDOMINAL ADJ PAIN
 00006 DELAYED ADJ BLADDER ADJ CONTROL OR DYSURIA OR (PAIN WITH URINATION)
 00007 PAINFUL ADJ URINATION OR DIFFICULT ADJ URINATION
 00008 URINALYSIS OR URINALYSES
 00009 URINE ADJ (ANALYSIS OR ANALYSES OR SAMPLE OR SAMPLES OR SAMPLING OR SPECIMEN OR SPECIMENS OR BAG OR BAGS OR MICROSCOPY OR CULTURE OR CULTURES)
 00010 URINE WITH (COLLECTION OR COLLECTING)
 00011 DIP ADJ (STICK OR STICKS)
 00012 REAGENT ADJ (STRIP OR STRIPS)
 00013 COLORIMETRIC ADJ (TEST OR TESTS OR TESTING)
 00014 GAS ADJ ANALYSIS
 00015 IMPEDANCE OR LUMINESCENCE
 00016 (IMMUNOLOGICAL OR ENZYME) ADJ (TEST OR TESTS OR TESTING)
 00017 ELISA OR BACTERIAL ADJ OXYGEN ADJ CONSUMPTION OR TURBIDIMETRY
 00018 BACTERIAL ADJ (CULTURE OR CULTURES)
 00019 DIPSLIDE OR DIPSLIDES OR ULTRASOUND OR ULTRASONOGRAPHY OR DIPSTICK OR DIPSTICKS OR PLANAR ADJ IMAGING OR RADIOGRAPHY OR UROGRAPHY OR PYELOGRAPHY OR KUB OR BLADDER ADJ IMAGING
 00020 CYSTOGRAPHY OR CYSTOURETHROGRAPHY OR SCINTIGRAPHY OR CYSTOGRAM OR CYSTOGRAMS

00021 DMSA OR UROGRAM OR UROGRAMS

00022 (PLANAR OR RENAL) ADJ (SCAN OR SCANS OR SCANNING)

00023 SPECT OR DIP ADJ SLIDE OR DIP ADJ SLIDES OR CYSTOSCOPY

00024 BLADDER ADJ ASPIRATION

00025 ACIDIFICATION ADJ (TEST OR TESTS OR TESTING) OR CORTICAL ADJ ECHOGENICITY

00026 WORKUP OR RADIOGRAPHIC OR CYSTOMANOMETRY SEARCH TERM 'CYSTOMANOMETRY' NOT IN THIS DATABASE

00027 BLADDER WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00028 KIDNEY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00029 URETHRA WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00030 RENAL WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00031 KIDNEYS WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00032 URINARY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00033 INFANT OR INFANTS OR PRESCHOOL OR PRESCHOOLER OR PRESCHOOLERS OR CHILD OR CHILDREN OR BABY OR BABIES OR PAEDIATRIC OR PAEDIATRICS OR PEDIATRIC OR PEDIATRICS

00034 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19
20 21 22 23 24 25 26 27 28 29 30 31

00035 3 AND 33 AND (32 OR 34)

NTIS

This database was searched on the Datastar system for the period 1970 to date.

00001 (URINARY ADJ TRACT ADJ (INFECTION OR INFECTIONS)) OR PYELONEPHRITIS OR BACTERIURIA OR CYSTITIS

00002 (URINARY OR URINE OR URETHRA OR URETER OR URETERS OR BLADDER OR KIDNEY OR KIDNEYS OR RENAL) WITH (BACTERIAL OR PSEUDOMONAS OR KLEBSIELLA OR GRAM ADJ NEGATIVE OR ESCHERICHIA ADJ COLI OR PROTEUS OR ENTEROCOCCUS OR STAPHYLOCOCCUS OR INFECTION OR INFECTIONS)

00003 1 OR 2

00004 RISK ADJ ASSESSMENT OR RISK ADJ ASSESSMENTS OR EXAM OR EXAMINATION OR FEEDING OR SLOW ADJ WEIGHT ADJ GAIN OR FEVER OR VOMITING OR DIARRHEA OR DIARRHOEA

00005 SEPSIS OR (FAILURE ADJ THRIVE) OR MALAISE OR FREQUENT ADJ URINATION OR ABDOMINAL ADJ DISCOMFORT OR ABDOMINAL ADJ PAIN

00006 DELAYED ADJ BLADDER ADJ CONTROL OR DYSURIA OR (PAIN WITH URINATION)

00007 PAINFUL ADJ URINATION OR DIFFICULT ADJ URINATION

00008 URINALYSIS OR URINALYSES

00009 URINE ADJ (ANALYSIS OR ANALYSES OR SAMPLE OR SAMPLES OR SAMPLING OR SPECIMEN OR SPECIMENS OR BAG OR BAGS OR MICROSCOPY OR CULTURE OR CULTURES)

00010 URINE WITH (COLLECTION OR COLLECTING)

00011 DIP ADJ (STICK OR STICKS)

00012 REAGENT ADJ (STRIP OR STRIPS)

00013 COLORIMETRIC ADJ (TEST OR TESTS OR TESTING)

00014 GAS ADJ ANALYSIS

00015 IMPEDANCE OR LUMINESCENCE

00016 (IMMUNOLOGICAL OR ENZYME) ADJ (TEST OR TESTS OR TESTING)

00017 ELISA OR BACTERIAL ADJ OXYGEN ADJ CONSUMPTION OR TURBIDIMETRY

00018 BACTERIAL ADJ (CULTURE OR CULTURES)

00019 DIPSLIDE OR DIPSLIDES OR ULTRASOUND OR ULTRASONOGRAPHY OR DIPSTICK OR DIPSTICKS OR PLANAR ADJ IMAGING OR RADIOGRAPHY OR UROGRAPHY OR PYELOGRAPHY OR KUB OR BLADDER ADJ IMAGING

00020 CYSTOGRAPHY OR CYSTOURETHROGRAPHY OR SCINTIGRAPHY OR CYSTOGRAM OR CYSTOGRAMS

00021 DMSA OR UROGRAM OR UROGRAMS

00022 (PLANAR OR RENAL) ADJ (SCAN OR SCANS OR SCANNING)

00023 SPECT OR DIP ADJ SLIDE OR DIP ADJ SLIDES OR CYSTOSCOPY

00024 BLADDER ADJ ASPIRATION

00025 ACIDIFICATION ADJ (TEST OR TESTS OR TESTING) OR CORTICAL ADJ ECHOGENICITY

00026 WORKUP OR RADIOGRAPHIC OR CYS TOMANOMETRY

00027 BLADDER WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00028 KIDNEY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00029 URETHRA WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00030 RENAL WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00031 KIDNEYS WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00032 URINARY WITH (INVESTIGATION OR INVESTIGATIONS OR DETECT OR DETECTION OR DETECTING OR

DETECTED OR DIAGNOSIS OR DIAGNOSES OR DIAGNOSTIC)

00033 INFANT OR INFANTS OR PRESCHOOL OR PRESCHOOLER OR PRESCHOOLERS OR CHILD OR CHILDREN OR BABY OR BABIES OR PAEDIATRIC OR PAEDIATRICES OR PEDIATRIC OR PEDIATRICES

00034 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31

00035 3 AND 33 AND (32 OR 34)

GreyLit network (<http://www.osti.gov/graylit/>)

The collections on this site were searched on 28 February 2003 using the following search terms:

Urinary tract infection* diagnos*
Uti diagnos*
Cystitis diagnos*
Pyelonephritis diagnos*
Bacteriuria

No relevant reports were found.

NHS EED

NHS EED was searched using the CRD internal search interface using the following search terms:

Urinary (W)tract(w)infection\$ or uti or cystitis or pyelonephritis

Eight records were downloaded.

EMBASE

EMBASE was searched on the Ovid Internet interface on 20 November 2002.

First draft search

- 1 exp urinary tract infection/ (6254)
- 2 bacterial infection/ or gram negative infection/ or enterobacter infection/ or staphylococcus infection/ or exp leukocyte/ (130521)
- 3 vesicoureteral reflux/ or exp pyelonephritis/ or bacteriuria/ or exp cystitis/ (4658)
- 4 exp urinary tract/ (30814)
- 5 (infection? adj3 (urinary or urine or urethra or bladder or ureter? or kidney or kidneys or renal)).ti,ab. (5824)
- 6 (2 or 3) and 4 (1827)
- 7 or/1,5-6 (10003)

- 8 exp infant/ or preschool child/ (63588)
 9 (infant? or baby or babies or toddler? or preschooler?).ti,ab. (41631)
 10 or/8-9 (87280)
 11 7 and 10 (1033)
 12 exp physical examination/ or weight gain/ or growth retardation/ or fever/ or vomiting/ (50693)
 13 exp feeding behavior/ (9803)
 14 exp diarrhea/ or sepsis/ or "failure to thrive"/ or malaise/ (29889)
 15 polyuria/ or abdominal discomfort/ or exp abdominal pain/ or dysuria/ or exp urinalysis/ (21153)
 16 test strip/ or colorimetry/ or exp gas analysis/ or impedance/ or exp luminescence/ (38765)
 17 exp immunoassay/ or enzyme assay/ or turbidimetry/ or urine culture/ or bacterium culture/ (67310)
 18 exp echography/ or ultrasound/ or planar chromatography/ or exp urologic examination/ or exp scintigraphy/ (93139)
 19 exp thiol derivative/ or ammonium chloride/ or "c reactive protein"/ or urodynamics/ (13118)
 20 bladder aspiration/ or exp computer assisted tomography/ or exp emission tomography/ (78550)
 21 (risk assessment? or exam or examination or feeding or slow weight gain or fever or vomiting or diarrh?).ti,ab. (126362)
 22 (sepsis or (failure adj2 thrive) or malaise or frequent urination or abdominal discomfort or abdominal pain).ti,ab. (18269)
 23 (delayed bladder control or dysuria or (pain adj3 urination) or painful urination or difficult urination).ti,ab. (564)
 24 (urinalysis or urine analysis or urine sample? or urine specimen? or (urine adj3 collect?).ti,ab. (7025)
 25 (urine bags or distick? or dip stick? or urine microscopy or reagent strip?).ti,ab. (151)
 26 (colorimetric test? or gas analysis or impedance or luminescence or immunological test?).ti,ab. (6559)
 27 (elisa or enzyme test? or bacterial oxygen consumption or turbidimetry or urine culture).ti,ab. (19582)
 28 (bacterial culture or dipslide? or ultrasonography or planar imag? or radiography or urography or pyelography or kub or bladder imag?).ti,ab. (18115)
 29 (cystography or cystourethrography or nuclear medicine or scintigraphy or cystogram?).ti,ab. (9129)
 30 (dmsa or urogram? or ultrasound or renal scan?).ti,ab. (26816)
 31 (spect or planar imag? or dip slide? or cystoscopy or bladder aspiration).ti,ab. (6227)
 32 (acidification test? or cortical echogenicity or workup? or radiographic or cystomanometry).ti,ab. (12375)
 33 (bladder adj3 (investigat? or detect?)).ti,ab. (126)
 34 (kidney adj3 (investigat? or detect?)).ti,ab. (100)
 35 (urethra adj3 (investigat? or detect?)).ti,ab. (4)
 36 (renal adj3 (investigat? or detect?)).ti,ab. (378)
 37 (kidneys adj3 (investigat? or detect?)).ti,ab. (23)
 38 (urinary adj3 (investigat? or detect?)).ti,ab. (191)
 39 or/12-38 (465240)
 40 11 and 39 (710)

The second and final search used a combination of the search structures in draft 1 above and MEDLINE draft 2 strategy (above).

Project registers

National Research Register (core NICE)

Issue 2003/1 of the NRR was searched on the Internet.

The search strategy was:

URINARY next (TRACT next INFECTION)
 UTI OR CYSTITIS
 PYELONEPHRITIS OR URINARY-TRACT-
 INFECTIONS*:ME
 BACTERIURIA OR BACTERIURIA:ME
 #1 or #2 or #3 or #4

Twenty-six records of ongoing and completed research were downloaded.

Controlled Clinical Trials

The meta-Register of controlled trials was searched on 24 February 2003 for 'urinary tract infection% or uti'. Of the 40 records identified, three were relevant. Two were on the NRR and the single non-NRR record was printed out.

Internet

This topic is very difficult to search for precisely on the Internet. Altavista (www.altavista.co.uk) was searched on 28 February 2003 using the search string:

(urinary near tract near infection near diagnosis) and (child or children or infant or infants or pediatric or paediatrics or baby or babies)

In total, 683 results were returned. The first 200 were scanned for relevance and to yield further papers.

Additional searches

Additional searches for product names associated with diagnostic tests for UTI were conducted.

MEDLINE

(1966 to April week 3 2003 (OVID
<http://gateway.uk.ovid.com/athens>)

- 1 exp Urinary Tract Infections/
- 2 testuria.ti,ab,sh
- 3 uricult.ti,ab,sh
- 4 bacturcult.ti,ab,sh
- 5 combi.ti,ab,sh
- 6 multistix.ti,ab,sh
- 7 nephur.ti,ab,sh
- 8 uristix.ti,ab,sh
- 9 uriglox.ti,ab,sh
- 10 urocomb.ti,ab,sh
- 11 (bac-up-dip or bac?u?dip or BUD).ti,ab,sh
- 12 chemstrip.ti,ab,sh
- 13 clinitek.ti,ab,sh
- 14 bm.ti,ab,sh
- 15 stat.ti,ab,sh
- 16 super.ti,ab,sh
- 17 combur.ti,ab,sh
- 18 (iris or international remote).ti,ab,sh
- 19 bactoscan.ti,ab,s
- 20 urotron.ti,ab,sh
- 21 filtracheck.ti,ab,sh
- 22 (automated urine analy?er).ti,ab,sh
- 23 euron.ti,ab,sh
- 24 urine collection pad.ti,ab,sh
- 25 uriscreen.ti,ab,sh

- 26 or /2-25
- 27 1 and 26

NB. Explosion of UTI selects its more specific terms Bacteriuria, Pyuria and Schistosomiasis haematobia.

EMBASE 1980 to 2003 Week 17 (OVID <http://gateway.uk.ovid.com/athens>)

- 1 exp Urinary Tract Infection/ or
BACTERIURIA/
- 2 testuria.ti,ab,sh
- 3 uricult.ti,ab,sh
- 4 bacturcult.ti,ab,sh
- 5 combi.ti,ab,sh
- 6 multistix.ti,ab,sh
- 7 nephur.ti,ab,sh
- 8 uristix.ti,ab,sh
- 9 uriglox.ti,ab,sh
- 10 (bac-up-dip or bac?u?dip or BUD).ti,ab,sh
- 11 chemstrip.ti,ab,sh
- 12 clinitek.ti,ab,sh
- 13 bm.ti,ab,sh
- 14 stat.ti,ab,sh
- 15 super.ti,ab,sh
- 16 combur.ti,ab,sh
- 17 labstix.ti,ab,sh
- 18 (iris or international remote).ti,ab,sh
- 19 bactoscan.ti,ab,sh
- 20 urotron.ti,ab,sh
- 21 filtracheck.ti,ab,sh
- 22 automated urine analy?er.ti,ab,sh
- 23 euron.ti,ab,sh
- 24 urine collection pad.ti,ab,sh
- 25 uriscreen.ti,ab,sh

NB. Explosion of UTI selects its associated terms Gonococcal Urethritis and Kidney Infection-Abscess, Tuberculosis, Pyonephrosis

Appendix 3

Search strategies for information used to populate the decision model

MEDLINE (1966 to April week 1 2003), using the Ovid Web interfaces, was searched for information to answer the following questions. The searches inevitably retrieved many duplicate records and records were deduplicated.

Decision model question	Total records retrieved	Unique records loaded into database
1.1 Incidence and prevalence of UTI in infants	242	242
1.3 Incidence and prevalence of UTI in preschool children	113	113
1.5 Incidence and prevalence of UTI in children over 5 years	414	152
1.2 Consequences of missed UTI in infants	364	347
1.4 Consequences of missed UTI in preschool children	111	108
1.6 Consequences of missed UTI in children over 5 years	203	184
2.1 What proportion of infants with symptoms of UTI do not see a GP?	22	14
2.4 What proportion of preschool children with symptoms of UTI do not see a GP?	33	11
2.7 What proportion of children over 5 with symptoms of UTI do not see a GP?	15	9
2.2 How many GP consultations per year are there for suspected UTI in infants?	4	4
2.5 How many GP consultations per year are there for suspected UTI in preschool children?	2	1
2.8 How many GP consultations per year are there for suspected UTI in children over 5?	3	1
2.3 What proportion of true-positive cases in infants are referred for imaging?	12	7
2.6 What proportion of true-positive cases in preschool children are referred for imaging?	2	1
2.9 What proportion of true-positive cases in children over 5 are referred for imaging?	9	1
3.1 What proportion of infants presenting with suspected UTI are treated without diagnostic test?	9	9
3.4 What proportion of preschool children presenting with suspected UTI are treated without diagnostic test?	5	4
3.7 What proportion of children over 5 presenting with suspected UTI are treated without diagnostic test?	7	6
3.2 What proportion of infants are given a diagnostic test before any treatment?	98	97
3.5 What proportion of preschool children are given a diagnostic test before any treatment?	24	24
3.8 What proportion of children over 5 are given a diagnostic test before any treatment?	51	23
3.3 If infants are treated for UTI without a diagnostic test what proportion of these are subsequently referred for imaging?	5	1
3.6 If preschool children are treated for UTI without a diagnostic test what proportion of these are subsequently referred for imaging?	0	0
3.9 If children over 5 are treated for UTI without a diagnostic test what proportion of these are subsequently referred for imaging?	2	0
4.1 What percentage of infants get which imaging technique?	5	4
4.5 What percentage of preschool children get which imaging technique?	0	0
4.9 What percentage of children over 5 get which imaging technique?	2	2
4.2, 4.3 and 4.4 What proportion of infants who have an imaging test have severe/mild/no reflux?	132	125
4.6, 4.7 and 4.8 What proportion of preschool children who have an imaging test have severe/mild/no reflux?	0	0
4.10, 4.11 and 4.12 What proportion of children over 5 who have an imaging test have severe/mild/no reflux?	22	18
5.1 and 5.2 What is the disutility associated with an acute or a missed UTI in infants?	15	12

continued

Decision model question	Total records retrieved	Unique records loaded into database
5.3 and 5.4 What is the disutility associated with an acute or a missed UTI in preschool children?	8	6
5.5 and 5.6 What is the disutility associated with an acute or a missed UTI in children over 5?	8	6
6.1 and 6.2 and 6.3 What is the disutility associated with each imaging technique in infants, preschool children and children over 5?	8	1

Answers to the following questions were also derived from other information sources:

Decision model question	Information source
2.3 How many GP consultations per year are there for suspected UTI in infants?	McCormick ²⁷⁸
2.6 How many GP consultations per year are there for suspected UTI in preschool children?	McCormick ²⁷⁸
2.9 How many GP consultations per year are there for suspected UTI in children over 5?	McCormick ²⁷⁸

The search strategies for the questions are shown below:

Questions 1.1 (set 4), 1.3 (set 6) and 1.5 (set 9)

- 1 exp *urinary tract infections/ep or exp *bacteriuria/ep
- 2 exp infants/
- 3 1 and 2
- 4 Child, Preschool/
- 5 1 and 5
- 6 6 not 3
- 7 child/
- 8 1 and 8
- 9 8 not (3 or 6)

Questions 1.2 (set 11), 1.4 (set 14) and 1.6 (set 17)

- 1 exp *urinary tract infections/co or exp *bacteriuria/co
- 2 exp *urinary tract infections/ or exp *bacteriuria/
- 3 missed diagnos\$.ti,ab.
- 4 untreated infection\$.ti,ab.
- 5 missed infection\$.ti,ab.
- 6 (consequences or effects).ti,ab.
- 7 or/3-6
- 8 2 and 7
- 9 1 or 8
- 10 exp infants/
- 11 9 and 10
- 12 children, preschool/
- 13 9 and 12
- 14 13 not 11
- 15 child/
- 16 9 and 15
- 17 16 not (11 or 14)

Questions 2.1 (set 7), 2.4 (set 10), and 2.7 (set 13)

- 1 exp *urinary tract infections/ or exp *bacteriuria/
- 2 exp infants/
- 3 (undiagnosed or untreated or missed diagnos\$.ti,ab.
- 4 (unrecognised or unrecognized).ti,ab.
- 5 (fail\$ adj4 (attend or present)).ti,ab.
- 6 or/3-5
- 7 1 and 2 and 6
- 8 children, preschool/
- 9 1 and 8 and 6
- 10 9 not 7
- 11 child/
- 12 1 and 11 and 6
- 13 12 not (7 or 10)

Questions 2.2 (set 15), 2.5 (set 18) and 2.8 (set 21)

- 1 exp urinary tract infections/
- 2 exp bacteriuria/
- 3 (consult or consultation or consultation or visit or visits or attendance\$.ti,ab.
- 4 "REFERRAL AND CONSULTATION"/
- 5 exp great britain/
- 6 (england or wales or scotland or ireland or uk or gb).in.
- 7 united kingdom.in.
- 8 great britain.in.
- 9 audit.ti,ab. or medical audit/
- 10 or/1-2
- 11 or/3-4,9
- 12 or/5-9
- 13 10 and 11 and 12
- 14 infants/
- 15 13 and 14
- 16 children, preschool/

- 17 13 and 16
 18 17 not 15
 19 child/
 20 13 and 19
 21 20 not (15 or 18)

Questions 2.3 (set 39), 2.6 (set 42) and 2.9 (set 45)

- 1 exp urinary tract infections/
 2 exp bacteriuria/
 3 "REFERRAL AND CONSULTATION"/
 4 exp great britain/
 5 (england or wales or scotland or ireland or uk or gb).in.
 6 united kingdom.in.
 7 great britain.in.
 8 infants/
 9 children, preschool/
 10 child/
 11 exp Diagnostic Imaging/
 12 (refer or referral or referrals or referred).ti,ab.
 13 imaging.ti,ab.
 14 scintigraphy.ti,ab.
 15 urography.ti,ab.
 16 (cystography or tomography or ct).ti,ab.
 17 (radiography or renography).ti,ab.
 18 (ultrasound or ultrasonography).ti,ab.
 19 dmsa.ti,ab.
 20 cystosonography.ti,ab.
 21 vcug.ti,ab.
 22 or/11,13-21
 23 or/1-2
 24 or/3,12
 25 or/4-7
 26 22 and 23 and 24 and 25
 27 26 and 8
 28 26 and 9
 29 28 not 27
 30 26 and 10
 31 30 not (27 or 29)
 32 positive.ti,ab.
 33 "sensitivity and specificity"/
 34 positives.ti,ab.
 35 or/32-34
 36 23 and 35
 37 36 and 25
 38 37 and 8
 39 27 or 38
 40 37 and 9
 41 40 or 29
 42 41 not 39
 43 37 and 10
 44 31 or 43
 45 44 not (39 or 42)

Questions 3.1 (set 15), 3.4 (set 18) and 3.7 (set 21)

- 1 exp *urinary tract infections/
 2 exp *urinary tract infections/di

- 3 primary care.mp. or exp Primary Health Care/
 4 family practice.mp. or Family Practice/
 5 (family practitioner\$ or general practi\$).ti,ab.
 6 or/3-5
 7 2 and 6
 8 (test or tests or result\$).ti,ab.
 9 1 and 8
 10 9 and 6
 11 ((untested or without) adj3 test\$).ti,ab.
 12 1 and 11
 13 or/7,10,12
 14 exp infants/
 15 13 and 14
 16 children, preschool/
 17 13 and 16
 18 17 not 15
 19 children/
 20 13 and 19
 21 20 not (15 or 18)

Questions 3.2 (set 23), 3.5 (set 25) and 3.8 (set 27)

- 1 exp *urinary tract infections/
 2 exp *urinary tract infections/di
 3 primary care.mp. or exp Primary Health Care/
 4 family practice.mp. or Family Practice/
 5 (family practitioner\$ or general practi\$).ti,ab.
 6 or/3-5
 7 2 and 6
 8 (test or tests or result\$).ti,ab.
 9 1 and 8
 10 9 and 6
 11 ((untested or without) adj3 test\$).ti,ab.
 12 1 and 11
 13 or/7,10,12
 14 exp infants/
 15 13 and 14
 16 children, preschool/
 17 13 and 16
 18 17 not 15
 19 children/
 20 13 and 19
 21 20 not (15 or 18)
 22 2 and 14
 23 22 not (15 or 18 or 21)
 24 2 and 16
 25 24 not (15 or 18 or 21 or 23)
 26 2 and 19
 27 26 not (15 or 18 or 21 or 23 or 25)

Questions 3.3 (set 10), 3.6 (set 12) and 3.9 (set 14)

- 1 exp *urinary tract infections/
 2 exp infants/
 3 children, preschool/
 4 children/
 5 investigation\$.ti,ab.
 6 primary care/ or primary care.ti,ab.
 7 family practice/ or family practice.ti,ab.

- 8 (family practitioner\$ or general practi\$).ti,ab.
- 9 or/6-8
- 10 1 and 2 and 5 and 9
- 11 1 and 3 and 5 and 9
- 12 11 not 10
- 13 1 and 4 and 5 and 9
- 14 13 not 11

Questions 4.1 (set 9), 4.5 (set 10), and 4.9 (set 11)

- 1 exp *urinary tract infections/
- 2 exp infants/
- 3 children, preschool/
- 4 children/
- 5 exp DIAGNOSTIC IMAGING/
- 6 exp great britain/
- 7 (england or ireland or wales or scotland or uk or united kingdom or gb or great britain).ti,ab,in.
- 8 or/6-7
- 9 1 and 2 and 5 and 8
- 10 (1 and 3 and 5 and 8) not 9
- 11 (1 and 4 and 5 and 8) not (9 or 10)

Questions 4.2 (set 10), 4.3 (set 10), and 4.4 (set 10)

- 1 exp infants/
- 2 children, preschool/
- 3 children/
- 4 exp DIAGNOSTIC IMAGING/
- 5 exp Vesico-Ureteral Reflux/
- 6 (vesicoureteral reflux or vesico ureteral reflux).ti,ab.
- 7 or/5-6
- 8 4 and 7
- 9 1 and 8
- 10 limit 9 to english language

Questions 4.6 (set 12), 4.7 (set 12), and 4.8 (set 12)

- 1 exp infants/
- 2 children, preschool/
- 3 children/
- 4 exp DIAGNOSTIC IMAGING/
- 5 exp Vesico-Ureteral Reflux/
- 6 (vesicoureteral reflux or vesico ureteral reflux).ti,ab.
- 7 or/5-6
- 8 4 and 7
- 9 1 and 8
- 10 limit 9 to english language
- 11 (2 and 8) not 10
- 12 limit 11 to english language

Questions 4.10 (set 14), 4.11 (set 14) and 4.12 (set 14)

- 1 exp infants/
- 2 children, preschool/
- 3 children/
- 4 exp DIAGNOSTIC IMAGING/

- 5 exp Vesico-Ureteral Reflux/
- 6 (vesicoureteral reflux or vesico ureteral reflux).ti,ab.
- 7 or/5-6
- 8 4 and 7
- 9 1 and 8
- 10 limit 9 to english language
- 11 (2 and 8) not 10
- 12 limit 11 to english language
- 13 (3 and 8) not (10 or 12)
- 14 limit 13 to english language

Questions 5.1 (set 21) and 5.2 (set 21); questions 5.3 (set 25) and 5.4 (set 25)

- 1 exp *urinary tract infections/
- 2 (sf36 or sf 36 or eq5d or eq 5d or euroqol or euro qol).ti,ab.
- 3 (short form or shortform or sf 36 or sf thirty six or sf36).ti,ab.
- 4 (hrql or hrqol or h qol or hql or hqol or hr qol).ti,ab.
- 5 (hye or hyes or health\$ year\$ equivalent\$ or utilit\$).ti,ab.
- 6 health related quality.ti,ab.
- 7 rosser.ti,ab.
- 8 (standard gamble\$ or time trade\$ or timetrade\$ or tto or willingness).ti,ab.
- 9 (disutilit\$ or daly or dalys or disability adjusted).ti,ab.
- 10 "quality of life"/
- 11 quality adjusted life year/
- 12 health status indicators/
- 13 (qaly\$ or quality adjusted life or life quality).ti,ab.
- 14 (qwb or qwbs or uwellbeing or factor analysis).ti,ab.
- 15 (preference\$ or state or states or hspv).ti,ab.
- 16 hui.ti,ab.
- 17 or/2-16
- 18 1 and 17
- 19 infant/
- 20 18 and 19
- 21 limit 20 to english language
- 22 child,preschool/
- 23 18 and 22
- 24 limit 23 to english language
- 25 24 not 21
- 26 child/
- 27 18 and 26
- 28 limit 27 to english language
- 29 28 not (21 or 25)

Questions 6.1 (set 21), 6.2 (set 21) and 6.3 (set 21)

- 1 exp diagnostic imaging/
- 2 exp urinary tract infections/
- 3 1 and 2

- | | |
|--|---|
| <p>4 (sf36 or sf 36 or eq5d or eq 5d or euroqol or euro qol).ti,ab.</p> <p>5 (short form or shortform or sf 36 or sf thirty six or sf36).ti,ab.</p> <p>6 (hrql or hrqol or h qol or hql or hqol or hr qol).ti,ab.</p> <p>7 (hye or hyes or health\$ year\$ equivalent\$ or utilit\$).ti,ab.</p> <p>8 health related quality.ti,ab.</p> <p>9 rosser.ti,ab.</p> <p>10 (standard gamble\$ or time trade\$ or timetrade\$ or tto or willingness).ti,ab.</p> <p>11 (disutilit\$ or daly or dalys or disability adjusted).ti,ab.</p> | <p>12 "quality of life"/</p> <p>13 quality adjusted life year/</p> <p>14 health status indicators/</p> <p>15 (qaly\$ or quality adjusted life or life quality).ti,ab.</p> <p>16 (qwb or qwbs or uwellbeing or factor analysis).ti,ab.</p> <p>17 (qwb or qwbs or wellbeing or factor analysis).ti,ab.</p> <p>18 (preference\$ or state or states or hspv).ti,ab.</p> <p>19 hui.ti,ab.</p> <p>20 or/4-19</p> <p>21 3 and 20</p> |
|--|---|

Appendix 4

QUADAS and details of criteria for scoring studies

1. Was the spectrum of patients representative of the patients who will receive the test in practice?	
<i>Yes</i>	UTI diagnosis: unselected or consecutive population of children suspected of having a UTI Further investigation: patients with confirmed UTI.
<i>No</i>	All other patient spectra including retrospectively selected patient spectra.
<i>Unclear</i>	If insufficient details are provided to make a judgement as to whether the patient spectrum would be scored as 'yes'.

2. Were selection criteria clearly described?	
<i>Yes</i>	Enough details are provided of how patients were selected so that the selection process could be replicated.
<i>No</i>	Insufficient details are presented.
<i>Unclear</i>	Not applicable.

3. Is the reference standard likely to correctly classify the target condition?	
<i>Yes</i>	UTI diagnosis: culture. Further investigation: diagnosis of APN: acute DMSA scan (<4 weeks) diagnosis of reflux: MCUG diagnosis of scarring: follow-up DMSA scan (at least 8 weeks).
<i>No</i>	All other reference standards.
<i>Unclear</i>	If details of the reference standard are not reported.

4. Is the time period between reference standard and index test short enough to be reasonably sure that the target condition did not change between the two tests?	
<i>Yes</i>	UTI diagnosis: should be performed on the same sample of urine. Further investigation: within 3 months.
<i>No</i>	If not as above.
<i>Unclear</i>	If details are not reported.

5. Did the whole sample or a random selection of the sample, receive verification using a reference standard of diagnosis?	
<i>Yes</i>	If the whole sample or a random selection of the sample received the same reference standard.
<i>No</i>	If only a selected sample received the reference standard.
<i>Unclear</i>	If it is not clear whether all the patients received the reference standard.

6. Did patients receive the same reference standard regardless of the index test result?	
<i>Yes</i>	If all patients received the same reference standard.
<i>No</i>	If some patients received a different reference standard.
<i>Unclear</i>	If it is not clear whether all patients received the same reference standard.

7. Was the reference standard independent of the index test (i.e. the index test did not form part of the reference standard)?	
<i>Yes</i>	If the index test and reference standard were independent.
<i>No</i>	If the index test formed part of the reference standard.
<i>Unclear</i>	If it is not clear if the index test and reference standard were independent.

8a. Was the execution of the index test described in sufficient detail to permit replication of the test?	
8b. Was the execution of the reference standard described in sufficient detail to permit its replication?	
<i>Yes</i>	If sufficient details of test execution are reported.
<i>No</i>	If sufficient details are not reported.
<i>Unclear</i>	Not applicable.

9a. Were the index test results interpreted without knowledge of the results of the reference standard?	
9b. Were the reference standard results interpreted without knowledge of the results of the index test?	
<i>Yes</i>	If the index test was interpreted without knowledge of the results of the reference standard and vice versa. If one test was clearly interpreted before the results of the other test were available then this should be scored as 'yes'.
<i>No</i>	If the person interpreting the index test was aware of the results of the reference standard or vice versa.
<i>Unclear</i>	If no information is provided regarding whether tests were interpreted blindly.

10. Were the same clinical data available when test results were interpreted as would be available when the test is used in practice?	
<i>Yes</i>	Dipstick or near patient tests: if clinical data were available. Culture or microscopy: if clinical data were not available. Further investigation: if clinical data were available.
<i>No</i>	If not as above.
<i>Unclear</i>	If details on the availability of clinical data are not reported.

11. Were uninterpretable/intermediate test results reported?	
<i>Yes</i>	If details are provided on uninterpretable/intermediate test results.
<i>No</i>	If there appear to be some uninterpretable/intermediate but the results of these are not reported.
<i>Unclear</i>	If it is not clear whether there were any uninterpretable/intermediate test results.

12. Were withdrawals from the study explained?	
<i>Yes</i>	If all patients recruited into the study were accounted for.
<i>No</i>	If there appear to be patients who were recruited into the study who are not accounted for.
<i>Unclear</i>	If it is not clear whether any withdrawals occurred.

Appendix 5

Included studies: diagnosis of UTI

TABLE 40 Included studies of tests for the diagnosis of UTI

Study details	Patient details	Reference standard	Index test
<p>Ahmad, 1991³⁸</p> <p>Study design Prospective cohort</p> <p>Country England</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum Infants with a wide range of symptoms seen in outpatient clinics or admitted to hospital</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: urine collection bags applied following cleaning of skin with chlorohexidine wipes. Disposable nappies put on by parents according to their normal routine with no specific instructions given about cleaning of the perineum. Nappies that had been on for longer than 4 hours or that were soiled by faeces were excluded</p> <p>Number of patients (number of girls): 45 (NR)</p> <p>Age: mean (range): NR (1–23 months)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Bag specimen</p> <p>Definition of a positive test result > 10⁵ cfu ml⁻¹ of the same single species. Mixed growth classed as negative</p>	<p>Test 1: Urine collection: pad specimen: urine was extracted by compressing nappy fibres in a disposable syringe. Nappies with ultra-absorbent gel-like material were not used in this study Half of each sample was sent for culture on MacConkey and blood agar plates</p> <p>Definition of positive result test 1: > 10⁵ cfu ml⁻¹ of the same single species. Mixed growth classed as negative</p>
<p>Anad, 2001³⁹</p> <p>Study design Prospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Secondary care (non-teaching)</p>	<p>Patient spectrum No details provided</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: except for two samples, all specimens were obtained by clean-catch in younger children, midstream in older children and SPA, catheter samples or Newcastle urine collection pad in infants and children who were still in nappies</p> <p>Number of patients (number of girls): 279 (490 urine specimens) (191)</p> <p>Age: mean (range): NR (4 days to 15 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution No details</p> <p>Definition of a positive test result > 10⁵ cfu ml⁻¹ of urine or any cfu in a urine sample obtained by SPA. Urine cultures with either mixed growth or unusual organisms were excluded from further analysis</p>	<p>Test 1: Dipstick (Nepheu-Test + Leuco (Boehringer Mannheim)) Definition of positive result test 1: LE and/or nitrite positive urine for plus cells</p> <p>Test 2: Microscopy: simple microscopy of fresh uncentrifuged urine for plus cells Definition of positive result test 2: Any WBC hpf⁻¹</p> <p>Test 3: Combination: dipstick and microscopy Definition of positive result test 3: Both tests positive</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Armengol, 2000⁴⁰</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (non-teaching)</p>	<p>Patient spectrum Outpatients aged <4 years with suspected UTI</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Catheter specimen</p> <p>Number of patients (number of girls): 260 (NR)</p> <p>Age: mean (range): NR (<4 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution No details</p> <p>Definition of a positive test result $\geq 10^5$ cfu ml⁻¹ of a uropathogen</p>	<p>Test 1: Dipstick</p> <p>Definition of positive result test 1: Positive nitrite and/or any LE, including trace</p> <p>Test 2: Microscopy for bacteriuria, no further details</p> <p>Definition of positive result test 2: Any bacteria, including 'rare' and 'occasional'.</p> <p>Test 3: Microscopy for pyuria</p> <p>Definition of positive result test 3: ≥ 5 WBC hpf⁻¹</p>
<p>Armengol, 2001²³⁰</p> <p>Study design Retrospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum Children aged <4 years seen as outpatients, identified using computerised records. Children were included if a urine sample had been obtained by catheterisation and a UA and culture performed</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Catheter specimen: no further details</p> <p>Number of patients (number of girls): 260 (NR)</p> <p>Age: mean (range): NR (<4 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Performed on 5% sheep blood-Trypticase agar and MacConkey agar. Organisms were speciated by an automatic method</p> <p>Definition of a positive test result $\geq 10^3$ cfu ml⁻¹ of a uropathogen</p>	<p>Tests 1–3: Dipstick: standard dipstick analysis</p> <p>Definition of positive result test 1: Nitrite positive</p> <p>Definition of positive result test 2: LE positive</p> <p>Definition of positive result test 3: Either or both positive</p> <p>Tests 4–6: Microscopy: urine centrifuged for 6 minutes at 2300 rpm. Sediment resuspended in drop of supernatant and examined either manually or with an automated method with a light microscope (Yellow Iris 900, (International Remote Imaging Systems, Chatsworth, CA, USA)</p> <p>Definition of positive result test 4: Presence of any bacteria</p> <p>Definition of positive result test 5: ≥ 5 WBC hpf⁻¹</p> <p>Definition of positive result test 6: Either or both positive</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Aronson, 1973 ⁴¹	Patient spectrum Children with suspected UTI (100), myelomeningocele (13) and glomerulonephritis (7)	Reference standard Culture or microscopy	Age < 18 months Test 1: Urine sampling: standard bacterial culture of CVU vs SPA culture Definition of positive result test 1: $> 10^5$ cfu ml ⁻¹
Study design Prospective cohort	Recurrent UTI: NR	Reference standard execution Standard bacterial culture or microscopy for pyuria of SPA urine	Test 2: Urine sampling: microscopy for pyuria using a counting chamber: CVU vs SPA microscopy Definition of positive result test 2: > 250 cells mm ⁻³
Country Sweden	Urine sampling methods Combination: CVU specimen obtained simultaneously with or immediately after performing a suprapubic bladder puncture. CVU specimens collected in sterile polyethylene bag (Hollister U-BAG) in infants	Definition of a positive test result $\geq 10^2$ cfu ml ⁻¹ in SPA urine > 10 cells mm ⁻³	Age 3–12 years Test 3: Urine sampling: standard bacterial culture of CVU vs SPA culture Definition of positive result test 3: $> 10^5$ cfu ml ⁻¹
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 120 (62)		Test 4: Urine sampling: microscopy for pyuria using a counting chamber: CVU vs SPA microscopy Definition of positive result test 4: > 250 cells mm ⁻³
	Age: mean (range): NR (0–12 years)		Age < 18 months Test 5: Microscopy: leucocyte counts from SPA urine vs SPA culture Definition of positive result test 5: > 10 cells mm ⁻³
			Test 6: Microscopy/urine sampling: leucocyte counts from CVU vs SPA culture Definition of positive result test 6: > 250 cells mm ⁻³
			Age 3–12 years Test 7: Microscopy: leucocyte counts from SPA urine vs SPA culture Definition of positive result test 7: > 10 cells mm ⁻³
			Test 8: Microscopy: leucocyte counts from CV urine vs SPA culture Definition of positive result test 8: > 250 cells mm ⁻³

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Arslian, 2002 ⁴²	Patient spectrum	Reference standard	Test 1: Microscopy: Gram stain alone
Study design	Children admitted to hospital with symptoms suggesting UTI. Inclusion criteria for infants were: fever with no apparent source, vomiting, decreased appetite and irritability; for toddlers: abdominal pain and voiding frequency with or without fever; and for older children: dysuria, frequency, urgency and abdominal/flank pain with or without fever	Culture	Definition of positive result test 1: Any organisms in 20 oifs
Country	Turkey	Reference standard execution	Test 2: Microscopy: pyuria plus Gram stain. Microscopy for pyuria performed on a centrifuged urine sample
Setting (teaching)	Secondary care (teaching)	Quantitative urine culture performed using a loop calibrated to deliver 0.01 ml to inoculate sheep blood agar and eosin methilen blue agar culture plates. All plates were incubated at 35°C and read at 24 and 48 hours for bacterial identification and colony count	Definition of positive result test 2: > 5 WBCs
		Definition of a positive test result	Test 3: Combination: overall UA. LE and nitrite performed in fresh and uncentrifuged urine using an automated urine analyser. (IRIS 500; IRIS Company, Los Angeles, CA, USA). Microscopy for bacteria and pyuria on centrifuged urine sample.
	Recurrent UTI: NR		Definition of positive result test 3: Positive if nitrite, LE, direct microscopy without Gram stain for bacteria or microscopy for WBC was positive
	Urine sampling methods		
	Combination: in infants an adhesive, sealed, sterile collecting bag was used to obtain a urine sample after disinfection of the skin of the genitals. In toilet-trained children, a midstream urine sample was taken		
	Number of patients (number of girls):		
	100 (52)		
	Age: mean (range): NR (2 days to 15 years)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Bachur, 2001 ⁴³	Patient spectrum Children aged <2 years with fever (temperature $\geq 38^{\circ}\text{C}$) who were seen in the emergency department, identified from medical records	Reference standard Culture	Test: Combination: overall UA: dipstick (Multistix; Bayer Corporation, Elkhart, IN, USA) and microscopy, urine centrifuged at 2000 rpm for 5 minutes
Study design Retrospective cohort		Reference standard execution No details	Definition of positive result test: Positive dipstick (LE and/or nitrite positive) and/or pyuria (≥ 5 WBC hpf ⁻¹)
Country USA	Recurrent UTI: NR	Definition of a positive test result $\geq 10^3$ cfu ml ⁻¹ from SPA, $\geq 10^4$ cfu ml ⁻¹ from catheterised specimens, $\geq 10^5$ cfu ml ⁻¹ from clean-voided specimens. Cultures were considered contaminated if more than one organism or non-pathogens were isolated. Cases with contaminated cultures were removed from further analysis	
Setting (teaching) Secondary care (NR)	Urine sampling methods Not clear: mixture of catheterisation, SPA and CVU		
	Number of patients (number of girls): 8815 (NR)		
	Age: mean (range): NR (<2 years)		
Baum, 1972 ⁴⁴	Patient spectrum Infants and children of both genders; no further details	Reference standard Culture	Test 1: Culture: dipslide with MacConkey medium on one side and nutrient agar on the other, inoculated immediately, dispatched together with clean urine sample to laboratory. Incubated for 18 hours at 37°C
Study design Prospective cohort		Reference standard execution Urine sample plated in the routine fashion. Incubated for 18 hours at 37°C	Definition of positive result test 1: Significant growth (no further details)
Country UK	Recurrent UTI: NR		
Setting (teaching) Secondary care, samples taken in the community	Urine sampling methods Fresh, clean urine samples; sampling details NR	Definition of a positive test result Negative: no significant growth or no growth; positive: significant growth	
	Number of patients (number of girls): 73 (NR)		
	Age: mean (range): NR		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Boreland, 1986 ⁴⁵ Study design Prospective cohort Country UK Setting (teaching) Not clear (NR)	Patient spectrum Paediatric inpatients and outpatients Recurrent UTI: NR Urine sampling methods Not clear Number of patients (number of girls): 700 (samples) (NR) Age: mean (range): NR (0–14 years)	Reference standard Culture Reference standard execution Blotting paper strip technique of Leigh and Williams ²⁷⁹ using both blood agar and MacConkey agar Definition of a positive test result $\geq 10^5$ cfu ml ⁻¹ , pure or mixed (not more than two species)	Index test Test 1: Dipstick: nitrite or blood or protein (N-Labstix SG strips, Ames; read visually, $n = 536$) Definition of positive result test 1: Any one or more of the three positive Test 2: Dipstick: nitrite or blood or protein (N-Multistix SG strips), read photometrically by Clinitek 200 reflectance photometer, Ames; $n = 164$) Definition of positive result test 2: Any one or more of the three positive Tests 3–5: Dipstick (N-Labstix SG strips, read visually, $n = 536$) Definition of positive result test 3: Nitrite positive Definition of positive result test 4: Blood positive Definition of positive result test 5: Protein positive Tests 6–8: Dipstick (N-Multistix SG strips, read photometrically by Clinitek 200 reflectance photometer, $n = 164$) Definition of positive result test 6: Nitrite positive Definition of positive result test 7: Blood positive Definition of positive result test 8: Protein positive
Braude, 1967 ⁴⁶ Study design Prospective cohort Country UK Setting (teaching) Secondary care (teaching)	Patient spectrum Symptomatic patients or (if asymptomatic) routine examination of the urine suggestive of infection Recurrent UTI: NR Urine sampling methods Combination: non-catheter specimens (midstream and/or bag) taken from 68 patients and compared with catheter Number of patients (number of girls): 68 (49) Age: mean (range): 3.2 years (9 days to 11 years)	Reference standard Catheter specimen Reference standard execution Polybactrim catheterisation specimen, assessment of infection by combination of culture and microscopy Definition of a positive test result $> 10^4$ cfu ml ⁻¹ and/or > 10 cells mm ⁻³	All children: Test 1: Urine sampling: non-catheter specimen (bag or midstream), assessment of infection by combination of culture and microscopy Definition of positive result test 1: one or more of the bacterial counts, cell counts or combined counts positive by previously described criteria Age ≤ 5 years Test 2: Urine sampling: as above Definition of positive result test 2: As above

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Bulloch, 2000 ⁴⁷	Patient spectrum Previously healthy patients aged <21 years presenting to an emergency department of a children's hospital who had a specimen collected for urine culture were eligible for the study. The decision to obtain a urine culture and the method of urine collection were made by the evaluating physician. A convenience sample was used: patients were involved when one of the investigators was available. Patients were excluded if they were referred for evaluation of an abnormal UA or positive urine culture, if they were receiving antibiotics, if they had an underlying medical problem requiring repeat catheterisation, if they were unable to obtain an adequate volume of urine or if they were known to have an underlying renal and/or genitourinary abnormality. All children aged <2 years (n = 51) were being evaluated for a fever without focus	Reference standard Culture	Test 1: Urine clarity: urine placed in red-topped blood tube, held against standard white background with black printed 1 l font text. If urine was not clear 0.1 ml of 0.1 mol l ⁻¹ acetic acid solution was added to dissolve any phosphates and visual inspection repeated. Two observers independently performed the inspection
Study design Prospective cohort		Reference standard execution Urine plated on MacConkey agar and sheep blood agar using a 0.01-ml calibrated loop. Incubated for 48 hours at 35°C and examined daily for growth	Definition of positive result test 1: To be considered clear (negative), printed text had to be legible through urine. If there was a discrepancy between observers the specimen was considered positive.
Country USA		Definition of a positive test result > 10 ⁴ cfu ml ⁻¹ from catheterised specimen, ≥ 10 ⁵ cfu ml ⁻¹ from midstream specimens. Considered contaminated if more than three organisms were isolated	Tests 2, 3: Dipstick: trained laboratory technician tested the unspun sample with a dipstick: LE read after 2 minutes classed as negative, trace, small, moderate or large. Nitrite read after 1 minute as positive or negative (Clinitek 200; Ames, Tarrytown, NJ, USA)
Setting (teaching) Secondary care (non-teaching)			Definition of positive result test 2: At least small LE Definition of positive result test 3: Positive nitrite
			Tests 4, 5: Microscopy: urine centrifuged for 5 minutes at 2200 rpm and microscopy performed. Volume of 0.5 ml used for resuspension of urinary sediment
			Definition of positive result test 4: ≥ 5 WBC hpf ⁻¹ Definition of positive result test 5: Any bacteria
			Test 6: Dipstick: abnormal dipstick Definition of positive result test 6: At least small LE or positive nitrite
			Test 7: Microscopy: Abnormal microscopy Definition of positive result test 7: ≥ 5 WBC hpf ⁻¹ or any bacteria
			Test 8: Combination Definition of positive result test 8: Abnormal dipstick or microscopy
	Recurrent UTI: NR		
	Urine sampling methods Combination: clean catch (n = 89) or catheterised specimens (n = 70)		
	Number of patients (number of girls): 159 (122)		
	Age: mean (range): 5.8 years (4 weeks to 19 years)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Caballero, 2001⁴⁸</p> <p>Study design Retrospective cohort</p> <p>Country Spain</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum Retrospective study of children aged <6 years in whom urinary sediment and culture were obtained in the emergency department</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Midstream: samples collected by midstream method from older children and from sterile pads in non-toilet-trained children</p> <p>Number of patients (number of girls): 511 (300)</p> <p>Mean age (range): 2 (<6 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution No details reported, two samples cultured from each child</p> <p>Definition of a positive test result > 10⁵ cfu ml⁻¹ of a single specimen.</p> <p>Considered contaminated when more than one species was present. UTI considered absent if two urine cultures were negative or one of them was negative and the other contaminated. In no case was one culture positive and the other negative in the same child</p>	<p>Tests 1 and 2: Microscopy: microscopy of sediment following centrifuging urine at 2000 rpm for 5 minutes. The number of leucocytes seen visually was counted</p> <p>Definition of positive result test 1: > 18 leucocytes in the urine sediment</p> <p>Definition of positive result test 2: Any leucocytes in the urine</p>
<p>Cervilla, 2001⁴⁹</p> <p>Study design Prospective cohort</p> <p>Country Spain</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum Children with fever > 38.1°C and/or other signs or symptoms related to UTI. All children were tested with dipsticks for LE, nitrite and blood</p> <p>Recurrent UTI: Combination</p> <p>Urine sampling methods Combination: culture samples: CVU or samples from older children, SPA or catheter for younger children</p> <p>Dipstick and microscopy samples: bag specimens</p> <p>Number of patients (number of girls): 102 (74)</p> <p>Mean age (range): 4.1 years (<1 month to 14 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Routine culture performed before administration of antibiotics. Performed where UTI was suspected (positive dipstick test or a negative test with previous history of UTI, renal abnormality, or neurogenic bladder disorder)</p> <p>Definition of a positive test result ≥ 10⁵ cfu ml⁻¹ for CVU samples; ≥ 10⁴ cfu ml⁻¹ for catheter samples; ≥ 100 cfu ml⁻¹ for SPA samples. Mixed culture results were repeated</p>	<p>Test 1: Clinical features: temperature</p> <p>Definition of positive result test 1: > 38.1°C</p> <p>Test 2: Microscopy: no details</p> <p>Definition of positive result test 2: Not clear</p> <p>Test 3: Dipstick</p> <p>Definition of positive result test 3: Blood, LE or nitrite positive</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Cid, 1997 ⁵⁰	Patient spectrum Febrile infants with suspicion of UTI	Reference standard Culture	Test 1: Culture: semi-quantitative urine culture comprised three agar plates: CLED (green) on one side, MacConkey (brown/red) and <i>Enterococcus</i> culture media (no colour). Urine added to test and incubated for 16–24 hours at 37°C (Uricult-plus) The removable section of the tube (containing the agar) was removed and the sensitivity of the colonies was compared with a standard model
Study design Prospective cohort	Recurrent UTI: NR	Reference standard execution Incubated for 18 hours at 37°C	Definition of positive result test 1: NR
Country Spain	Urine sampling methods Not clear: sterile urine samples obtained from urine collection bags	Definition of a positive test result ≥ 10 ⁵ cfu ml ⁻¹ of the same organism in two consecutive urine cultures	
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 100 (61)		
	Mean age (range): NR (1–24 months)		
Cohen, 1997 ⁵¹	Patient spectrum Infants presenting with fever, irritability and no signs of toxicity and no obvious focus of infection on clinical exam. Not sufficiently sick to justify hospitalisation	Reference standard Culture	Test 1: Urine sampling: culture of sample extracted from lining of disposable nappy under sterile conditions
Study design Prospective cohort		Reference standard execution Culture of samples obtained by SPA or catheter on blood agar and MacConkey plates	Definition of positive result test 1: ≥ 10 ⁵ cfu ml ⁻¹
Country Israel	Recurrent UTI: NR	Definition of a positive test result: ≥ 10 ⁵ cfu ml ⁻¹	
Setting (teaching) Primary care (not secondary care)	Urine sampling methods Combination: urine samples were collected from each infant either by catheter or by SPA, and from disposable nappy. Nappies contaminated with faeces or that had been on the infant for > 3 hours were excluded.		
	Number of patients (number of girls): 38 (24)		
	Age: mean (range): NR (1–24 months)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Craver, 1997 ⁵²	Patient spectrum	Reference standard	Tests 1 and 2: Dipstick: unspun urine tested. At 1 minute nitrite was reported as negative or positive, blood was reported as negative, trace, 1+, 2+ or 3+, and protein was interpreted as negative, trace, 1+, 2+ or 3+. Proteinuria of trace or more was confirmed by mixing equal parts of 3% sulfosalicylic acid with centrifuged urine and evaluating the precipitation as none, trace, 1+, 2+ or 3+.
Study design	All children examined at a children's hospital emergency room and suspected of having a UTI were studied. All tests were performed on the basis of clinical necessity determined by the emergency room physician	Reference standard execution	At 2 minutes the LE was reported as negative, trace, 1+, 2+ or 3+ (Ames Multistix 10SG, Miles, Elkhart, IN, USA)
Country	USA	Blood and MacConkey agar plates were each inoculated with 0.01 and 0.001 ml of urine and incubated at 35°C, observed daily for 48 hours.	Definition of positive result test 1: LE at least trace, nitrite positive with blood at least trace
Setting (teaching)	Recurrent UTI: NR	For any growth of one or two organisms the child's symptoms and response to antibiotics were further investigated to determine whether the culture represented infection	Definition of positive result test 2: LE at least trace, nitrite positive with blood at least 1+
Secondary care (NR)	Urine sampling methods	Definition of a positive test result	Test 3: Microscopy: performed on 10 ml of urine spun at 1800 rpm for 5 minutes. Supernatant decanted and sediment pellet resuspended in 1 ml of urine and examined on a 6.6- μ l slide chamber (Kova slide chamber; Hycor Biomedical, Irvine, CA, USA)
	Combination: catheterisation, midstream clean catch or a urine bag	>50,000 cfu of one or two pathogens from a clean-catch or urine bag or > 10 ³ cfu ml ⁻¹ from catheterised samples	Definition of positive result test 3: WBC > 10, any bacteria
	Number of patients (number of girls): 236 (136)		Test 4: Combination
	Age: mean (range): NR (3 weeks to 21 years)		Definition of positive result test 4: LE at least trace, nitrite positive with blood at least trace, any WBC > 10
			Test 5: Combination
			Definition of positive result test 5: LE at least trace, nitrite positive with blood at least 1+, any WBC > 10

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Dayan, 2000 ⁵⁴ Study design Prospective cohort Country USA Setting (teaching) Secondary care (non-teaching)	Patient spectrum Convenience sample of children aged < 4 years undergoing bladder catheterisation to screen for an acute UTI. Patients were being evaluated for UTI mainly secondary to an unexplained fever. Patients were recruited from an urban paediatric emergency department. Patients were excluded if their final diagnosis was a condition that was associated with sterile pyuria Recurrent UTI: NR Urine sampling methods Catheter specimen: early- and late-stream catheterised specimens were obtained Number of patients (number of girls): 87 (36) Age: median (range): median 98 days (3–1330 days)	Reference standard Culture Reference standard execution Later stream samples used: 0.01 ml of urine inoculated via a calibrated loop onto plates containing sheep-blood agar and MacConkey agar. Incubated at 35°C and examined at 24 and 48 hours Definition of a positive test result Single organism growth of $\geq 50,000$ cfu ml ⁻¹ . Equivocal culture defined as single organism growth ≥ 1000 but $< 50,000$ cfu ml ⁻¹ . Negative defined as 'not positive'; included equivocal findings	Early sample Test 1: Culture: as reference standard Definition of positive result test 1: As reference standard: positive or equivocal finding. Test 2: Microscopy: Gram stain: unspun urine completed using automated preparation system (MIDAS 2) Definition of positive result test 2: Any organisms per oif Late sample Test 3: Microscopy: Gram stain. Unspun urine completed using automated preparation system (MIDAS 2) Definition of positive result test 3: Any organisms per oif Early sample Tests 4–6: Microscopy: UA. Completed with centrifuged urine (Yellow IRIS) Definition of positive result test 4: Any bacteria present Definition of positive result test 5: ≥ 5 WBC hpf ⁻¹ Definition of positive result test 6: ≥ 10 WBC hpf ⁻¹ Late sample Tests 7–9: Microscopy: UA. Completed with centrifuged urine (Yellow IRIS) Definition of positive result test 7: Any bacteria present Definition of positive result test 8: ≥ 5 WBC hpf ⁻¹ Definition of positive result test 9: ≥ 10 WBC hpf ⁻¹

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Dayan, 2002 ⁵³ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching)	Patient spectrum Consecutive febrile infants (rectal temperature $\geq 38^{\circ}\text{C}$) aged 1–60 days. Infants were excluded if they received antibiotics within 48 hours of evaluation, if urine collection was attempted but not obtained, or if a Gram stain was not completed secondary to laboratory unavailability Recurrent UTI: Initial UTI Urine sampling methods Combination: Samples only included if obtained by urethral catheterisation ($n = 225$) or SPA ($n = 7$) Number of patients (number of girls): 246 (232 included in analysis) (118) Age: mean (range): 36 days (2–60 days)	Reference standard Culture Reference standard execution 0.001 ml of urine inoculated using a calibrated loop onto MacConkey agar and Columbia agar with 5% sheep blood, incubated at 35°C and examined at 24 and 48 hours Definition of a positive test result $\geq 10,000$ cfu ml^{-1} for catheter samples and ≥ 1000 cfu ml^{-1} for SPA samples of a single urinary pathogen. All other cultures considered negative	Index test Test 1: Microscopy: Gram stain prepared by automated system using uncentrifuged urine read manually (Midas II) Definition of positive result test 1: Identification of any organism Tests 2–5: Dipstick: tests for nitrite and LE, uncentrifuged urine (Super UA automated urine analyser) Definition of positive result test 2: Any positive nitrite Definition of positive result test 3: Any positive LE Definition of positive result test 4: Nitrite and LE positive Definition of positive result test 5: Nitrite or LE positive Tests 6 and 7: Microscopy: automated microscopy of uncentrifuged urine (Yellow IRIS) Definition of positive result test 6: ≥ 5 WBC hpf ⁻¹ Definition of positive result test 7: ≥ 10 WBC hpf ⁻¹
Demi, 1993 ⁵⁵ Study design Prospective cohort Country Italy Setting (teaching) Secondary care (teaching)	Patient spectrum Neonatal inpatients Recurrent UTI: NR Urine sampling methods Urine bag Number of patients (number of girls): 247 (NR) Age: mean (range): NR (neonates; follow-up testing at intervals over 3 year)	Reference standard Culture Reference standard execution Urine culture where dipstick positive or patient symptomatic. No details of technique Definition of a positive test result NR	Index test Tests 1 and 2: Dipstick: 2–3 second immersion (Boehringer Mannheim) Definition of positive result test 1: LE positive Definition of positive result test 2: Nitrite positive

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Dosa, 1973 ⁵⁶	Patient spectrum Two groups of children were investigated: Group A: 422 healthy newborn infants (274 male, 148 female) aged 3–10 days old Group B: 35 infants < 12 months old suspected of having a UTI because earlier routine urine cultures had yielded > 10 ⁵ bacteria ml ⁻¹ of urine. 22 of these infants had a meningomyelocele that may have led to some stasis of urine within the bladder	Reference standard Culture Reference standard execution No details Definition of a positive test result: ≥ 10 ⁵ cfu ml ⁻¹	Group A Tests 1–3: Dipstick: test for glucose: protruding tip changes to blue-green when the urinary glucose concentration exceeds 2–3 mg 100 ml ⁻¹ (Uriglox) Definition of positive result: No colour change Test 1: Initial examination only Test 2: Initial and second examinations combined Test 3: Initial, second and third examination combined Group B Test 4: Dipstick: group B: test for glucose: protruding tip changes to blue-green when the urinary glucose concentration exceeds 2–3 mg 100 ml ⁻¹ (Uriglox) Definition of positive result test 4: No colour change
Study design Prospective cohort			
Country UK			
Setting (teaching) Secondary care (NR)			
	Recurrent UTI: NR Urine sampling methods Combination: Group A: adhesive bag. Second sample of midstream urine collected without the bag; if any test (reference standard or index) was positive in initial sample (<i>n</i> = 66). Third sample collected by SPA if any of the second samples were positive by either test (<i>n</i> = 9) Group B: all samples SPA Number of patients (number of girls): 422 + 35 (NR) Age: mean (range): NR (< 12 months)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Farrell, 2002 ⁵⁷	Patient spectrum	Reference standard	Test 1: Urine sampling: microscopy of pad sample (Euron Urine Collection Pad; Ontex; Buggenhout)
Study design	A convenience, non-consecutive sample from children attending a children's hospital either as outpatients (12) or as inpatients (8) was recruited. In 12 urine samples were required as part of routine surveillance care, while in eight microscopy was requested to confirm either the existence or absence of UTI in the presence of signs of acute illness.	Microscopy and culture	Definition of positive result test 1: WBC present
Country	UK	Reference standard execution	Test 2: Urine sampling: culture of pad sample (Euron Urine Collection Pad)
Setting (teaching)	Children treated with antibiotics within 7 days of the proposed date of urine specimen collection were excluded. However, children receiving long-term prophylactic antibiotic treatment were eligible for inclusion	Urine collected via urine specimen bag (Biotrol Urinocol, B.B. Braun Medical, Sheffield, UK). Urine was retrieved from the bag and decanted into a sterile specimen container. Within 30 minutes of collection samples were sent to the laboratory for standard hospital microbiological analysis	Definition of positive result test 2: Bacteria present
Secondary care (NR)		Definition of a positive test result	
	Recurrent UTI: NR	WBC present (test 1), bacteria present (test 2)	
	Urine sampling methods		
	Combination: urine specimens collected by bag and pad from all children. The bag was applied, having first made a large-bore perforation in the bag with a sterile needle to allow some draining of urine into the pad. The pad was then placed over the bag and then the child's nappy was secured. This ensured that the bag and pad contained the same urine sample		
	Number of patients (number of girls):		
	20 (16)		
	Age: mean (range): 16 months (2–27 months)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Feasey, 1999 ⁵⁸	Patient spectrum	Reference standard	Test 1: Urine sampling: culture: pad specimen
Study design	Convenience sample of 50 children taken from the population of children presenting on the acute services	Culture	Definition of positive result test 1: $\geq 10^5$ cfu ml ⁻¹ of a single organism
Country	UK	Reference standard execution	
Setting (teaching)	paediatric unit of a district general hospital in whom urine microscopy and culture were indicated. Children who were able to cooperate with the collection of a midstream specimen of urine were excluded and included children had to provide at least two specimens of urine. Children were excluded if the specimen was contaminated by faeces, or if the child commenced a course of antibiotics before both specimens were collected	Bag specimen, no further details	
Secondary care (non-teaching)	Recurrent UTI: NR	Definition of a positive test result	
	Urine sampling methods	$\geq 10^5$ cfu ml ⁻¹ of a single organism. All other results (including contaminated specimens) classed as negative	
	Combination: two urine specimens collected from each child. One specimen was collected using a Newcastle urine collection pad and the other by the method normally used in the unit, usually a sterile adhesive bag. Half the children had pad specimens collected first and half had bag specimens collected first		
	Number of patients (number of girls):		
	50 (NR)		
	Age: mean (range): NR (1 day to 5 years)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Fennell, 1977 ⁵⁹	Patient spectrum	Reference standard	Test 1: Culture: following dipstick test, urine was poured into a roll-
Study design	Children were selected from a clinic population known to have recurrent bacteriuria on the basis of willingness and ability to learn the home culturing methods. 20/50 children had an associated structural abnormality of the urinary tract	Culture	tube; after 10 seconds of exposure to the nutrient agar coating was discarded by inverting the tube and the tube was loosely sealed (not airtight). Incubated at 37°C for 24 hours in home incubator. Number of colonies within a unit circle on a transparent flexible plastic strip was counted (Bacturcult; Wampole Laboratories)
Country	USA	Reference standard execution	Definition of positive result 1: Classified by parents after 24 hours as < 10 colonies, 10–50 colonies or > 50 colonies per unit area
Setting (teaching)	Community	Children followed regularly at clinic at bimonthly intervals. If home testing indicated recurrent bacteriuria child was reassessed at clinic as soon as possible. In the clinic another urine specimen was procured by the clean-catch method. A calibrated loop was used to streak on blood agar. When bacteriuria was suspected (by symptoms, home testing or UA) two additional urine specimens were cultured by the standard pour plate technique. Colony counts were also obtained on all first voided specimens brought to clinic on ice	Test 2: Dipstick: dipped into urine and withdrawn immediately [Bac-U-Dip (BUD); Warner-Chilcott]
	Recurrent UTI: Recurrent UTI		Definition of positive result test 2: Pink or red colour change
	Urine sampling methods		
	Clean catch: first urine specimen of the morning		
	Number of patients (number of girls):		
	50 (49)		
	Age: median (range): 9 years (NR, boy was aged 2 years)		
		Definition of a positive test result	
		> 10 ⁵ cfu ml ⁻¹ in both specimens	

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Fernandez, 1996⁶¹</p> <p>Study design Case-control</p> <p>Country Spain</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum Previously healthy children who presented to the emergency department who required a urine culture: previous positive culture ($n = 19$), evaluation of febrile illness ($n = 13$), other clinical presentation ($n = 8$), due to a previously contaminated urine culture specimen ($n = 8$)</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Two consecutive urine samples were collected: one in a perineal bag followed by SPA</p> <p>Number of patients (number of girls): 61 (48 completed) (20 completed)</p> <p>Age: mean (range): ≤ 12 months</p>	<p>Reference standard Reference standard Culture</p> <p>Reference standard execution Culture of SPA urine samples (no further details)</p> <p>Definition of a positive test result Pure growth of > 1000 cfu ml⁻¹ in the SPA specimen</p>	<p>Test 1: Culture of perineal bag specimens (no further details)</p> <p>Definition of positive result test 1: $> 10^5$ cfu ml⁻¹</p>
<p>Fernandez, 2000⁶⁰</p> <p>Study design Prospective cohort</p> <p>Country Spain</p> <p>Setting (teaching) Emergency department (NR)</p>	<p>Patient spectrum Children aged 1–24 months, presence of fever ($> 38^\circ\text{C}$)</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: urine specimen collected by urethral catheterisation (57%) or from a bag attached to the perineum (43%). If bag specimens were positive for leucocyturia or nitrituria, or patients were deemed to be at high risk of UTI, then specimens were also collected by urethral catheterisation (done for 75 patients)</p> <p>Number of patients (number of girls): 175 (115)</p> <p>Age: mean (range): 9.8 months (median 8 months) (1–24 months)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Urine cultured on blood agar and examined in 24–48 hours</p> <p>Definition of a positive test result $> 50,000$ cfu ml⁻¹</p>	<p>Test 1: Dipstick: Nitrite analysis (Multistix; Bayer)</p> <p>Definition of positive result test 1: Nitrite</p> <p>Test 2: Microscopy:</p> <p>Definition of positive result test 2: Presence of leucocytes</p> <p>Test 3: Microscopy: uncentrifuged urine Gram stained and drop placed on slide, dried and examined under microscope</p> <p>Definition of positive result test 3: Gram stain positive</p> <p>Test 4: Combination</p> <p>Definition of positive result test 4: Leucocyturia or nitrituria</p> <p>Test 5: Combination</p> <p>Definition of positive result test 5: Leucocyturia or Gram stain positive</p> <p>Test 6: Combination</p> <p>Definition of positive result test 6: Leucocyturia and nitrituria</p> <p>Test 7: Combination</p> <p>Definition of positive result test 7: Leucocyturia and Gram stain positive</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Giraldez, 1998 ⁶² Study design Prospective cohort Country Venezuela Setting (teaching) Secondary care (NR)	Patient spectrum Children with fever of unknown source, and/or symptoms of UTI who were seen at a paediatric emergency department Recurrent UTI: NR Urine sampling methods Combination: bag specimens and 'Urolab' Number of patients (number of girls): 50 (22) Age: mean (range): 3.9 years (0.4–12 years)	Reference standard Culture Reference standard execution Incubated for 48 hours Definition of a positive test result: $\geq 10^5$ cfu ml ⁻¹	Index test Test 1: Dipstick: nitrite Definition of positive result test 1: Positive Test 2: Dipstick: LE Definition of positive result test 2: Positive Test 3: Dipstick: nitrite and LE Definition of positive result test 3: Both positive Test 4: Microscopy: urine sediment, assume that specimen was centrifuged Definition of positive result test 4: ≥ 10 WBC hpf ⁻¹
Godard, 1979 ⁶³ Study design Prospective cohort Country Switzerland Setting (teaching) Community (not secondary care)	Patient spectrum Girls aged 1–5 years. 86 consulted for acute illnesses, and the remainder for vaccinations or health control Recurrent UTI: NR Urine sampling methods Combination: either by midstream urine (n = 57) or by voiding in a usual pot previously disinfected by alcohol. For non-midstream urine a 5-ml sample was taken from the pot by a plastic syringe. Sample collection method was the same in office in 278 cases and different in 19 cases Number of patients (number of girls): 309 (309) Age: mean (range): NR (1–5 years)	Reference standard Culture Reference standard execution Office culture, no details reported. Patients with colony counts $\geq 10^5$ ml ⁻¹ in both dipslides submitted to a third urine culture using an ideal method of collection (bladder puncture, or midstream urine after local cleaning) Definition of a positive test result $\geq 10^5$ cfu ml ⁻¹ sent for a third culture. No details of how this was considered positive	Index test Test 1: Culture: urine specimen inoculated immediately on dipslide. Dipslide kept in warm place (kitchen or living room) and returned to medical office within several to 24 hours. Colony counts estimated by comparison with photographic standards after 24 hours (NR) Definition of positive result test 1: $\geq 10^5$ cfu ml ⁻¹

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Gorelick, 2000 ⁶⁴	Patient spectrum Girls aged <2 years presenting to the emergency department with fever ($\geq 38.3^{\circ}\text{C}$). Children with definitive sources of fever (e.g. confirmed bacterial infection, specific viral infection and recognisable febrile disease) were excluded. Children with possible but not definitive sources of fever (e.g. upper respiratory tract infection, gastroenteritis, otitis media or non-specific viral syndrome) were included	Reference standard Culture	Index test Test 1: Clinical features: the clinical decision rule developed from various demographic, historical and physical examination results. Developed using multiple logistic regression to identify variables associated with UTI. Following variables were selected: <12 months old, white race, temperature $\geq 39^{\circ}\text{C}$, fever for ≥ 2 days, absence of another source of fever on examination Definition of positive result test 1: Presence of two or more of these five variables. Cut-off selected to maximise sensitivity
Study design Prospective cohort		Reference standard Culture	
Country USA		Reference standard execution No further details	
Setting (teaching) Secondary care (teaching)		Definition of a positive test result Pure growth of $\geq 10^4$ cfu ml ⁻¹ of a pathogenic species of bacteria	
	Recurrent UTI: NR		
	Urine sampling methods All specimens obtained by urethral catheterisation		
	Number of patients (number of girls): 1469 (1469)		
	Age: mean (range): 11 months (<2 years)		
Hardy, 1976 ⁶⁵	Patient spectrum Routine screening for UTI of all inpatient admissions. Children in whom bacterial growth was obtained from first bag specimen. Patients whose condition demanded immediate antimicrobial therapy where delay caused by screening could not be justified were also entered. In the latter group all three specimens were taken without prior screening. Most common reasons for admission were neonatal jaundice, poor feeding, poor weight gain, vomiting, diarrhoea, fever and febrile convulsions	Reference standard SPA culture	Test 1: Urine sampling: culture: bag vs SPA Definition of positive result test 1: $\geq 10^4$ cfu ml ⁻¹ pure growth; mixed growth classed as negative
Study design Prospective cohort		Reference standard execution SPA by Pyles' method ²⁸⁰ Culture on CLED media for 18 hours, WBCs counted unstained on centrifuged urine	Test 2: Urine sampling: culture: clean-catch vs SPA Definition of positive result test 2: $\geq 10^4$ cfu ml ⁻¹ pure growth; mixed growth classed as negative
Country UK		Definition of a positive test result: $\geq 10^4$ cfu ml ⁻¹ pure growth	Test 3: Microscopy: SPA vs SPA culture Definition of positive result test 3: > 10 WBC mm ⁻³
Setting (teaching) Secondary care (teaching)			Test 4: Microscopy: bag vs SPA culture Definition of positive result test 4: > 10 WBC mm ⁻³
	Recurrent UTI: NR		Test 5: Microscopy: clean-catch vs SPA culture Definition of positive result test 5: > 10 WBC mm ⁻³
	Urine sampling methods Combination: all children provided bag, clean-catch and SPA specimens		
	Number of patients (number of girls): 30 (10)		
	Age: mean (range): 16 weeks (2 days to 3 years)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Hiraoka, 1995 ⁶⁶ Study design Prospective cohort Country Japan Setting (teaching) Secondary care (teaching)	Patient spectrum 41 of 89 samples were from children without previous UTI Recurrent UTI: Combination Urine sampling methods Combination: 55 midstream, 20 catheter and 14 indwelling catheter Number of patients (number of girls): 53 (89 samples) (29) Age: mean (range): 6.7 years (1 month to 15 years)	Reference standard Culture Reference standard execution Standard culture, 1- μ l calibrated loop, 48-hour culture Definition of a positive test result 10^5 cfu ml ⁻¹ midstream specimens, 10^3 cfu ml ⁻¹ catheter specimens	Index test Test 1: Microscopy: bacterial count ml ⁻¹ (Kova Slide 10 Grid, Miles-Sankyo) Definition of positive result test 1: NS Test 2: Microscopy: leucocyte count μ l ⁻¹ (Kova Slide 10 Grid) Definition of positive result test 2: ≥ 10 WBC μ l ⁻¹
Hoberman, 1993 ⁶⁸ Study design Prospective cohort Country USA Setting (teaching) Secondary care (NR)	Patient spectrum Urine specimens obtained by bladder catheterisation in the emergency department of a children's hospital. No further details Recurrent UTI: NR Urine sampling methods Catheter specimens Number of patients (number of girls): 698 (NR) Age: mean (range): NR (<2 years)	Reference standard Culture Reference standard execution Quantitative urine culture performed using loop calibrated to deliver 0.001 ml to inoculate plates containing sheep blood agar, Columbia CNA agar and MacConkey agar. Plates incubated at 35-37°C and examined at 24 and 48 hours Definition of a positive test result Growth of single pathogen at concentration of >50,000 cfu ml ⁻¹	Index test Test 1: Microscopy: standard: specimens > 1 ml centrifuged at 2000 rpm for 10 minutes, samples < 1 ml examined uncentrifuged. Unstained specimens examined microscopically for pyuria and bacteriuria Definition of positive result test 1: Pyuria ≥ 5 wbc hpf ⁻¹ and bacteriuria of any bacteria per hpf Test 2: Microscopy: enhanced UA: performed on uncentrifuged urine. Urine drawn into Neubauer haemocytometer and WBCs counted on each side of the chamber. Smears prepared using two drops of uncentrifuged urine on sterile slide and Gram-stained Definition of positive result test 2: Pyuria of ≥ 10 wbc mm ⁻³ and bacteriuria of any bacteria per 10 oifs

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Hoberman, 1994 ⁶⁹ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching)	Patient spectrum Children aged <2 years, seen in the emergency department, from whom a urine sample was obtained by catheterisation were eligible for inclusion. 95% of specimens obtained during diagnostic evaluation for fever Recurrent UTI: NR Urine sampling methods Catheter specimen Number of patients (number of girls): 2181 (NR) Age: mean (range): NR (<2 years)	Reference standard Culture Reference standard execution Calibrated loop (0.001 ml) used to inoculate plates containing sheep's blood agar, Columbia CNA agar and MacConkey agar. All plates incubated at 35–37°C and examined at 24 and 48 hours Definition of a positive test result > 50,000 cfu ml ⁻¹ . Samples with multiple organisms were considered to have contaminated specimens, classed as negative	Index test Test 1: Microscopy: uncentrifuged urine, leucocyte count on a Neubauer haemocytometer Definition of positive result test 1: > 10 WBC mm ⁻³ Test 2: Microscopy: presence of bacteria, by smear (air dried and Gram stained), oif (NS) Definition of positive result test 2: any bacteria Test 3: Microscopy: combination of above Definition of positive result test 3: > 10 WBC mm ⁻³ and any bacteria
Hoberman, 1996 ⁶⁷ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching)	Patient spectrum Children seen in a hospital emergency department from whom a urine specimen for UA and culture was obtained by bladder catheterisation. Urine specimens were obtained as part of a diagnostic evaluation of fever from 95% of children and of sepsis in 5% Recurrent UTI: NR Urine sampling methods Catheter specimens Number of patients (number of girls): 4253 (NR) Age: mean (range): NR (<24 months)	Reference standard Culture Reference standard execution Quantitative urine culture using loop calibrated to deliver 0.001 ml to inoculate plates containing sheep blood agar and MacConkey agar, incubated at 35–37°C and examined at 24 and 48 hours for a colony count and bacterial identification Definition of a positive test result ≥ 50,000 cfu ml ⁻¹ of a single urinary pathogen	Index test Test 1: Microscopy: presence of pyuria. Uncentrifuged urine drawn into Neubauer haemocytometer by capillary action Definition of positive result test 1: ≥ 10 WBC mm ⁻³ Tests 2, 3: Microscopy: uncentrifuged urine drawn into Neubauer haemocytometer by capillary action. WBC counted. Smears were prepared using uncentrifuged urine on a slide within a standardised marked area, air dried and Gram stained Definition of positive result test 2: Presence of pyuria and bacteriuria: ≥ 10 WBC mm ⁻³ and any bacteria per hpf Definition of positive result test 3: Presence of pyuria or bacteriuria: ≥ 10 WBC mm ⁻³ or any bacteria per hpf

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Holland, 1968 ⁷⁰ Study design Prospective cohort Country Ireland Setting (teaching) Secondary care (teaching)	Patient spectrum NR Recurrent UTI: NR Urine sampling methods Midstream: all midstream for tests 1 and 3, for test 2 mainly midstream, but also routine collections (42%) and catheter collections (5%) Number of patients (number of girls): Test 1: 70 (51) Test 2: 372 (206) Test 3: 126 (49), age 1 month to 15 years Age: mean (range): NR (Not clear at least 2.5–7 years)	Reference standard Reference standard Culture Reference standard execution Pour plate colony count method Definition of a positive test result > 10 ⁵ cfu ml ⁻¹	Index test Test 1: Chemical: Greiss nitrate reduction test with the addition of nitrate to the urine before testing. Reported in 4 hours Definition of positive result test 1: Pink or red colour change Test 2: Chemical: TTC test. Reported in 4 hours Definition of positive result test 2: Formation of red precipitate Test 3: Dipstick: Stat-Test (Mallinckrodt Pharmaceuticals), also nitrate reduction test Definition of positive result test 3: Production of red colour
Kohler, 1970 ⁷¹ Study design Prospective cohort Country Sweden Setting (teaching) Community (not secondary care)	Patient spectrum Study was part of a general healthy control, partly of 4-year-old children (n = 948) and partly of schoolgirls (n = 1065) Recurrent UTI: NR Urine sampling methods Midstream: midstream morning samples collected at home under standardised conditions. The sample was chilled with ice cubes in a plastic bag and delivered during the day to the laboratory Number of patients (number of girls): 2016 (1514) Age: mean (range): NR (4–19 years)	Reference standard Culture Reference standard execution No details reported Definition of a positive test result ≥ 10 ⁵ cfu ml ⁻¹	Index test Test 1: Dipstick: Test for glucose (Uriglox) Definition of positive result test 1: No colour change

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Kunin, 1977 ⁷²	Patient spectrum Girls with previous bacteriuria were tested at 3-month intervals for periods of 3 months to 6 years. Some girls were included more than once in the analysis	Reference standard Culture	Mothers given three test strips every 3 months: Test 1: Dipstick: nitrite dipstick performed on three consecutive first morning urine specimens (Microstix-Nitrite; Ames, Elkhart, IN, USA) Definition of positive result test 1: Any shade of pink appearing within 30 seconds after wetting strip classed as positive
Study design Cohort	Recurrent UTI: Recurrent UTI	Reference standard execution Dipsticks taken to laboratory incubated and read at 24–48 hours (Uricult, Orion Laboratories, Helsinki, Finland)	
Country USA	Urine sampling methods Clean catch; CVU collected by mother	Definition of a positive test result > 10 ⁵ cfu ml ⁻¹ . If nitrite tests negative a positive slide culture was confirmed by a second culture	
Setting (teaching) Community (not secondary care)	Number of patients (number of girls): 21 (21) Age: mean (range): preschool		
Labbe, 1982 ⁷³	Patient spectrum Home-based study: children followed at a paediatric UTI clinic who presented with at least two UTIs in the previous 12 months. Patients who received prophylactic antibiotics were excluded	Reference standard Culture	Test 1: Dipstick: home-based study: test performed by mothers, who had received detailed instructions on how to perform it. Mothers tested morning urine samples on Mondays and Thursdays and if child was symptomatic (N-Uristix; Ames) Definition of positive result test 1: Colour change from beige to pink
Study design Prospective cohort	Recurrent UTI: Recurrent UTI	Reference standard execution If nitrite test was positive then the child was taken to the clinic for a urine culture. Children also had regular 3-monthly appointments at the clinic, during which time a sample of urine was taken for culture.	
Country Canada	Urine sampling methods Not clear: no details on method of collection for urine culture samples, assume dipstick samples were collected by mother	Definition of a positive test result > 10 ⁵ cfu ml ⁻¹ of a single organism, intermediate cultures (10 ⁴ –10 ⁵) were repeated	
Setting (teaching) Community (non-teaching)	Number of patients (number of girls): 32 (13 samples) (31) Age: mean (range): 6 years (23 months to 15 years)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Lagos, 1994 ⁷⁴	Patient spectrum	Reference standard	Tests 1, 2: Dipstick: interpreted according to manufacturers' instructions (Combur-9R; Boehringer Mannheim)
Study design	Children presenting suspected of UTI.	Culture and microscopy	Definition of positive result test 1: LE positive
Prospective cohort	Group A: children divided into three groups:	Reference standard execution	Definition of positive result test 2: Nitrite positive
Country	Group A: children with some symptoms of urinary tract inflammation (559)	Cultured on selective (Levine) and non-selective agar (blood).	Test 3: Clinical features: naked eye examination of urine in tube
Chile	Group B: febrile children with urinary tract symptoms (252)	Culture followed by microscopy without centrifugation	Definition of positive result test 3: Not limp
Setting (teaching)	Group C: children with nutritional or gastrointestinal disorders, alterations of urine appearance or previous history of UTI without further follow-up (169)	Definition of a positive test result:	Test 4: Combination
Secondary care (teaching)	Children who had received antibiotics with systematic effects that acted against Gram-negative bacteria in the previous 4 days were excluded	$\geq 10^5$ cfu ml ⁻¹ of a single bacterial isolates with leucocyturia ≥ 10 cells mm ⁻³ or two urine cultures with the same characteristics	Definition of positive result test 4: LE or nitrite positive
	Recurrent UTI: NR		Test 5: Combination
	Urine sampling methods		Definition of positive result test 5: LE positive or not limp
	Combination: midstream samples, for children who could not produce midstream SPA samples were obtained.		Test 6: Combination
	Analysis only included first sample of urine in each patient		Definition of positive result test 6: LE or nitrite positive or not limp
	Number of patients (number of girls): 990 (737)		
	Age: mean (range): (<15 years; includes <24 months)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Lejeune, 1991⁷⁵</p> <p>Study design Prospective cohort</p> <p>Country France</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum Consecutive urine samples from infants aged < 18 months</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Not clear</p> <p>Number of patients (number of girls): 243 (NR)</p> <p>Age: mean (range): NR (21 days to 18 months)</p>	<p>Reference standard Combination</p> <p>Reference standard execution Microscopy and culture, details not reported</p> <p>Definition of a positive test result >25 x 10⁹ WBC l⁻¹ for boys or >50 x 10⁹ WBC l⁻¹ for girls aged < 8 days; >10⁹ WBC l⁻¹ for those aged > 8 days and a bacterial count of >10⁵ cfu ml⁻¹ with a maximum of two bacterial species. Samples with <10⁴ cfu ml⁻¹ or with more than two bacterial species were considered contaminated. Classified as negative</p>	<p>Tests 1–6: Dipstick: (Multistick 8, SG; Ames) read by Clinitek System photometer (Ames)</p> <p>Definition of positive result test 1: LE positive</p> <p>Definition of positive result test 2: Nitrite positive</p> <p>Definition of positive result test 3: Protein positive</p> <p>Definition of positive result test 4: LE and nitrite positive</p> <p>Definition of positive result test 5: LE and protein positive</p> <p>Definition of positive result test 6: LE and nitrite and protein positive</p>
<p>Lin, 2000⁷⁷</p> <p>Study design Prospective cohort</p> <p>Country Taiwan</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum All febrile infants (rectal temperature >38°C) admitted to the paediatric department aged < 12 months. Patients were excluded if they had received antibiotics or SPA or bladder catheterisation within 24 hours</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods All specimens obtained by SPA</p> <p>Number of patients (number of girls): 230 (104)</p> <p>Age: median (range): 1 month (3 days to 12 months)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Quantitative urine cultures using loop calibrated to deliver ± 0.01 ml used to inoculate plates containing sheep blood agar, Columbia CNA agar and MacConkey agar. Plates incubated at 35–37°C and examined at 24–48 hours for colony count and bacterial identification</p> <p>Definition of a positive test result Growth of a single urinary pathogen at a concentration of ≥ 1000 cfu ml⁻¹ considered significant. Cultures with growth of mixed organisms or non-pathogenic Gram-positive cocci were considered contaminated</p>	<p>Test 1: Microscopy: standard microscopy. Specimens centrifuged at 2000 rpm for 10 minutes and examined microscopically for pyuria, reported as number of leucocytes per hpf (NR)</p> <p>Definition of positive result test 1: ≥ 5 WBC hpf⁻¹</p> <p>Test 2: Microscopy: Haemocytometer WBC counts, uncentrifuged urine examined microscopically using counting chamber (Kova slide chamber)</p> <p>Definition of positive result test 2: ≥ 10 WBC μl⁻¹</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Lin, 2000 ⁷⁶ Study design Prospective cohort Country Taiwan Setting (teaching) Secondary care (teaching)	Patient spectrum All febrile (rectal temperature > 38°C) infants aged < 8 weeks who presented to the emergency department or paediatric clinic. All infants were hospitalised. Infants who had received an antibiotic agent or who had undergone SPA or bladder catheterisation within 24 hours were excluded. Children had to provide >5 ml of urine on a single aspiration Recurrent UTI: NR Urine sampling methods All specimens obtained by SPA Number of patients (number of girls): 162 (68) Age: mean (range): NR (<8 weeks)	Reference standard Culture Reference standard execution Quantitative culture performed. Loop calibrated to deliver ±0.01 ml used to inoculate plates containing sheep blood agar, Columbia CNA agar and MacConkey agar. Plates incubated at 35–37°C and examined at 24–48 hours for colony count and bacterial identification Definition of a positive test result Growth of a single urinary pathogen at a concentration of ≥ 100 cfu ml ⁻¹ . Cultures with growth of mixed organisms or non-pathogenic Gram-positive cocci were considered contaminated and classed as negative	Index test Test 1: Microscopy: standard UA. Samples centrifuged at 2000 rpm for 10 minutes and examined microscopically for pyuria, reported as number of leucocytes per hpf Definition of positive result: test 1: ≥ 5 WBC hpf ⁻¹ Test 2: Microscopy: haemocytometer WBC counts. Uncentrifuged urine specimens examined microscopically on a slide chamber. Average leucocyte number per small grid counted and transformed to total cells per µl (Kova slide 10; Hycor Biomedical, Irvine, CA, USA), glass disposable cell counting chamber) Definition of positive result: test 2: ≥ 10 WBC µl ⁻¹ Test 3: CRP: no details, part of full evaluation of sepsis. Definition of positive result: test 3: > 20 mg l ⁻¹ Test 4: ESR: no details, part of full evaluation of sepsis. Definition of positive result: test 4: > 30 mm h ⁻¹ Test 5: Peripheral WBC: no details, part of full evaluation of sepsis. Definition of positive result: test 5: > 15,000 µl ⁻¹
Liptak, 1993 ⁷⁸ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching)	Patient spectrum Children with neurogenic bladder secondary to spinal cord disease or injury Recurrent UTI: NR Urine sampling methods Catheter specimens Number of patients (number of girls): 141 (329 specimens) (NR) Age: mean (range): NR (2 weeks to 21 years)	Reference standard Culture Reference standard execution Performed using 0.01 ml calibrated loop to inoculate MacConkey agar and blood plate Definition of a positive test result ≥ 10 ⁵ cfu ml ⁻¹ of a single organism or of the predominant organism in a mixed culture where there were three or fewer total organisms	Tests 1–3: Dipstick (Chemstrip 9; Boehringer Mannheim) Definition of positive result: test 1: LE positive Definition of positive result: test 2: Nitrite positive Definition of positive result: test 3: Either LE or nitrite or both positive Test 4: Clinical features Definition of positive result: test 4: Signs and symptoms including: urinary odour and cloudiness, fever, malaise, haematuria, pain (NA) Test 5: Microscopy: WBCs and/or bacteria Definition of positive result: test 5: ≥ 10 WBC hpf ⁻¹ or any bacteria

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Littlewood, 1977⁷⁹</p> <p>Study design Prospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum Children attending a general paediatric outpatient clinic. None was receiving antibiotics</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: suprapubic, midstream, clean-catch or bag</p> <p>Number of patients (number of girls): 189 (NR)</p> <p>Age: mean (range): NR (2 weeks to 14 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Culture on well-dried CLED plates (Oxoid), incubation at 37°C for 18 hours</p> <p>Definition of a positive test result ≥ 10⁵ organisms ml⁻¹</p>	<p>Test 1: Microscopy: centrifuged urine, bacteria count</p> <p>Definition of positive result test 1: ≥ 10 organisms per microscopic field</p>
<p>Lockhart, 1995⁸⁰</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Emergency department (teaching)</p>	<p>Patient spectrum Infants (aged <6 months) who presented to a paediatric emergency department over an 8-month period, and from whom urine was obtained for culture using a sterile technique. Patients were excluded if insufficient urine was obtained for both Gram stain and UA</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: SPA (three specimens) or catheterisation (remainder)</p> <p>Number of patients (number of girls): 207 (NR)</p> <p>Age: mean (range): NR (≤ 6 months)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Quantitative urine culture using a 0.1-ml calibrated loop to inoculate blood agar and MacConkey agar culture plates. Plates were incubated and read at 24 and 48 hours</p> <p>Definition of a positive test result Pure growth of more than 1000 cfu ml⁻¹</p>	<p>Test 1: Microscopy: Gram stains on 25 µl of uncentrifuged urine. Stains were read by microbiology technicians and results were verified by supervising microbiologists</p> <p>Definition of positive result test 1: Any organisms in 20 oifs</p> <p>Test 2: UA: qualitative studies performed using an automated urine analyser (Clinitek 200 Plus; Miles): LE, nitrite, microscopy for bacteria and microscopy for pyuria on uncentrifuged sample</p> <p>Definition of positive result test 2: Any one of the following: more than trace LE, nitrite positive, >5 WBC on unstained microscopy, at least 'slight' bacteriuria</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Lohr, 1993⁸¹</p> <p>Study design Retrospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum Urine specimens, from patients whose physicians requested the urine studies to assist in ruling out or documenting a UTI, which were subjected to both UA and culture in an on-site outpatient clinic laboratory were reviewed for a 3-year period. Urine specimens collected for follow-up evaluations of previous UTI were excluded. Children with neurogenic bladders or other urinary tract abnormalities or who were known to have persistent bacteriuria were excluded</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: midstream void ($n = 604$), catheterisation ($n = 76$) and SPA ($n = 9$)</p> <p>Number of patients (number of girls): 689 (NR)</p> <p>Age: mean (range): NR (1 month to 16 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution 0.001-ml calibrated wire loop to inoculate sheep blood agar and MacConkey agar plates</p> <p>Definition of a positive test result > 1000 cfu ml⁻¹ for SPA, > 10⁴ cfu ml⁻¹ for catheter specimens and > 10⁵ cfu ml⁻¹ for midstream voided specimens of a single urinary tract pathogen</p>	<p>Tests 1–3: Dipstick: non-centrifuged sample (Ames Multistix reagent strip read with a Clinitek 10 Analyser; Miles, Elkhart, IN, USA)</p> <p>Definition of positive result test 1: LE: at least trace</p> <p>Definition of positive result test 2: Nitrite: positive</p> <p>Definition of positive result test 3: LE and/or nitrite</p> <p>Test 4: Microscopy: drop of urine taken from sediment of a 10/12-ml sample centrifuged at 1800 rpm for 6 minutes</p> <p>Definition of positive result test 4: > 5 WBC hpf⁻¹</p> <p>Tests 5, 6: Microscopy: drop of urine taken from sediment of a 10/12-ml sample centrifuged at 1800 rpm for 6 minutes. Examined with Gram stain to detect bacteria if they could not be detected in the unstained sediment and if any of the other tests were positive</p> <p>Definition of positive result test 5: any bacteria per hpf or oif</p> <p>Definition of positive result test 6: > 5 WBC hpf⁻¹ and/or any bacteria per hpf or oif</p> <p>Tests 7–9: Combination</p> <p>Definition of positive result test 7: LE and nitrite and/or > 5 WBC hpf⁻¹</p> <p>Definition of positive result test 8: LE and nitrite and/or any bacteria per hpf or oif</p> <p>Definition of positive result test 9: Combination of all tests</p>
<p>Manson, 1985⁸²</p> <p>Study design Prospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Community (not secondary care)</p>	<p>Patient spectrum Primary schoolgirls</p> <p>Recurrent UTI: Other</p> <p>Urine sampling methods Midstream: collection by trained children's nurse</p> <p>Number of patients (number of girls): 2234 (2234)</p> <p>Age: mean (range): NR (4–12 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Standard calibrated loop (1 µl) plated onto CLED agar and MacConkey agar</p> <p>Definition of a positive test result ≥ 10⁵ cfu ml⁻¹</p>	<p>Test 1: Microscopy: protocol 1: undiluted urine + borate buffer + acridine orange (Bactoscan)</p> <p>Definition of positive result test 1: 10⁵ counts ml⁻¹</p> <p>Test 2: Microscopy: protocol 6: urine diluted (1:2) with distilled water + borate buffer + acridine orange (Bactoscan)</p> <p>Definition of positive result test 2: ≥ 10⁵ counts ml⁻¹</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Marret, 1995 ⁸³	Patient spectrum Children with suspected UTI	Reference standard Culture	Tests 1–4: Dipstick: test for bacteria (FiltracCheck-UTI) Definition of positive result test 1: No details
Study design Prospective cohort	Recurrent UTI: NR	Reference standard execution No details	Test 2: Dipstick: detection of LE and nitrites (Combur-9) Definition of positive result test 2: Not clear
Country Singapore	Urine sampling methods Combination: sterile bag, midstream voided specimens, catheterisation and SPA	Definition of a positive test result No details	
Setting (teaching) Not clear (NR)	Number of patients (number of girls): 69 (NR)		
	Age: mean (range): NR (2 weeks to 14 years)		
Marsik, 1986 ⁸⁴	Patient spectrum NR	Reference standard Culture	Tests 1–4: Dipstick: (Chemstrip LN; Biodynamics)
Study design Prospective cohort	Recurrent UTI: NR	Reference standard execution Accepted techniques (Barry <i>et al.</i> , 1975 ²⁸¹)	Definition of positive result test 1: LE positive Definition of positive result test 2: Nitrite positive Definition of positive result test 3: LE or nitrite positive Definition of positive result test 4: LE and nitrite positive
Country USA	Urine sampling methods Combination: clean-catch, catheterised or SPA samples (numbers by each method NR)	Definition of a positive test result Clean-catch: single isolate of Gram-positive $\geq 10^4$ cfu ml ⁻¹ , or Gram-negative $\geq 10^5$ cfu ml ⁻¹ Catheterised: $\geq 10^4$ cfu ml ⁻¹ any organism SPA: any number of bacteria per ml	
Setting (teaching) Secondary care (NR)	Number of patients (number of girls): 402 (601 samples) (NR)		
	Age: mean (range): NR (0–21 years)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Matthai, 1995⁸⁵</p> <p>Study design Prospective cohort</p> <p>Country India</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum Children with suspected UTI; none of the children had received antibiotics in the 24 hours before the study</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Clean catch: freshly voided midstream clean-catch</p> <p>Number of patients (number of girls): 376 (NR)</p> <p>Age: mean (range): NR (6 months to 5 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Cultured on blood agar and MacConkey agar and observed for 48 hours</p> <p>Definition of a positive test result > 10⁵ cfu ml⁻¹</p>	<p>Test 1: Urine albumin: proteinuria: quantitatively estimated by the sulphosalicylic acid method</p> <p>Definition of positive result test 1: Not clear</p> <p>Tests 2–4: Microscopy: pyuria: unstained centrifuged samples and examined for leucocytes and erythrocytes</p> <p>Definition of positive result test 2: > 5 WBC hpf⁻¹</p> <p>Definition of positive result test 3: > 10 WBC hpf⁻¹</p> <p>Definition of positive result test 4: > 20 WBC hpf⁻¹</p> <p>Test 5: Microscopy: bacteriuria: performed on unstained and Gram-stained sediment</p> <p>Definition of positive result test 5: Not clear</p> <p>Test 6: Combination: proteinuria, pyuria and bacteriuria</p> <p>Definition of positive result test 6: > 10 WBC hpf⁻¹ for pyuria, not clear for others</p> <p>Test 7: Pyuria and bacteriuria positive</p> <p>Definition of positive result test 7: > 10 WBC hpf⁻¹ for pyuria, not clear for others</p> <p>Test 8: Combination: pyuria and proteinuria positive</p> <p>Definition of positive result test 8: > 10 WBC hpf⁻¹ for pyuria, not clear for others</p> <p>Test 9: Combination: proteinuria and bacteriuria positive</p> <p>Definition of positive result test 9: Not clear</p> <p>Test 10: Combination: any two positive</p> <p>Definition of positive result test 10: > 10 WBC hpf⁻¹ for pyuria, not clear for others</p> <p>Test 11: Combination: any one positive</p> <p>Definition of positive result test 11: > 10 WBC hpf⁻¹ for pyuria, not clear for others</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Mongeau, 1972 ⁸⁶ Study design Prospective cohort Country Canada Setting (teaching) Secondary care (teaching)	Patient spectrum Children selected at random from a renal clinic to provide a urine specimen Recurrent UTI: NR Urine sampling methods Not clear Number of patients (number of girls): 100 (65) Age: mean (range): NR (3–16 years)	Reference standard Culture Reference standard execution Regular urine culture Definition of a positive test result ≥ 10 ⁵ cfu ml ⁻¹ or urinary pathogens. Growth of 104 considered doubtful, classed as negative	Test 1: Culture: miniature culture plate smeared with a dipstick after its immersion in the urine. After 19–24 hours incubation at 37°C. Read after 24 hours (Testuria) Definition of positive result test 1: No growth (0–2 colonies), suspicious growth (3–25 colonies) positive growth (>25 colonies). Suspicious classed as negative Test 2: Culture: semi-quantitative culture medium. Slide covered with nutrient agar on one side and MacConkey agar on the other dipped into freshly voided urine. Incubated at 37°C for 18 hours, results read semi-quantitatively. Read after 24 hours (Uricult) Definition of positive result test 2: ≥ 10 ⁵ cfu ml ⁻¹ , doubtful classed as negative
Morton, 1982 ⁸⁷ Study design Prospective cohort Country Nigeria Setting (teaching) Secondary care (teaching)	Patient spectrum Children suspected of having UTI and who had not taken antimicrobials in the previous 7 days Recurrent UTI: NR Urine sampling methods Combination: SPA performed about 20 minutes after drinking. If the aspiration failed one further attempt was made after a further drink. Following cleaning of the external genitalia midstream urine specimens were collected Number of patients (number of girls): Not clear (NR) Age: mean (range): NR (<10 years; includes infants)	Reference standard Culture Reference standard execution Samples plated out by loop onto MacConkey medium or blood agar for 18 hours at 37°C. Performed by SPA for comparison of test 1, urine specimen unclear for other tests Definition of a positive test result Virtually any growth from SPA, ≥ 10 ⁵ cfu ml ⁻¹ from midstream samples	Test 1: Culture: performed on MSU Definition of positive result test 1: > 10 ⁵ cfu ml ⁻¹ Tests 2, 3: Microscopy: uncentrifuged urine for WBCs: not stained Definition of positive result test 2: > 10 cells mm ⁻³ Definition of positive result test 3: > 15 organisms hpf ⁻¹ Test 4: Microscopy: uncentrifuged urine for bacteria: Gram stained Definition of positive result test 4: > 15 organisms hpf ⁻¹ Test 5: Microscopy: urine sediment, obtained by centrifugation at 2500 rpm for 5 minutes, for bacteria: Gram stained Definition of positive result test 5: > 15 organisms hpf ⁻¹

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Navarrete, 1966⁸⁸</p> <p>Study design Prospective cohort</p> <p>Country Not clear</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum Urine samples were obtained from hospitalised children. Reasons for urine samples were: children presenting with general symptoms and fever, and for gastrointestinal symptoms including nausea, vomiting and diarrhoea or for control of a previously positive sample</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Not clear: external genitalia cleaned with sterile solution, sterile bag was positioned with sticky tape</p> <p>Number of patients (number of girls): 140 (60)</p> <p>Age: mean (range): NR, 91% <2 years (1 month to 6 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Quantitative culture was performed</p> <p>Definition of a positive test result ≥ 10⁵ cfu ml⁻¹</p>	<p>Tests 1 and 2: Culture: 2 cm³ of urine was used for the test. Readings were taken every 30 minutes for 9 hours (Uriscreen)</p> <p>Definition of positive result test 1: Results after 4 hours</p> <p>Definition of positive result test 2: Results after 9 hours</p>
<p>Ordonez, 1994⁸⁹</p> <p>Study design Prospective cohort</p> <p>Country Spain</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum Children presenting to the service with fever and/or a suspicion of UTI</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Bag specimens</p> <p>Number of patients (number of girls): 100 (40)</p> <p>Age: mean (range): NR (1–24 months)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Sample incubated for 18 hours at 37°C</p> <p>Definition of a positive test result ≥ 10⁵ cfu ml⁻¹ of the same organism in two consecutive urine cultures</p>	<p>Test 1: Clinical features: semi-quantitative urine culture comprised three agar plates: CLED (green) on one side, MacConkey (brown/red) and <i>Enterococcus</i> culture media (no colour). Urine added to test and incubated for 16–24 hours at 37°C (Uricult-plus)</p> <p>Definition of positive result test 1: The removable section of the tube (containing the agar) was removed and the sensitivity of the colonies was compared with a standard model</p> <p>Tests 2, 3: Dipstick (Multistix; Ames)</p> <p>Definition of positive result test 2: Nitrite</p> <p>Definition of positive result test 3: Nitrite positive</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Palmer, 1997 ⁹⁰ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching)	Patient spectrum Consecutive children undergoing urodynamic evaluation for various indications. Indications for urodynamic evaluation included various aetiologies of neurogenic and non-neurogenic bladder dysfunction. All children were asymptomatic for UTI at the time of the study. Children on suppressive or therapeutic antibiotic regimens were maintained on that agent Recurrent UTI: No Urine sampling methods Catheter specimens Number of patients (number of girls): 200 (NR) Age: mean (range): NR (0–15 years)	Reference standard Culture Reference standard execution Uncentrifuged urine was mixed and inoculated onto plates of eosin-methylene blue and Columbia CNA soy agar supplemented with 5% sheep blood. Incubated overnight at 35–37°C with 5–10% carbon dioxide Definition of a positive test result: ≥ 50,000 cfu ml ⁻¹	Test 1: An aliquot of urine was tested immediately according to manufacturer's specifications. A 1.5–2-ml urine sample was transferred into a test-tube containing the reagent powder; four drops of 10% hydrogen peroxide were added and mixed (Uriscreeen; no further details) Definition of positive result test 1: Formation of foam sufficient to form a complete ring or layer on the surface of the liquid within 1–2 minutes
Parmington, 1989 ⁹¹ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching)	Patient spectrum All patients presenting to the paediatric emergency department, in whom both a urine culture was ordered and a nitrite dipstick result recorded Recurrent UTI: NR Urine sampling methods Not clear Number of patients (number of girls): 305 (214) Age: mean (range): NR (includes <2 years, age range NR)	Reference standard Culture Reference standard execution NS Definition of a positive test result NS	Total population Test 1: Dipstick Definition of positive result test 1: Nitrite

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Pryles, 1965 ⁹² Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching)	Patient spectrum Infants and children attending consecutive paediatric renal clinics. One urine specimen per patient was obtained at each clinic; patients could contribute several samples over several visits Recurrent UTI: NR Urine sampling methods Clean-catch Number of patients (number of girls): 136 (275 urine specimens) (114) Age: mean (range): NR (infants and children)	Reference standard Culture Reference standard execution No further details Definition of a positive test result $\geq 10^5$ cfu ml ⁻¹	Tests 1 and 2: Microscopy: well-mixed aliquot was examined in a Neubauer counting chamber and a cell count performed Definition of positive result test 1: ≥ 10 WBC mm ⁻³ Definition of positive result test 2: ≥ 100 WBC mm ⁻³ Test 3: Microscopy: aliquot centrifuged at 3000 rpm for 3 minutes, supernatant fluid poured off and sediment diluted to 0.5 ml with more uncentrifuged urine and mixed well Definition of positive result test 3: ≥ 5 WBC mm ⁻³
Purwar, 1972 ⁹³ Study design Prospective cohort Country India Setting (teaching) Secondary care (teaching)	Patient spectrum NR Recurrent UTI: NR Urine sampling methods Not clear: urine specimens for routine examination were collected in clean test tubes or bottles. For culture samples were clean-voided midstream samples with strict aseptic precautions in sterile test-tubes, in very few cases catheter specimens were taken Number of patients (number of girls): 232 (62) Age: mean (range): NR (2 months to 14 years)	Reference standard Culture Reference standard execution Colony counts performed 24–48 hours after incubating sterilised urine inoculated agar plate Definition of a positive test result $> 10^4$ cfu ml ⁻¹	Test 1: Microscopy: Gram-stained film of a loopful of uncentrifuged urine Definition of positive result test 1: Any bacteria

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Pylkkanen, 1979⁹⁴</p> <p>Study design Prospective cohort</p> <p>Country Finland</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum Patients seen at a general outpatient clinic with symptoms suggestive of UTI or with a high leukocyte count or an abnormal quantitative bacterial culture from a CVU specimen</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 477 (not reported)</p> <p>Age: mean (range): not reported (not reported, 164 infants included)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Suprapubic aspiration sample cultured on Uricult dipslides and blood agar plates</p> <p>Definition of a positive test result Both cultures showed consistent bacterial growth</p>	<p>Tests 1 and 2: Leukocyte count by microscopy of CVU specimen</p> <p>Definition of positive result test 1: ≥ 200 leukocytes mm^{-3}</p> <p>Definition of positive result test 2: ≥ 11 leukocytes mm^{-3}</p> <p>Tests 3 and 4: Bacterial count by microscopy of CVU specimen</p> <p>Definition of positive result test 3: ≥ 30 bacteria/microscope field</p> <p>Definition of positive result test 4: > 1 bacteria/microscope field</p> <p>Tests 5 and 6: Quantitative bacterial culture of CVU</p> <p>Definition of positive result test 5: $\geq 10^5$ bacteria ml^{-1}</p> <p>Definition of positive result test 6: $\geq 10^4$ bacteria ml^{-1}</p>
<p>Ramage, 1999⁹⁵</p> <p>Study design Prospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum Infants in whom urine culture was indicated, with paired urine cultures obtained by SPA and clean-catch within a 48-hour period, in the absence of prior or intervening antibiotic 58 specimen pairs from 49 patients</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: clean-catch specimens and paired SPA specimens</p> <p>Number of patients (number of girls): 49 (58 specimens) (27)</p> <p>Age: mean (range): 0.36 years (<24 months)</p>	<p>Reference standard Culture of SPA specimen</p> <p>Reference standard execution Specimen obtained by SPA as described by Nelson and Peters.²⁸² Culture on 'Till-U-Test' dipslide plate incubated at 37°C for 16 hours</p> <p>Definition of a positive test result Presence of growth of any organism</p>	<p>Test 1: Urine sampling: as for reference standard, but sample obtained by clean-catch by nursing staff or parents supervised by nursing staff</p> <p>Definition of positive result test 1: $\geq 10^5$ cfu ml^{-1}</p>
<p>Rich, 1976⁹⁶</p> <p>Study design Prospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Community (not secondary care)</p>	<p>Patient spectrum Population screening in six primary schools and one secondary school</p> <p>Recurrent UTI: Other</p> <p>Urine sampling methods Midstream</p> <p>Number of patients (number of girls): 1329 (1329)</p> <p>Age: mean (range): NR (4–16 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Supervised collection, dipslide</p> <p>Definition of a positive test result Three supervised midstream sample with $\geq 10^5$ cfu ml^{-1}</p>	<p>Tests 1 and 2: Culture: home dipslide, after explanatory letter to parents and explanation to children by nurse</p> <p>Definition of positive result test 1: $\geq 10^5$ cfu ml^{-1}. Unsatisfactory samples ($n = 213$) classed as negative</p> <p>Definition of positive result test 2: $\geq 10^5$ cfu ml^{-1} (negative includes unreturned dipslides)</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Santos, 1982⁹⁷</p> <p>Study design Prospective cohort</p> <p>Country Brazil</p> <p>Setting (teaching) Not clear (NR)</p>	<p>Patient spectrum Children with possible UTI</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: bag samples from infants, midstream from older children</p> <p>Number of patients (number of girls): 2000 (1314)</p> <p>Age: mean (range): NR (<12 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Urine diluted and plated on MacConkey and blood agar. Incubated for 24 hours at 37°C</p> <p>Definition of a positive test result Not clear</p>	<p>Test 1: Microscopy: quantitative bacterioscopy: urine Gram stained, area was standardised and reading of homogenised urine smears was used to calculate the quantity of bacteria per ml and their morpho-tinctorial condition Definition of positive result test 1: Not clear</p> <p>Test 2: Microscopy for the presence of leucocyturia: urine was centrifuged at 2000 rpm for 5 minutes, stained with the dye of Sterheimer-Malbin, and number of leucocytes counted Definition of positive result test 2: Not clear</p>
<p>Saxena, 1975⁹⁸</p> <p>Study design Prospective cohort</p> <p>Country India</p> <p>Setting (teaching) Not clear (NR)</p>	<p>Patient spectrum Children in whom a clinical diagnosis of UTI was made and who were referred for laboratory confirmation. All children were ambulatory and continent</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Midstream samples</p> <p>Number of patients (number of girls): 70 (45)</p> <p>Age: mean (range): NR (4–12 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Semi-quantitative urine culture performed within 1 hour of collection on MacConkey agar and blood agar plates</p> <p>Definition of a positive test result NR</p>	<p>Test 1: Microscopy: urine sample was shaken well and a drop placed on a Neubaur counting chamber Definition of positive result test 1: > 10 WBC mm⁻³</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Scherstein, 1968 ⁹⁹ Study design Prospective cohort Country Sweden Setting (teaching) Community (not secondary care)	Patient spectrum Screening of 4-year-old children and schoolgirls; no further details reported Recurrent UTI: Combination Urine sampling methods Midstream: samples were clean-voided, midstream specimens from first morning voiding, after overnight fast Number of patients (number of girls): 719 (592) Age: mean (range): NR (4 to 18 years)	Reference standard Culture Reference standard execution Culture as previously described ²⁸³ Definition of a positive test result Positive when two or more consecutive samples of $> 10^5$ organisms ml ⁻¹	Test 1: Dipstick: test paper A, prepared in the laboratory: for the adsorbent part, 2-diethylaminoethanol-substituted paper, washed with phosphate buffer, was used. Glucose oxidase, peroxidase and o-tolidine were used for the colour reagent. Reagents were fixed on a plastic stick. Test performed by immersing adsorbent part in urine (max. 0.5 cm) and allowing urine to pass to colour reaction part by capillary action. Reaction evaluated 3 minutes after urine reached colour reaction part Definition of positive result: test 1: If no colour after 3 minutes, test negative for glucose and therefore positive for bacteriuria
Schreiter, 1971 ¹⁰⁰ Study design Prospective cohort Country Germany Setting (teaching) Secondary care (NR)	Patient spectrum Children admitted to the paediatric department with suspected or confirmed UTI. 349 children aged 0–1 years and 660 1–14 years. Children with confirmed glomerulonephritis were excluded. Some children were receiving chemotherapy ($n = 188$), the rest were not ($n = 82$) Recurrent UTI: NR Urine sampling methods Not clear: no details reported Number of patients (number of girls): 1009 (614) Age: mean (range): NR (0–14 years)	Reference standard Culture Reference standard execution Urine streaked on blood agar and an endoplate. Urine centrifuged and sediment streaked on fructose bouillon. Incubated for 24 hours at 37°C and colonies counted Definition of a positive test result $> 10^5$ cfu ml ⁻¹	Test 1: Microscopy: leucocytes counted. no details. Aged < 1 year. Children who did not receive chemotherapy Definition of positive result: test 1: > 20 LE μ l ⁻¹ Test 2: Microscopy: leucocytes counted. No details. Aged 1–14 years. Children who did not receive chemotherapy Definition of positive result: test 2: > 20 LE μ l ⁻¹ Test 3: Microscopy: leucocytes counted. No details. Aged < 1 year. Children who did chemotherapy Definition of positive result: test 3: > 20 LE μ l ⁻¹ Test 4: Microscopy: leucocytes counted. No details. Aged 1–14 years. Children who did receive chemotherapy Definition of positive result: test 4: > 20 LE μ l ⁻¹

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Sharief, 1998 ¹⁰¹	Patient spectrum	Reference standard	All tests: Dipstick (Multistix 8 SG; (Bayer), read on a Clinitek 10 reflectance meter (Bayer)
Study design	Children in whom UTI was a possibility; unselected children with fever, no urinary tract symptoms	Culture	
Country	Recurrent UTI: NR	Reference standard execution	Total study population:
Setting (teaching)	Urine sampling methods	Definition of a positive test result	Definition of positive result test 1: LE positive
Secondary care (non-teaching)	Combination: 188 clean-catch, 167 bag	UTI defined as pure growth of $\geq 10^5$ cfu ml ⁻¹ , inconclusive culture defined as 10^4 – 10^5 cfu ml ⁻¹ , and pyuria, $< 10^4$ cfu ml ⁻¹ , and mixed growth were taken to be negative	Definition of positive result test 2: Nitrite positive
	Number of patients (number of girls): 325 (131)		Definition of positive result test 3: LE and nitrite positive
	Age: mean (range): 39.5 months (2 days to 16 years)		Infants < 1 year
			Definition of positive result test 4: LE positive
			Definition of positive result test 5: Nitrite positive
			Definition of positive result test 6: LE and nitrite positive
			Total study population:
			Definition of positive result test 7: LE or nitrite positive
			Infants < 1 year
			Definition of positive result test 8: LE or nitrite positive

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Shaw, 1991¹⁰³</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum All children who were examined in the paediatric emergency department who had a urine specimen collected for culture between 08.00 h and 24.00 h (when a laboratory technician was available). The evaluating physician made the decision to order a urine culture and chose the method of specimen collection</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: urine collected in a urine bag or by midstream, clean-catch techniques were cultured using dipslide (Uricult). Urine obtained by urethral catheterisation cultured in laboratory</p> <p>Number of patients (number of girls): 491 (309)</p> <p>Age: mean (range): 6 years (14 days to 19 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Either dipslide (Uricult, Medical Technology Corp., Somerset, NJ, USA) or inoculated onto blood and MacConkey agar plates with a 0.01-ml calibrated loop. Cultures and dipslides incubated at 35°C and examined for growth daily for 2 days</p> <p>Definition of a positive test result ≥ 10⁵ cfu ml⁻¹ of one or two urinary pathogens for clean-catch specimens or ≥ 10³ cfu ml⁻¹ for catheterised specimens</p>	<p>All children</p> <p>Tests 1, 2: Dipstick: LE measurement, read after 2 minutes as negative, trace, small (1+), moderate (2+) or large (3+); nitrite read at 60 seconds and recorded as negative or positive. Performed by trained laboratory technician (Multistix 10 SG)</p> <p>Definition of positive result test 1: at least small LE or positive nitrite</p> <p>Definition of positive result test 2: at least trace LE or positive nitrite</p> <p>Definition of positive result test 3: at least small LE and positive nitrite</p> <p>Tests 4–6: Microscopy: centrifuged at 2200 rpm for 5 minutes</p> <p>Definition of positive result test 4: at least 10 WBC hpf⁻¹ or moderate bacteria</p> <p>Definition of positive result test 5: at least 5 WBC hpf⁻¹ or few bacteria</p> <p>Definition of positive result test 6: at least 10 WBC hpf⁻¹ and moderate bacteria</p> <p>Aged <2 years</p> <p>Tests 7 and 8: Dipstick: as above</p> <p>Definition of positive result test 7: at least trace LE or positive nitrite</p> <p>Definition of positive result test 8: at least small LE and positive nitrite</p> <p>Tests 9 and 10: Microscopy: as above</p> <p>Definition of positive result test 9: at least 5 WBC hpf⁻¹ or few bacteria</p> <p>Definition of positive result test 10: at least 10 WBC hpf⁻¹ and moderate bacteria</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Shaw, 1998¹⁰²</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (teaching)</p>	<p>Patient spectrum Patients presenting to a children's hospital emergency department with unexplained fever $\geq 38.3^{\circ}\text{C}$ or symptoms of UTI. It was standard practice in the department to obtain urine specimens on such patients</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: in 1% of patients urine was obtained, after sterile preparation, by midstream clean-catch. The remainder of specimens were obtained by catheterisation, as per standard practice in the unit</p> <p>Number of patients (number of girls): 3873 (2363)</p> <p>Age: mean (range): 9 months (<2 years girls, <1 year boys)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Quantitative cultures performed in the microbiology laboratory. Urine inoculated onto blood and MacConkey agar plates with a 0.1-ml calibrated loop, incubated at 35°C, and examined daily for growth for 2 days</p> <p>Definition of a positive test result: $\geq 10^4$ cfu ml^{-1} of a urinary tract pathogen. Mixed organisms of non-pathogens were considered contaminated and classed as negative</p>	<p>Tests 1, 2: Dipstick: performed in the haematology laboratory, results interpreted visually according to standard colour charts. LE read after 2 minutes and recorded as negative, trace, small (+), moderate (+2) or large (+3). Nitrate read at 60 seconds and recorded as positive or negative (Multistix 10SG 228; Bayer, Elkhart, IN, USA)</p> <p>Definition of positive result test 1: At least trace LE or positive nitrite ($n = 3394$)</p> <p>Definition of positive result test 2: At least moderate LE or positive nitrite ($n = 3394$)</p> <p>Test 3: Microscopy: cell count using uncentrifuged urine (Neubauer haemocytometer, Reichert)</p> <p>Definition of positive result test 3: ≥ 10 WBC mm^{-3} ($n = 2193$)</p> <p>Tests 4, 5: Microscopy: Gram stain, smears were prepared using uncentrifuged urine (NR)</p> <p>Definition of positive result test 4: Any bacteria ($n = 2305$)</p> <p>Definition of positive result test 5: Single organism ($n = 2305$)</p> <p>Tests 6, 7: Microscopy: cell count using uncentrifuged urine. Gram stain, smears were prepared using uncentrifuged urine (Neubauer haemocytometer, Reichert)</p> <p>Definition of positive result test 6: Cell count ≥ 10 WBC mm^{-3} or positive Gram stain ($n = 2016$)</p> <p>Definition of positive result test 7: Cell count ≥ 10 WBC mm^{-3} and positive Gram stain ($n = 2016$)</p> <p>Tests 8, 9: Combination: dipstick analysis as before. Urine centrifuged for 5 minutes at 2200 rpm and number of leucocytes and bacteria per hpf recorded. Microscopy only performed if any component of the dipstick was positive (protein, blood, glucose, LE, nitrite, ketones)</p> <p>Definition of positive result test 8: Dipstick positive or ≥ 5 WBC hpf^{-1} or any bacteria per hpf ($n = 3394$)</p> <p>Definition of positive result test 9: Dipstick positive and ≥ 5 WBC hpf^{-1} and bacteraemia per hpf</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Struthers, 2003 ¹⁰⁴ Study design Prospective cohort Country UK Setting (teaching) Secondary care (non-teaching)	Patient spectrum Children aged <6 years, whose urine was being collected, in a paediatric admissions department. There were no specific exclusions, but children were only included when study team was on duty. Unit policy was that all unwell or febrile young children should have a urine sample collected; this will include children with symptoms of UTI and children with non-specific symptoms Recurrent UTI: NR Urine sampling methods Combination: clean-catch ($n = 105$), SPA ($n = 3$), midstream ($n = 2$) Number of patients (number of girls): 110 samples (NR) Age: mean (range): 23 months (2 days to 62 months)	Reference standard Culture Reference standard execution No details Definition of a positive test result > 10^5 cfu ml ⁻¹ , mixed growths were classed as negative	Test 1: Clinical features: questionnaire on urine odour given to parents at time of sample collection Definition of positive result test 1: One or more positive replies
Tahirovic, 1988 ¹⁰⁵ Study design Prospective cohort Country Yugoslavia Setting (teaching) Secondary care (teaching)	Patient spectrum Unselected patients Recurrent UTI: NR Urine sampling methods Midstream Number of patients (number of girls): 306 (NR) Age: mean (range): NR (0–14 years)	Reference standard Culture Reference standard execution Quantitative culture Definition of a positive test result ≥ 10^5 cfu ml ⁻¹	Test 1: Dipstick (Urocomb 8 test Pliva; Boehringer Mannheim) Definition of positive result test 1: Nitrite positive Test 2: Dipstick: nitrite test after sterile incubation for 4 hours at 37°C, for samples initially nitrite negative (Urocomb 8 test Pliva) Definition of positive result test 2: Nitrite positive Test 3: Dipstick: nitrite test after sterile incubation with NaNO ₃ for 4 hours at 37°C, for samples initially nitrite negative (Urocomb 8 test Pliva) Definition of positive result test 3: Nitrite positive

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Todd, 1974 ⁰⁶ Study design Prospective cohort Country USA Setting (teaching) Community (non-teaching)	Patient spectrum Children with recurrent urinary tract infections. 13 of the children had underlying anatomical or physiological abnormalities of the urinary tract, 12 were receiving suppressant medications Recurrent UTI: Recurrent UTI Urine sampling methods Combination: not clearly reported for home specimen collection. For reference standard culture, a midflow catheter specimen or clean-catch midstream sample was required Number of patients (number of girls): 25 (22) Age: mean (range): NR (3–14 years)	Reference standard Culture Reference standard execution Midflow catheter quantitative urine culture performed according to the calibrated loop method. Culture performed if home tests indicated UTI positive, and samples from all children were cultured at 1–2-month intervals. Reference standard was not clearly reported Definition of a positive test result > 10 ³ cfu ml ⁻¹ or if two clean-catch midstream urine cultures (one for boys) > 10 ⁵ cfu ml ⁻¹ of a single organism	Test 1: Dipstick: urine specimens taken after overnight fasting tested weekly at home. The first morning specimen was tested with Uriglox Strips (Kabi, Stockholm, Sweden). If glucose was present infection was assumed to be absent. If glucose was absent on two consecutive specimens the children were seen in the clinic to confirm the presence of infection Definition of positive result test 1: Absence of glucose (no blue colour)
Vangone, 1985 ⁰⁷ Study design Prospective cohort Country Italy Setting (teaching) Secondary care (teaching)	Patient spectrum No further details Recurrent UTI: NR Urine sampling methods Combination: sterile bag, catheter Midstream: no further details Number of patients (number of girls): 561 (NR) Age: mean (range): NR (15 days to 15 years)	Reference standard Culture Reference standard execution No details reported Definition of a positive test result > 10 ⁵ cfu ml ⁻¹	Tests 1 and 2: Microscopy: fresh, unstained, uncentrifuged urine, bacterial count using a Neubauer haemocytometer Definition of positive result test 1: > 5 bacteria ml ⁻¹ Definition of positive result test 2: > 10 bacteria ml ⁻¹

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Vickers, 1991 ¹⁰⁸ Study design Prospective cohort Country UK Setting (teaching) Secondary care (NR)	Patient spectrum Consecutive children who attended an outpatient clinic (many because of previous UTI diagnosis) or who were admitted to hospital with a wide range of diagnoses and symptoms and signs that included unexplained pyrexia, abdominal pain, haematuria, malaise and weight loss. A few had known chronic renal failure Recurrent UTI: Combination Urine sampling methods Combination: midstream specimens or sterile urine bags in younger children Number of patients (number of girls): 357 (NR) Age: mean (range): NR (6 weeks to 18 years)	Reference standard Combination Reference standard execution As with index tests Definition of a positive test result Patients were considered positive if urine microscopy and culture results were both positive, and to have uninfected urine when the culture result was negative, irrespective of the microscopic findings. If the initial urine specimen showed uncertain results on microscopy or culture, a repeat specimen was taken as soon as possible. Urine samples were obtained repeatedly until a specimen fulfilled criteria for a UTI or no infection	Test 1: Microscopy: samples examined three times at x400 magnification. Samples examined for red and white blood cells, epithelial cells, crystals, debris and bacteria. Neubauer counting chamber with a mirrored surface was used in all examinations Definition of positive result test 1: $\geq 10^7$ bacteria of the same morphology per ml; negative if no bacteria seen; uncertain for any other result. For 2×2 analysis uncertain classed as negative Test 2: Culture: urine plated onto MacConkey and blood agar and the interval between the collection and plating noted Definition of positive result test 2: $> 10^5$ cfu ml ⁻¹ of one species; negative if $< 10^4$ cfu ml ⁻¹ ; uncertain if growth of more than one species was observed. For 2×2 analysis uncertain classed as negative

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Waisman, 1999¹⁰⁹</p> <p>Study design Prospective cohort</p> <p>Country Israel</p> <p>Setting (teaching) Secondary care (NR)</p>	<p>Patient spectrum A random sample of children presenting to the emergency department of a children's medical centre with symptoms suggesting UTI. Inclusion criteria were: for infants, fever with no apparent source, vomiting, decreased appetite and irritability; for toddlers, abdominal pain and voiding frequency with or without fever; and for older children, dysuria, frequency, urgency and abdominal flank pain with or without fever. Children receiving antibiotics were excluded from the study</p> <p>Recurrent UTI: NR</p> <p>Urine sampling methods Combination: SPA for neonates, catheterisation for girls ≤ 3 years and urine bags for boys until they were toilet trained, clean-catch for older children</p> <p>Number of patients (number of girls): 121 (82)</p> <p>Age: mean (range): NR (1 month to 17 years)</p>	<p>Reference standard Culture</p> <p>Reference standard execution Commercial Diaslide methods (Diathec Diagnostica, Rehovot, Israel)</p> <p>Hinged case containing two opposing agar media, both agar surfaces inoculated with a streaking dilution</p> <p>Definition of a positive test result > 10^5 cfu ml⁻¹ for clean-catch or urine bag, 10^3 for catheterised and SPA samples</p>	<p>Test 1: Uriscreeen (catalase test): aliquot of urine placed in a test-tube containing Uriscreeen reagent powder, four drops of 10% hydrogen peroxide added and shaken gently (Uriscreeen; Diathec Diagnostica, Rehovot, Israel)</p> <p>Definition of positive result test 1: Formation of foam sufficient to form a complete ring or layer on the surface of the liquid within 1–2 minutes of the addition of the hydrogen peroxide</p> <p>Test 2: Dipstick: aliquot of uncentrifuged urine tested for presence of nitrite or LE (Multistix; Bayer, Elkhart, IN, USA)</p> <p>Definition of positive result test 2: Positive if the dipstick turned pink or red for nitrites or purple (from trace to +2) for leucocytes within 2 minutes of contact with the urine</p> <p>Test 3: Microscopy: uncentrifuged urine screened initially with automated urine analyser, specimens found to be positive underwent centrifugation and the sediment was examined by standard microscopy (Urotron RL9; Boehringer Mannheim)</p> <p>Definition of positive result test 3: > 10 leucocytes hpf⁻¹</p>

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Wammanda, 2000 ¹⁰	Patient spectrum Consecutive children presenting at a paediatric hospital unit with features suggestive of urinary tract infection as well as other clinical conditions necessitating infection screen, in whom urine culture was carried out as part of the infection screen	Reference standard Culture Reference standard execution Blood agar and MacConkey agar plated simultaneously. Incubated for 18–24 hours. Definition of a positive test result $\geq 10^5$ cfu ml ⁻¹	Test 1: Microscopy: for leucocyturia: 10 ml of urine centrifuged, supernatant fluid decanted and remainder shaken to mix; drop placed on slide and examined under high power Definition of positive result test 1: ≥ 10 cells hpf ⁻¹ Test 2: Dipstick: nitrite using Multistix 10 SG test strip according to manufacturer's instructions (Multistix; Bayer) Definition of positive result test 2: Manufacturer's instructions
Study design Prospective cohort			
Country Nigeria			
Setting (teaching) Secondary care (teaching)	Recurrent UTI: NR Urine sampling methods Combination: clean-catch midstream urine collected in a sterile container in older children. For infants, collected in a sterile urine bag attached to their perineum after cleaning with dilute Savlon		
	Number of patients (number of girls): 185 (67) Age: mean (range): NR (2 days to 12 years)		

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Weinberg, 1991 ¹¹¹	Patient spectrum	Reference standard	Tests 1–4: Microscopy: drop of well-mixed uncentrifuged urine transferred to slide and stained using Gram stain reagents. Five oifs were examined and the averaged results reported as number of microorganisms seen.
Study design	Children who had a complete UA, quantitative urine Gram-stained smear and urine culture performed as part of an evaluation for urinary tract infection or acute febrile illness of uncertain cause.	Reference standard execution	Definition of positive result test 1: Any
Country	Follow-up specimens were excluded from the analysis. Children came from a clinic serving children from birth to aged 18 years	0.001-ml calibrated loop was used to plate and streak the urine on MacConkey's and CNA agar. Organism identification and quantification were by standard microbiological techniques	Definition of positive result test 2: ≥ 1 oif ⁻¹
Setting (teaching)	Recurrent UTI: NR	Definition of a positive test result	Definition of positive result test 3: ≥ 2 oif ⁻¹
Secondary care (teaching)	Urine sampling methods	$\geq 10^5$ cfu ml ⁻¹	Definition of positive result test 4: ≥ 5 oif ⁻¹
	Combination: the method of urine collection was at the discretion of the physician caring for the patient: catheter ($n = 261$), bag ($n = 186$), SPA ($n = 253$) and clean-catch ($n = 319$).		Tests 5–7: Dipstick: used in accordance with manufacturer's instructions (Multistix; Ames)
	Number of patients (number of girls): 1019 (NR)		Definition of positive result test 5: Nitrite positive
	Age: mean (range): NR		Definition of positive result test 6: LE positive
			Definition of positive result test 7: Nitrite or LE positive
			Tests 8, 9: Microscopy: urine centrifuged for 5 minutes at 2000 rpm, resuspended after supernatant was decanted. Placed in disposable slide chamber and examined microscopically (Kova slide chamber; ICL Scientific)
			Definition of positive result test 8: ≥ 5 WBC hpf ⁻¹
			Definition of positive result test 9: ≥ 10 WBC hpf ⁻¹

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Wiggelinkhuizen, 1988 ^{11,12}	Patient spectrum Children attending medical casualty and outpatient follow-up clinics were screened	Reference standard Culture	Tests 1–6: Dipstick (Combur-9 test; Boehringer Mannheim)
Study design Prospective cohort			Definition of positive result test 1: LE positive
Country South Africa	Recurrent UTI: NR	Reference standard execution Semi-quantitative culture using Bacteruritest (Mast Laboratories) filterstrip imprints on CLED agar	Definition of positive result test 2: Nitrite positive
Setting (teaching) Secondary care (teaching)	Urine sampling methods Combination: clean-catch midstream ($n = 668$), bag ($n = 400$), catheter ($n = 44$), suprapubic aspiration ($n = 6$), other (ostomy samples) ($n = 19$)	Definition of a positive test result $> 10^5$ cfu ml ⁻¹ of one predominant organism. Lower counts were considered significant if samples obtained by bladder catheterisation or SPA	Definition of positive result test 3: LE or nitrite positive
	Number of patients (number of girls): 1137 (522)		Definition of positive result test 4: LE and nitrite positive
	Age: mean (range): NR (0 to > 12 years)		Definition of positive result test 5: LE, nitrite or protein positive
			Definition of positive result test 6: LE, nitrite and protein positive
			Tests 7–12: Dipstick (Multistix 9; Ames Bayer-Miles)
			Definition of positive result test 7: LE positive
			Definition of positive result test 8: Nitrite positive
			Definition of positive result test 9: LE or nitrite positive
			Definition of positive result test 10: LE and nitrite positive
			Definition of positive result test 11: LE, nitrite or protein positive
			Definition of positive result test 12: LE, nitrite and protein positive
			Note: For the analysis for dipstick positive for both LE and nitrite the authors appear to have omitted test results that were positive for LE or positive for nitrite and negative for the other. The reviewers used the results for dipstick positive for either test to deduce these and have added these to the 'negative' cells of the 2×2 table of the data

continued

TABLE 40 Included studies of tests for the diagnosis of UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Woodward, 1993 ¹¹³	Patient spectrum Patients in paediatric surgical department. Patients were selected using pre-existing criteria for urine testing (NS); most had acute abdominal pain	Reference standard Culture and microscopy	Test 1: Dipstick: LE and nitrite positive (Multistix 10SG; Bayer) Definition of positive result test 1: more than a trace of LE, any colour change for nitrites
Study design Prospective cohort	Recurrent UTI: NR	Reference standard execution NS	Test 2: Dipstick: either test positive (Multistix 10SG) Definition of positive result test 2: more than a trace of LE detected, any colour change for nitrites
Country UK	Urine sampling methods Combination: 86% of samples were taken midstream	Definition of a positive test result > 20 WBC μl^{-1} and > 10^8 cfu l^{-1} (10^5 cfu ml^{-1})	
Setting (teaching) Secondary care (non-teaching)	Number of patients (number of girls): 133 (71)		
	Age: mean (range): 8.1 years (1 month to 15 years)		
CNA, colistin-nalidixic acid; CLED, cystine lactose electrolyte-deficient; UA, urinalysis.			

Appendix 6

Included studies: further investigation

TABLE 41 Included studies of tests for the further investigation of confirmed UTI

Study details	Patient details	Reference standard	Index test
Alon, 1986 ¹¹⁴	Patient spectrum Confirmed UTI, further investigation	Reference standard IVP: examination preceded by plain abdominal and pelvic radiographs. 60% urographin injected i.v. at 1 ml kg ⁻¹ body weight in children and 2–3 ml kg ⁻¹ body weight in small infants. Supine radiographs of the kidneys and bladder were obtained at 5 and 10 minutes; if necessary, prone radiographs were added. Additional oblique and postvoiding radiographs or tomography were used when indicated	Test 1: MCUG: bladder filled to capacity with 30% Urographin under fluoroscopic visualisation. If passive reflux was observed during filling a radiograph was obtained. The full bladder was radiographed routinely. After catheter removal, the patient voided, and, if reflux was noted during voiding, a radiograph was taken, if not a postvoiding radiograph of the kidneys, ureters, bladder, and urethral areas was obtained Time between infection and test 1: 5–7 weeks Definition of positive result test 1: Reflux was graded 1–5 according to the international system ²⁸⁴
Study design Prospective cohort	Further details All children hospitalised or seen at an outpatient clinic, over approx. 11 months, because of confirmed symptomatic UTI, in whom radiological evaluation was indicated.		Test 2: Ultrasound: ultrasonography was performed first. Examination with a grey-scale static scanner and 3.5–5-MHz transducers, and a real-time sector scanner with a 5-MHz transducer. Longitudinal supine and prone scans, and transverse prone scans were performed to evaluate position, size and anatomy of the kidneys. For detection of renal scars the parenchymal width was measured along the length of each kidney and compared bilaterally. In addition, serial longitudinal scans were carried out with 0.5–1.0-cm intervals, with accentuation on the central echographic pattern, to detect double collecting systems Time between infection and test 2: <4 days Definition of positive result test 2: NR
Country Israel	Indications for radiological evaluation were: first documented UTI in boys; first documented UTI in girls aged <5 years; first infection presenting as severe clinical pyelonephritis in older girls; second infection in girls aged >5 years	Time between infection and reference standard: 5–7 weeks Definition of a positive test result NR	Test 3: Combination: VCUG and ultrasonography, as above Time between infection and test 3: As above Definition of positive result test 3: VCUG or ultrasound positive
Setting (teaching) Secondary care (teaching)	Recurrent UTI: NR		
Aim of imaging Detection of scarring	Number of patients (number of girls): 81 (68) (33 outpatients, 48 inpatients) Age: mean (range): 4 years (2 weeks to 12 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Alzen, 1994 ¹⁵ Study design Prospective cohort Country Germany Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux	Patient spectrum Mixed, some UTI Further details Consecutive children. Indications for imaging were: initial UTI ($n = 14$), recurrent UTI ($n = 2$), known reflux ($n = 4$), pathological ultrasound ($n = 15$), others ($n = 17$) Recurrent UTI: Combination Number of patients (number of girls): 52 (21) Age: mean (range): NR (<2 years)	Reference standard Reference standard MCUG: performed immediately after ultrasound. Bladder filled with contrast agent and standard MCUG performed Time between infection and reference standard: NR Definition of a positive test result Reflux graded according to international recommendations	Index test Test 1: Ultrasound: child catheterised and bladder emptied. Children lay on front. Bladder filled with air. Kidneys examined dorsally by standard ultrasound Time between infection and test 1: NR Definition of positive result test 1: NR
Andrich, 1992 ¹⁶ Study design Retrospective cohort Country USA Setting (teaching) Secondary care (NR) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children referred for evaluation with a clinical diagnosis of pyelonephritis. None had either obstructive uropathy or neuropathic bladder Recurrent UTI: NR Number of patients (number of girls): 72 (50) Age: mean (range): 1.6 years (1 month to 5.4 years)	Reference standard ^{99m} Tc-DMSA, no further details Time between infection and reference standard: NR Definition of a positive test result: NR	Index test Test 1: Ultrasound: renal ultrasound, no further details Time between infection and test 1: NR Definition of positive result test 1: NR Test 2: Cystography: no details Time between infection and test 2: NR Definition of positive result test 2: NR

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Bagni, 1997 ¹¹⁷	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA, planar: imaging 2–3 hours after i.v. 20–74 MBq ^{99m} Tc-DMSA. Image acquisition using standard posterior, right posterior oblique left posterior oblique and anterior views	Test 1: DMSA, SPECT: data obtained over 360° as 120 projections Time between infection and test 1: NR Definition of positive result test 1: Reduced uptake or well-defined scars
Study design Prospective cohort	Further details Patients with a history of recent ($n = 37$) or previous ($n = 11$) UTI, 80% of patients aged <5 years	Time between infection and reference standard: NR	
Country Italy	Recurrent UTI: Combination	Definition of a positive test result Reduced uptake or well-defined scars	
Setting (teaching) Secondary care (NR)	Number of patients (number of girls): 48 (36)		
Aim of imaging Detection of scarring	Age: mean (range): 3.2 years (0.1–17 years)		
Barnett, 1978 ¹¹⁸	Patient spectrum Confirmed UTI, further investigation	Reference standard Combination: IJU and MCUG	Test 1: Microscopy: antibody-coated bacteria tested for by immunofluorescence. Urine specimen centrifuged for 10 minutes, sediment washed with phosphate-buffered saline. 0.05 ml of undiluted FITC-conjugated rabbit anti-human IgA, IgG and IgM polyvalent serum added to 0.2 ml washed urine. Mixed and incubated at 27°C for 20 minutes. Sediment washed and examined under microscope. Blocking antibody and negative controls carried out
Study design Prospective cohort	Further details Children with Gram-negative UTI attending the paediatric genitourinary clinic and wards. Where more than one UTI was tested for in a patient the results from the first episode were used in the analysis	Time between infection and reference standard: NR	Time between infection and test 1: NR Definition of positive result test 1: Any fluorescence observed significant if both controls were negative
Country New Zealand	Recurrent UTI: Combination	Definition of a positive test result Detection of radiological abnormalities, no details provided	
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 69 (97 UTIs) (NR)		
Aim of imaging Localisation of UTI	Age: mean (range): NR (<1 month to >1 year)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Baronciani, 1986¹¹⁹</p> <p>Study design Retrospective cohort</p> <p>Country Italy</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children admitted to a paediatric department with UTI</p> <p>Recurrent UTI: Initial UTI</p> <p>Number of patients (number of girls): 98 (71)</p> <p>Age: mean (range): (0–14 years)</p>	<p>Reference standard</p> <p>Reference standard MCUG: no details reported</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result Reporting by criteria of Dwoskin <i>et al.</i>²⁸⁵</p>	<p>Test 1: Ultrasound: real-time ultrasound with 2.5–3.5-MHz probe, longitudinal and transverse scans with subject in prone position</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: classified as normal, moderate dilatation, hydronephrosis, severe hydronephrosis</p>
<p>Barry, 1998¹²⁰</p> <p>Study design Retrospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details In children following UTI (no details given)</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 300 (648 kidneys) (176)</p> <p>Age: mean (range): 3.89 years (18 days to 13.6 years)</p>	<p>Reference standard</p> <p>DMSA: scanning 3 hours post-injection, dose was fraction by body weight of 80 MBq (min. 18 MBq), posterior and left and right posterior oblique projections</p> <p>Time between infection and reference standard: > 3 months</p> <p>Definition of a positive test result NR</p>	<p>Test 1: Ultrasound: using 3.5, 5 or 7.5-MHz microconvex transducers. Prone and supine scans unless lack of cooperation indicated other approaches</p> <p>Time between infection and test 1: 1–3 months</p> <p>Definition of positive result test 1: Scarring assessed by following criteria: major: (1) proximity of sinus echoes to cortical surface; intermediate: (2) loss of pyramids, (3) irregularity of outline; minor: (4) loss of definition of capsular echo, (5) calyceal dilatation. The major criterion plus at least one intermediate criterion were necessary for the diagnosis of scarring</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Benador, 1994 ¹²¹ Study design Prospective cohort Country Switzerland Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children hospitalised with a clinical diagnosis of pyelonephritis Recurrent UTI: Combination Number of patients (number of girls): 111 (220 kidneys) (51) Age: median (range): 5.5 months (1 week to 16 years)	Reference standard Reference standard ^{99m} Tc-DMSA, dose 75 μ Ci kg^{-1} (min. 400 μ Ci, max. 2 mCi). Scans 3–4 hours postinjection. Six views (one posterior, two posterior oblique, one anterior, two anterior oblique) Time between infection and reference standard: Mean 3 days Definition of a positive test result Abnormal defined as focal or diffuse decrease or absence of DMSA uptake	Index test Test: Ultrasound: anterior, posterior, longitudinal and transverse views obtained of each patient Time between infection and test: Mean 3 days Definition of positive test result: Increase in renal size, loss of cortico-medullary differentiation, focal or generalised hyperchogenicity or hyposechogenicity, and parenchymal reduction were considered signs of pyelonephritis Test 1: For first or multiple UTI in patients of all ages Test 2: For first UTI in patients of all ages Test 3: For multiple UTI in patients of all ages Test 4: For first or multiple UTI in patients aged < 1 year Test 5: For first UTI in patients aged < 1 year Test 6: For multiple UTI in patients aged < 1 year Test 7: For first or multiple UTI in patients aged \geq 1 year Test 8: For first UTI in patients aged \geq 1 year Test 9: For multiple UTI in patients aged \geq 1 year
Benigno, 1986 ¹²² Study design Prospective cohort Country Italy Setting (teaching) Secondary care (teaching) Aim of imaging Anatomical	Patient spectrum Confirmed UTI, further investigation Further details Patients admitted with UTI over a 4-year period Recurrent UTI: NR Number of patients (number of girls): 49 (34) Age: mean (range): NR (15 days to 10 years)	Reference standard IVU and MCUG: no details reported Time between infection and reference standard: Cystography 8 weeks after urine becomes sterile Definition of a positive test result NR	Index test Test 1: ESR Time between infection and test 1: NR Definition of positive result test 1: 25 (no units reported) Test 2: CRP, semi-quantitative, latex agglutination test Time between infection and test 2: NR Definition of positive result test 2: Positive threshold 2+ Test 3: Leucocyte differential Time between infection and test 3: NR Definition of positive result test 3: Not clear Test 4: Body temperature Time between infection and test 4: NR Definition of positive result test 4: 38.5°C Test 5: Biochemical: renal concentrating capacity Time between infection and test 5: NR Definition of positive result test 5: 1025 (no units reported)

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Bergius, 1990 ¹²³ Study design Prospective cohort Country Finland Setting (teaching) Secondary care (NR) Aim of imaging Detection of reflux	Patient spectrum Mixed, some UTI Further details 124 consecutive infants and children referred for urological work-up, usually because of relapsing febrile UTI. Main indications for urological work-up were: recurrent UTI ($n = 106$), postoperative follow-up ($n = 10$), enuresis ($n = 5$), other ($n = 3$) Recurrent UTI: NR Number of patients (number of girls): 124 (248 ureters) (59) Age: mean (range): NR (NR; 101 <5 years)	Reference standard Reference standard MCUG: standard dynamic MCUG, immediately after ultrasound with the same catheter. Fluoroscopy was not used. To image the urethra and trigonum, one film was taken during voiding with the patient leaning forwards Time between infection and reference standard: NR Definition of a positive test result Reflux graded on international scale ²⁸⁴	Index test Test: Ultrasound: bladder emptied by catheter before examination. Both renal areas imaged with a 5-MHz transducer, special emphasis on thickness and structure of the renal parenchyma and on the shape of the pelvis and calyces. Bladder filled, according to size of patient, with Isopaque contrast medium and monitored during filling and voiding Time between infection and test: NR Definition of positive test result: The presence of air bubbles in the ureters was considered indicative of reflux and findings were graded according to the five-degree classification of Hofmann and Beyer ²⁸⁶ Definition of positive result test 1: Reflux grade \geq II Definition of positive result test 2: High-grade reflux: (grade \geq III)
Berrocal, 2001 ¹²⁴ Study design Prospective cohort Country Spain Setting (teaching) Secondary care (teaching) Aim of imaging Diagnosis of reflux	Patient spectrum Mixed, some UTI Further details Consecutive patients referred for suspected reflux. Patients unable to void spontaneously (e.g. because of spinal cord or CNS lesions) or who had neurogenic bladder were excluded. Clinical diagnoses included: UTI ($n = 88$), APN ($n = 120$), known reflux ($n = 64$), urinary incontinence ($n = 2$), structural abnormalities ($n = 33$), other ($n = 18$) Recurrent UTI: NR Number of patients (number of girls): 247 (216 patients and 440 kidneys included) (116) Age: mean (range): 3 years (3–18 years)	Reference standard Reference standard MCUG: performed immediately after ultrasound using the same catheter. The contrast medium Plenigraf (iodine concentration 16%) was infused. Intermittent fluoroscopy was performed, and the following abdominal radiographs were obtained: half and completely filled bladder, anteroposterior and oblique voiding, postvoiding, and voiding lateral in males Time between infection and reference standard: NR Definition of a positive test result Interpreted by one radiologist and later reinterpreted by consensus. Reflux graded by international scale ²⁸⁴	Index test Test: Ultrasound: preliminary ultrasound followed by cystosonography. Baseline ultrasound included transverse and longitudinal scanning of both kidneys and bladder, using 3.5 or 5-MHz sector array or 7.5-MHz linear transducer. Bladder filled, by infusion through catheter, to half volume with saline. 10 ml of SH U 508A enhancer (300 mg ml ⁻¹) was then injected and further saline infused until the bladder was full. Each kidney and bladder were imaged sequentially at approximately 15-second intervals during filling. Scans were obtained before filling, during filling and during voiding Time between infection and test: NR Definition of positive test result: reflux was diagnosed at cystosonography when microbubbles appeared in a ureter or renal pelvis. Reflux grading: grade 1: echo contrast in only the ureter grade 2: echo contrast in the pelvicalyceal system with no dilatation grade 3: mild to moderate dilatation of the pelvicalyceal system grade 4: moderate to severe dilatation of the pelvicalyceal system grade 5: gross dilatation of the pelvicalyceal system with total or partial atrophy Test 1: Results by renal unit Test 2: Results by patient

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Biggi, 2001 ¹²⁵			
Study design Prospective cohort	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: performed after i.v. injection of ^{99m} Tc-technetium, schedule based on body surface area (max. adult dose 110 MBq, min. dose 20 MBq). After injection (2–3 hours) three views (one posterior and two posterior oblique) were obtained	Test 1: CRP: no further details Time between infection and test 1: NR Definition of positive result test 1: >88 mg dl ⁻¹
Country Italy	Further details Children referred following a documented UTI associated with alterations in the urinary sediment (leucocyturia, bacteriuria, haematuria). Only children with first symptomatic UTI, in whom the interval between onset of symptoms and diagnosis of UTI was <6 days and in whom a DMSA scan was performed no later than 15 days following presentation were included	Time between infection and reference standard: <15 days Definition of a positive test result One or more areas of focal decreased cortical uptake, or diffuse areas of diminished cortical uptake of DMSA with no evidence of cortical loss. Involvement of each kidney graded as mild/moderate/severe	Test 2: ESR: no further details Time between infection and test 2: >68 mm h ⁻¹ Definition of positive result test 2: >68 mm h ⁻¹
Setting (teaching) Secondary care (NR)	Recurrent UTI: Initial UTI		Test 3: Microscopy: no further details: WBC count Time between infection and test 3: NR Definition of positive result test 3: >14,601 WBC mm ⁻³
Aim of imaging Localisation of UTI	Number of patients (number of girls): 101 (60) Age: mean (range): 18 months (1 month to 13.5 years)		Test 4: Microscopy: no further details: granulocytes Time between infection and test 4: NR Definition of positive result test 4: >52% Test 5: Clinical features: maximum rectal temperature Time between infection and test 5: NR Definition of positive result test 5: ≥39.1°C
			Test 6: Ultrasound: performed on computerised sonographic units equipped with 3.5 and 5-MHz transducers, results for number of kidneys Time between infection and test 6: NR Definition of positive result test 6: Abnormal echogenicity or dilatation of collecting system
Bircan, 1995 ¹²⁶			
Study design Prospective cohort	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: performed 2–3 hours after i.v. injection of 2 MBq kg ⁻¹ ^{99m} Tc-DMSA; anterior, posterior, right lateral and left lateral images were obtained	Test 1: Ultrasound: abdominal ultrasound using 5-MHz linear probe Time between infection and test 1: <48 hours Definition of positive result test 1: Kidneys examined for presence of congenital anomalies, renal scarring and for changes related to acute infection (changes in renal parenchymal echogenicity)
Country Turkey	Further details 18 patients had a previous UTI or an episode of "fever of unknown origin"	Time between infection and reference standard: <48 hours Definition of a positive test result Focal areas of diminished uptake in the renal parenchyma	Test 2: IVU: sodium meglumine diatrizoate (Urografin) used as contrast material Time between infection and test 2: <48 hours Definition of positive result test 2: Anatomical pathologies related to the urinary system were evaluated
Setting (teaching) Secondary care (teaching)	Recurrent UTI: Combination		
Aim of imaging Localisation of UTI	Number of patients (number of girls): 63 (56) Age: mean (range): 5 years (3 months to 12 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Boudailliez, 1998¹²⁷</p> <p>Study design Prospective cohort</p> <p>Country France</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children with their first episode of UTI; temperature > 38.5°C, CRP > 20 mg l⁻¹, no urinary tract malformation and no reflux grade ≥ 3 as assessed by mode B sonography and cystography (to exclude previous lesions of renal parenchyma)</p> <p>Recurrent UTI: Initial UTI</p> <p>Number of patients (number of girls): 49 (32)</p> <p>Age: mean (range): 4.2 years (0.2–14 years)</p>	<p>Reference standard</p> <p>Reference standard DMSA: 50 µCi kg⁻¹, min. dose 350 µCi</p> <p>Time between infection and reference standard: < 5 days</p> <p>Definition of a positive test result NR</p>	<p>Test 1: Ultrasound: power Doppler renal sonography (5–7.5-MHz probe)</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: NR</p>
<p>Bower, 1985¹²⁸</p> <p>Study design Prospective cohort</p> <p>Country Australia</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Children referred to a nuclear medicine department for assessment of renal and ureteric function as part of the follow-up of reflux, or recurrent UTI</p> <p>Recurrent UTI: Recurrent UTI</p> <p>Number of patients (number of girls): 30 (27 children, 54 ureters included in analysis) (22)</p> <p>Age: mean (range): NR (4 months to 12 years)</p>	<p>Reference standard</p> <p>Reference standard Direct radionuclide cystography: ^{99m}Tc-DTPA renal scan and a delayed voiding cystogram. Voiding for cooperative children was allowed in the upright sitting position viewed posteriorly. Bolus of ^{99m}Tc-sulfur colloid (37 MBq) was injected in the catheter at the start of bladder filling. Analogue and digital acquisition was in 15-second frames for 14 frames followed by 30-second analogue image when the bladder was full before catheter removal and voiding. 60 5-second voiding frames were collected and a 30-second postvoid analogue image was obtained</p> <p>Time between infection and reference standard: Day after indirect cystogram</p> <p>Definition of a positive test result Images scored for reflux, no attempt to grade reflux</p>	<p>Test 1: Indirect radionuclide cystography: performed when 80% of the intravenously administered ^{99m}Tc-DTPA from renal study was in the bladder and the child expressed a strong desire to void. The study was then identical to that of the direct cystogram</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: Indirect playback loops read independently by three observers unaware of patients' identity or other investigation results. A consensus score for each ureter as positive or negative for reflux was given</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Buyan, 1993 ¹²⁹ Study design Prospective cohort Country Turkey Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details None of the children had had previous UTI or "fever of unknown origin" Recurrent UTI: Initial UTI Number of patients (number of girls): 24 (20) Age: mean (range): 6 years (13 months to 12 years)	Reference standard Reference standard DMSA: performed 2–3 hours following i.v. injection of 2 MBq kg ⁻¹ of ^{99m} Tc-DMSA; anterior, posterior, right lateral and left lateral images were obtained. 3D images also obtained using SPECT Time between infection and reference standard: Before treatment Definition of a positive test result Focal areas in the renal parenchyma showing no radioactivity	Test 1: Clinical features: flank pain, chills, nausea, vomiting, high fever (> 38°C), tenderness of the costovertebral angle diagnosed as upper UTI. Those with dysuria, enuresis, pollakiuria, abdominal pain, haematuria, fever (≤38°C) diagnosed as lower UTI Time between infection and test 1: On admission Definition of positive result test 1: Upper UTI Test 2: WBC: no details reported Time between infection and test 2: On admission Definition of positive result test 2: > 15,000 WBC cm ⁻³ Test 3: ESR: no details reported Time between infection and test 3: On admission Definition of positive result test 3: > 25 mm h ⁻¹ Test 4: CRP: single radial immunodiffusion, Behring, LC-Partigen Time between infection and test 4: On admission Definition of positive result test 4: > 20 µg l ⁻¹ Test 5: ACB: prepared and detected as described by Thomas <i>et al.</i> ²⁸⁷ Time between infection and test 5: On admission Definition of positive result test 5: > 2 ACB per 200 fields or in 5 minutes of scanning
Capa Kaya, 2001 ¹³⁰ Study design Prospective cohort Country Turkey Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children with symptoms of upper UTI and positive urine culture Recurrent UTI: Initial UTI Number of patients (number of girls): 100 (82) Age: mean (range): 5 years (0–12 years)	Reference standard DMSA: 0.03–0.05 µCi kg ⁻¹ ^{99m} Tc-DMSA administered intravenously; anterior, posterior and right and left posterior oblique views were obtained Time between infection and reference standard: 3 days Definition of a positive test result Presence of focal or diffuse decreased uptake, segmental cortical thinning and cortical irregularity for upper UTI	Test 1: NAG activity was measured with a kit (Boehringer Mannheim) using a spectrophotometric method, NAG/creatinine ratios Time between infection and test 1: Performed at time of infection Definition of positive result test 1: 5 U l ⁻¹ for NAG and 7 U g ⁻¹ for NAG creatinine ratio for upper UTI

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Cavanagh, 1983 ¹³¹	Patient spectrum Confirmed UTI, further investigation	Reference standard MCUG: bladder catheterisation without sedation immediately before X-ray, bladder filling using contrast medium sufficient to induce voiding	Test: Early IVU films available for study of the renal parenchyma on the nephrogram. Contrast medium was sodium meglumine diatrizoate Time between infection and test: within 3 weeks of MCUG
Study design Prospective cohort	Further details Children referred to X-ray department for first MCUG and IVU because of provisional or proven diagnosis of UTI.	Time between infection and reference standard: not clear	Definition of positive result test 1: Graded as normal or suspect: duplex, scarred moiety, dilated ureter, scarred kidney Definition of positive result test 2: Grade 3 reflux on reference standard considered positive, all others considered negative
Country England	41/62 had proved UTIs, the remainder were suspected UTI but had no laboratory confirmation. 11 had suffered one attack of symptoms and the remainder had multiple infections.	Definition of a positive test result Reflux graded as follows: grade 1: slight – incomplete filling of the upper urinary tract without dilatation grade 2: moderate – complete filling of the upper urinary tract with slight dilatation but no ballooning of calices grade 3: gross – complete filling of the upper urinary tract with marker dilatation and obvious ballooning of calices. Any reflux considered abnormal	
Setting (teaching) Secondary care (teaching)	Children aged <6 months were excluded. Children with an obvious anomaly of renal drainage on the IVU, e.g. ectopic ureterocele, congenital megaureter, were excluded. Referrals came from paediatric and urological clinics		
Aim of imaging Detection of reflux	Recurrent UTI: Combination Number of patients (number of girls): 62 (48) Age: mean (range): 5 years (6 months to 13 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Chan, 1999 ¹³² Study design Prospective cohort Country Hong Kong Setting (teaching) Secondary care (teaching) Aim of imaging Detection of scarring	Patient spectrum Confirmed UTI, further investigation Further details Children with increased susceptibility to UTI due to spinal dysraphism and neurogenic bladder ($n = 18$), anorectal anomaly ($n = 6$). Interval between last UTI and current UTI varied from 2 months to 2 years. No patient had a history of APN. Ten children had documented previous UTI Recurrent UTI: Recurrent UTI Number of patients (number of girls): 24 (48 kidneys) (11) Age: mean (range): 5.3 years (1–13 years)	Reference standard Reference standard DMSA, ^{99m} Tc-DMSA scans performed 2–3 hours after i.v. injection in dose range 38–120 MBq by body weight. Simultaneous planar anterior and posterior, right posterior oblique and left anterior oblique, left posterior oblique and right anterior oblique images obtained Time between infection and reference standard: 2 months to 2 years Definition of a positive test result Presence or absence of renal scarring (not defined)	Index test Test: MRI: 1.5 T with spine coil, field of view 300–375 mm, matrix 256 × 256, slice thickness 4 mm Time between infection and test: 2 months to 2 years Definition of positive test result: Presence or absence of renal scarring (not defined) Test 1: Coronal, postgadolinium STIR sequence (TR/TE/TI, 2000/17/160) Test 2: Coronal, fat-saturated, spin-echo T1-W sequence (TR/TE, 550/15) Test 3: MRI: STIR or T1-W
Clarke, 1990 ¹³³ Study design Prospective cohort Country England Setting (teaching) Secondary care (teaching) Aim of imaging Detection of scarring	Patient spectrum Confirmed UTI, further investigation Further details Children with known UTI Recurrent UTI: NR Number of patients (number of girls): 22 (NR) Age: mean (range): NR (4–12 years)	Reference standard IVU, no details Time between infection and reference standard: NR Definition of a positive test result No details	Index test Test 1: DMSA: planar ^{99m} Tc-DMSA imaging Time between infection and test 1: NR Definition of positive result test 1: NR Test 2: DMSA: SPECT ^{99m} Tc-DMSA imaging Time between infection and test 2: NR Definition of positive result test 2: NR

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Dacher, 1996 ¹³⁴ Study design Prospective cohort Country France Setting (teaching) Secondary care (NR) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children referred because of upper UTI. Criteria for inclusion were: confirmed UTI, high fever (> 38°C), abdominal or lumbar fossa pain, and presence of urinary nitrites and LE as determined by dipstick. Uncooperative children, children with obstructed urinary tracts or pyelonephrosis, and those who had undergone surgery were excluded Recurrent UTI: NR Number of patients (number of girls): 30 (24) Age: mean (range): 7.5 years (7 months to 15 years)	Reference standard Enhanced CT: 10-mm slices without interslice intervals were obtained immediately after infusion of ionic iodinated contrast medium (1 ml kg ⁻¹ body weight) Time between infection and reference standard: Within 24 hours of admission Definition of a positive test result Sharply bordered triangular areas of decreased attenuation in the renal parenchyma that involved both the medulla and the cortex or rounded masses of low attenuation that modified renal contours were considered indicative of APN	Test 1: Ultrasound: power Doppler sonography performed with a 5-MHz curved array with patients in the prone position. Axial and longitudinal scans were obtained for each renal pole Time between infection and test 1: within 24 hours of admission Definition of positive result test 1: Abnormality was defined by the presence of a triangular zone of decreased or absent flow in the parenchyma. Each kidney was divided into two poles by a vertical line perpendicular to the major axis
De Sadeleer, 1994 ¹³⁵ Study design Retrospective cohort Country Belgium Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux and scarring	Patient spectrum Confirmed UTI, further investigation Further details Patients with UTI referred for imaging who underwent indirect radionuclide cystography during the acute phase of the UTI and MCUG 6 weeks later Recurrent UTI: NR Number of patients (number of girls): 40 (80 kidneys) (24) Age: mean (range): median 4.5 years (2 weeks to 11 years)	Reference standard test 1 MCUG: performed after catheterisation. Bladder emptied and 18% iodinated contrast material run into bladder until bladder capacity reached. Some images obtained during this phase. Spontaneous micturition occurred when the bladder was full. Images obtained by intermittent fluoroscopy Time between infection and reference standard: 6 weeks for MCUG, acute or postacute phase for DMSA Definition of a positive test result: Severity of reflux graded according to the international classification ²⁸⁴ Reference standard tests 2, 3 DMSA scintigraphy performed 2 hours after i.v. injection of tracer at dose of 0.75 MBq kg ⁻¹ (minimum dose 17 MBq); one posterior (supine position) and two oblique posterior views performed	Test 1: Indirect radionuclide cystography: performed after ^{99m} Tc-MAG3 renography, the tracer dose was scaled according to body surface area. In cooperative children above 3–4 years old voiding cystography was performed approximately 2 hour after i.v. injection of the tracer. A dynamic series of images using a 5-second frame rate was acquired using a gamma camera at least 30 seconds before the child was allowed to void. Uncooperative children only imaged if there was enough computer time and gamma camera time. Analogue images were reviewed one by one. Time between infection and test 1: Acute phase Definition of positive result test 1: Any increase in ureteral or renal activity Test 2: MCUG: as for reference standard 1 Time between infection and test 2: 6 weeks Definition of positive result test 2: Severity of reflux graded according to international classification ²⁸⁴ Test 3: As for test 1 Time between infection and test 3: Acute phase Definition of positive result test 3: Any increase in ureteral or renal activity

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Ditchfield, 1994 ¹³⁶	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: performed according to standard technique including high-resolution tomography. Either ^{99m} Tc-gluconate or ^{99m} Tc-DMSA was administered. Images were obtained after 1 hour for gluconate and 3 hours for DMSA. One posterior view delayed image of the kidneys was acquired, followed by posterior tomographic acquisition of 32 steps each for 30 seconds	Test: MCUG; no details Time between infection and test: Performed within 15 days of start of treatment Definition of positive test result: Classified according to international classification. Reflux grades 0–1 considered negative, 2–5 considered positive
Study design Prospective cohort	Further details Consecutive patients aged <5 years with a first proven UTI	Time between infection and reference standard: Performed within 15 days of start of treatment	Test 1: Result: per kidney Test 2: Result per patient
Country Australia	Recurrent UTI: Initial UTI	Definition of a positive test result Graded on scale of 0–3: grade 0: no defect grade 1: possible defect grade 2: probably defect grade 3: definite defect	
Setting (teaching) Secondary care (NR)	Number of patients (number of girls): 150 (300 kidneys) (82)	Scans of grades 2–3 considered positive. When more than one defect present, grade represented the most severe defect	
Aim of imaging Detection of scarring	Age: mean (range): NR (<5 years)		
Drachman, 1984 ¹³⁷	Patient spectrum Confirmed UTI, further investigation	Reference standard MCUG, NR	Test 1: IVP; no details reported Time between infection and test 1: 4–6 weeks Definition of positive result test 1: NR
Study design Prospective cohort	Further details Children who underwent IVP and MCUG 4–6 weeks after UTI	Time between infection and reference standard: 4–6 weeks	
Country Israel	Recurrent UTI: Initial UTI	Definition of a positive test result Reflux, graded as follows: grade I: reflux limited to the ureter grade II: reflux reaching the pelvis and calyces without dilatation of the collecting system grade III: complete reflux with some distension of ureter, pelvis and calyces grade IV: reflux associated with massive dilatation of ureter, pelvis and calyces.	
Setting (teaching) Secondary care (NR)	Number of patients (number of girls): 191 (177)	All grades of reflux considered abnormal. Also abnormal if ureteropelvic junction stenosis, atrophic kidney or anatomy of the collecting system	
Aim of imaging Detection of reflux	Age: mean (range): NR (6 months to 12 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Elison, 1992¹³⁸</p> <p>Study design Prospective cohort</p> <p>Country Australia</p> <p>Setting (teaching) Secondary care (non-teaching)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Patients referred by a paediatric nephrologist and urologist. Patients with bacteriological evidence of recent UTI were excluded. Patients had experienced their last document UTI at least 3 months previously</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 208 (415 kidneys) (139)</p> <p>Age: mean (range): 3.7 years (3 months to 12 years)</p>	<p>Reference standard</p> <p>Reference standard IVU: performed with non-ionic contrast medium (Omnipaque 240) at a dose of 1.5–2 ml kg⁻¹ (min. 10 ml, max. 40 ml). Each study comprised four plain films with no additional tomography of the kidneys</p> <p>Time between infection and reference standard: NR (min. 3 months)</p> <p>Definition of a positive test result Reflux graded according to internal criteria.²⁸⁴ Renal scarring reported as occurring in the upper, middle and/or lower poles of each kidney. Criteria for scarring included: disruption of the cortical rim of uptake seen in the pinhole images, clearly defined defects in uptake on emission computed scans; significant variation in the left:right relative function was also considered. Evaluated by three nuclear medicine physicians, consensus of at least two required for classification, otherwise image discarded</p>	<p>Test: DMSA: children who were unable to cooperate or who were aged <18 months old were sedated. DMSA dose dependent on age (min. 40 MBq). Imaging performed 3 hours after injection of ^{99m}Tc-DMSA in the supine position. Normal planar images were acquired in the anterior and posterior projections for 5 minutes per image using a high-resolution collimator. SPECT was undertaken in 76% of patients</p> <p>Time between infection and test: NR (min. 3 months)</p> <p>Definition of positive test result: Renal scarring reported as occurring in the upper, middle and/or lower poles of each kidney, no further details</p> <p>Test 1: Results by number of patients Test 2: Results by number of kidneys</p>
<p>Evans, 1999¹³⁹</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Children known to be undergoing both renal ultrasound and MCUG on the same day were eligible for inclusion. Only children with a full bladder who voided during the course of the ultrasound examination were included. Indications for referral were: UTI (n = 51), antenatal hydronephrosis (n = 3), incontinence (n = 1), unknown (n = 2)</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 57 (113 kidneys) (48)</p> <p>Age: mean (range): 5 years (1 month to 16 years)</p>	<p>Reference standard MCUG: fluoroscopic voiding cystourethrogram, performed on the same day as the preceding ultrasound</p> <p>Time between infection and reference standard: No details</p> <p>Definition of a positive test result VUR graded on international scale²⁸⁴</p>	<p>Test 1: Ultrasound: measurements made on widest section of each renal pelvis on transverse mid-kidney images. Patients were imaged before voiding, either supine or prone, and after voiding in the same position</p> <p>Time between infection and test 1: No details</p> <p>Definition of positive result test 1: A difference of 1 mm or more on pre- and postvoiding images was considered a change in pelvic diameter, and evidence of reflux</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Everaert, 1998¹⁴⁰</p> <p>Study design Retrospective cohort</p> <p>Country Belgium</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children referred to the clinic for a UTI, had received a kidney ultrasound and DMSA scan within 4 days after onset of treatment and could be administered a second DMSA scan at least 3 months later for control. Children were excluded if they were aged <3 months, had proven hydronephrosis, if DMSA lesions compatible with scars were seen on a previous DMSA scan or on the follow-up scan, if it was impossible to decide from scans whether acute lesions were present, or if overt renal failure was present</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 62 (43)</p> <p>Age: mean (range): 4 years (NR)</p>	<p>Reference standard</p> <p>Reference standard DMSA: performed after hyperhydration of the patients. A salt and glucose solution was infused intravenously at 20 ml kg⁻¹ over 30 minutes. Patients were injected with 185 mBq 1.73 m⁻² of freshly prepared ^{99m}Tc-DMSA. Renal activity on both kidneys 24 hours after the tracer injection was investigated</p> <p>Time between infection and reference standard: Within 4 days of therapy</p> <p>Definition of a positive test result To exclude pre-existing scars the ultimate diagnosis of APN was made retrospectively by comparing the DMSA images during and after the acute disease</p>	<p>Index test</p> <p>Test 1: Urinary α_1-MG: urinary α_1-MG detected by latex enhanced nephelometry. Concentration expressed as the urinary α_1-MG: creatinine ratio</p> <p>Time between infection and test 1: On admission</p> <p>Definition of positive result test 1: > 10 mg g⁻¹</p> <p>Test 2: Clinical features: clinical symptoms</p> <p>Time between infection and test 2: On admission</p> <p>Definition of positive result test 2: Symptoms of APN positive, symptoms of cystitis or atypical symptoms negative</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Farnsworth, 1991⁴¹</p> <p>Study design Prospective cohort</p> <p>Country Australia</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Consecutive patients undergoing urological assessment for: UTI ($n = 86$), family history of reflux ($n = 18$), mild urinary tract dilatation detected on antenatal ultrasound ($n = 4$), imperforate anus ($n = 2$), haematuria ($n = 1$), retention ($n = 1$), painful voiding ($n = 1$). The indications for performing the DMSA study were UTI associated with high ($> 38.5^{\circ}\text{C}$) and persistent (> 2 days) fever and/or severe systemic upset, or grade 3–5 reflux, according to international classification. Only infants with primary reflux were included</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 113 (224 kidneys) (53)</p> <p>Age: mean (range): NR (< 1 year)</p>	<p>Reference standard Excretory urography (IVP)</p> <p>Reference standard execution IVP after 4 hours prep of clear fluids. Non-ionic contrast medium (Omnipaque 240), 10 mL kg⁻¹ in children < 4 kg and 1.5–2 kg⁻¹ in children > 4 kg. Films taken at end of injection, 10 minutes in prone position and after micturition</p> <p>Time between infection and reference standard: Not clear</p> <p>Definition of a positive test result Decrease in parenchymal thickness as defined by a focal decrease in the distance from the renal outline to the renal calyces and a generalised decrease in renal size and/or abnormal calyces, particularly medial deviation of the upper pole calyx. Accessory signs such as renal tract dilatation and striation of the renal pelvis were not considered</p>	<p>Test 1: DMSA 3 hours after i.v. injection of 40 MBq ^{99m}Tc-DMSA. Posterior planar view</p> <p>Time between infection and test 1: < 1 month to > 3 months</p> <p>Definition of positive result test 1: Reduction in kidney function and/or decreased or absent uptake of tracer in the renal cortex causing distortion or indentation of the normal renal outline</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Foresman, 2001 ¹⁴² Study design Retrospective cohort Country USA Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux	Patient spectrum Confirmed UTI, further investigation Further details Children hospitalised with a clinical diagnosis of APN, in whom renal and bladder ultrasound had been performed and whose charts were available for review. Children with known genitourinary pathology, neurological disease or concomitant pregnancy were excluded from the analysis Recurrent UTI: NR Number of patients (number of girls): 209 (123) Age: mean (range): 2.4 years (2.1 days to 18 years)	Reference standard Reference standard MCUG including fluoroscopic control, gravity filling, and filling and voiding views to assess the urethra, bladder wall and evidence of reflux Time between infection and reference standard: NR Definition of a positive test result NR	Index test Test 1: Ultrasound: Duplex ultrasound with 5 or 7.5-MHz probes. Sequential examination of kidneys and bladder before and after voiding if feasible Time between infection and test 1: NR Definition of positive result test 1: Renal size, presence of hydronephrosis and hydroureter, renal scarring, parenchymal character, calculi and bladder wall thickening assessed. Radiologist reimaged questionable areas and provided an official report
Fretzayas, 2000 ¹⁴³ Study design Prospective cohort Country Greece Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children with a first symptomatic confirmed UTI were included regardless of the presence of fever. Children were included if there was no previous history of urinary tract obstruction, other chronic inflammatory condition or other current infectious disease Recurrent UTI: Initial UTI Number of patients (number of girls): 83 (54) Age: mean (range): 3.8 years (2.5 days to 14 years)	Reference standard DMSA: dose of 5 MBq kg ⁻¹ ^{99m} Tc-DMSA (min. 10 MBq) administered i.v. At least 4 hours after injection images were obtained in anterior, posterior and oblique views. Pinhole views used to image areas of particular interest Time between infection and reference standard: <96 hours Definition of a positive test result APN diagnosed when scintiscan revealed focal or diffuse areas of diminished uptake with an intact or slightly bulging contour	Index test Test 1: Body temperature Time between infection and test 1: On admission Definition of positive result test 1: ≥38°C Test 2: CRP: no details Time between infection and test 2: On admission Definition of positive result test 2: 20 mg dl ⁻¹ Test 3: ESR: no details Time between infection and test 3: Measured on admission. No details Definition of positive result test 3: 30 mm h ⁻¹ Test 4: Polymorphonuclear elastase-α ₁ -antitrypsin complex: measured in EDTA-plasma and in urine supernatant, both obtained by centrifugation at 2000 g for 10 minutes within 1 hour of collection. Measurement was performed with an ELISA according to the manufacturers instructions Time between infection and test 4: On admission Definition of positive result test 4: 95th percentile of reference range (figure not reported) Test 5: MCUG: obtained by standard radiographic technique Time between infection and test 6: median 17 days after treatment Definition of positive result test 6: reflux graded according to international recommendations

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Frutos, 2000¹⁴⁴</p> <p>Study design Prospective cohort</p> <p>Country Spain</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Children suspected of reflux. Children who did not receive both imaging modalities were excluded. 68 children had previously diagnosed reflux</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 169 (144 included in the analysis, 293 renal units) (78)</p> <p>Age: mean (range): 3 years 9 months (3 days to 18 years)</p>	<p>Reference standard</p> <p>Reference standard MCUG: contrast media infused using catheter used for ultrasound, same volume of contrast agent was used. The complete urinary tract was imaged using a frontal projection. Images were taken of the full bladder, during micturition, after micturition and, in boys, lateral plates were taken during micturition</p> <p>Time between infection and reference standard: Not clear, performed during same diagnostic session as ultrasound</p> <p>Definition of a positive test result Graded according to international classification²⁸⁴</p>	<p>Index test</p> <p>Test 1: Ultrasound: standard ultrasound followed by cystosonography. Bladder filled with Levograf (an echo-enhancing agent). Ultrasonographic images were obtained during bladder filling and micturition</p> <p>Time between infection and test 1: Not clear</p> <p>Definition of positive result test 1: On standard ultrasound kidneys were evaluated for position, size and morphology, and for changes in the parenchyma or dilatation of excretory paths. Special attention was paid to dilatation of the distal ureter and the ureterovesical junction. Reflux was diagnosed on cystosonography whenever bubbles of Levograf were found above the bladder, outside the ureter or in the renal pelvis</p>
<p>Gervais, 2001¹⁴⁵</p> <p>Study design Prospective cohort</p> <p>Country Switzerland</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children attending the emergency unit at a children's hospital with fever (rectal temperature > 38°C), signs and symptoms suggestive of a UTI and/or a positive urine dipstick analysis (positive LE and/or nitrite) were consecutively enrolled. Children who received antibiotics in the previous week were excluded from the study. Children with a positive urine culture were included</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 54 (36)</p> <p>Age: mean (range): NR (1 week to 16 years)</p>	<p>Reference standard DMSA, no further details</p> <p>Time between infection and reference standard: 3–5 days</p> <p>Definition of a positive test result APN diagnosed if focal or diffuse ^{99m}Tc-DMSA uptake was noted on at least two projections</p>	<p>Test 1: PCT: Lumitest PCT (Brahms Diagnostica, Berlin, Germany) and Brahms PCT-Q (Brahms Diagnostica). Measured quantitatively in EDTA-blood samples by an immunoluminometric assay. PCT values were also determined by a rapid semi-quantitative immunochromatographic test. Both performed according to manufacturer's instructions</p> <p>Time between infection and test 1: On admission</p> <p>Definition of positive result test 1: PCT-Q ≥ 0.5 ng ml⁻¹ (cut-off determined by ROC analysis)</p> <p>Test 2: CRP: NyoCard CRP (Nycomed Pharma, Oslo, Norway) CRP measured in EDTA-blood samples by a rapid immunometric method following the manufacturer's instructions</p> <p>Time between infection and test 2: On admission</p> <p>Definition of positive result test 2: CRP ≥ 40 mg dl⁻¹</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Girona, 1995 ¹⁴⁶ Study design Prospective cohort Country Spain Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children aged <2 years with symptomatic UTI. Patients with obstructive uropathy or secondary reflux were excluded Recurrent UTI: Initial UTI Number of patients (number of girls): 85 (170 kidneys) (46) Age: mean (range): NR (15 days to 2 years)	Reference standard ^{99m} Tc-DMSA i.v. administered at a dose calculated according to standard guidelines, min. dose 15 MBq, 3 hours after administration; posterior, left oblique posterior and right oblique posterior images were obtained Time between infection and reference standard: Not clear Definition of a positive test result APN diagnosed if differential renal function was <48–52%, non-symmetrical renal size, defined as focal or multifocal perfusion defects	Test 1: Ultrasound: renal ultrasound performed with 5-MHz transducers. Longitudinal and transverse images were obtained with the patient in prone and supine positions Time between infection and test 1: NR Definition of positive result test 1: Kidney size was compared with tables of age-standardised kidney Test 2: MCUG: no details Time between infection and test 2: NR Definition of positive result test 2: Graded from I to V according to international classification. Grade ≥ II positive Test 3: CRP: no details Time between infection and test 3: no details Definition of positive result test 3: > 2 mg dl ⁻¹
Gordon, 1992 ¹⁴⁷ Study design Prospective cohort Country UK Setting (teaching) Secondary care (teaching) Aim of imaging Detection of scarring	Patient spectrum Confirmed UTI, further investigation Further details Children referred 12–14 weeks after a UTI Recurrent UTI: NR Number of patients (number of girls): 59 (110 kidneys) (31) Age: median (range): 5.3 years (0.1–14 years)	Reference standard DMSA scintigraphy 6 hours after ^{99m} Tc-DMSA injection (dose scaled by body surface area to adult dose of 100 MBq). Posterior and two posterior oblique views acquired by gamma camera in 256 × 256 matrix Time between infection and reference standard: 12–14 weeks Definition of a positive test result A kidney was considered abnormal if differential function on ^{99m} Tc-DMSA was <43% and/or if a focal defect was seen	Test 1: Scintigraphy: ^{99m} Tc-MAG3 study carried out with child supine on gamma camera dynamic 20-minute acquisition in 10-second frames. Dose scaled on body surface area to adult dose of 80 or 112 MBq Time between infection and test 1: 12–14 weeks Definition of positive result test 1: NR

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Guermazi, 1993¹⁴⁸</p> <p>Study design Retrospective cohort</p> <p>Country Belgium</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Children who underwent DMSA, including those with a doubtful or negative urine culture. The locations of the infection before DMSA was made based on clinical and biological features. 60/166 children were classed as having a lower UTI and 27 as having an upper UTI; in the remaining 79 the diagnosis was 'non-specific', the urine culture was negative or doubtful</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 166 (92)</p> <p>Age: mean (range): 5 years (15 days to 17 years)</p>	<p>Reference standard</p> <p>Reference standard DMSA: performed 2 hours after i.v. injection of ^{99m}Tc-DMSA at a dose of 0.74 MBq kg⁻¹. One posterior and two obliques posterior: images were taken</p> <p>Time between infection and reference standard: Not clear</p> <p>Definition of a positive test result Images were classed as abnormal based on two scintigraphic features: acute lesions or chronic lesions (definitions provided)</p>	<p>Index test</p> <p>Test 1: Ultrasound: 5-MHz convex</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: Suspicion of inflammation (any type): focal hypoechoic lesion, size anomaly, irregularity in the contours or aberrant differentiation of the corticomedulla</p>
<p>Haberlik, 1997¹⁴⁹</p> <p>Study design Prospective cohort</p> <p>Country Austria</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details 102 children in whom radiological evaluation by MCUG was indicated. Indications for MCUG included: UTI (<i>n</i> = 37), follow-up of previous reflux (<i>n</i> = 21), enuresis (<i>n</i> = 13), UTI and enuresis (<i>n</i> = 3), dysuria (<i>n</i> = 2), haematuria (<i>n</i> = 1)</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 102 (77 included in analysis, 154 kidneys (59))</p> <p>Age: mean (range): 7.3 years (7 months to 14 years)</p>	<p>Reference standard</p> <p>Reference standard MCUG: performed immediately after second conventional ultrasound following emptying of bladder via catheter</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result Reflux graded according to standard criteria²⁸⁴</p>	<p>Index test</p> <p>Test 1: Ultrasound: colour Doppler imaging-mode cystography with 5 and 3.5-MHz linear and sector transducers used. Catheterisation performed using standard aseptic procedures. Residual urine removed from bladder, patient placed supine for rest of study. Bladder filled by drip infusion using room-temperature saline solution. Both distal ureters and ureterovesical junction were alternately scanned by sagittal and oblique sections. A second conventional scan of the kidneys at the end of filling was made for detection of possible pelvicalyceal dilatation or urine-filled pyelon</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: Reflux defined as visualisation of a blue-coloured jet from the ureterovesical junctions to the distal ureter during the course of bladder filling or of pelvicalyceal dilatation or urine-filled pyelon on conventional ultrasound</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Hajjar, 2002 ¹⁵⁰	Patient spectrum Confirmed UTI, further investigation	Reference standard ^{99m} Tc-DMSA scan 6 hours postinjection (50 μ Ci kg^{-1} body weight)	Test 1: Ultrasound: B-mode sonography with 7.5-MHz transducer used for baseline examination to measure renal size and identify uropathies, 5–10-MHz linear transducer to study renal structure. Doppler sonography to map the intrarenal vessels. Each kidney studied with the patient supine and prone, in the transverse and sagittal planes
Study design Prospective cohort	Further details Patients admitted to a paediatric nephrology unit, over an 18-month period, with clinical and biological suspicion of APN	Time between infection and reference standard: <2 days	Time between infection and test 1: < 2 days
Country France	Recurrent UTI: NR	Definition of a positive test result Evidence of images characteristic of APN	Definition of positive result test 1: criteria for positive ultrasound: increased global renal volume in relation to diameter (ratio 1.4 significant); focal hyper- or hypoechogenicity, with changes in renal parenchymal echogenicity and triangular forms; diffuse hyperechogenicity, differential echogenicity between kidneys; abnormal renal vasculature on Doppler examination, evidence of triangular zones of hypovascularisation on more than two occasions
Setting (teaching) Secondary care (NR)	Number of patients (number of girls): 49 (NR)		
Aim of imaging Localisation of UTI	Age: median (range): 3 years (3 months to 15 years)		
Hanbury, 1989 ¹⁵¹	Patient spectrum Mixed, some UTI	Reference standard MCUG and IVU	Test 1: Combination: ultrasound and KUB X-ray
Study design Prospective cohort	Further details Children investigated for suspected UTI who received ultrasound and plain X-ray of the kidneys, ureters and bladder as the initial investigation. Presenting symptoms included dysuria, frequency, abdominal pain, enuresis, daytime wetting, fevers, loin pain, haematuria, urgency, vaginal soreness, difficult voiding and balanitis.	Reference standard execution No details reported. Only performed if child had abnormal ultrasound/X-ray or if child presented with a further UTI (parents followed up after 1–8 years to enquire about further symptoms of UTI)	Time between infection and test 1: Not clear
Country England	80% of children had two or more symptoms; 39% had firm evidence of UTI (10^5 cfu ml^{-1} and >50 pus cells mm^{-3})	Time between infection and reference standard: NR	Definition of positive result test 1: No details
Setting (teaching) Secondary care (teaching)	Recurrent UTI: NR	Definition of a positive test result No details	
Aim of imaging Detection of reflux and scarring	Number of patients (number of girls): 332 (309 included) (173)		
	Age: mean (range): 6 years (2–16 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Hedman, 1978 ¹⁵² Study design Prospective cohort Country Sweden Setting (teaching) Secondary care (NR) Aim of imaging Detection of reflux	Patient spectrum Confirmed UTI, further investigation Further details "In a few cases the investigation was performed as part of periodic control of earlier proven reflux." All children had recurrent UTI Recurrent UTI: Recurrent UTI Number of patients (number of girls): 51 (102 kidneys) (44) Age: mean (range): 8 years (2–15 years)	Reference standard Reference standard MCUG: technique of Shopfner ²⁸⁸ Time between infection and reference standard: NR Definition of a positive test result NR	Test 1: Scintigraphy: 15 μCi ^{99m} Tc-DTPA kg^{-1} body weight injected i.v. after voiding. 2–5 hours postinjection, sequential images recorded at 5-second intervals for approx. 30 seconds, patient then allowed to void and images recorded during voiding Time between infection and test 1: NR Definition of positive result test 1: NR
Hellerstein, 1978 ¹⁵³ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details The group included children with their second or third UTI as well as some with a history of many UTIs. Several children were known to have reflux. A few of the children had ureteral reimplantations performed ≥ 1 years before these studies Recurrent UTI: Combination Number of patients (number of girls): 45 (49 studies) (42) Age: mean (range): NR (3–17 years)	Reference standard Bladder washout test: bladder filled sterile water containing neomycin and Elase. After 45 minutes the bladder was drained through the catheter. Quantitative cultures were done on final few millilitres of washout solution. Three consecutive urine samples were collected at 20-minute intervals and each of these samples was cultured quantitatively on two blood agar plates. Plates were incubated for 15–18 hours at 37°C and colonies on each plate counted Time between infection and reference standard: NR Definition of a positive test result Growth of ≥ 1000 cfu ml^{-1} in one of the three final urine specimens taken as upper UTI. Counts of 100–1000 cfu ml^{-1} considered indeterminate	Test 1: ACB: catheter urine samples obtained for culture, sensitivity studies and enumeration of ACB. Urinary sediments and saline controls were incubated with fluorescein-conjugated antihuman globulin of goat origin and rewashed twice in phosphate-buffered saline. Smears were prepared and examined microscopically for fluorescence. A minimum of 20 bacteria, identified by phase-contrast microscopy, were examined for fluorescence. Bacteria showing bright apple-green fluorescence were recorded as antibody coated Time between infection and test 1: On admission Definition of positive result test 1: $< 1\%$ of bacteria in urinary sediment with antibody coating classed as negative

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Hellstrom, 1989 ¹⁵⁴ Study design Prospective cohort Country Sweden Setting (teaching) Secondary care (NR) Aim of imaging Detection of scarring	Patient spectrum Confirmed UTI, further investigation Further details All children with a first known episode of confirmed febrile UTI diagnosed at the children's hospital were enrolled Recurrent UTI: Initial UTI Number of patients (number of girls): 88 (84 included in the analysis) (65) Age: mean (range): median 10 months (2 months to 6 years)	Reference standard Urography: performed according to standardised procedure. ²⁸⁹ Appears that follow-up urography was also performed after several years (2.3–4.1 years) Time between infection and reference standard: Not clear Definition of a positive test result Renal damage (scarring) defined as reduction in parenchymal thickness with or without calyceal deformity	Test 1: MCUG; performed according to standardised procedure adhering to the recommendations of the International Reflux Study in Children. ²⁸⁴ Drip infusion of contrast medium by gravity from a standardised height. Intermittent fluoroscopy with an image intensified and 70-mm film for documentation used during bladder filling and voiding Time between infection and test 1: Not clear Definition of positive result test 1: Reflux graded according to international criteria ²⁸⁴
Hitzel, 2000 ¹⁷⁷ Study design Prospective cohort Country France Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI and prediction of scarring	Patient spectrum Confirmed UTI, further investigation Further details Children with clinically defined APN (fever >38°C, abdominal pain and positive urinary culture). Infants who had high-grade reflux and obstruction were excluded Recurrent UTI: NR Number of patients (number of girls): 56 (111 kidneys) (NR) Age: mean (range): 5 years (NR)	Reference standard DMSA: no further details Time between infection and reference standard: Performed during acute phase and at 7 months Definition of a positive test result NR	Test: Ultrasound: B-mode and colour/power Doppler sonography performed by an experienced radiologist in both prone and supine positions with 5–7.5-MHz transducer Time between infection and test: NR Definition of positive result test: NR Test 1: Compared with acute-phase DMSA reference standard Test 2: Compared with follow-up DMSA reference standard at 7 months

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Hitzel, 2002¹⁵⁵</p> <p>Study design Prospective cohort</p> <p>Country France</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Imaging, other</p> <p>Further details To be included a child had to have clinical findings consistent with APN, be aged > 6 months, and have undergone Doppler ultrasound and DMSA at the acute stage of pyelonephritis.</p> <p>APN was diagnosed if the patient had abdominal or lumbar fossa pain, fever > 38°C, and positive urine culture results. Children were excluded if they had urinary tract obstruction, > grade III reflux, or breakthrough infection between inclusion and follow-up</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 57 (50)</p> <p>Age: mean (range): 5 years (6 months to 15.5 years)</p>	<p>Reference standard</p> <p>DMSA: scintigraphy by standard protocol, 4 hours after i.v. injection of ^{99m}Tc-DMSA. Planar posterior, left posterior and right posterior oblique views obtained</p> <p>Time between infection and reference standard: During acute phase</p> <p>Definition of a positive test result If the criteria for inflammation were met: slightly bulging or normal contour; single or multiple, local or diffuse areas of decreased activity in parenchyma, which are diffuse or, rarely, spheric, in at least two projections; mild to severe degree of photopenia or, rarely, complete absence of activity; no volume loss</p>	<p>Test: Ultrasound: sonography included grey-scale and Doppler ultrasound examination of both kidneys with a variable-frequency 5–7-MHz curved transducer. Axial and longitudinal scans were obtained. Performed by an experienced radiologist</p> <p>Time between infection and test: During acute phase</p> <p>Definition of positive test result: Presence of a triangular zone of decreased or absent flow in the parenchyma. Grey-scale abnormalities: thickening of pelvic or ureteral wall, renal sinus hyperechogenicity, mild dilatation of pelvis or ureter, increased renal size, triangular hyperechogenicity</p> <p>Test 1: Results by patient</p> <p>Test 2: Results by kidney</p>
<p>Ilyas, 2002¹⁵⁶</p> <p>Study design Retrospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children presenting with a febrile UTI. Patients also had to have had an initial DMSA renal scan. Children were excluded if they had had a renal transplant, obstructive uropathy, neurogenic bladder, renal calculi or recurrent chronic pyelonephritis. The children were divided in three age groups: < 2 years (<i>n</i> = 85), 2–8 years (<i>n</i> = 91), > 8 years (<i>n</i> = 46)</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 222 (175)</p> <p>Age: median (range): 4.6 years (2 months to 19 years)</p>	<p>Reference standard</p> <p>DMSA: performed by the SPECT method within 48 hours of admission. Weight-adjusted adult-equivalent dose of 37 MBq ^{99m}Tc-DMSA was injected i.v. and SPECT acquisition was performed for 20 minutes 2 hours after isotope administration. Coronal and 3D surface rendering images were reviewed</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result Presence of any of the following: change in relative size of the functional renal mass, an area of either decreased or absent activity within the cortex, a cortical defect with central extension to the collecting system</p>	<p>Test 1: MCUG: no further details</p> <p>Time between infection and test 1: Within 6 weeks of UTI</p> <p>Definition of positive result test 1: Grade based on international classification²⁸⁴</p> <p>Test 2: Ultrasound: renal ultrasound using 5 and 7.5-MHz probes.</p> <p>Time between infection and test 2: Performed during admission</p> <p>Definition of positive result test 2: Presence of renal volume enlargement, hyper- or hypoechogenicity (focal or diffuse) and renal pelvic dilatation measured from anterior and posterior projection and related to body weight</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Jakobsson, 1992 ¹⁵⁷	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: children examined in supine position. ^{99m} Tc-DMSA given at dose of 0.5 MBq kg ⁻¹ (min. 10 MBq). At least 3 hours after injection single anterior and posterior images were taken. In 60 children the DMSA uptake was also determined by computer-designed method that compared results with population of normal kidneys from age-matched control group	Test 1: Ultrasound: performed with 3.5 or 5-MHz sector scanner. Renal volume calculated and expressed as a percentage of normal kidney volume for body weight. Compared with initial DMSA scan Time between infection and test 1: <5 days Definition of positive result test 1: Changes in ultrasound considered present if renal volume exceeded 120% of normal kidney volume or if changes in parenchymal echogenicity and/or dilatation of pelvices were observed
Study design Prospective cohort	Further details Children admitted to hospital with clinically defined APN: fever ≥ 38.5°C, ESR > 20 mm h ⁻¹ or CRP > 20 mg l ⁻¹ , and a positive urine culture	Time between infection and reference standard: < 5 days for test 1 (initial scan), 6–20 weeks later for tests 3 and 5	Test 2: IVU: performed according to standard procedures Time between infection and test 3: 6–20 weeks (median 8 weeks) Definition of positive result test 3: Changes considered present if parenchymal reduction, deformities of calyces, hydronephrosis, ureteral dilatation or malformations were observed. Renal size and parenchymal thickness evaluated according to Claesson criteria. A parenchymal thickness < 2 SD of the normal considered a parenchymal reduction
Country Sweden	Recurrent UTI: NR	Definition of a positive test result Each kidney judged subjectively to be normal, borderline or with definite changes. For computerised results kidneys with a DMSA uptake below 2 SD of the control kidney were defined as abnormal	Test 3: MCUG: no further details Time between infection and test 5: 6–20 weeks (median 8 weeks) Definition of positive result test 5: Presence of reflux noted on MCUG and graded according to international criteria ²⁸⁴
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 72 (59)		
Aim of imaging Localisation of UTI and detection of scarring	Age: median (range): 1 year (0.1 to 15 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Jantausch, 1994 ¹⁵⁸	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: all patients received a renal scan to localise the site of UTI. No further details reported	Test 1: β_2 M concentration: β_2 M determined according to the manufacturer's procedure. Micro-particle Enzyme Immunoassay (Abbott Laboratories) on the Abbott IMX analyser
Study design Prospective cohort	Further details Children admitted to a children's medical centre over a 1.5-year period with a febrile UTI	Time between infection and reference standard: NR	Time between infection and test 1: On admission Definition of positive result test 1: $\geq 0.5 \mu\text{g mg}^{-1}$ of CR
Country USA	Recurrent UTI: NR	Definition of a positive test result Patients with a positive scan (no details) classified as having pyelonephritis, patients with a negative scan as having cystitis	Test 2: NAG: 50 μl of urine incubated with 100 μl of 4-methylumbelliferyl β -D-N-acetylglucosaminide substrate at 37°C for 60 minutes. The reaction was stopped by the addition of 2.5 ml of 0.17 M glycinecarbonate buffer, pH 10.3, and read in a spectrofluorometer with excitation at 365 nm and emission at 450 nm. Fluorescence compared with that of standard solutions of 4-methylumbelliferone. Sample blanks were prepared for each specimen to compensate for non-specific fluorescence
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 24 (NR)		Time between infection and test 2: On admission Definition of positive result test 2: $\geq 40 \mu\text{mol h}^{-1} \text{mg}^{-1}$ of CR
Aim of imaging Localisation of UTI	Age: mean (range): NR (2 weeks to 10 years)		Test 3: Combination: no details Time between infection and test 3: No details Definition of positive result test 3: Two tests combined (not clear whether both positive or either positive)

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Jequier, 1998¹⁶⁰</p> <p>Study design Prospective cohort</p> <p>Country Switzerland</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI and prediction of scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children with clinically suspected pyelonephritis: rectal temperature $\geq 38.5^{\circ}\text{C}$, positive culture; and abdominal or flank pain or, in younger children unable to complain of pain, irritability, vomiting, feeding problems and a CRP level $> 10\text{ mg l}^{-1}$</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 290 (199)</p> <p>Age: median (range): 394 days (4 days to 15 years)</p>	<p>Reference standard</p> <p>DMSA: scintigraphy performed 3–4 hours after i.v. injection of $^{99\text{m}}\text{Tc-DMSA}$ (2.78 MBq kg^{-1} body weight to a maximum of 74 MBq). One posterior, two posterior oblique, one anterior and two anterior oblique images obtained. Global DMSA uptake of each kidney was measured and the two sides were compared</p> <p>Time between infection and reference standard: < 3 days then 60–90 days if initial scan positive</p> <p>Definition of a positive test result Focal or diffuse decreased or absent uptake seen on two projections or more in a normal-sized or swollen kidney</p>	<p>Comparison with initial DMSA scan</p> <p>Test 1: Ultrasound: grey-scale ultrasound, with scan head adapted to patient size and varying from 7 to 4 MHz</p> <p>Time between infection and test 1: < 3 days</p> <p>Definition of positive result test 1: Focal swelling of the renal parenchyma, loss of corticomedullary differentiation, focal hyper- or hypoechogenicity, thickening of the mucosa of the renal pelvis</p> <p>Test 2: Ultrasound: Doppler ultrasound, with scan head adapted to patient size and varying from 7 to 4 MHz</p> <p>Time between infection and test 2: < 3 days</p> <p>Definition of positive result test 2: Focal swelling of the renal parenchyma, loss of corticomedullary differentiation, focal hyper- or hypoechogenicity, thickening of the mucosa of the renal pelvis, increased or decreased blood flow</p> <p>Comparison with follow-up DMSA scan</p> <p>Test 3: As test 1</p> <p>Test 4: As test 2</p>
<p>Jequier, 1985¹⁵⁹</p> <p>Study design Prospective cohort</p> <p>Country Canada</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Anatomical and detection of reflux and scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details UTI diagnosis made according to the criteria of Forbes <i>et al.</i>²⁹⁰</p> <p>Recurrent UTI: Initial UTI</p> <p>Number of patients (number of girls): 240 (121)</p> <p>Age: mean (range): NR (2 days to 15 years)</p>	<p>Reference standard IVU and/or MCUG: no procedural details reported. Pertinent echographic findings such as obstructive lesions were confirmed immediately by IVU and/or MCUG. Suspected reflux or normal sonograms were followed by IVU and/or MCUG 3 weeks to 5 months later. All patients had IVU</p> <p>Time between infection and reference standard: < 5 months after acute phase (usually 3–6 weeks)</p> <p>Definition of a positive test result NR (positive: abnormal MCUG or IVU)</p>	<p>Test 1: Ultrasound: ultrasound using various machines with probes varying from 3.5 to 7.5 MHz. No further details reported</p> <p>Time between infection and test 1: All patients examined during the acute phase of UTI</p> <p>Definition of positive result test 1: NR</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Johnson, 1985 ¹⁶¹	Patient spectrum Confirmed UTI, further investigation	Reference standard Radiological evaluation: All children had a renal and pelvic ultrasound, IVP and MCUG. Boys had MCUG done under fluoroscopic direction: bladder filled with contrast material and spot film taken at time of voiding and postvoid. IVP was done with 50% sodium diatrizoate injection. Ultrasound performed with real-time sector scanner	Test 1: Body temperature: rectal/oral temperature recorded after onset of symptoms before antibiotic therapy was started Time between infection and test 1: On admission Definition of positive result test 1: >38°C rectally or >37°C orally Test 2: Age Definition of positive result test 2: <5 years Test 3: Fever, ESR, CRP, DDAVP-RCA: ESR measured by the Westergen method; serum CRP determined quantitatively by nephelometry; DDAVP renal concentrating ability determined by the method of Monnen <i>et al.</i> ²⁹¹ DDAVP laced in anterior nares at dose of 20 µg in children aged > 1 year and 10 µg in infants. Urine collected at 3 and 5 hours and osmolality determined by freezing point depression using a Fiske osmometer. Values compared with normal values for maximum renal concentrating ability Time between infection and test 3: On admission Definition of positive result test 3: ERP ≥ 25 mm h ⁻¹ ; CRP ≥ 1.0 µg dl ⁻¹ ; Any two of the four positive: both samples > 2 SD below mean Test 4: IVP: methods as reference standard Time between infection and test 4: 4–6 weeks Definition of positive result test 4: as reference standard Test 5: Ultrasound: methods as reference standard Time between infection and test 5: 4–6 weeks Definition of positive result test 5: as reference standard Test 6: MCUG: methods as reference standard Time between infection and test 6: 4–6 weeks Definition of positive result test 6: As reference standard
Study design Prospective cohort	Further details Patients admitted for pyelonephritis were eligible for inclusion as long as they were not referral cases. Children were excluded if they had asymptomatic bacteriuria or myelodysplasia, were attending the urology clinic or had a previous cystogram and intravenous pyelogram	Time between infection and reference standard: 4–6 weeks following start of antibiotic therapy Definition of a positive test result Reflux graded I–V according to international classification (not referenced). Identification of a treatable medical problem – one requiring either medical or surgical intervention in addition to the standard 10-day course of antimicrobial therapy – considered positive. Medical problems identified were: reflux and ectopic ureteroceles with associated upper pole hydronephrosis	
Country USA	Recurrent UTI: Combination		
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 75 (69 included) (NR)		
Aim of imaging Identification of treatable urological problems	Age: mean (range): NR (>2 weeks to <13 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Johnson, 1990¹⁶²</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Further details of inclusion criteria reported previously^{292,293}</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 147 (NR)</p> <p>Age: mean (range): NR (3 months to 13 years)</p>	<p>Reference standard NR: presence of reflux. Children received renal ultrasonogram, excretory urogram and voiding cystogram, in latter part of study children only received excretory urogram if reflux was found. Assume that these tests were used to diagnose reflux, but this is not clearly reported</p> <p>Time between infection and reference standard: No details</p> <p>Definition of a positive test result Reflux grades II–IV considered positive</p>	<p>Index test Test: NAG: assays of NAG were done by the spectrophotometric method of Lockwood and Bosmann.²⁹⁴ Creatinine concentration determined by standard methods and NAG values standardised to creatinin</p> <p>Time between infection and test: On admission</p> <p>Definition of positive test result: Normal values for NAG: creatinine ratios taken from Horak et al.²⁹⁵ Values > 1 SD from the mean were classified as elevated</p> <p>Test 1: Children with cystitis (determined according to criteria of Jodal)²⁹⁶</p> <p>Test 2: Children with pyelonephritis (determined according to criteria of Jodal)²⁹⁶</p>
<p>Kenda, 1989¹⁶³</p> <p>Study design Prospective cohort</p> <p>Country Yugoslavia</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Hospitalised children with confirmed UTI. 73/101 children were presenting with first UTI, remaining 28/101 had recurrent UTI. 56 aged < 1 year</p> <p>Recurrent UTI: Combination</p> <p>Number of patients (number of girls): 101 (67)</p> <p>Age: mean (range): NR (2 weeks to 6 years)</p>	<p>Reference standard Excretory urography: no further details</p> <p>Time between infection and reference standard: < 3 months</p> <p>Definition of a positive test result NR</p>	<p>Test 1: Ultrasound: no details reported</p> <p>Time between infection and test 1: < 3 months</p> <p>Definition of positive result test 1: NR</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Kenda, 2000¹⁶⁴</p> <p>Study design Prospective cohort</p> <p>Country Slovenia</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details The indications for cystography were UTI, follow-up of a previously detected reflux or screening of siblings of children with reflux. Children aged < 1 year excluded</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 99 (198 kidneys) (78)</p> <p>Age: mean (range): 4.6 years (1–12 years)</p>	<p>Reference standard</p> <p>Reference standard Direct radionuclide voiding cystography: performed with child in supine position. Catheter inserted and urine allowed to drain. Bladder filled with 20 MBq of ^{99m}Tc pertechnetate-labelled colloid. When the bladder was full an echo-enhancing agent (Levovist) was administered intravesically. Child was asked to void or allowed to void spontaneously. Images acquired</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result Reflux graded as follows: grade 1: radiotracer reaching the ureter only grade 2: radiotracer reaching the pelvis grade 3: radiotracer reaching the pelvis, which seemed dilated</p>	<p>Test 1: Ultrasound voiding cystography: performed using real-time scanner with a 3.7 and 7-MHz transducer. Baseline ultrasound image obtained and kidneys evaluated through ventral and dorsal approach. When bladder filled to predicted volume, Levovist was administered through catheter. Bladder and proximal parts of the ureters were examined, further Levovist was administered and kidneys were scanned ventrally and dorsally in longitudinal and transverse sections while awaiting micturition. Child asked to void or voided spontaneously. Procedure continued until voiding complete</p> <p>Time between infection and test 1: Performed at same time as reference standard</p> <p>Definition of positive result test 1: Reflux identified when hyperchoegenic microbubbles detected in ureter and/or renal pelvis. Same grading scale used as used for reference standard</p>
<p>Kessler, 1982¹⁶⁵</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Children referred for MCUG to evaluate known VUR or for work-up of UTI</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 32 (55 kidneys) (30)</p> <p>Age: mean (range): NR (1–10 years)</p>	<p>Reference standard MCUG: no details reported</p> <p>Time between infection and reference standard: No details</p> <p>Definition of a positive test result Reflux was graded as follows: grade I: lower ureteral filling grade IIA: ureteral and pelvocalyceal filling without other changes grade IIB: ureteral and pelvocalyceal filling with mild calyceal blunting, but without clubbing and without dilatation of the pelvis or tortuosity or the ureter grade III: ureteral and pelvocalyceal filling, calyceal clubbing, and minor to moderate pelvic dilation with slight tortuosity of the ureters grade IV: massive hydronephrosis and hydroureter²⁸⁵</p> <p>Grades II–IV were considered positive for reflux</p>	<p>Test 1: Ultrasound: ultrasound with 5-MHz transducer used to scan both kidneys. Representative sagittal images obtained with the subject prone. Patient catheterised and placed prone for the remainder of study, residual urine removed via catheter. Cysto-Conray (20%) used as the contrast agent. Kidneys scanned simultaneously with injection of contrast material. The end-point for filling was determined according to patient age or by the child's inability to retain further contrast material. Reflux was not evaluated during voiding</p> <p>Time between infection and test 1: Immediately before MCUG</p> <p>Definition of positive result test 1: Visualisation of moving microbubbles in the renal collecting system and/or dilatation of the collecting system during the course of bladder filling</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Krzemien, 2002 ¹⁶⁶ Study design Prospective cohort Country Poland Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children with UTI referred to a paediatric hospital. 22 newborns, seven aged > 1 year. No other illnesses detected among the children tested Recurrent UTI: Initial UTI Number of patients (number of girls): 29 (58 kidneys) (18) Age: mean (range): 8.2 months (1–24 months)	Reference standard DMSA: no further details Time between infection and reference standard: Within 3 weeks Definition of a positive test result Disorders of uptake of the indicator associated with presence of inflammatory processes in the kidneys. Normal if no differences in the uptake of dye, borderline images classed as normal	Test 1: Ultrasound: power Doppler ultrasound Time between infection and test 1: Within 7 days of presentation Definition of positive result test 1: Hyperfusion in the kidneys
La Cava, 1990 ²⁰⁰ Study design Prospective cohort Country Italy Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children affected by UTI according to clinical, bacteriological and instrumental data Recurrent UTI: Combination Number of patients (number of girls): 162 (321 kidneys) (88) Age: median (range): 3.5 years (20 days to 15 years)	Reference standard Contrast urography performed according to the standard technique of the radiological department Time between infection and reference standard: ≥ 4 weeks Definition of a positive test result Scarring was classified according to the International Reflux Study Committee guidelines ³⁰¹	Test 1: Scintigraphy: renal sequential scintigraphy with ¹²³ I-hippuran, single-kidney ERPF calculated from ¹²³ I-hippuran clearance Time between infection and test 1: ≥ 4 weeks Definition of positive result test 1: Abnormal ERPF 270 ml minute ⁻¹

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Landau, 1994 ¹⁶⁸ Study design Retrospective cohort Country USA Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Mixed, some UTI Further details Medical records of all infants aged <4 months who underwent a DMSA scan to rule out APN were reviewed. Infants were evaluated for fever of undetermined source or suspected sepsis. Patients excluded if no UA was available from admission or if the urine specimen was not SPA or catheter sample. Those with known immunodeficiency, indwelling urinary catheter or a neuropathic bladder were also excluded. Associated clinical findings (age/temperature/peripheral WBC count/additional source of infection) were recorded to classify the infants as high or low risk for serious infection Recurrent UTI: NR Number of patients (number of girls): 186 (128 included in analysis) (38) Age: mean (range): 8 weeks (<16 weeks)	Reference standard Reference standard DMSA: renal cortical scans were done during the acute phase of the disease and obtained 2–3 hours after an i.v. injection of ^{99m} Tc-DMSA dose of 50 μ Ci kg^{-1} (min. 300 μ Ci). High-resolution magnified images of the kidneys in posterior and posterior-oblique projections were obtained Time between infection and reference standard: NR Definition of a positive test result The finding of at least one discrete or diffuse photopenic lesion in the renal parenchyma, without loss in the cortex volume in that area	Test 1: Microscopy: urine samples centrifuged and unstained sediment examined under the microscope for WBC using $\times 400$ magnification Time between infection and test 1: NR Definition of positive result test 1: ≥ 5 WBC hpf^{-1} for APN Test 2: Clinical features: low-risk criteria for a serious bacterial infection Time between infection and test 2: NR Definition of positive result test 2: No focal bacterial infection on physical examination, previously healthy, blood WBC $< 15 \times 10^9 \text{ l}^{-1}$, band forms $< 15 \times 10^9 \text{ l}^{-1}$, UA < 5 WBC hpf^{-1} , when diarrhoea present: < 5 WBC hpf^{-1} in stool
Landau, 1994 ¹⁶⁷ Study design Retrospective cohort Country USA Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Infants aged <16 weeks who underwent a DMSA scan during episodes of febrile UTI. Infants had to have a positive urine culture and available UA from admission. Exclusion criteria included a known obstructive uropathy, immunodeficiency or an indwelling urinary catheter Recurrent UTI: NR Number of patients (number of girls): 142 (NR) Age: mean (range): NR (<16 weeks)	Reference standard DMSA: no details Time between infection and reference standard: During acute phase Definition of a positive test result No details	Test 1: UA: WBCs Time between infection and test 1: On admission Definition of positive result test 1: ≥ 5 WBC hpf^{-1}

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Lavocat, 1997 ¹⁶⁹	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: performed 6 hours after i.v. injection of 2 MBq kg ⁻¹ (min. 15 MBq) of ^{99m} Tc-DMSA. One posterior, one anterior and two posterior oblique images were obtained. The relative contribution of each kidney to renal function was determined by using the geometric average from anterior and posterior plane data. The isofixation curves were used to locate cortical scars	Test 1: Ultrasound: carried out using a 5-MHz transducer for children aged > 3 months and a 7.5-MHz transducer for children aged < 3 months Time between infection and test 1: < 48 hours, repeated within 10 days if required Definition of positive result test 1: Increase in parenchymal volume (confirmed by a decrease in volume on a second ultrasound examination 10 days later); alteration of parenchymal echogenicity (hypoechoic or hyperechoic foci). Isolated dilatation and hypotonicity of the renal pelvis not considered abnormal
Study design Prospective cohort	Further details All children aged > 15 days admitted to a paediatric unit with febrile symptomatic UTI were included. Presented with symptoms of upper UTI and positive urine culture. All patients with uropathy, with the exception of reflux, were excluded from the study. UTI prophylaxis was given until the time of MCUG. Nine children had a history of pyelonephritis, and 24 had upper urinary tract symptoms (upper abdominal or lumbar pain)	Time between infection and reference standard: < 1 week Definition of a positive test result For each kidney, uptake of < 45%; single or multiple peripheral cortical defects; localised or diffuse hypoactivity in one or both kidneys; large defect located at one pole. Small kidney volume and/or deformation of the outlines was considered as evidence of previous parenchymal involvement in APN	Test 2: CT scan: scans were obtained before and after i.v. injection of 2 ml kg ⁻¹ of 38% sodium- meglumine ioxitalamate (Telebrix 38 R) or an ioversol (Optray 350) bolus of 1 mg kg ⁻¹ . A plain film of the urinary tract was taken approximately 20 minutes after iodine injection Time between infection and test 2: < 1 week Definition of positive result test 2: Round, ill-defined and non-enhancing areas of low attenuation exhibiting a mass effect reflected 'acute lobular nephronia'. Wedge-shaped and sharply delimited enhanced areas of reduced parenchyma reflected focal or multifocal lesions of pyelonephritis
Country France			Test 3: MCUG; no details Time between infection and test 3: No details Definition of positive result test 3: Graded according to international classification ²⁸⁴
Setting (teaching) Secondary care (non-teaching)			
Aim of imaging Localisation of UTI	Recurrent UTI: NR Number of patients (number of girls): 55 (110 kidneys) (32) Age: mean (range): 33.4 months (3 weeks to 15 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Leonidas, 1985¹⁷⁰</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Abnormal urinary tract</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children referred for excretory urography as part of the diagnostic investigation of UTI</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 71 (59)</p> <p>Age: mean (range): 5.8 (1 month to 28 years)</p>	<p>Reference standard</p> <p>Reference standard IVU: i.v. injection of 2–3 ml sodium diatrizoate 50% kg⁻¹ body weight; films exposed at 3, 5 and 10 minutes postinjection, with additional exposures as necessary. 59/71 (89%) of patients also had MCUG</p> <p>Time between infection and reference standard: No details</p> <p>Definition of a positive test result Comments on the various components of the urinary tract were recorded separately (length of kidney, renal parenchymal thickness, and degree of dilatation of the collecting system) and a final diagnosis was indicated</p>	<p>Index test</p> <p>Test 1: Ultrasound: carried out using a 5 or 3.5-MHz transducer. Sonograms usually obtained with the patient supine and in the right and left decubitus positions as required. Kidneys scanned in sagittal, coronal and transverse planes</p> <p>Time between infection and test 1: No details</p> <p>Definition of positive result test 1: Comments on the various components of the urinary tract were recorded separately (length of kidney, renal parenchymal thickness and degree of dilatation of the collecting system) and a final diagnosis was indicated</p>
<p>LeQuesne, 1986¹⁷¹</p> <p>Study design Retrospective cohort</p> <p>Country Australia</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children who were investigated with both ultrasound and DMSA scan. 29/33 children had proven reflux at the time of examination</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 33 (66 kidneys) (NR)</p> <p>Age: mean (range): NR (5 months to 14 years)</p>	<p>Reference standard DMSA: no details reported</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result NR</p>	<p>Index test</p> <p>Test 1: Ultrasound: no details reported</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: Ultrasonic signs of possible reflux nephropathy included: focal cortical thinning, focal caliectasis, segmental or generalised scarring, changes in echogenicity of renal tissue, renal size below normal range for height, disparity in renal size > 10%</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Lindsell, 1986 ¹⁷²			
Study design Prospective cohort	Patient spectrum Confirmed UTI, further investigation	Reference standard IVU performed using Iopamidol (Niopam 300) (dose 2 ml kg ⁻¹). The number of films varied; the minimum necessary to show renal outlines clearly, an opacified pelvicalyceal system and a reasonably full bladder. Performed immediately after ultrasound	Test 1: Ultrasound: all examinations by one radiologist using 5-MHz probes. If possible the pelvis was scanned when the bladder was full Time between infection and test 1: NR
Country UK	Further details Age distribution: <1 year (n = 11), 1–5 years (n = 48), >5 years (n = 41)		Definition of positive result test 1: Kidneys assessed for congenital abnormalities and for evidence of renal scarring or dilatation of the distal ureters suggestive of reflux
Setting (teaching) Secondary care (teaching)	Recurrent UTI: NR		
Aim of imaging Detection of scarring	Number of patients (number of girls): 100 (71) Age: mean (range): NR (children, including under 5 years)	Time between infection and reference standard: NR Definition of a positive test result Abnormality defined by: structural abnormalities of the renal tract; evidence of renal scarring; changes, such as minor calyceal blunting and ureteric dilatation, suggestive of reflux	
Lonergan, 1998 ¹⁷³			
Study design Prospective cohort	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: scintigraphy using ^{99m} Tc-DMSA (1.85 MBq kg ⁻¹ , min. 18.5 MBq) or ^{99m} Tc-glucosheptonate (7.4 MBq kg ⁻¹ , min. 92.5 MBq). Scanning approx. 3 hours postinjection, one posterior and one posterior oblique view of each kidney	Test: MRI: 1.5 T, pelvic coil. Gadolinium-enhanced inversion recovery imaging (TR/TE/TI, 2000/16/160), slice thickness 3–8 mm, axial and/or coronal planes Time between infection and test: NR Definition of positive test result: Positive or negative for pyelonephritis (not defined)
Country USA	Further details Children presenting with a fever and positive culture were considered for inclusion. Exclusion criteria: children aged <2 years, elevated serum creatinine, allergy to gadopentetate dimeglumine or to scintigraphic agent, presence of MR-incompatible device, contraindications to sedation (if sedation deemed necessary)		Test 1: By patient Test 2: By kidney
Setting (teaching) Secondary care (NR)	Recurrent UTI: NR	Time between infection and reference standard: NR Definition of a positive test result Positive or negative for pyelonephritis (not defined)	
Aim of imaging Localisation of UTI	Number of patients (number of girls): 37 (70 kidneys) (NR) Age: mean (range): 5.2 years (2–16 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
MacKenzie, 1994 ¹⁷⁴ Study design Prospective cohort Country UK Setting (teaching) Secondary care (NR) Aim of imaging Detection of scarring	Patient spectrum Confirmed UTI, further investigation Further details All children presenting to hospital over a 5-year period, with a first documented, symptomatic, bacteriologically proven <i>E. coli</i> UTI were entered into the study. Symptoms, duration and child's temperature at presentation were recorded Recurrent UTI: Initial UTI Number of patients (number of girls): 175 (112 analysed) (112) Age: mean (range): NR (1 month to 13 years)	Reference standard Reference standard DMSA: posterior, right and left posterior oblique views were obtained 1.5–2 hours after injection of a weight-related dose of DMSA Time between infection and reference standard: <28 days Definition of a positive test result Showing photon-deficient areas, scarring (persistent loss of normal reniform outline), showing both photon-deficient areas and scarring, showing a significant difference in size, showing divided function outside the normal range (45–55%)	Test 1: Ultrasound: ultrasound examinations performed using a 3.5 or 5-MHz probe. Images of the kidneys were obtained with the patient in supine and prone positions, images of the bladder also obtained. Time between infection and test 1: <48 hours Definition of positive result test 1: Ultrasound examinations were reported as normal, showing scarring, showing dilatation or a suspected obstruction, or showing a difference in size. The presence of a duplex kidney was looked for. Swelling of the kidney, generalised or localised changes in echogenicity, prominence of the pyramids and loss of the cortico-medullary differentiation were also commented on
Mage, 1989 ¹⁷⁵ Study design Prospective cohort Country France Setting (teaching) Secondary care (NR) Aim of imaging Detection of reflux	Patient spectrum Confirmed UTI, further investigation Further details Children presenting over a 7-year period. 77 children aged <2 years, 12 children aged <2 months Recurrent UTI: Initial UTI Number of patients (number of girls): 122 (85) Age: mean (range): NR (16 days to 11 years)	Reference standard Reference standard MCUG: examinations during filling, with full bladder, during voiding and postvoiding. Always retrograde in girls, retrograde or suprapubic in boys Time between infection and reference standard: >3 weeks Definition of a positive test result Not clear	Test 1: Ultrasound: ultrasound examination with 5-MHz transducer. Kidneys, ureteral junctions and bladder were examined. The anteroposterior renal pelvis diameter was systematically measured in transverse sections Time between infection and test 1: Before MCUG Definition of positive result test 1: Not clear

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Mahant, 2002 ¹⁷⁶			
Study design Retrospective cohort	Patient spectrum Confirmed UTI, further investigation	Reference standard MCUG using Hypaque 18% contrast material. Intermittent fluoroscopy during the filling stage and a spot image of the filled bladder obtained. Spot images obtained of the bladder and urethra during voiding without catheter, with girls in a supine position and boys left anterior oblique. A postvoid image of the bladder and renal fossae was also obtained	Test 1: Ultrasound: ultrasound scans, with sector curved array and linear high-resolution transducers Time between infection and test 1: Median 2 (IQR 1–3) following start of treatment Definition of positive result test 1: Renal ultrasound considered suggestive of reflux if: dilatation of the pelvicalyces, dilatation of the ureters, dilatation of the collecting system reported (for one or both kidneys)
Country Canada	Further details The charts of all children aged <5 years admitted with UTI were reviewed. Children with known genitourinary abnormalities or with previous history of UTI were excluded		
Setting (teaching) Secondary care (teaching)	Recurrent UTI: Initial UTI		
Aim of imaging Detection of reflux	Number of patients (number of girls): 170 (162 included) (71)	Time between infection and reference standard: NR	
	Age: mean (range): median 3 months (3 days to 4 years)	Definition of a positive test result Reflux graded using international scale and reported by staff radiologists ²⁸⁴	
McLorie, 1989 ¹⁷⁸			
Study design Prospective cohort	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: ^{99m} Tc-DMSA injected i.v. at a dose not exceeding 5 µCi 70 kg ⁻¹ 1.73 m ⁻² body surface area. Scans obtained 3 hours after injection in the sagittal, coronal and transverse planes	Test 1: IVP after i.v. injection of 50% diatrizoate meglumine and diatrizoate sodium, (injection volume 1 ml kg ⁻¹ body weight). A minimum of two radiographs taken at 5 and 15 minutes after injection were obtained Time between infection and test 1: NR Definition of positive result test 1: Criteria for diagnosis of renal scarring were diffuse or focal parenchymal thinning and adjacent calyceal clubbing or deformity
Country Canada	Further details Patients with documented primary reflux, presenting with UTI. Criteria for participation in the study were: progression in degree of reflux, previous documentation of renal scars by IVP or ^{99m} Tc-DMSA, recurrent UTI or contemplation of surgical correction of reflux. Patients with renal insufficiency, hydronephrosis, neurovesical dysfunction or outflow obstruction were excluded		
Setting (teaching) Secondary care (NR)	Recurrent UTI: NR	Time between infection and reference standard: NR	
Aim of imaging Detection of scarring	Number of patients (number of girls): 32 (64 renal units) (26)	Definition of a positive test result Diagnosis of focal scarring was based on identification of areas of reduced radioactivity in the parenchyma in at least two tomographic cuts in one axis	
	Age: mean (range): NR (2–14 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Mentzel, 2002 ¹⁷⁹ Study design Prospective cohort Country Germany Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux	Patient spectrum Mixed, some UTI Further details All patients were referred for the evaluation of reflux. The indications for investigation were: UTI ($n = 67$), follow-up of previously diagnosed reflux ($n = 15$), sonographically diagnosed dilatation of the urinary pelvicalyceal system or megaureter ($n = 20$), malformation syndromes with high probability of renal involvement ($n = 10$), enuresis ($n = 6$). Patients were excluded for: galactosemia, ongoing UTI, unsuccessful sonographic examination caused by restlessness, boys aged < 1 year Recurrent UTI: NR Number of patients (number of girls): 118 (234 uretero-renal units) (90) Age: median (range): 4.5 years (3 weeks to 16 years)	Reference standard Reference standard MCUG: saline solution and X-ray contrast medium (Peritrat 400; Kohler-Chemie, Germany) were drip infused into the bladder. Procedure monitored by fluoroscopy and spot film using one anterior-posterior spot film during filling and one in the Lauenstein position during voiding Time between infection and reference standard: NR Definition of a positive test result Reflux graded according to international scale ²⁸⁴ (positive: reflux of any grade; negative: no reflux)	Index test Test 1: Ultrasound: ultrasound with 3–5-MHz curved array and 5–10-MHz linear transducers. Baseline sonograms were obtained in two perpendicular planes of the kidney, ureters and bladder, with the patient in the supine and prone positions. The patient was then catheterised and the bladder emptied and afterwards filled with saline solution. Ultrasonographic contrast medium (Levovist 300 mg ml ⁻¹ solution) was then added. Examination in the longitudinal and transverse sections of each flank in prone and supine positions was conducted before voiding. Patients were then returned to the supine position and asked to void (uncooperative, young children allowed to void spontaneously). The retrovesical parts of the ureters and the renal pelvis were scanned during voiding Time between infection and test 1: NR Definition of positive result test 1: Reflux graded according to international scale ²⁸⁴ (positive: reflux of any grade; negative: no reflux)
Merrick, 1980 ¹⁸⁰ Study design Retrospective cohort Country Scotland Setting (teaching) Secondary care (NR) Aim of imaging Detection of scarring	Patient spectrum Confirmed UTI, further investigation Further details All children referred for renal investigations were identified from files. Patients had to receive both DMSA and IVU within 3 months of each other, or IVU in the interval between two identical DMSA scans, or a DMSA scan in the interval between two identical IVUs. All children had bacteriologically proven UTI Recurrent UTI: NR Number of patients (number of girls): 79 children (155 kidneys) (NR) Age: mean (range): NR (appears to be 1 to > 10 years)	Reference standard Reference standard: Scintigraphy: performed 1–4 hours after administration of ^{99m} Tc-glucoheptonate at a dose of 7.4 MBq kg ⁻¹ or of ^{99m} Tc-DMSA at a dose of 3.7 MBq kg ⁻¹ . Analogue images without enhancement or any form of data processing were recorded simultaneously in the posterior, left posterior, oblique and right posterior oblique positions Time between infection and reference standard: Not clear Definition of a positive test result No details	Index test Test 1: IVU: technique of IVU determined by the radiologist. Minimum of three films were taken: a plain film immediately before administration of contrast, a coned view of the renal areas at 5 minutes and a full-length film at 15 minutes. Additional views and tomograms were obtained as deemed necessary to obtain adequate visualisation of both the renal outlines, the calyces and the ureters Time between infection and test 1: Not clear Definition of positive result test 1: No details

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Misselwitz, 1971 ¹⁸¹	Patient spectrum Mixed, some UTI	Reference standard IVU: NR	Test 1: Renogram: performed 30 minutes after 10 ml Tee kg ⁻¹ body weight in lying position. 2.5 µC kg ⁻¹ body weight of ¹³¹ I-o-hippurat injected Time between infection and test 1: NR Definition of positive result and test 1: Positive for 'harnstauung'. Qualitative assessment: subjective assessment of the secretion curve
Study design Retrospective cohort	Further details Children who had received both renography and IVU. 37 children with APN, 66 with recurrent or chronic pyelonephritis without retention on the IVU and without reflux, 64 children with recurrent or chronic pyelonephritis with retention on IVU or reflux. 167 had confirmed pyelonephritis, 13 had retention without pyelonephritis	Time between infection and reference standard: NR Definition of a positive test result Positive for 'harnstauung'	Test 2: Renogram: performed 30 minutes after 10 ml Tee kg ⁻¹ body weight in lying position. 2.5 µC kg ⁻¹ body weight of ¹³¹ I-o-hippurat injected Time between infection and test 2: NR Definition of positive result and test 2: Positive for 'harnstauung'. Semi-quantitative assessment according to method of Zum Winkel ²⁷
Country Germany			
Setting (teaching) Secondary care (teaching)			
Aim of imaging Detection of reflux	Recurrent UTI: NR Number of patients (number of girls): 180 (NR) Age: mean (range): NR (6 months to 16 years)		
Montplaisir, 1981 ¹⁸²	Patient spectrum Confirmed UTI, further investigation	Reference standard Not clear: on the basis of clinical, radiological and cystoscopic investigations children were classified as having kidney infection (n = 94), bladder/urethra infection (n = 102), or asymptomatic bacteriuria (n = 86)	Test 1: ACB: direct immunofluorescence performed on urine sediment as previously described ²⁸ and on stool and vaginal secretions smeared on glass slides, air-dried and subsequently fixed with acetone. Antisera used were fluorescein-conjugated anti-human-IgG, -IgA and -IgM and anti-IgG containing only G(ab'). As a negative control for each sample giving a positive result the test was repeated in the same manner. Smears were examined with a fluorescence microscope at a magnification of x250 Time between infection and reference standard: NR Definition of positive result and test 1: NR Definition of positive result test 1: > 3 fluorescent bacteria in 200 microscopic fields
Study design Prospective cohort	Further details 53 children aged <2 years		
Country Canada	Recurrent UTI: NR		
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 282 (216) Age: mean (range): NR (0–16 years)	Time between infection and reference standard: NR Definition of a positive test result Lower UTI or asymptomatic bacteria considered negative. Kidney infection considered positive	
Aim of imaging Localisation of UTI			

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Morin, 1999 ¹⁸³			
Study design Prospective cohort	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: children examined in supine position. ^{99m} Tc-DMSA given at dose of 0.5 MBq kg ⁻¹ body weight. 6 hours after injection, posterior and posterior oblique images obtained	Compared with reference standard 1 Test 1: Ultrasound: renal ultrasound performed using 3.5, 5.0, and 9.0-MHz transducers. Anterior, posterior, longitudinal and transverse views were obtained for each kidney Time between infection and test 1: On admission Definition of positive result test 1: Considered abnormal if one of the following features was found: parenchymal hyperechogenicity, focal lesion with hyper/hypoechogenicity, thickening of the renal pelvic wall, significant enlargement of both kidney length and width Test 2: MCUG: conducted as for reference standard 2 Time between infection and test 2: < 4 weeks Definition of positive result test 2: As reference standard 2
Country France	Further details Patients referred for a first episode of APN. Clinical criteria for diagnosing APN were: fever, CRP > 20 mg l ⁻¹ and a positive urine culture. Children with features of dilated uropathy were excluded	Time between infection and reference standard: < 5 days Definition of a positive test result Considered abnormal if one or more areas of decreased DMSA uptake were noted. Care was taken not to consider central defects located over the pelivcalyceal system as abnormal	Compared with reference standard 2 Test 3: Ultrasound: conducted as for test 1 Time between infection and test 3: As for test 1 Definition of positive result test 3: As for test 1
Setting (teaching) Secondary care (NR)	Recurrent UTI: Initial UTI	Reference standard 2 MCUG: performed within 4 weeks of admission according to standard procedure	
Aim of imaging Localisation of UTI and detection of reflux	Number of patients (number of girls): 70 (43) Age: median (range): 2 years (1 month to 17 years)	Time between infection and reference standard 2: < 4 weeks Definition of positive result reference standard 2 Reflux graded according to international criteria ²⁸⁴	

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Mucci, 1994 ¹⁸⁴	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: no details reported	Test 1: Ultrasound: ultrasound, no details given Time between infection and test 1: NR Definition of positive result test 1: NR
Study design Retrospective cohort	Further details All included children had undergone at least DMSA and ultrasound	Time between infection and reference standard: NR	
Country UK	Recurrent UTI: Combination	Definition of a positive test result NR	
Setting (teaching) Secondary care (non-teaching)	Number of patients (number of girls): 193 (141)		
Aim of imaging Detection of scarring	Age: mean (range): 4.8 years (5 months to 17 years)		
Muensterer, 2002 ¹⁸⁵	Patient spectrum Mixed, some UTI	Reference standard MCUG: no further details	Test: Ultrasound: renal size on ultrasound examination by a standard protocol using curvilinear or linear probes of the highest frequency possible (7, 5, or 3 MHz for infants, children and adolescents, respectively). Length measurements performed on screen using an integrated electronic calliper Time between infection and test: >2 weeks
Study design Retrospective cohort	Further details Patients who underwent renal ultrasound over a 6-month period were retrospectively reviewed. Patients were included if they had had MCUG and ultrasound examinations within 1 week.	Time between infection and reference standard: >2 weeks	
Country USA	Indications for the studies were as follows: UTI, dysuria, frequency ($n = 101$, 42 first UTI, 59 recurrent), lower genitourinary malformations, ($n = 28$), dilatation on prenatal ultrasound ($n = 31$), neurological dysfunction ($n = 26$), haematuria, ($n = 7$). Patients with previous urinary tract surgery, kidney malformations or solitary kidney were excluded	Definition of a positive test result Reflux graded according to international reflux study in children ²⁸⁴	Compared with any reflux on MCUG Definition of positive result test 1: Abnormal kidney size (defined as one outside +2 SD of normal for age) Definition of positive result test 2: Any dilatation ²⁹⁹
Setting (teaching) Secondary care (teaching)			
Aim of imaging Detection of reflux	Recurrent UTI: Combination		Compared with high-grade reflux on MCUG (grades 3–5) Definition of positive result test 3: Abnormal kidney size (defined as one outside of +2 SD of normal for age) Definition of positive result test 4: Any dilatation Definition of positive result test 5: Abnormal kidney size or any dilatation
	Number of patients (number of girls): 193 (386 kidneys) (111)		
	Age: median (range): 3.6 years (1 day to 18.3 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont d)

Study details	Patient details	Reference standard	Index test
Oostenbrink, 2000 ¹⁸⁶	Patient spectrum Confirmed UTI, further investigation	Reference standard MCUG: no further details	Compared with what? Test 1: Ultrasound: no details
Study design Prospective cohort	Further details Children aged <5 years presenting to the paediatric outpatient or emergency department. Patients with neurological bladder dysfunction were excluded	Time between infection and reference standard: NR	Time between infection and test 1: NR
Country Netherlands		Definition of a positive test result Presence of reflux. Grade I-IV according to international classification ²⁸⁴	Definition of positive result 1: Presence of dilatation of urinary tract, graded subjectively by radiologist as 0 if no dilatation; 1 in presence of pyelectasis, 2 in the case of mild dilatation, 3 if severe dilatation was present. Classed as positive in the case of dilatation grade ≥ 2
Setting (teaching) Secondary care (teaching)	Recurrent UTI: Initial UTI		
Aim of imaging Detection of reflux	Number of patients (number of girls): 140 (89) Age: mean (range): 1 year (<5 years)		Compared with any grade reflux Test 2: combination: combined risk score calculated from following model: score = 6 × male gender + 7 × positive family history - 1 × age + 1 × CRP + 14 × ultrasound dilatation. Model produced by regression analysis of prespecified variables thought to be associated with reflux. Points assigned to age: <1 year = 0; 1-2 year = 1; 2-3 year = 2, etc., with max. of 5 points for children aged 5 years. Points assigned to CRP: 0-9 mg l ⁻¹ = 1; 10-19 mg l ⁻¹ = 2 points, etc., up to max. of 20 points for CRP > 200 mg l ⁻¹
			Time between infection and test 2: NR Definition of positive result test 2: ≥ 1 Definition of positive result test 3: ≥ 6 Definition of positive result test 4: ≥ 11 Definition of positive result test 5: ≥ 16 Definition of positive result test 6: > 25
			Compared with reflux grade ≥ 3 Definition of positive result test 7: ≥ 11 Definition of positive result test 8: ≥ 16 Definition of positive result test 9: > 25

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Piaggio, 2003¹⁸⁷</p> <p>Study design Prospective cohort</p> <p>Country Italy</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Detection of reflux</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Children undergoing MCUG and ultrasound over a 1-year period. Indications for examinations were: clinical signs, including recurrent UTI, pyelonephritis and significant micturition disorders; ultrasound data of moderate to severe or worsening hydronephrosis; urinary tract malformations with or without clinical signs; clinical signs (UTI, hydronephrosis) in transplanted kidneys</p> <p>Recurrent UTI: Combination</p> <p>Number of patients (number of girls): 158 (305 kidney-ureter units) (61)</p> <p>Age: mean (range): 3.9 years (0.1–12.7 years)</p>	<p>Reference standard MCUG</p> <p>Reference standard execution Performed after cystoscopy with same catheter, no further details reported</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result Reflux appears to be classified according to the international scale²⁸⁴</p>	<p>Test 1: Ultrasound: ultrasound with convex 2–4 and 4–7-MHz multiple-frequency transducers. Levovist contrast medium (2.5 g at 300 micro-particles/mg ml⁻¹) was used. Half the dose was administered after mild bladder distension, the remainder retained in case of further need. Saline solution was continued until spontaneous micturition in younger children, and until urgent need in older ones</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: NR</p>
<p>Pickworth, 1992¹⁸⁸</p> <p>Study design Retrospective cohort</p> <p>Country England</p> <p>Setting (teaching) Secondary care (non-teaching)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details The notes, radiological investigation and radionuclide scans of all patient who received ^{99m}Tc-MAG3 scans over a 2-year period were reviewed. Reasons for referral included: UTI ($n = 57$), pre-/postsurgical assessment ($n = 21$), further assessment of antenatally diagnosed renal tract anomaly ($n = 13$), multiple congenital defects ($n = 4$), screening ($n = 3$), loin pain ($n = 2$)</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 100 (NR)</p> <p>Age: mean (range): 5.6 years (6 days to 17 years)</p>	<p>Reference standard 1 DMSA: no details reported</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result: NR</p> <p>Reference standard 2 MAG3: ^{99m}Tc-MAG3 scans were performed following oral hydration based on an adult dose of 100 MBq scaled according to body surface area. Patients were studied supine on a gamma camera. Micturating studies were performed at the end of the 30-minute renography phase with the patient sitting on a modified commode. Two frames were acquired during micturition</p> <p>Time between infection and reference standard 2: NR</p> <p>Definition of positive result reference standard 2 Kidneys were classified as showing focal scarring if there were discrete photon-deficient areas on the 2.5-minute images with or without underlying dilatation of the collecting system. Global scarring defined as uniform loss of cortical thickness</p>	<p>Compared with DMSA Test 1: MAG3: conducted as for reference standard 2</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: as for reference standard 2</p> <p>Compared with MAG-3 Test 2: IVU: no details</p> <p>Time between infection and test 2: NR</p> <p>Definition of positive result test 2: No details</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Piepsz, 1992¹⁸⁹</p> <p>Study design Retrospective cohort</p> <p>Country Belgium</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Retrospectively selected on the basis of clinical suspicion of APN</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 30 (60 kidneys) (NR)</p> <p>Age: mean (range): 7.2 years (4 months to 18 years)</p>	<p>Reference standard</p> <p>Reference standard DMSA: ^{99m}Tc-DMSA scintigraphy 2 hours after i.v. injection. Dose calculated as a fraction of the adult dose, by child's weight and body surface area (min. 18 MBq). Posterior and two oblique posterior views</p> <p>Time between infection and reference standard: within 48 hours</p> <p>Definition of a positive test result Routine written report used for study. Abnormal: small kidney with or without irregular kidney shape or cortical defects, single or multiple cortical defects, diffuse hypoactivity of one kidney (<45% of total activity)</p>	<p>Index test</p> <p>Test 1: Scintigraphy: MAG3 renography after i.v. injection of a fraction of the adult dose determined by child's weight and body surface area (min. 17 MBq). Sixty 20-second frames were recorded and the three frames acquired during the second minute of the test were added</p> <p>Time between infection and test 1: Within 48 hours</p> <p>Definition of positive result test 1: Abnormal: small kidney with or without irregular kidney shape or cortical defects; single or multiple cortical defects; diffuse hypoactivity of one kidney (<45% of total activity)</p>
<p>Pylkkanen, 1978¹⁹⁰</p> <p>Study design Prospective cohort</p> <p>Country Finland</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Symptomatic (n = 114) and asymptomatic (n = 14) children</p> <p>Recurrent UTI: Initial UTI</p> <p>Number of patients (number of girls): 128 (NR)</p> <p>Age: mean (range): NR (0–14 years)</p>	<p>Reference standard Clinical: Jodal criteria²⁹⁶</p> <p>Time between infection and reference standard: On admission</p> <p>Definition of a positive test result High fever, increased ESR, abnormally high CRP level, decreased renal concentrating capacity. Patient classed as upper UTI if at least 3 criteria fulfilled</p>	<p>Test 1: ACB: performed according to method of Thomas <i>et al.</i>²⁸⁷ Fluorescein-conjugated goat antiserum to human globulin was used</p> <p>Time between infection and test 1: On admission</p> <p>Definition of positive result test 1: ≥2 brightly fluorescent bacteria</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Radmayr, 2002 ¹⁹¹ Study design Prospective cohort Country Austria Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux	Patient spectrum Confirmed UTI, further investigation Further details Patients transferred to a radiology department for urological investigation because of a recent UTI. Patients with renal duplications were excluded from the study. None of the children had acute UTI at the time of the investigation Recurrent UTI: NR Number of patients (number of girls): 104 (208 renal and ureter units) (81) Age: mean (range): 5.4 years (14 days to 19 years)	Reference standard MCUG: using intermittent digital fluoroscopy. Filling volume kept comparable to that used in ultrasound, prewarmed X-ray contrast medium injected directly into the bladder by the transurethral route Time between infection and reference standard: NR Definition of a positive test result International classification for reflux grade ²⁸⁴	Test 1: Ultrasound: performed using a 3.5–7.5-MHz curved array transducer attached to a Doppler ultrasound unit. An initial standard ultrasound of the whole urinary tract was performed with the patient prone and supine. After transurethral catheterisation, the bladder was completely emptied and filled with saline, administered until the patient had the urge to void or until the operator felt slight back pressure. 15% of the filling volume was then replaced with galactose-based, microbubble containing echo-enhancer agent. The bladder, retrovesical space and kidneys were monitored during injection of agent. Both kidneys were scanned during voiding Time between infection and test 1: NR Definition of positive result test 1: Reflux was diagnosed when microbubbles appeared in the ureter or renal pelvis. The international classification of VUR was used for grading ²⁸⁴
Redman, 1984 ¹⁹² Study design Retrospective cohort Country USA Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux	Patient spectrum Confirmed UTI, further investigation Further details Girls who received an IVP and MCUG as part of the evaluation for UTI. The evaluation was recommended primarily on the basis of a strong history of UTI or the UA and culture obtained by referring physicians previously. No studies were performed when the child was acutely infected Recurrent UTI: NR Number of patients (number of girls): 200 (200) Age: mean (range): NR (<1 year to 14 years)	Reference standard MCUG: performed in a retrograde fashion using a no. 8 feeding tube. Contrast medium was instilled by gravity using image intensification to observe filling. Cystography during voiding was also done Time between infection and reference standard: Not acute Definition of a positive test result No details reported	Test 1: IVP: performed using 1 ml lb ⁻¹ body weight 60% diatrizoate. Radiographs were made immediately following injections, and supine and prone radiographs were obtained 10 minutes after injections Time between infection and test 1: Not acute Definition of positive result test 1: No details reported

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Rehling, 1989 ¹⁹³ Study design Prospective cohort Country Denmark Setting (teaching) Secondary care (teaching) Aim of imaging Detection of scarring	Patient spectrum Mixed, some UTI Further details Children with UTI and/or congenital hydronephrosis. 12 had UTI, six had hydronephrosis. 69 children presented with UTI and/or congenital hydronephrosis, but only 20 included in analysis Recurrent UTI: NR Number of patients (number of girls): 69 (20 included in analysis) (9) Age: median (range): 5 years (6 days to 13 years)	Reference standard Reference standard IVU: standard IVU performed starting with control film of kidneys and bladder. Maximum 40 ml of Omnipaque 300 mg I ml ⁻¹ or Telebrix 380 mg I ml ⁻¹ was injected in children under and over 15 kg, respectively. On average seven films taken Time between infection and reference standard: NR, not during infection Definition of a positive test result Urinary tract evaluated as normal (0), moderately abnormal (1), severely abnormal (2) or impossible to evaluate. 0 considered negative, all other results positive	Index test Test 1: Scintigraphy: performed with the patient supine. Data were stored from 0 to 21 minutes postinjection of 4 MBq kg ⁻¹ body weight (min. 50 MBq) of ^{99m} Tc-DTPA. Regions of interest of the kidneys and the left ventricle of the heart were defined and time-activity curves computed during infection Time between infection and test 1: NR, not during infection Definition of positive result test 1: Urinary tract evaluated as normal (0), moderately abnormal (1), severely abnormal (2) or impossible to evaluate; (0) considered negative, all other results positive
Rickwood, 1992 ¹⁹⁴ Study design Prospective cohort Country England Setting (teaching) Secondary care (non-teaching) Aim of imaging Detection of scarring, reflux or anatomical abnormality	Patient spectrum Confirmed UTI, further investigation Further details Consecutive children presenting to one of three hospitals with a symptomatic, bacteriologically proven UTI. Outpatients (<i>n</i> = 153), emergency admissions (<i>n</i> = 47) Recurrent UTI: NR Number of patients (number of girls): 200 (59) Age: mean (range): NR (<10 years, included infants)	Reference standard Reference standard Combination: all children received IVU, further investigations determined by the ultrasound findings. Cystography and static DMSA were performed when ultrasound and IVU gave normal results or showed abnormalities in renal size, outline or echxture. Cystography also performed when distal ureteric dilatation was detected and was followed by DMSA in the presence of reflux or by dynamic diuresis renography otherwise. Cases of pure pelvic hydronephrosis were examined by dynamic renography only. Static DMSA was always deferred until at least a month after the UTI had cleared Time between infection and reference standard: Static DMSA performed a minimum of 1 month postinfection, no further details reported Definition of a positive test result Combination of all these methods, no further details	Index test Test 1: Ultrasound: no details Time between infection and test 1: NR Definition of positive result test 1: NR

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Rohden, 1995 ¹⁹⁵ Study design Cohort, not clear if prospective Country Germany Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux	Patient spectrum Mixed, some UTI Further details Children after a UTI, children with suspicion for urinary tract malformations, enuresis diurnal and nocturnal, pathological ultrasound after birth, and children followed up after long-term reflux Recurrent UTI: NR Number of patients (number of girls): 26 (16) Age: mean (range): 3.5 years (3 days to 13 years)	Reference standard MCUG: none reported Time between infection and reference standard: Not clear Definition of a positive test result NR	Test 1: Ultrasound: contrast-enhanced ultrasound with Echovist. Standard ultrasound performed initially following micturition, children catheterised, 10–30 ml of saline infused, depending on bladder volume 5–30 ml of contrast solution was infused. Children imaged in dorsal position. Bladder filled until needed to micturate, extra contrast infused and imaged during micturition Time between infection and test 1: Not clear Definition of positive result test 1: NR: subjective assessment
Rossleigh, 1990 ¹⁹⁶ Study design Prospective cohort Country Australia Setting (teaching) Secondary care (NR) Aim of imaging Detection of scarring	Patient spectrum Mixed, some UTI Further details Children undergoing urological assessment. 49 presented with UTI, in 14 the assessment was carried out for other reasons Recurrent UTI: NR Number of patients (number of girls): 63 (NR) Age: mean (range): NR (<1 year)	Reference standard IVU: no details Time between infection and reference standard: > 1 month Definition of a positive test result No details	Test: DMSA: performed 3 hours after injection of 40 MBq ^{99m} Tc-DMSA, standard posterior planar views were collected for 5 minutes. Posterior pinhole views were also performed. Some patients also underwent SPECT examination. The study was reconstructed in coronal, sagittal and transaxial planes with slices of 1 pixel width. Differential renal function was calculated from the posterior planar data Time between infection and test: > 1 month Definition of positive test result: Reduction in kidney function and/or decreased or absent uptake of tracer in the renal cortex causing distortion or indentation of the normal renal outline Test 1: UTI patients only Test 2: Reflux, no UTI patients

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Salih, 1994 ¹⁹⁷	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children who were investigated by standard MCUG and were shown to have reflux in at least one kidney</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 21 (42 kidneys) (14)</p> <p>Age: mean (range): 6 (2–16 years)</p>	<p>Reference standard MCUG: no further details</p> <p>Time between infection and reference standard NR</p> <p>Definition of a positive test result Graded according to classification of Dwoskin²⁸⁵ For 2 × 2 analysis any reflux classed as positive</p>	<p>Test 1: Colour-flow Doppler sonography: conventional ultrasound and colour-flow Doppler sonography performed using 3.75 and 5-MHz phased array transducers. Patient initially examined by conventional ultrasound. Renal dimensions, parenchymal thickness, scars and parenchymal echo patterns were evaluated during sonography examinations while the bladder was empty. Patients were asked to drink water until they had a strong sensation of bladder fullness or to the point of reflex voiding in infants. Patients were examined in supine position. Both ureterovesical junctions and distal ureters were scanned by sagittal and oblique sections one after the other. Patients were asked to urinate and examination repeated to find out whether there was a back-flow into the ureters. Residual volume of the bladder also examined</p> <p>Time between infection and test 1: On same day as MCUG</p> <p>Definition of positive result test 1: Reflux of urine into the ureter depicted as blue colour jet. Collecting systems examined and compared with prior sonography findings to grade reflux as follows: I: low-grade reflux; II: reflux with lower ureteral filling only, kidney is normal; III: reflux with minor to moderate ureteral and pelvic dilatation; IV: reflux with massive hydronephrosis and hydroureter. For 2 × 2 analysis any reflux classed as positive</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Scherz, 1994 ¹⁹⁸ Study design Prospective cohort Country USA Setting (teaching) Secondary care (teaching) Aim of imaging Detection of scarring	Patient spectrum Imaging, other Further details 51 children presenting with febrile UTI, 24 asymptomatic (presenting with non-febrile UTI or for sibling screening). All children had reflux, demonstrated by MCUG Recurrent UTI: NR Number of patients (number of girls): 75 (150 kidneys) (53) Age: mean (range): NR (3 weeks to 12 years)	Reference standard Reference standard DMSA from a commercial kit, dose determined by body weight (min. 0.3 µCi, max. 3.0 µCi). Imaging 2–6 hours postinjection, posterior and right and left posterior images obtained Time between infection and reference standard: > 3 months Definition of a positive test result DMSA abnormality defined as: reduction in renal function (<43% overall function) and/or decreased or absent uptake of tracer in the renal cortex causing distortion or indentation of the normal renal outline	Index test Test 1: Ultrasound: ultrasound, 3-MHz transducers. 14 images per kidney, including longitudinal, transverse and coronal Time between infection and test 1: Not clear Definition of positive result test 1: Abnormal defined as: changes in parenchymal echogenicity and/or dilation of pelvis; scar, defined as cortical thickening with or without depression of margin; renal size discrepancy (> 1 cm) Test 1: Symptomatic patients Test 2: Asymptomatic patients Test 3: All patients
Schneider, 1984 ¹⁹⁹ Study design Prospective cohort Country Germany Setting (teaching) Secondary care (teaching) Aim of imaging Detection of reflux	Patient spectrum Mixed, some UTI Further details Children presenting for examination of the urinary tract over a 7-month period. Indication for MCUG was: UTI (n = 88), follow-up after surgery or prior pathology (n = 15), malformation syndromes (n = 4), haematuria (n = 2), 'persistent dribbling' (n = 1) Recurrent UTI: NR Number of patients (number of girls): 110 (219 renal units) (75) Age: mean (range): NR (6 days to 14 years)	Reference standard Reference standard MCUG: standard MCUG using fluoroscopy and a spot-film technique. No further details reported Time between infection and reference standard: NR Definition of a positive test result Reflux graded according to the classification of Heikel and Parkkula ³⁰⁰	Index test Test 1: Ultrasound: ultrasound performed with a real-time sector-scanner using a 5-MHz transducer. The bladder was catheterised and completely emptied, and a baseline scan of the kidneys in supine and prone positions measured in transverse and longitudinal sections performed. The bladder was filled with contrast medium Conray FL, and the ureterovesical junction investigated for reflux. Air contrast studies were also conducted: small amounts of air (2–20 cm ³) were injected into the full bladder and the ureter was surveyed for migration of air bubbles associated with reflux Time between infection and test 1: NR Definition of positive result test 1: Reflux graded according to the classification of Heikel and Parkkula ³⁰⁰ Positive: any grade reflux vs any grade reflux by MCUG Definition of positive result test 2: Increased separation in the central renal echo complex vs grade ≥ II reflux by MCUG

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Sfakianakis, 1989 ²⁰¹	Patient spectrum Confirmed UTI, further investigation	Reference standard Scintigraphy: ^{99m} Tc-glucosheptonate triple phase (flow, 10 × 2 minutes early and delayed)	Test 1: Ultrasound: NR Time between infection and test 1: NR Definition of positive result test 1: NR
Study design Prospective cohort	Further details Infants with clinical and bacteriological proof of first UTI	Time between infection and reference standard: NR	
Country USA	Recurrent UTI: Initial UTI	Definition of a positive test result NR	
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 104 (48 ultrasound) (NR)		
Aim of imaging Localisation of UTI	Age: mean (range): NR (< 2 years)		
Siampilis, 1996 ²⁰²	Patient spectrum Mixed, some UTI	Reference standard MCUG: standard MCUG with urography, under fluoroscopy including the voiding phase. 2–5 days after ultrasound	Test 1: Ultrasound: fluid-cystorenography, 3.5-MHz sector and 10-MHz linear transducers. Normal saline administered dropwise by catheter (40–200 ml, according to capacity). Sonographic study of the renal collecting system performed during filling Time between infection and test 1: NR Definition of positive result test 1: NR
Study design Prospective cohort	Further details First episode of UTI (<i>n</i> = 51), recurrent UTI (<i>n</i> = 19), follow-up after past recurrent UTI (<i>n</i> = 5), follow-up of a familial reflux (<i>n</i> = 6), other disorders of the urinary tract (e.g. malformations or stone) (<i>n</i> = 6), other (<i>n</i> = 3)	Time between infection and reference standard: NR	Test 2: Ultrasound: air-cystorenography, conducted 30 minutes after removal of fluid. Air administered by same catheter (70–180 ml according to capacity). Sonographic examination of the renal pelvicalyceal system via a dorsal approach, in sitting position or upright in the arms of parent Time between infection and test 2: NR Definition of positive result test 2: NR
Country Greece	Recurrent UTI: Combination	Definition of a positive test result NR	
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 90 (180 kidneys) (35)		
Aim of imaging Detection of reflux	Age: mean (range): 2.8 years (2.5 days to 17 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Smellie, 1995 ²⁰³ Study design Prospective cohort Country England Setting (teaching) Secondary care (teaching)	Patient spectrum Confirmed UTI, further investigation Further details Children aged <14 years who had at least one bacteriologically proven UTI. 42 primary/secondary referrals and 16 tertiary referrals from other hospitals Recurrent UTI: Combination	Reference standard Combination: MCUG, IU and DMSA performed according to standard methods. IU and MCUG each limited to four films at one of the two hospitals Time between infection and reference standard: <12 months Definition of a positive test result More than one investigation abnormal. Children in whom DMSA study was initially abnormal but later resolved after treatment were classed as normal for DMSA	Test 1: Ultrasound: renal tract ultrasonography Time between infection and test 1: <12 months Definition of positive result test 1: NR
Aim of imaging Detection of renal scarring and reflux	Number of patients (number of girls): 58 (46) Age: median (range): 3 years (1 month to 13 years)		
Smolkin, 2002 ²⁰⁴ Study design Prospective cohort Country Israel Setting (teaching) Secondary care (NR) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children admitted to the paediatric department with febrile UTI Recurrent UTI: NR Number of patients (number of girls): 64 (60 included in analysis) (44) Age: mean (range): 16.7 months (2 weeks to 3 years)	Reference standard DMSA, no further details Time between infection and reference standard: <7 days Definition of a positive test result Renal pathology defined as focal or multifocal perfusion defects, or as split renal uptake of <45%. If the first DMSA was abnormal a repeat scan was performed ≥6 months after the initial one. Diagnosis of APN was confirmed only in patients with totally or partially reversible lesions on scintigraphy	Test 1: PCT: clotted blood (2 ml) was centrifuged, serum separated and frozen at -70°C before analysis. All patients with high PCT on admission had second PCT determination 72 hours after treatment Time between infection and test 1: <3 days Definition of positive result test 1: ≥0.5 µg/l Test 2: CRP: clotted blood (2 ml) was centrifuged, serum separated and frozen at -70°C before analysis. Performed on admission and 72 hours after treatment. Determined by semi-quantitative immunometric test Time between infection and test 2: <3 days Definition of positive result test 2: ≥20 mg l ⁻¹

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Sreenarasimhaiah, 1995²⁰⁵</p> <p>Study design Prospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (NR)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Consecutive series of patients; all children hospitalised with the clinical diagnosis of APN and who required radiological evaluation, over a 9-month period. Indications for radiological investigation published previously^{114,302}</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 50 (42)</p> <p>Age: mean (range): NR (2 months to 15 years)</p>	<p>Reference standard ^{99m}Tc-glucuheptonate renal scintigraphy</p> <p>Reference standard execution Performed by injecting tracer intravenously in dose proportional to body surface area, (min. 1.0 μCi, max. 7.0 μCi). Images recorded every 15 seconds for 1–6 minutes. After 2 hours, posterior, right and left posterior oblique images were taken</p> <p>Time between infection and reference standard: 48–96 hours</p> <p>Definition of a positive test result The criteria for the diagnosis of APN were renal scarring and anatomical and functional abnormalities of the urinary system, published elsewhere.^{303–308}</p>	<p>Test 1: Ultrasound: standard techniques for renal ultrasound,¹¹⁴ no further details reported</p> <p>Time between infection and test 1: 48–96 hours</p> <p>Definition of positive result test 1: published elsewhere.^{114,302} Results by renal unit</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Stokland, 1998 ²⁰⁹	Patient spectrum Confirmed UTI, further investigation	Reference standard DMSA: performed approximately 4 hours after injection of 0.5 MBq kg ⁻¹ body weight ^{99m} Tc-DMSA (minimum 10 MBq), posterior view with 250,000 counts acquired on a gamma camera with a general purpose collimator	Test 1: DMSA: as for reference standard Time between infection and test 1: Median 13 days (range 1–66 days) Definition of positive result test 1: As for reference standard
Study design Prospective cohort	Further details All children aged 0–6 years who were treated for culture-verified acute first symptomatic UTI over a 2-year period were included in the study. Children aged ≥ 1 year had to have a body temperature $\geq 38.5^\circ\text{C}$ during the index UTI, children below this age were included irrespective of body temperature. Children with urinary tract obstruction were excluded. All children were treated immediately with appropriate antibiotics	Time between infection and reference standard: 1.1 (range 0.5–2.5 years) Definition of a positive test result Focal or generalised uptake defect with or without indentation of the renal contour considered abnormal. Split renal function was calculated based on background subtracted kidney uptake. Split renal function $< 45\%$ considered abnormal irrespective of any uptake defects at visual examination. Each kidney classified as normal, equivocal or abnormal. Equivocal classed as normal	Test 2: MCUG: no details Time between infection and test 3: No details Definition of positive result test 3: Reflux graded according to international recommendations ²⁸⁴
Country Sweden			
Setting (teaching) Secondary care (teaching)			
Aim of imaging Prediction and detection of scarring	Recurrent UTI: Initial UTI Number of patients (number of girls): 157 (314 renal units) (72) Age: median (range): 0.4 years (5 days to 5 years)		Test 3: Urography: initial urography Time between infection and test 4: NR Definition of positive result test 4: Renal damage defined as reduction of parenchymal thickness with or without calyceal deformation. In the presence of calyceal deformation, parenchymal thickness below 2 SD was considered abnormal. In the absence of calyceal deformation parenchymal thickness below 2.5 SD was considered abnormal. The parenchymal damage was classified according to a three-grade scale like that used for DMSA. Equivocal classed as normal
			Test 4: Urography: follow-up urography Time between infection and test 6: Within mean 1 day (range 1 day to 1.2 years) of follow-up DMSA Definition of positive result test 6: As for test 4

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Stokland, 1996 ²⁰⁷	Patient spectrum	Reference standard	Test 1: CRP; highest values used for analyses
Study design	Confirmed UTI, further investigation	DMSA: performed approximately 4 hours after injection of 0.5 MBq kg ⁻¹ body weight ^{99m} Tc-DMSA (minimum 10 MBq), posterior view with 250,000 counts acquired on a gamma camera with a general purpose collimator	Time between infection and test 1: Determined at the index UTI
Country	Further details		Definition of positive result test 1: > 20 mg l ⁻¹
Sweden	All children aged 0–6 years who were treated for culture-verified acute first symptomatic UTI over a 2-year period were included in the study. Children aged ≥ 1 year had to have a body temperature ≥ 38.5°C during the index UTI, children below this age were included irrespective of body temperature. Children with urinary tract obstruction were excluded. All children were treated immediately with appropriate antibiotics. Children who were not followed up for 1 year or who did not receive a follow-up DMSA scan were excluded		Test 2: Body temperature
Setting (teaching)			Time between infection and test 2: Determined at index UTI, highest values used for analyses
Secondary care (teaching)			Definition of positive result test 2: ≥ 38.5°C
Aim of imaging		Time between infection and reference standard: 1.1 (range 0.5–2.5 years)	Test 3: MCUG; no details
Prediction of scarring		Definition of a positive test result	Time between infection and test 3: 18 days (range 1–123 days)
		Focal or generalised uptake defect with or without indentation of the renal contour considered abnormal. Split renal function was calculated based on background subtracted kidney uptake. Split renal function below 45% considered abnormal irrespective of any uptake defects at visual examination. Each kidney classified as normal, equivocal or abnormal. Equivocal classed as normal	Definition of positive result test 3: Reflux graded 1–5 according to international recommendations ²⁸⁴
	Recurrent UTI: Initial UTI		
	Number of patients (number of girls): 157 (72)		
	Age: median (range): 0.4 years (5 days to 5.8 years)		

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Stokland, 1994 ²⁰⁶	Patient spectrum Confirmed UTI, further investigation	Reference standard Urography: no further details	Test: Ultrasound: 3–7.5-MHz sector, vector, linear or convex transducers. Kidneys classified as normal, probably normal, uncertain, probably abnormal or abnormal
Study design Prospective cohort	Further details Children recently examined with urography because of UTI	Time between infection and reference standard: NR	Time between infection and test: NR
Country Sweden	Recurrent UTI: NR	Definition of a positive test result Equivocal findings were excluded.	Tests 1–4: Focal renal scarring used as the diagnostic criteria
Setting (teaching) Secondary care (NR)	Number of patients (number of girls): 25 (240 kidney examinations) (NR)	Abnormal defined as calyceal deformation plus parenchymal reduction. Classified by three radiologists	Definition of positive result test 1: Positive: abnormal or probably abnormal Definition of positive result test 2: Positive: abnormal or probably abnormal Definition of positive result test 3: Positive: abnormal or probably abnormal, or uncertain
Aim of imaging Detection of scarring	Age: median (range): 6.5 years (2–16 years)		Definition of positive result test 4: Positive: abnormal or probably abnormal, or uncertain, or probably normal Tests 5–8: Focal renal scarring and/or reduced renal size used as the diagnostic criteria Definition of positive result test 5: Positive: abnormal Definition of positive result test 6: Positive: abnormal or probably abnormal Definition of positive result test 7: Positive: abnormal or probably abnormal, or uncertain Definition of positive result test 8: Positive: abnormal or probably abnormal, or uncertain, or probably normal

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Stokland, 1996 ²⁰⁸ Study design Prospective cohort Country Sweden Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children aged <6 years presenting with their first recognised bacteriologically proven symptomatic UTI were included. In the age group 1–6 years a body temperature of $\geq 38.5^{\circ}\text{C}$ was required, children aged <1 year included regardless of temperature. No patient had recurrent infection before DMSA was performed Recurrent UTI: Initial UTI Number of patients (number of girls): 186 (175 included in analysis) (78) Age: median (range): 0.9 years (10 days to 5 years)	Reference standard Reference standard DMSA: performed 4 hours after i.v. injection of 0.5 MBq kg^{-1} of $^{99\text{m}}\text{Tc-DMSA}$. Posterior view acquired using gamma camera Time between infection and reference standard: 1–66 days Definition of a positive test result Localised or generalised uptake defects with or without indentation in the renal contour were regarded as abnormal. Relative renal function calculated based on background subtracted kidney uptake. A split renal side distribution <45% considered abnormal irrespective of any uptake defects at visual evaluation. All defects classified as normal, equivocal or abnormal. Equivocal cases analysed as negative	Index test Test 1: Urography: no details; results by kidneys Time between infection and test 1: NR Definition of positive result test 1: Renal damage defined as a reduction in parenchymal thickness with or without calyceal deformation. In the presence of calyceal deformation a parenchymal thickness below 2 SD was considered abnormal. In the absence of calyceal deformity a parenchymal thickness below 2.5 SD was considered abnormal. The parenchymal damage was classified according to a three-graded scale like that used for DMSA Test 2: MCUG: no details Time between infection and test 3: No details Definition of positive result test 3: Graded according to International recommendations ²⁸⁴ Test 3: CRP: no details Time between infection and test 4: On admission Definition of positive result test 4: >20 mg l ⁻¹
Tan, 1988 ²¹⁰ Study design Prospective cohort Country Singapore Setting (teaching) Secondary care (NR) Aim of imaging Detection of reflux	Patient spectrum Confirmed UTI, further investigation Further details Children referred by paediatrics department with confirmed ($n = 66$) or strongly suspected ($n = 34$) UTI Recurrent UTI: NR Number of patients (number of girls): 100 (55 MCUG) (67) Age: mean (range): 1.76 years (0–11 years)	Reference standard Reference standard MCUG: by infusing dilute contrast into the bladder by catheter. Under fluoroscopic screening reflux looked for in both ureters before, during and after micturition Time between infection and reference standard: NR Definition of a positive test result Reflux classified on the Smellie scale, grades I–IV	Index test Test 1: Ultrasound: first examination carried out. Real-time sector scanning with 3.5 or 5-MHz transducer. Supine oblique and prone scans obtained in both longitudinal and transverse planes. Younger children were sedated Time between infection and test 1: NR (first investigation carried out) Definition of positive result test 1: NR

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Study design Retrospective cohort</p> <p>Country USA</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Localisation of UTI</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children who had undergone scintigraphy, over a 2-year period, for the evaluation of possible UTI</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 55 (39 included) (31)</p> <p>Age: median (range): 3 years 11 months (3 days to 19 years)</p>	<p>Reference standard</p> <p>Reference standard Clinical and laboratory findings: children classified as APN, cystitis or no UTI on clinical grounds.</p> <p>Time between infection and reference standard: On admission</p> <p>Definition of a positive test result <i>Pyelonephritis:</i> temperature $\geq 38^{\circ}\text{C}$, costovertebral angle tenderness, chills, urine culture $\geq 10^5$ cfu ml⁻¹, ESR > 25 mm h⁻¹</p> <p><i>Cystitis:</i> temperature $< 38^{\circ}\text{C}$, dysuria, frequency, urine culture $\geq 10^5$ cfu ml⁻¹, ESR < 25 mm h⁻¹</p> <p><i>No UTI:</i> temperature $\leq 38.5^{\circ}\text{C}$, no costovertebral angle tenderness, no chills, no dysuria, no frequency, urine culture $< 10^5$ cfu ml⁻¹</p> <p>Positive: clinical evidence of APN</p>	<p>Test 1: Scintigraphy: studies performed using a dual-isotope method. Angiographic and cortical images were derived using ^{99m}Tc-glucroheptonate and functional measures with ¹³¹I-o-hippurate. Initial and early 5-minutes static images using ^{99m}Tc-glucroheptonate (100 Ci lb⁻¹). Delayed images at 1.5–2.5 hours. Posterior and posterior oblique views of each kidney recorded</p> <p>Time between infection and test 1: Usually < 1 week</p> <p>Definition of positive result test 1: NR</p> <p>Test 2: Ultrasound: using a 3.5 or 5-MHz transducer</p> <p>Time between infection and test 2: Usually < 1 week</p> <p>Definition of positive result test 2: NR</p> <p>Test 3: IVP: injection of 2 ml kg⁻¹ Renographin 60. Imaging sequences: preliminary AP radiograph, AP radiograph at 1 minute, AP and both oblique radiographs of the abdomen at 10 minutes. Tomograms or delayed radiographs, as determined by the radiologist</p> <p>Time between infection and test 3: NR</p> <p>Definition of positive result test 3: NR</p> <p>Test 4: Cystography: no details reported</p> <p>Time between infection and test 4: NR</p> <p>Definition of positive result test 4: NR</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Trave, 1997¹²</p> <p>Study design Prospective cohort</p> <p>Country Spain</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of scarring and reflux</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children with a diagnosis of APN. Patients were studied by means of clinical and laboratory assessments in addition to the imaging methods discussed below</p> <p>Recurrent UTI: Initial UTI</p> <p>Number of patients (number of girls): 33 (19)</p> <p>Age: mean (range): 4 years (0.2–12.3 years)</p>	<p>Reference standard</p> <p>Reference standard DMSA: performed 3–4 hours after administration of i.v. 25–74 MBq of ^{99m}Tc-DMSA. Images were taken in the anterior, posterior and oblique posteriors (left and right) planes</p> <p>Time between infection and reference standard: Mean 3.9 days after admission</p> <p>Definition of a positive test result Scans were considered abnormal when kidney function was <45% and/or there was visualisation of areas of decreased uptake</p> <p>Reference standard 2 MCUG: no further details</p> <p>Time between infection and test 4: 3–4 weeks after infection</p> <p>Definition of positive result test 4 NR</p>	<p>Index test</p> <p>Compared with DMSA for detection of scarring</p> <p>Test 1: Ultrasound: renal ultrasound, no details</p> <p>Time between infection and test 1: Mean 2.6 days following recruitment</p> <p>Definition of positive result test 1: No details</p> <p>Compared with MCUG for diagnosing reflux</p> <p>Test 2: Ultrasound: no details</p> <p>Time between infection and test 2: As above</p> <p>Definition of positive result test 2: No details</p> <p>Test 3: DMSA: as for reference standard</p> <p>Time between infection and test 3: As for reference standard</p> <p>Definition of positive result test 3: As for reference standard</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Valentini, 2001 ²¹³	Patient spectrum Mixed, some UTI	Reference standard MCUG: conventional MCUG was performed using a standardised technique.	Test 1: Ultrasound: contrast-enhanced ultrasound performed using a 3.75-MHz convex-array transducer. The bladder was filled to its estimated capacity with saline, by catheter. The bladder, ureteral orifices and both kidneys were evaluated using conventional grey-scale sonography. Levovist was introduced through the catheter (300 mg ml ⁻¹ , to 8% of bladder volume), and the bladder and ureteral orifices were reevaluated by axial and oblique scans with the patient in the supine position. Both kidneys were re-examined in two orthogonal planes. The voiding phase was attempted with the catheter in place, but was not always achieved
Study design Prospective cohort	Further details Consecutive patients admitted because of recurrent UTI, a history of reflux, or myelomeningocele spina bifida	An ionic, water-soluble contrast agent (36% iodamine, Uromiro) was introduced, bladder volume was kept comparable to that used in ultrasound examinations. Examination of the voiding phase was attempted by removing the catheter, but was not always successful. Radiographs were taken during the filling and voiding phases	Time between infection and test 1: Immediately before MCUG
Country Italy	Recurrent UTI: Recurrent UTI		Test 2: Ultrasound: Doppler sonography, using a 4-kHz pulse repetition frequency, with colour filters and colour gain adjusted to optimise imaging. Contrast-enhanced ultrasound performed as test 1
Setting (teaching) Secondary care (teaching)	Number of patients (number of girls): 74 (60 patients and 14 girls) included in the analysis) (NR)	Time between infection and reference standard: NR	Time between infection and test 2: Immediately before MCUG
Aim of imaging Detection of reflux	Age: mean (range): 5 years (3 weeks to 16 years)	Definition of a positive test result Reflux was graded in accordance with the international system. ²⁸⁴	Definition of positive result tests 1 and 2: The diagnostic criterion for reflux was the presence of microbubbles in the collecting system appearing as hyperechoic dots on grey-scale sonograms. Reflux was graded as follows: grade I: hyperechoic dots in the ureter above the ureteral orifice grade II: hyperechoic dots extending up to the renal pelvis without ureteral dilatation grade III: hyperechoic dots extending up to the renal pelvis and calices (non-dilated) with dilatation of the ureter grade IV: hyperechoic dots in mildly dilated ureter, pelvis and calyces grade V: hyperechoic dots in markedly dilated ureter, calyces and pelvis

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
<p>Verber, 1988^{2,14}</p> <p>Study design Prospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Secondary care (teaching)</p> <p>Aim of imaging Detection of reflux or renal scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children aged <5 years with symptomatic confirmed UTI</p> <p>Recurrent UTI: NR</p> <p>Number of patients (number of girls): 115 (80)</p> <p>Age: mean (range): 1.7 years (<5 years)</p>	<p>Reference standard MCUG or IVU:</p> <p>MCUG: performed after catheterisation. Hypaque fed into the bladder by gravity. Bladder screened intermittently during filling, catheter removed before continuous screening during micturition</p> <p>IVU: Performed after laxative the previous evening and restricted fluid for 5–8 hours pre-examination (dependent on age). Examinations comprised: a full-length control film, films of the renal area at 5–10 minutes after injection of 20–40 ml Hypaque or Niopam, full-length films before and after micturition</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result MCUG: reflux recorded on a scale of 1–3 (grade 1: confined to the ureter; grade 2: reaching the renal calyces without distension; grade 3: with distension)</p> <p>IVU: renal scarring and renal size were recognised by the criteria of Hodson;³⁰⁹ significant dilatation of the collecting system or the ureter on the postmicturition film and other features associated with reflux, such as dilatation of the renal pelvis, were recorded</p>	<p>Compared with MCUG</p> <p>Test 1: Scintigraphy: DMSA scans performed 2–3 hours after i.v. injection of 2 MBq kg⁻¹ ^{99m}Tc-DMSA (min. 20 MBq, max. 160 MBq). Images acquired in posterior and both posterior oblique projections, with the patient supine, using a gamma camera (<i>n</i> = 130 kidneys)</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: Defects reported if there was a clear interruption in the renal outline on any projection. Relative renal uptake outside the range of 45–55% was considered abnormal. Discrepancies in renal size were noted</p> <p>Test 2: IVU: as for reference standard</p> <p>Time between infection and test 2: NR</p> <p>Definition of positive result test 2: As for reference standard</p> <p>Test 3: Ultrasound: renal ultrasound in the anterior and posterior sagittal and transverse planes using a 5-MHz transducer and a static or real-time (coronal planes also included) scanner (<i>n</i> = 62 kidneys)</p> <p>Time between infection and test 3: NR</p> <p>Definition of positive result test 3: Transverse scans of the full bladder were used to visualise any ureteral dilatation. Renal length was measured and any focal cortical abnormality recorded. Separation of the renal sinus echoes by >0.5 cm was recorded as a dilated upper tract</p> <p>Compared with IVU</p> <p>Test 4: As for test 1 (<i>n</i> = 92 kidneys)</p> <p>Time between infection and test 4: NR</p> <p>Definition of positive result test 4: As for test 1</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont'd)

Study details	Patient details	Reference standard	Index test
Verboven, 1990 ²¹⁵ Study design Prospective cohort Country Belgium Setting (teaching) Secondary care (teaching) Aim of imaging Localisation of UTI	Patient spectrum Confirmed UTI, further investigation Further details Children admitted to a hospital with symptomatic UTI. Symptoms included fever, chills, anorexia, nausea, weight loss, dysuria, enuresis, abdominal pain and low back pain Recurrent UTI: NR Number of patients (number of girls): 24 (18) Age: mean (range): 4 years (3 weeks to 12 years)	Reference standard Reference standard Clinical: patients classified according to clinical criteria of Pylkkänen; ³¹⁰ fever, ESR, serum level of CRP and renal concentrating capacity Time between infection and reference standard: At presentation Definition of a positive test result Refer to Pylkkänen; ³¹⁰ no further details	Test 1: DMSA: performed 2 hours after injection of dose of ^{99m} Tc-DMSA adjusted for the child's height and weight. One posterior and two posterior oblique images of 300 counts were obtained. When necessary SPECT was performed immediately after the planar imaging Time between infection and test 1: During acute infection Definition of positive result test 1: No details
Volti, 1991 ²¹⁶ Study design Prospective cohort Country Italy Setting (teaching) Secondary care (teaching) Aim of imaging Abnormal urinary tract	Patient spectrum Confirmed UTI, further investigation Further details Patients with VUR on MCUG were excluded ($n = 10$) Recurrent UTI: NR Number of patients (number of girls): 56 (37) Age: mean (range): NR (2 months to 12 years)	Reference standard Reference standard IVU: non-ionic contrast medium Iopamidol (dose 1.5–3 ml kg ⁻¹). Supine radiographs of kidney and bladder 5–10 minutes after i.v. injection, 4–6 films Time between infection and reference standard: 7–10 days after treatment Definition of a positive test result NR	Test 1: Ultrasound: renal ultrasound using 5 and 3.5-MHz probes. Longitudinal, supine, prone and transverse scans performed. Kidneys evaluated for position, size, morphology and echogenicity. Bladder and degree of distension also observed. Time between infection and test 1: 7–10 days after treatment Definition of positive result test 1: NR

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont d)

Study details	Patient details	Reference standard	Index test
<p>Whitear, 1990²¹⁷ Study design Retrospective cohort</p>	<p>Patient spectrum Mixed, some UTI</p> <p>Further details Children who received both IVU and DMSA were included. These were a heterogeneous group. The largest diagnostic groups were UTI and reflux (42%), hypertension (8%) and nephritis (6%)</p> <p>Recurrent UTI: Other</p> <p>Number of patients (number of girls): 205 (388 kidneys) (97)</p> <p>Age : Mean (range): NR (3 days to 16 years)</p>	<p>Reference standard IVU: majority consisted of a control film, 5-minute renal area and 15-minute full-length film</p> <p>Time between infection and reference standard: No details</p> <p>Definition of a positive test result: NR</p>	<p>Test 1: Scintigraphy: ^{99m}Tc-DMSA scans on gamma camera, posterior and both posterior-oblique projections obtained 6 hours post-injection</p> <p>Time between infection and test 1: no details</p> <p>Definition of positive result test 1: NR</p>

continued

TABLE 41 Included studies of tests for the further investigation of confirmed UTI (cont d)

Study details	Patient details	Reference standard	Index test
<p>Wujanto, 1987²¹⁸</p> <p>Study design Prospective cohort</p> <p>Country UK</p> <p>Setting (teaching) Secondary care (non-teaching)</p> <p>Aim of imaging Detection of scarring</p>	<p>Patient spectrum Confirmed UTI, further investigation</p> <p>Further details Children referred for radionuclide assessment of renal function and scarring. All had previous IVU, 36 had radiographically demonstrated reflux, 30 had ureteric reimplantation to correct the reflux, and 16 were treated conservatively</p> <p>Recurrent UTI: Recurrent UTI</p> <p>Number of patients (number of girls): 46 (92 kidneys) (28)</p> <p>Age: mean (range): 5.6 years (2 months to 11 years)</p>	<p>Reference standard 1 IVU: assessment of renal scarring: dose 1 MBq kg⁻¹ body weight. After 2–3 hours analogue images were acquired in anterior, posterior and both posterior oblique projections. Relative renal function calculated. Performed previously, no details reported</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result 1 Judged according to radiologist's report and opinion of referring urologist, no further details. Equivocal classed as normal</p> <p>Reference standard 2 DMSA: assessment of renal scarring, performed 2 days after hippuran renogram. Dose 1 MBq kg⁻¹ body weight. After 2–3 hours analogue images were acquired in anterior, posterior and both posterior oblique projections. Relative renal function calculated</p> <p>Time between infection and reference standard: NR</p> <p>Definition of a positive test result 2 Criteria used for reporting renal scarring were: reduction in kidney size, calyceal dilatation with thinning of cortex, decreased uptake of tracer in the cortex causing distortion or indentation of the normal renal outline. Equivocal classed as normal</p>	<p>Compared with IVU Test 1: Renography: Assessment of renal function: 1123 hippuran at a dose of 0.2 MBq kg⁻¹ body weight (min. 2.5 MBq). The scan was performed with the patient either sitting in front of or lying supine over a gamma camera. Computerised image acquisition every 20 seconds for at least 20 minutes. Relative renal function calculated</p> <p>Time between infection and test 1: NR</p> <p>Definition of positive result test 1: Criteria used for reporting renal scarring were: reduction in kidney size, calyceal dilatation with thinning of cortex, decreased uptake of tracer in the cortex causing distortion or indentation of the normal renal outline. Equivocal classed as normal</p> <p>Test 2: DMSA: as for reference standard 2</p> <p>Time between infection and test 2: NR</p> <p>Definition of positive result test 2: As for reference standard 2</p> <p>Compared with DMSA Test 3: As for test 1</p> <p>Time between infection and test 3: NR</p> <p>Definition of positive result test 3: As for test 1</p>

AP, anterior–posterior; ELISA, enzyme-linked immunosorbent assay; FITC, fluorescein isothiocyanate; IgA, IgG, IgM, immunoglobulin A, G, M; IMLA, ???; 3D, three-dimensional.

Appendix 7

Studies excluded from the review

TABLE 42 Studies excluded from the review

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Aathithan, 2001	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Abbott, 1972	Other				No	
Abbott, 1978	Other				No	
Abdou, 1977	Other				No	
Abdul Mujeeb, 1994	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Abdurrahman, 1992	Other				No	
Abraham, 1970	Other				No	
Abraham, 1979	Other				No	
Abyholm, 1979	Other				No	
Addor, 1984	Other				No	
Aggarwal, 1989	Other				No	
Aggarwal, 1991	Other				No	
Akdilli, 1999	Other				No	
Akhundova, 1980	Other				No	
Akoua-Koffi, 1994	Other				No	
Aladjem, 1997	Other				No	
Alam, 1981	Other				No	
Albert, 1982	Other				No	
Alexander, 1972	Other				No	
Allen, 1974	Other				No	
Almeida, 1993	Further investigation	Other	Yes	≥20	No	
Almeida, 1994	Other				No	
Alon, 1988	Further investigation	Other	<18	≥20	No	
Alon, 1999	Other				No	
Al-Orifi, 2000	Other				No	
Altemaier, 1996	Other				No	
Altman, 1969	Other				No	
Altman, 1971	Other				No	
Al Umran, 1994	Other				No	
Alur, 2000	Further investigation	Diagnostic accuracy	Yes	≥20	No	
American Academy of Pediatrics, 1999	Other				No	
Amir, 1993	Other				No	
Anders, 1974	Other				No	
Anderson, 1995	Other				No	
Andersz, 1966	Other				No	
Andrich, 1992	Other				No	
Anonymous, 1968	Other				No	
Anonymous, 1979	Other				No	
Anonymous, 1984	Other				No	
Anonymous, 2000	Other				No	
Anonymous, 2002	Other				No	
Ansell, 2002	Other				No	
Anttila, 1970	Other				No	
Arap, 1986	Other				No	
Ariceta Iraola, 1991	Other				No	
Arrieta, 1994	Other				No	
Artibani, 1990	Other				No	
Arya, 2002	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Ascenti, 2000	Further investigation	Other	Yes	≥20	No	
Asscher, 1970	Other				No	
Aubert, 1983	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Augusidou-Savopoulou, 1993	Other				No	
Austin, 1999	Other				No	
Avni, 1997	Other				No	
Bacharach, 1982	UTI diagnosis	Diagnostic accuracy	None <5	≥20	No	Culture
Bachelard, 2001	Other				No	
Bachur, 2001	Other				No	
Bailey, 1995	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Bakakos, 2002	Other				No	
Balsamo, 1976	Other				No	
Balzar, 1991	Other				No	
Banzo, 1999	Other				No	
Baraff, 1992	Other				No	
Barbin, 1978	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Barkemeyer, 1993	Other				No	
Barta, 1969	Other				No	
Basso, 1981	Other				No	
Batinic, 1988	Other				No	
Batista, 1998	Other				No	
Beeram, 1991	Other				No	
Behrendt, 1985	Other				No	
Belli, 1996	Further investigation	Other	<18	≥20	No	
Benador, 1998	Other				No	
Ben-Ami, 1984	Other				No	
Ben-Ami, 1989	Further investigation	Other	Yes	≥20	No	
Bendall, 1994	Other				No	
Bengtsson, 1970	Other				No	
Benigno, 1990	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Benitz, 1998	Other				No	
Bensman, 1978	UTI diagnosis	Other	Yes	≥20	No	
Bensman, 1993	Other				No	
Bensman, 2002	Other				No	
Berg, 1971	Other				No	
Bergman, 1999	Other				No	
Bergstrom, 1972	Other				No	
Bergstrom, 1973	Other				No	
Berro, 2000	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Bhatnagar, 2002	Further investigation	Other	Yes	≥20	No	
Biasini, 1974	UTI diagnosis	Diagnostic accuracy	Yes	≥20	No	Culture
Bixler-Forell, 1985	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Biggi, 2001	Other				No	
Bjerkklund Johansen, 2002	Other				No	
Bjorgvinsson, 1991	Further investigation	Other	<12	≥20	No	
Blane, 1993	Further investigation	Other	None <5	≥20	No	
Bleeker, 2001	Other				No	
Blickman, 1984	Other				No	
Blickman, 1985	Further investigation	Other	Yes	≥20	No	
Blickman, 1991	Other				No	
Bliznakova, 1998	Other				No	
Block, 1986	Other				No	
Blondeau, 1995	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Bodaghi, 1978	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Bonadio, 1987	Other				No	
Bonadio, 1990	Other				No	
Bonadio, 1991	Other				No	
Bonadio, 1991	Other				No	
Bonadio, 1992	Other				No	
Bonadio, 1993	Other				No	
Bonadio, 1993	Other				No	
Borrelli, 1983	Other				No	
Bos, 1975	Other				No	
Bosio, 1998	Other				No	
Bosio, 2002	Other				No	
Boudailliez, 1989	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Boudry, 1970	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Bouissou, 1988	Other				No	
Bouissou, 1994	Further investigation	Other	<18	≥20	No	
Braae, 1971	Other				No	
Bradley, 1971	Other				No	
Brendstrup, 1983	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Breunung, 1969	Other				No	
Breunung, 1970	Other				No	
Breunung, 1971	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Breunung, 1973	Other				No	
Bridji, 1999	Further investigation	Diagnostic accuracy	<18	≥20	No	
Brindle, 1990	Other				No	
Brindle, 1994	Other				No	
Brock, 1977	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Broderick, 1995	Other				No	
Broner, 1990	Other				No	
Brook, 1981	Other				No	
Brouhard, 1997	Other				No	
Bruna, 1997	Other				No	
Brundtland, 1973	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Bruns, 1969	Other				No	
Buchanan, 1971	Other				No	
Buchanec, 1974	Other				No	
Buchino, 1988	Other				No	
Bueschen, 1974	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Buffatti, 1977	Other				No	
Buiuc, 1980	Other				No	
Burko, 1970	Other				No	
Busch, 1984	Other				No	
Busch, 1985	Other				No	
Busch, 1985	Other				No	
Buys, 1994	Other				No	
Cakar, 1998	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Calado, 2002	Other				No	
Calver, 1988	Other				No	
Campos, 1996	Other				No	
Canarelli, 1979	Further investigation	Diagnostic accuracy	None <5	<20	No	
Cannon, 1986	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Capdevila Cogul, 2001	Other				No	
Cardiff-Oxford Bacteriuria Study Group, 1978	Other				No	
Castellanos, 1985	UTI diagnosis	Other	<18	<20	No	Culture
Cejkova, 1979	Other				No	
Chapman, 1988	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Chatys-Gorska, 1973	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Chatys-Gorska, 1974	Other				No	
Cheskis, 1972	Other				No	
Cheskis, 1973	Other				No	
Chiou, 2001	Further investigation	Diagnostic accuracy	Yes	≥20	No	
Chiou, 2001	Further investigation	Diagnostic accuracy	Yes	≥20	No	
Cho, 1998	Other				No	
Christen, 1974	Other				No	
Christian, 1998	Other				No	
Christophe, 1983	Other				No	
Chu, 2002	Other				No	
Chuang, 1989	Other				No	
Cielak-Puchalska, 2000	Other				No	
Cigna, 1992	Other				No	
Cohen, 1972	Other				No	
Conn, 1970	Other				No	
Conway, 1972	Other				No	
Cook, 1995	Further investigation	Other	None <5	≥20	No	
Cooper, 1993	Further investigation	Diagnostic accuracy	<18	≥20	No	
Corman, 1982	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Corso, 1989	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Coulthard, 1999	Other				No	
Coulthard, 2001	Other				No	
Courteau, 1980	Other				No	
Craig, 1997	Other				No	
Craig, 1998	Other				No	
Craig, 1999	Other				No	
Craig, 2000	Other				No	
Cruickshank, 1967	Other				No	
Cusins, 1973	UTI diagnosis	Other	Yes	≥20	No	Culture
Cuzzolin, 2001	Other				No	
Dacher, 1993	Other				No	
Dajani, 1980	Other				No	
Daly, 1986	UTI diagnosis	Diagnostic accuracy	None <5	≥20	No	Culture
Darge, 1996	Further investigation	Diagnostic accuracy	<12	<20	Yes	
Darge, 1997	Other				No	
Darge, 1999	Other				No	
Darge, 2001	Other				No	
Darge, 2001	Other				No	
Darge, 2002	Other				No	
Darge, 2002	Other				No	
Daschner, 1976	Other				No	
Daschner, 1976	Other				No	
Daschner, 1977	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Davey, 1997	Further investigation	Diagnostic accuracy	<12	≥20	No	
David, 1998	Other				No	
de Man, 1991	Other				No	
Defrenne, 1979	Other				No	
Deflandre, 1987	Further investigation	Other	<18	≥20	No	
Denis, 1978	Other				No	
Dennis, 1979	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Desnottes, 1978	Other				No	
Díaz Alvarez, 1998	Other				No	
Dick, 1996	Other				No	
Dikshit, 1993	Further investigation	Diagnostic accuracy	Other	≥20	Yes	
Dillon, 1984	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 x 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Dinkel, 1986	Further investigation	Other			No	
Ditchfield, 1993	Other				No	
Ditchfield, 1995	Other				No	
Ditchfield, 1998	Further investigation	Other	<18	≥20	No	
Dittrich, 1986	Other				No	
Djojohadipringgo, 1976	Other				No	
Dodge, 1969	Other				No	
Dodge, 1971	Other				No	
Dolezel, 2000	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Domic, 1991	Other				No	
Dosa, 1972	Other				No	
Downs, 1999	UTI diagnosis	SR	Other	≥20	Yes	Culture
Dubnickova, 1970	Other				No	
Dudley, 1996	Other				No	
Durbin, 1993	Other				No	
Eckstein, 1966	Other				No	
Edwards, 1975	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Edwards, 1997	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture + microscopy
Eggert, 1971	Other				No	
Eggli, 1992	Further investigation	Diagnostic accuracy	<18	<20	Yes	
Eichenwald, 1986	Other				No	
El-Hatw, 1999	Other				No	
Eliakim, 1997	UTI diagnosis	Other			No	
El-Ridi, 1986	Other				No	
Endsjo, 1967	Other				No	
Enerback, 1987	Other				No	
Erasmie, 1989	Other				No	
Ergenekon, 2000	Other				No	
Ericsson, 1971	Other				No	
Espinosa, 1991	Other				No	
Everaert, 1998	Other				No	
Eyrick, 1969	Other				No	
Falcao, 1999	Other				No	
Farhat, 2002	Other				No	
Farkas, 1969	Other				No	
Fass, 1973	Other				No	
Faunce, 1992	Other				No	
Fennell, 1976	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Fernbach, 1989	Other				No	
Ferrario, 1970	UTI diagnosis	Other	None <5		No	
Ferrera, 1997	Other				No	
Fettich, 1989	Other				No	
Fettich, 1992	Other				No	
Fischman, 1982	Further investigation	Other	Other		No	
Fletcher, 1975	Other				No	
Fletcher, 1980	Other				No	
Fliedner, 1986	Other				No	
Fliegel, 1983	Other				No	
Fowlis, 1994	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Foxman, 2002	Background				No	
Franczak, 1968	Other				No	
Franz, 1999	Background				No	
Frauscher, 1999	Other				No	
Freitag, 1973	Other				No	
Fuchs, 1969	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Furness, 1999	Other				No	
Gacinovic, 1996	Further investigation	Other			No	
Ganapathy, 1997	Other				No	
García, 1997	UTI diagnosis	Case-control	< 18	≥ 20	Yes	Culture
García Cañete, 2001	Other				No	
García, 2002	UTI diagnosis	Other	<5 only	≥ 20	No	Culture
García Muñoz, 1996	Other				No	
Gazal, 1989	Other				No	
Gbadegesin, 2002	Other				No	
Gelfand, 1999	Other				No	
Gelfand, 2000	Background				No	
Genster, 1972	UTI diagnosis	Diagnostic accuracy	None <5	≥ 20	Yes	Culture
Gentelet, 2001	UTI diagnosis	Diagnostic accuracy	Other	≥ 20	Yes	Culture
Giddens, 1998	Other				No	
Gil Salom, 1989	Other				No	
Gillenwater, 1975	Other				No	
Gillenwater, 1975	UTI diagnosis	Diagnostic accuracy	None <5	≥ 20	Yes	Culture
Gillenwater, 1976	UTI diagnosis	Diagnostic accuracy	None <5	≥ 20	Yes	Culture
Ginevri, 1992	Other				No	
Ginocchi, 1968	Other				No	
Girardet, 1970	Further investigation	Diagnostic accuracy	None <5	<20	No	
Girardet, 1970	Other				No	
Girardet, 1971	Other				No	
Girardet, 1980	UTI diagnosis	Diagnostic accuracy	None <5	≥ 20	Yes	Culture
Glazier, 1997	Other				No	
Gleason, 1974	Other				No	
Gleeson, 1991	Other				No	
Glezerov, 1975	Other				No	
Godard, 1977	Other				No	
Goldman, 2000	Other				No	
Goldraich, 1989	Other				No	
Goldsmith, 1990	UTI diagnosis	Diagnostic accuracy	None <5	≥ 20	Yes	Culture
Golebiowska, 1977	Other				No	
Goosens, 1985	UTI diagnosis	Diagnostic accuracy	<5 only	≥ 20	No	Other
Gordon, 1986	Other				No	
Gorelick, 1999	UTI diagnosis	SR	Other	≥ 20	Yes	Culture
Goszczyk, 2000	Other				No	
Graborn, 1968	Other				No	
Granados, 1998	Other				No	
Grechi, 1980	Other				No	
Grechi, 1980	Other				No	
Grechi, 1989	Other				No	
Grisaru-Soen, 2000	Other				No	
Grob, 1978	Other				No	
Groshar, 1989	Other				No	
Grunberg, 1991	Other				No	
Guareschi, 1976	Other				No	
Guidoni, 1995	Other				No	
Gupta, 1993	Other				No	
Gutierrez, 1974	Other				No	
Haahr, 1971	UTI diagnosis	Other	Other	≥ 20	No	Culture
Haahr, 1971	Other				No	
Hachey, 1992	Other				No	
Hallett, 1974	UTI diagnosis	Diagnostic accuracy	Not clear	≥ 20	No	Culture
Hamburger, 1986	Other				No	
Hamoudi, 1986	UTI diagnosis	Diagnostic accuracy	Not clear	≥ 20	Yes	Microscopy

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 x 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Hanbury, 1990	Further investigation	Diagnostic accuracy	Other	≥20	Yes	
Hansen, 1995	Further investigation	Other	<18	≥20	No	
Hansen, 1996	Further investigation	Other	<18	≥20	No	
Hansson, 1997	Other				No	
Hansson, 1998	Other				No	
Hansson, 1999	Other				No	
Haran, 1967	Other				No	
Haraoka, 1996	Other				No	
Hassan, 1998	UTI diagnosis	Diagnostic accuracy	Other		No	
Hawatmeh, 1980	Other				No	
Haycock, 1986	Background				No	
Haycock, 1991	Other				No	
Hayden, 1986	Other				No	
Haznedaroglu, 1996	Other				No	
Heldrich, 1992	Further investigation	Diagnostic accuracy	Other	≥20	Yes	
Heldrich, 2000	Other				No	
Hellerstein, 1981	Other				No	
Hellerstein, 1982	Other				No	
Hellerstein, 1984	Other				No	
Hellerstein, 1988	Other				No	
Hellerstein, 1992	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Helwig, 1978	Other				No	
Hernandez Aguado, 2001	Other				No	
Herr, 2001	Other				No	
Herrera Labarca, 1995	Other				No	
Herrera Labarca, 1997	Other				No	
Hertz, 1970	Other				No	
Hervas, 2001	Other				No	
Hesse, 1991	Other				No	
Hida, 1995	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Hilderbrand, 1981	Other				No	
Hiraga, 1985	Other				No	
Hiraoka, 1994	UTI diagnosis	Diagnostic accuracy	<18	≥20	Yes	Microscopy
Hiraoka, 1994	Other				No	
Hiraoka, 1996	Further investigation	Diagnostic accuracy	<5 only	≥20	No	
Hiraoka, 1997	Further investigation	Other	<12	<20	No	
Hiraoka, 1999	Other				No	
Hjalmas, 1967	Other				No	
Hobday, 1966	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Hoberman, 1993	UTI diagnosis	Diagnostic accuracy	<5 only	≥20	No	
Hodson, 1966	Other				No	
Hoebeke, 1996	Other				No	
Hofmann, 1985	Further investigation	Other	<18	≥20	No	
Hogg, 1987	Other				No	
Hole, 1975	Other				No	
Holemans, 1994	Other				No	
Holland, 1969	Other				No	
Holland, 1995	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Holloway, 1982	Other				No	
Holt, 1983	Other				No	
Holy, 1971	Other				No	
Honkinen, 1986	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Honkinen, 1999	Other				No	
Houston, 1969	Other				No	
Houston, 1969	UTI diagnosis	Diagnostic accuracy	Other	≥20	No	Culture
Howman-Giles, 1998	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Huang, 1992	Further investigation	Other	Not clear	≥20	No	
Huicho, 2002	UTI diagnosis	SR	Other	≥20	Yes	Culture
Huland, 1984	Other				No	
Hulbert, 1973	Other				No	
Hurlbut, 1991	UTI diagnosis	SR	Other	≥20	No	Culture
Huttunen, 1970	Other				No	
Iitaka, 1990	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Ingels, 1989	Other				No	
Isaacman, 2002	Other				No	
Itoh, 1991	Further investigation	Diagnostic accuracy	Other	≥20	No	
Itoh, 1995	Further investigation	Diagnostic accuracy	Other	≥20	Yes	
Jakobsson, 1991	Other				No	
Jakobsson, 1992	Further investigation	Diagnostic accuracy	<18	≥20	No	
Jakobsson, 1994	Other				No	
Jakobsson, 1997	Other				No	
Jakobsson, 2000	Other				No	
Jaya, 1996	Other				No	
Jequier, 1983	Other				No	
Jequier, 1983	Other				No	
Jerkins, 1988	Other				No	
Jewkes, 1990	Other				No	
Jimenez-Cruz, 1981	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Jodal, 1975	Further investigation	Diagnostic accuracy	None <5	≥20	No	
Jodal, 2000	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Jodal, 2002	Other				No	
Johnson, 1986	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Jojart, 1977	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Joseph, 1990	Other				No	
Jothilakshmi, 2001	Other				No	
Kal'tianis, 1988	Other				No	
Kal'tyanis, 1979	Other				No	
Kang, 1989	Other				No	
Kangaroo, 1985	Other				No	
Kaniewska, 1966	Other				No	
Karakatsani, 1997	Other				No	
Kass, 1992	Further investigation	Diagnostic accuracy	<12	≥20	No	
Kass, 2000	Further investigation	Other	<18	≥20	No	
Kassim, 1989	Other				No	
Kaufman, 1971	Other				No	
Kavukcu, 1998	Other				No	
Kawata, 1981	Further investigation	Other		<20	No	
Kellogg, 1992	Other				No	
Kelly, 1995	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Kenda, 1983	Other				No	
Kenda, 1998	Other				No	
Kenney, 1991	Other				No	
Kernen, 1999	Other				No	
Khan, 1984	UTI diagnosis	Other	Other	≥20	No	Culture
Kholin, 1973	Other				No	
Kienitz, 1970	Other				No	
Kiliku, 1991	Other				No	
Kjaerulff, 1979	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Klauber, 1984	Other				No	
Kobayasi, 1982	Other				No	
Koenig, 1988	Other				No	
Kohli, 1993	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Konda, 1984	Other				No	
Konda, 1989	Other				No	
Korpal-Szczyrska, 1989	Further investigation	Diagnostic accuracy	None <5	≥20	No	
Koskimies, 1995	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	
Kostiala, 1980	UTI diagnosis	Other	<18	≥20	No	Culture
Kostro, 2002	Other				No	
Krepler, 1969	Other				No	
Krepler, 1977	Other				No	
Kroigaard, 1967	Other				No	
Kumar, 1996	Other				No	
Kunin, 1975	Background				No	
Kunin, 1975	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Kunin, 1976	Other				No	
Kunin, 1976	Other				No	
Kurol, 1988	Other				No	
Kuzmanovska, 1995	Other				No	
Kuzmanovska, 1996	Translate				No	
Labbe, 1983	Other				No	
Lachner, 1984	UTI diagnosis	Other	<18	≥20	No	
Ladron De Guevara, 2001	Other				No	
Lafave, 1979	Other				No	
Laguna, 1998	Other				No	
Lagutina, 1980	Other				No	
Lahde, 1981	Further investigation	Other	Other		No	
Lamabadusuriya, 2000	Other				No	
Lanning, 1979	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Larcombe, 2002	Background				No	
Latorre, 2001	Other				No	
Latorre, 2001	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Lavocat, 1998	Further investigation	Other	<18		No	
Leanos-Miranda, 1996	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Leape, 1974	UTI diagnosis	Other	None <5	≥20	No	Culture
Lechi, 1976	Other				No	
Lee, 1991	Other				No	
Lefcoe, 1970	Other				No	
Lejeune, 1969	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Leong, 1976	Other				No	
Leonidas, 1983	Other				No	
Leonidas, 1984	Further investigation	Diagnostic accuracy	<18	≥20	No	
Leonidas, 1984	Further investigation	Diagnostic accuracy	<18	≥20	No	
Leonidas, 1995	Other				No	
Leonidas, 1999	Other				No	
Levitt, 1977	Other				No	
Lewis, 1998	UTI diagnosis	Other	<5 only	<20	No	
Li, 2002	Other				No	
Liaw, 2000	Other				No	
Lidefelt, 1989	Other				No	
Lin, 1998	Other				No	
Lines, 1970	Other				No	
Linne, 1994	Further investigation	Other	<12	≥20	No	
Lip, 1992	Other				No	
Lipshultz, 1972	Other				No	
Lirenman, 1969	UTI diagnosis	Other	<5 only	≥20	No	
Littlewood, 1972	Other				No	
Livsey, 1994	Other				No	
Lloyd, 1993	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Lohr, 1991	Other				No	
Loirat, 1978	Other				No	
Lorentz, 1979	Other				No	
Loutfi, 1998	Other				No	
Loutfi, 1999	Other				No	
Lowe, 1985	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Luders, 1966	Other				No	
Lue, 1982	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	
Lythgoe, 1992	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
MacGregor, 1966	Other				No	
MacKenzie, 1990	Other				No	
MacKenzie, 1990	Other				No	
MacKenzie, 1991	Other				No	
MacKenzie, 1992	Other				No	
Macpherson, 1986	Other				No	
Madon, 1973	Other				No	
Mafe, 1997	Other				No	
Mage, 1989	Further investigation	Other	<12	≥20	No	
Magill, 1986	Other				No	
Mahant, 2001	Other				No	
Maizels, 1979	Other				No	
Maizels, 1987	Other				No	
Majd, 1979	Other				No	
Majd, 1986	Further investigation	Diagnostic accuracy	<12	≥20	No	
Majd, 1992	Background				No	
Malaga, 1978	Other				No	
Malviya, 2000	Background				No	
Mantel, 1970	Other				No	
Margileth, 1976	Other				No	
Marild, 1982	Other				No	
Marild, 1989	Other				No	
Marosvari, 1975	Other				No	
Marr, 1975	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Marshall, 1990	Further investigation	Other	<12	≥20	No	
Martin, 1985	Further investigation	Diagnostic accuracy	<18	≥20	No	
Martin, 2001	Other				No	
Martin Aguado, 2000	Other				No	
Martin Caballero, 1983	Other				No	
Martin Puerto, 1999	Other				No	
Mason, 1984	Other				No	
Masters, 1966	Other				No	
Mastin, 1995	Further investigation	Diagnostic accuracy	Other	≥20	No	
Matajc, 1965	Other				No	
Matsaniotis, 1971	Other				No	
Mattsby-Baltzer, 1981	Other				No	
Mazzoleni, 1984	Further investigation	Other	<12	≥20	No	
McAfee, 1979	Other				No	
McAlister, 1974	Other				No	
McCarthy, 1982	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
McCauley, 1984	Further investigation	Diagnostic accuracy	<18	≥20	No	
McCormick, 1978	Other				No	
McDonald, 2000	Other				No	
McGillivray, 2002	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Meadows, 1967	Other				No	
Medhat, 1997	Further investigation	Other	Other		No	
Melis, 1992	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 x 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Mendygarina, 1983	Other				No	
Merguerian, 1997	Other				No	
Merguerian, 1999	Other				No	
Merrick, 1979	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Merrick, 1995	Further investigation	Other			No	
Merrick, 1995	Further investigation	Other			No	
Middleton, 1980	Other				No	
Mitchell, 1990	Other				No	
Miyata, 1969	Other				No	
Mok, 1979	Other				No	
Molyneux, 1995	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Mongeau, 1971	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Montplaisir, 1976	Other				No	
Moosmann, 1977	Other				No	
Morey, 1999	Background				No	
Morin de Finfe, 1989	Other				No	
Morris, 1978	Other				No	
Motoiro, 1975	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Mouratidis, 1993	Further investigation	Other	<18	≥20	No	
Mrhac, 1988	Other				No	
Mucke, 1971	Other				No	
Muller Suur, 1992	Further investigation	Diagnostic accuracy	<18	≥20	No	
Murahovschi, 1988	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Murphy, 1989	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Murphy, 1989	Other				No	
Mussarakis, 1994	Other				No	
Nakaarai, 1971	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Nanayakkar, 1988	Other				No	
Nangia, 1998	Other				No	
Narbutowicz, 1975	Other				No	
Nauschuetz, 1993	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Need, 2000	Other				No	
Nelson, 1990	Other				No	
Newman, 1967	Other				No	
Newman, 1997	Other				No	
Ng, 1971	Other				No	
N'Goran, 1989	Other				No	
Nicolet, 1983	Further investigation	Diagnostic accuracy	<18	≥20	No	
Nicolet, 1984	Further investigation	Diagnostic accuracy	<18	≥20	No	
Nicolet, 1984	Other				No	
Nitz, 1980	Other				No	
Nowak, 2000	Other				No	
Nunan, 1997	Other				No	
Nuutinen, 2001	Other				No	
Nwaorgu, 1992	Other				No	
Oak, 1999	Other				No	
Oehme, 1972	Other				No	
O'Hara, 1996	Other				No	
Okolo, 1988	Other				No	
Oldak-Skvirsky, 1994	Other				No	
Orellana, 2002	Other				No	
Ort, 1965	Other				No	
Ozturk, 1991	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Padelt, 1967	Other				No	
Padhy, 1989	Other				No	
Pahl, 1992	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Pai, 1991	Other				No	
Palmer, 1996	Other				No	
Paltiel, 1992	Other				No	
Panaretto, 1999	Other				No	
Panikratov, 1975	Other				No	
Panuel, 1984	Further investigation	Other	<18	≥20	No	
Panuel, 1984	Further investigation	Diagnostic accuracy	<18	≥20	No	
Panuel, 1992	Other				No	
Papadopoulou, 2002	Other				No	
Parkkulainen, 1971	Other				No	
Parsons, 1974	Other				No	
Patel, 1993	Other				No	
Pead, 1994	Other				No	
Pecile, 1995	Further investigation	Diagnostic accuracy	<18	≥20	No	
Peng, 1999	Further investigation	Other	<18	≥20	No	
Peng, 2001	Further investigation	Other	<18	≥20	No	
Penza, 1981	Other				No	
Perri, 1989	Other				No	
Perri, 1992	Other				No	
Pezzlo, 1988	Other				No	
Pfaller, 1987	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Piepsz, 1990	Further investigation	Diagnostic accuracy	None <5		Yes	
Piepsz, 1999	Other				No	
Piepsz, 2002	Other				No	
Poli-Merol, 1998	Further investigation	Diagnostic accuracy	<18	≥20	No	
Pollet, 1980	Further investigation	Other	Not clear	≥20	No	
Pollet, 1981	Further investigation	Other	<21	≥20	No	
Polmear, 1999	Other				No	
Porter, 1978	Other				No	
Prassopoulos, 1995	Other				No	
Preston, 1994	Other				No	
Preuss, 1969	Other				No	
Principi, 1969	Other				No	
Pronicka, 1967	Other				No	
Proesmans, 1980	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Proesmans, 1981	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Proesmans, 1992	UTI diagnosis	Diagnostic accuracy	Other	<20	No	Culture
Pryles, 1959	Other				No	
Pulatov, 1973	Other				No	
Pulliam, 2001	Other				No	
Putto, 1986	Other				No	
Pylkkanen, 1981	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Raes, 1997	Other				No	
Rahman, 1997	Other				No	
Ramzy, 1993	Other				No	
Randolph, 1970	Other				No	
Randolph, 1971	Other				No	
Randolph, 1974	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Randolph, 1979	UTI diagnosis	Other	<5 only	≥20	No	
Randolph, 1981	UTI diagnosis	Other	<5 only	≥20	No	
Raymond, 1998	Other				No	
Redman, 2001	Background				No	
Rees, 1996	Other				No	
Riccabona, 1991	Further investigation	Diagnostic accuracy	<5 only	≥20	No	
Riccabona, 2001	Other				No	
Riccabona, 2002	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 x 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Ring, 1989	Other				No	
Roberts, 2000	Background				No	
Robins, 1975	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Robson, 1996	Other				No	
Roch, 1980	Other				No	
Roderick, 1997	Other				No	
Roebuck, 1999	Further investigation	SR	None <5	≥20	Yes	
Roelants, 2001	Other				No	
Ronchetti, 1976	Other				No	
Rosenberg, 1990	Other				No	
Rosenberg, 1992	Other				No	
Rosenberg, 2001	Background				No	
Rosenberg, 2002	Other				No	
Rosenkranz, 1971	UTI diagnosis	Diagnostic accuracy	Other	≥20	No	Culture
Rossleigh, 1990	Other				No	
Rossmueller, 2001	Other				No	
Roy, 1986	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Rushton, 1992	Other				No	
Sabbag Chauvez, 1988	Other				No	
Sackey, 2000	Other				No	
Santos, 1980	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Saraga, 1996	Further investigation	Diagnostic accuracy	<12	≥20	No	
Saxton, 1995	Other				No	
Scheifele, 1978	Other				No	
Scherf, 1977	Other				No	
Schersten, 1967	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Schiff, 1993	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Schlager, 2001	Other				No	
Schreiter, 1970	Further investigation	Diagnostic accuracy	<5 only	≥20	No	
Schreiter, 1971	Other				No	
Scully, 1990	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Seigle, 1995	Other				No	
Sellin, 1982	Other				No	
Servadio, 1974	Further investigation	Diagnostic accuracy	Other	≥20	No	
Sfakianakis, 1992	Other				No	
Sfakianakis, 2000	Further investigation	Diagnostic accuracy	Other	≥20	Yes	
Shah, 1989	Other				No	
Shanon, 1992	Further investigation	Diagnostic accuracy	<18	≥20	No	
Shapiro, 1988	Further investigation	Other	<18	≥20	No	
Sharief, 2001	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Sharma, 1970	Other				No	
Shaw, 1997	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Sherwood, 1984	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Shrestha, 1973	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Siafarikas, 1977	Other				No	
Siegel, 1975	Other				No	
Sigmund, 1991	Other				No	
Sikund, 1968	Other				No	
Silbert, 1992	UTI diagnosis	Case-control	<18	≥20	Yes	Culture
Silvestro, 1983	Other				No	
Simon, 1972	Other				No	
Sinaniotis, 1978	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Sitzmann, 1970	Other				No	
Skelton, 1977	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Slosky, 1977	Other				No	
Smellie, 1989	Other				No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 × 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Smellie, 1994	Other				No	
Smellie, 1995	Other				No	
Smith, 1973	Other				No	
Smith, 1998	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Snyder, 1987	Follow-up	Case-control	Not clear	<20	Yes	
Sorantin, 1997	Other				No	
South Bedfordshire Practitioners' Group, 1990	Other				No	
South Bedfordshire Practitioners' Group, 1990	Other				No	
Sreenarasimhaiah, 1998	Background				No	
Stake, 1979	Other				No	
Stake, 1983	Other				No	
Stark, 1980	Other				No	
Steiner, 1988	Other				No	
Stepanov, 1976	Other				No	
Stokland, 1992	Other				No	
Strife, 1988	Further investigation	Diagnostic accuracy	<18	≥20	No	
Strife, 1996	Other				No	
Sty, 1987	Other				No	
Subat-Dezulovic, 1998	Other				No	
Subat-Dezulovic, 1998	Further investigation	Cross-sectional	<18	≥20	No	
Tabuki, 1981	Other				No	
Taffinder, 1984	Other				No	
Takano, 1983	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Takeda, 1994	Other				No	
Tamminen, 1978	Other				No	
Tarp, 2000	Other				No	
Tasker, 1993	Further investigation	Diagnostic accuracy	<12	≥20	No	
Taylor, 1986	Other				No	
Tereshchenko, 1987	Other				No	
Thabet, 2002	UTI diagnosis	Diagnostic accuracy	<18	≥20	Yes	Microscopy
Thayyil-Sudhan, 2000	UTI diagnosis	Diagnostic accuracy	<18	≥20	No	Culture
Thayyil-Sudhan, 2000	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Thelle, 1985	Further investigation	Diagnostic accuracy	<18	≥20	No	
Thornbury, 1965	Other				No	
Tobiansky, 1998	UTI diagnosis	RCT/CCT	<5 only	≥20	No	
Traisman, 1987	Further investigation	Diagnostic accuracy	<18	≥20	No	
Troell, 1984	Other				No	
Troger, 1999	Other				No	
Tsai, 1998	Other				No	
Tsukamoto, 1999	Other				No	
Tsukamoto, 1999	Other				No	
Tucci, 1985	Other				No	
Ugarte, 1986	Other				No	
Uhari, 1976	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Utikalova, 1974	Other				No	
Vallee, 1999	Other				No	
Van de Vijver, 2001	Other				No	
van der Voort, 1997	Other				No	
van der Voort, 1997	Other				No	
Vanderfaellie, 1998	Other				No	
Van Herreweghe, 1980	Other				No	
Varela Salgado, 1994	UTI diagnosis	Diagnostic accuracy	Other		No	

continued

TABLE 42 Studies excluded from the review (cont'd)

Study details	Does the study address diagnosis or further investigation of UTI?	Is it a diagnostic cohort study or an RCT?	Does the study include at least some children aged <5 years?	Minimum 20 participants included?	Data to construct 2 x 2 tables reported?	For studies of diagnosis, does the reference standard include culture?
Vargas Rosendo, 1984	Further investigation	Diagnostic accuracy	<18	<20	Yes	
Vazquez, 1984	Other				No	
Vegni, 1969	Other				No	
Vela Navarrete, 1969	Other				No	
Veleminsky, 1975	Other				No	
Venturoli, 1982	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Vernon, 1997	Other				No	
Verrier Jones, 1993	Other				No	
Vilka, 1980	Other				No	
Vlahakis, 1971	Further investigation	Diagnostic accuracy	Other		No	
Von Rohren, 1981	Other				No	
Wagle, 1989	UTI diagnosis	Diagnostic accuracy	Not clear	≥20	Yes	Culture
Wallin, 2001	Further investigation	Other	<5 only	≥20	No	
Walten, 1971	Other				No	
Wassner, 1992	Other				No	
Weber, 1966	Other				No	
Weingartner, 1969	Other				No	
Weitzel, 1980	Other				No	
Whitaker, 1969	Other				No	
White, 2001	Background				No	
Whyte, 1988	Other				No	
Wichmann, 1970	Other				No	
Wiebel, 1972	Other				No	
Wientzen, 1979	Other				No	
Wille, 1969	Other				No	
Wilson, 1993	Other				No	
Winnicki, 1975	Other				No	
Winqvist, 1997	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Wlodarczak, 1974	Other				No	
Wlodarczak, 1975	Other				No	
Wlodarczak, 1975	Other				No	
Wong, 1998	UTI diagnosis	Diagnostic accuracy	<5 only	≥20	No	Culture
Woodard, 1976	Other				No	
Wyszynska, 1967	Other				No	
Yen, 1994	Further investigation	Diagnostic accuracy	<18	≥20	No	
Yen, 1996	Further investigation	Other	<12	≥20	No	
Yen, 1998	Further investigation	Diagnostic accuracy	<18	≥20	No	
Yen, 1999	Other				No	
Yen, 1999	Other				No	
Yuen, 2001	UTI diagnosis	Diagnostic accuracy	<5 only	≥20	Yes	Microscopy
Yunes-Zarraga, 1997	Other				No	
Zacchello, 1966	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Zainal, 1996	UTI diagnosis	Diagnostic accuracy	None <5	≥20	Yes	Culture
Zaki, 1996	Other				No	
Zaman, 1998	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Zaman, 2001	UTI diagnosis	Diagnostic accuracy	Other	≥20	Yes	Culture
Zhilinskaia, 1973	Other				No	
Zilleruelo, 1995	Further investigation	Diagnostic accuracy	<12	≥20	No	
Zocchi, 1988	Further investigation	Diagnostic accuracy	None <5	≥20	Yes	
Zoch-Zwierz, 1979	Other				No	
Zoppardo, 1988	Further investigation	Diagnostic accuracy	<18	≥20	No	
Zwolinska, 1999	Other				No	

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Appendix 8

Downs analysis tables

TABLE 43 Quality assessment of the Downs study¹⁰

Study question	Comments
<i>General</i>	
1. Costs and effects examined	✓
2. Alternatives compared	✓
3. The viewpoint(s)/perspective of the analysis is clearly stated (e.g. NHS, society)	✗ Can be assumed to be US third party payer
<i>Selection of alternatives</i>	
4. All relevant alternatives are compared (including do-nothing if applicable)	✗ A range of additional strategies could, in principle, have been assessed, including combination and sequential strategies
5. The alternatives being compared are clearly described (who did what, to whom, where and how often)	✓
6. The rationale for choosing the alternative programmes or interventions compared is stated	✓
<i>Form of evaluation</i>	
7. The choice of form of economic evaluation is justified in relation to the questions addressed	✗ There was no clear explanation of how the results of the analysis could be used for decision-making
8. If a cost-minimisation design is chosen, have equivalent outcomes been adequately demonstrated?	NA
<i>Effectiveness data</i>	
9. The source(s) of effectiveness estimates used are stated (e.g. single study, selection of studies, systematic review, expert opinion)	✓
10. Effectiveness data from RCT or review of RCTs	✓
11. Potential biases identified (especially if data not from RCTs)	✓
12. Details of the method of synthesis or meta-analysis of estimates are given (if based on an overview of a number of effectiveness studies)	✗ A number of systematic reviews were undertaken some of which culminated in pooling, but the methods were not detailed
<i>Costs</i>	
13. All the important and relevant resource use included	✓ Assuming a payer perspective
14. All the important and relevant resource use measured accurately (with methodology)	✗ Estimates of cost from the literature
15. Appropriate unit costs estimated (with methodology)	✓
16. Unit costs reported separately from resource-use data	✗ Resource-use data not presented
17. Productivity costs treated separately from other costs	NA
18. The year and country to which unit costs apply are stated, with appropriate adjustments for inflation and/or currency conversion	✗ Country but not year
<i>Benefit measurement and valuation</i>	
19. The primary outcome measure(s) for the economic evaluation are clearly stated (cases detected, life-years, QALYs, etc.)	✓
20. Methods to value health states and other benefits are stated (e.g. time trade-off)	NA Outcomes presented in natural units
21. Details of the individuals from whom valuations were obtained are given (patients, members of the public, healthcare professionals, etc.)	NA

continued

TABLE 43 Quality assessment of the Downs study¹⁰ (cont'd)

Study question	Comments
<i>Decision modelling</i>	
22. Details of any decision model used are given (e.g. decision tree, Markov model)	✓
23. The choice of model used and the key input parameters on which it is based are adequately detailed and justified	✓
24. All model outputs described adequately.	✓
<i>Discounting</i>	
25. Discount rate used for both costs and benefits	× No discounting appears to have been undertaken for health benefits
26. Do discount rates accord with NHS guidance?	NA 3% for costs and 0% for health benefits. Not the same as NHS guidance
<i>Allowance for uncertainty</i>	
<i>Stochastic analysis of patient-level data</i>	
27. Details of statistical tests and confidence intervals are given for stochastic data	× No patient-level data NA
28. Uncertainty around cost-effectiveness expressed (e.g. confidence interval around incremental cost-effectiveness ratio, cost-effectiveness acceptability curves)	NA
29. Sensitivity analysis used to assess uncertainty in non-stochastic variables (e.g. unit costs, discount rates) and analytical decisions (e.g. methods to handle missing data)	NA
<i>Probabilistic analysis of decision models</i>	
30. Are all appropriate input parameters included with uncertainty?	×
31. Is second order uncertainty (uncertainty in means) included rather than first order (uncertainty between patients)?	NA
32. Are the probability distributions adequately detailed and appropriate?	NA
33. Sensitivity analysis used to assess variability (e.g. unit costs, discount rates) and analytical decisions (e.g. methods to handle missing data)	×
<i>Deterministic analysis</i>	
34. The approach to sensitivity analysis is given (e.g. univariate, threshold analysis)	✓ ✓ One and two way, including threshold analysis. Mostly diagrammatically
35. The choice of variables for sensitivity analysis is justified	✓
36. The ranges over which the variables are varied are stated	✓
<i>Presentation of results</i>	
37. Incremental analysis is reported using appropriate decision rules	✓
38. Major outcomes are presented in a disaggregated as well as aggregated form	✓
39. Applicable to the NHS setting	× US based and not relevant in UK setting

Appendix 9

Additional economics tables

TABLE 44 Long-term costs and QALYs lost (girls) taken from the long-term model

Distribution	Girl < 1 year		Girl 1-2 years		Girl 2-3 years		Girl > 3 years	
	Mean	SE	Mean	SE	Mean	SE	Mean	SE
Cost long-term pathway 1	£1022.735	£53.480	£980.980	£36.492	£963.149	£28.198	£956.166	£24.107
Cost long-term pathway 2	£988.162	£640.574	£616.785	£496.336	£426.143	£421.172	£220.438	£335.570
Cost long-term pathway 3	£117.774	£17.704	£103.305	£11.295	£97.539	£8.902	£92.547	£8.765
Cost long-term pathway 4	£221.768	£140.056	£138.623	£107.033	£109.429	£105.696	£56.700	£84.120
Cost long-term pathway 5	£153.299	£95.847	£50.532	£36.845	£109.429	£105.696	£28.489	£37.434
Cost long-term pathway 6	£153.299	£95.847	£50.532	£36.845	£109.429	£105.696	£28.489	£37.434
Cost long-term pathway 7	£0	£0	£0	£0	£0	£0	£0	£0
Cost long-term pathway 8	£0	£0	£0	£0	£0	£0	£0	£0
QALYs lost long-term pathway 1	0.171	0.102	0.094	0.071	0.061	0.055	0.049	0.048
QALYs lost long-term pathway 2	0.736	0.500	0.649	0.523	0.443	0.422	0.237	0.347
QALYs lost long-term pathway 3	0.047	0.031	0.025	0.019	0.016	0.015	0.008	0.014
QALYs lost long-term pathway 4	0.235	0.168	0.133	0.115	0.114	0.112	0.061	0.089
QALYs lost long-term pathway 5	0.094	0.066	0.051	0.044	0.114	0.112	0.012	0.011
QALYs lost long-term pathway 6	0.094	0.066	0.051	0.044	0.114	0.112	0.012	0.011
QALYs lost long-term pathway 7	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
QALYs lost long-term pathway 8	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000

TABLE 45 Long-term costs and QALYs lost (boys) taken from the long-term model

Distribution	Boy < 1 year		Boy 1-2 years		Boy 2-3 years		Boy > 3 year	
	Mean	SE	Mean	SE	Mean	SE	Mean	SE
Cost long-term pathway 1	£967.934	£27.802	£957.931	£26.883	£950.585	£27.723	£944.887	£27.407
Cost long-term pathway 2	£476.160	£419.396	£380.752	£397.239	£262.298	£349.554	£164.145	£300.980
Cost long-term pathway 3	£98.413	£8.421	£95.576	£8.204	£92.843	£6.974	£91.582	£7.963
Cost long-term pathway 4	£118.069	£100.803	£92.372	£94.729	£59.281	£77.327	£43.805	£83.276
Cost long-term pathway 5	£76.802	£68.023	£65.642	£65.170	£41.502	£53.913	£30.133	£55.432
Cost long-term pathway 6	£76.802	£68.023	£65.642	£65.170	£41.502	£53.913	£30.133	£55.432
Cost long-term pathway 7	£0	£0	£0	£0	£0	£0	£0	£0
Cost long-term pathway 8	£0	£0	£0	£0	£0	£0	£0	£0
QALYs lost long-term pathway 1	0.061	0.048	0.044	0.045	0.032	0.047	0.023	0.047
QALYs lost long-term pathway 2	0.426	0.384	0.349	0.347	0.239	0.318	0.145	0.252
QALYs lost long-term pathway 3	0.015	0.013	0.011	0.012	0.008	0.010	0.006	0.011
QALYs lost long-term pathway 4	0.106	0.095	0.084	0.090	0.047	0.062	0.040	0.077
QALYs lost long-term pathway 5	0.040	0.036	0.034	0.034	0.018	0.024	0.016	0.029
QALYs lost long-term pathway 6	0.040	0.036	0.034	0.034	0.018	0.024	0.016	0.029
QALYs lost long-term pathway 7	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
QALYs lost long-term pathway 8	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000

TABLE 46 The optimal strategy (and probability ranked first) for each subgroup of girls as a function of the threshold willingness to pay for an additional QALY, including glucose tests

Threshold value (£)	Girl over 3	
	Optimal	Prob. ranked 1st
0	2	0.67
1,000	2	0.757
2,000	2	0.816
3,000	2	0.854
4,000	2	0.885
5,000	2	0.902
6,000	2	0.909
7,000	2	0.91
8,000	2	0.897
9,000	2	0.873
10,000	2	0.862
11,000	2	0.846
12,000	2	0.829
13,000	2	0.805
14,000	2	0.787
15,000	2	0.77
16,000	2	0.758
17,000	2	0.745
18,000	2	0.728
19,000	2	0.714
20,000	2	0.702
21,000	2	0.691
22,000	2	0.683
23,000	2	0.662
24,000	2	0.652
25,000	29	0.306
26,000	29	0.312
27,000	29	0.32
28,000	29	0.324
29,000	29	0.335
30,000	29	0.348
31,000	29	0.351
32,000	29	0.357
33,000	29	0.362
34,000	29	0.369
35,000	29	0.373
36,000	29	0.379
37,000	29	0.382
38,000	29	0.389
39,000	29	0.393
40,000	29	0.398
41,000	29	0.401
42,000	29	0.405
43,000	29	0.408
44,000	29	0.41
45,000	29	0.417
46,000	29	0.423
47,000	29	0.425
48,000	29	0.431
49,000	29	0.434
50,000	29	0.439

Details of the strategies are given in *Table 37* of the main text.

TABLE 47 The optimal strategy (and probability ranked first) for each subgroup of boys as a function of the threshold willingness to pay for an additional QALY, including glucose tests

Threshold value	Boy over 3	
	Optimal	Prob. ranked 1st
0	2	0.685
1,000	2	0.754
2,000	2	0.807
3,000	2	0.839
4,000	2	0.876
5,000	2	0.906
6,000	2	0.917
7,000	2	0.918
8,000	2	0.915
9,000	2	0.909
10,000	2	0.906
11,000	2	0.898
12,000	2	0.884
13,000	2	0.861
14,000	2	0.849
15,000	2	0.835
16,000	2	0.815
17,000	2	0.802
18,000	2	0.786
19,000	2	0.776
20,000	2	0.76
21,000	2	0.751
22,000	2	0.74
23,000	2	0.727
24,000	2	0.72
25,000	2	0.71
26,000	2	0.7
27,000	2	0.691
28,000	2	0.682
29,000	2	0.673
30,000	2	0.668
31,000	2	0.66
32,000	2	0.654
33,000	2	0.649
34,000	2	0.64
35,000	2	0.63
36,000	2	0.623
37,000	2	0.616
38,000	2	0.606
39,000	2	0.595
40,000	2	0.586
41,000	29	0.269
42,000	29	0.274
43,000	29	0.28
44,000	29	0.282
45,000	29	0.286
46,000	29	0.293
47,000	29	0.297
48,000	29	0.299
49,000	29	0.3
50,000	29	0.302

Details of the strategies are given in *Table 37* of the main text.

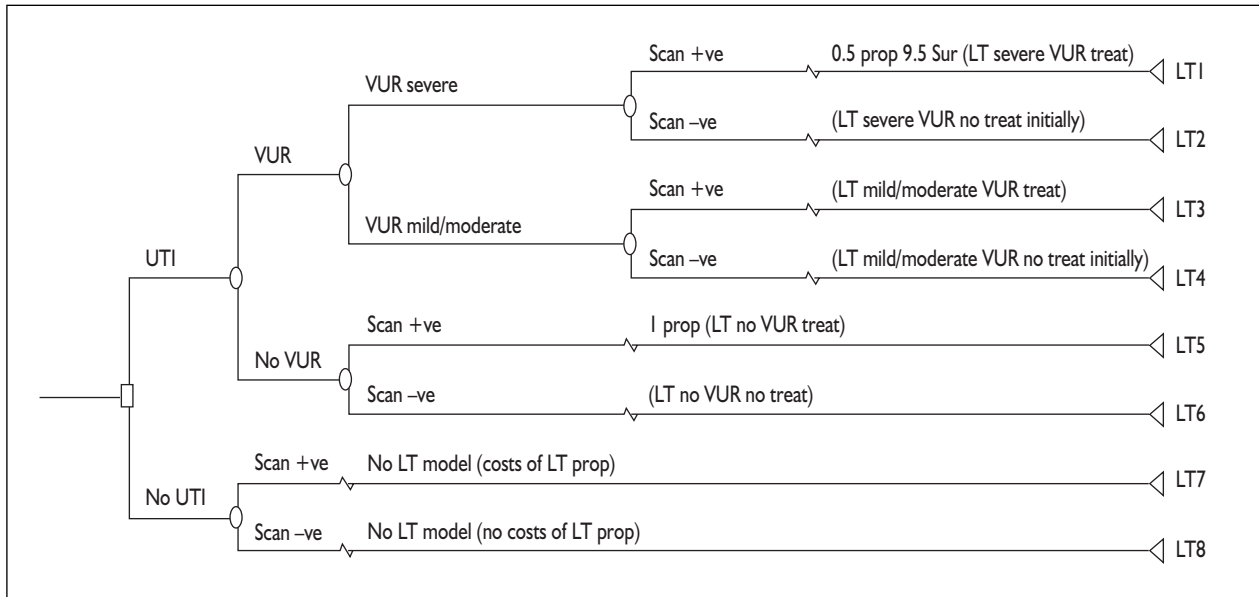


FIGURE 34 Diagnostic tree for no diagnostic tests (treat all and image all). LT, long-term; prop, prophylaxis; Sur, surgery.

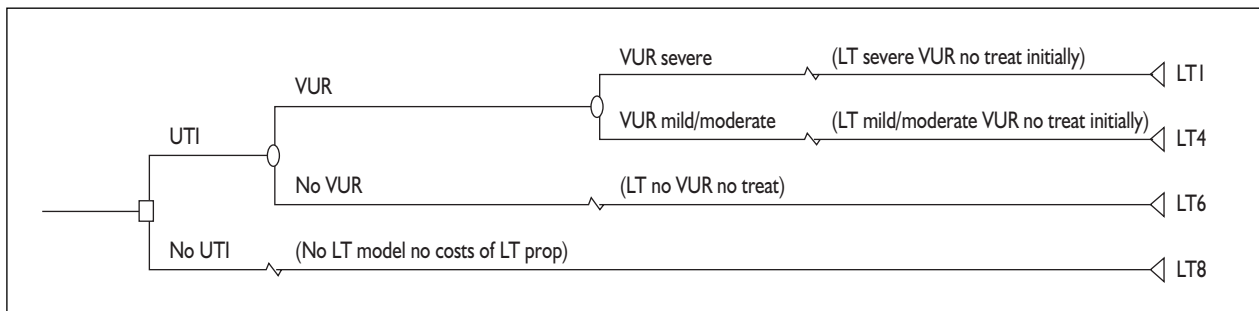


FIGURE 35 Diagnostic tree for no diagnostic tests (treat all)

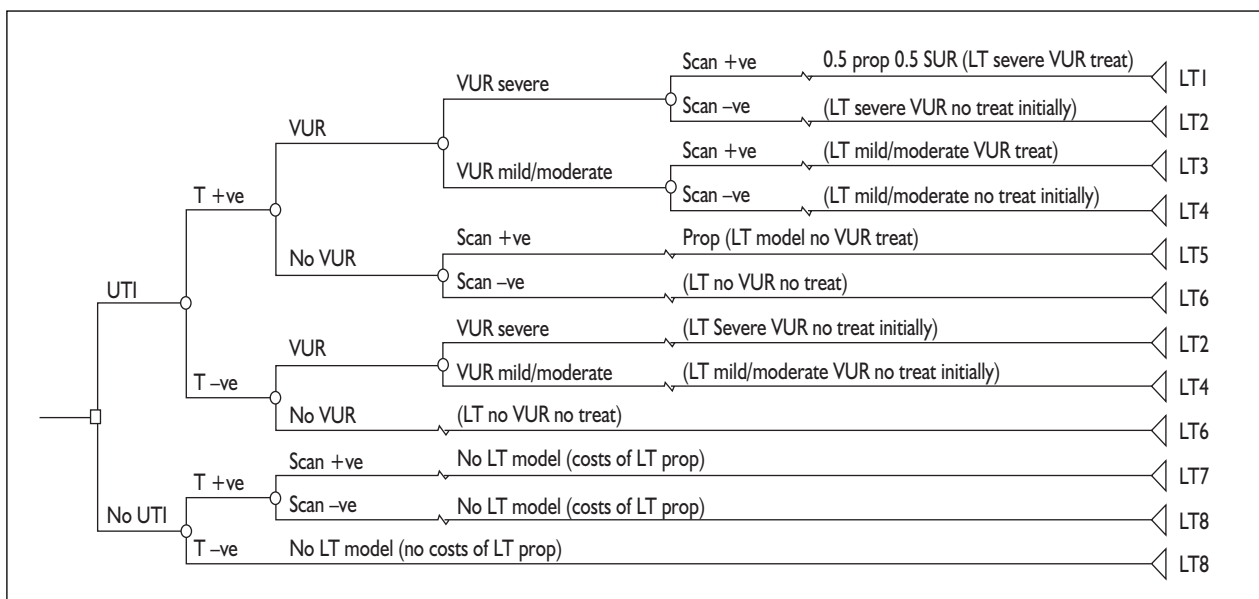


FIGURE 36 Diagnostic tree for single diagnostic test

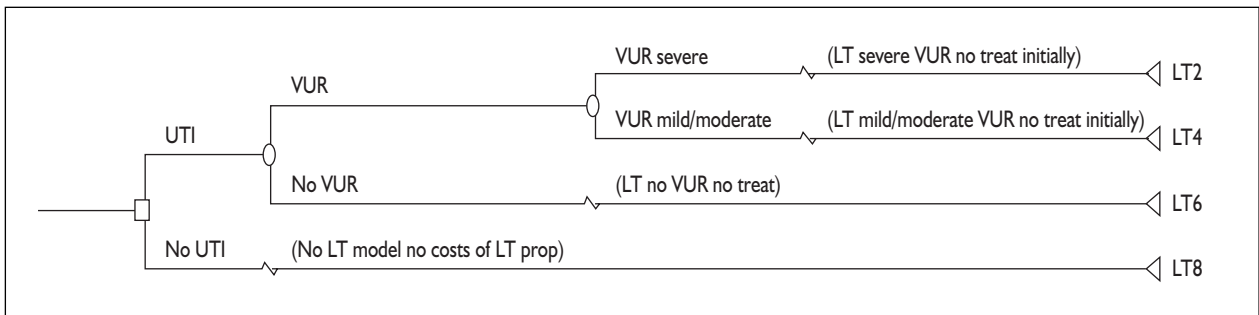


FIGURE 37 Diagnostic tree for no diagnostic tests (treat no one)

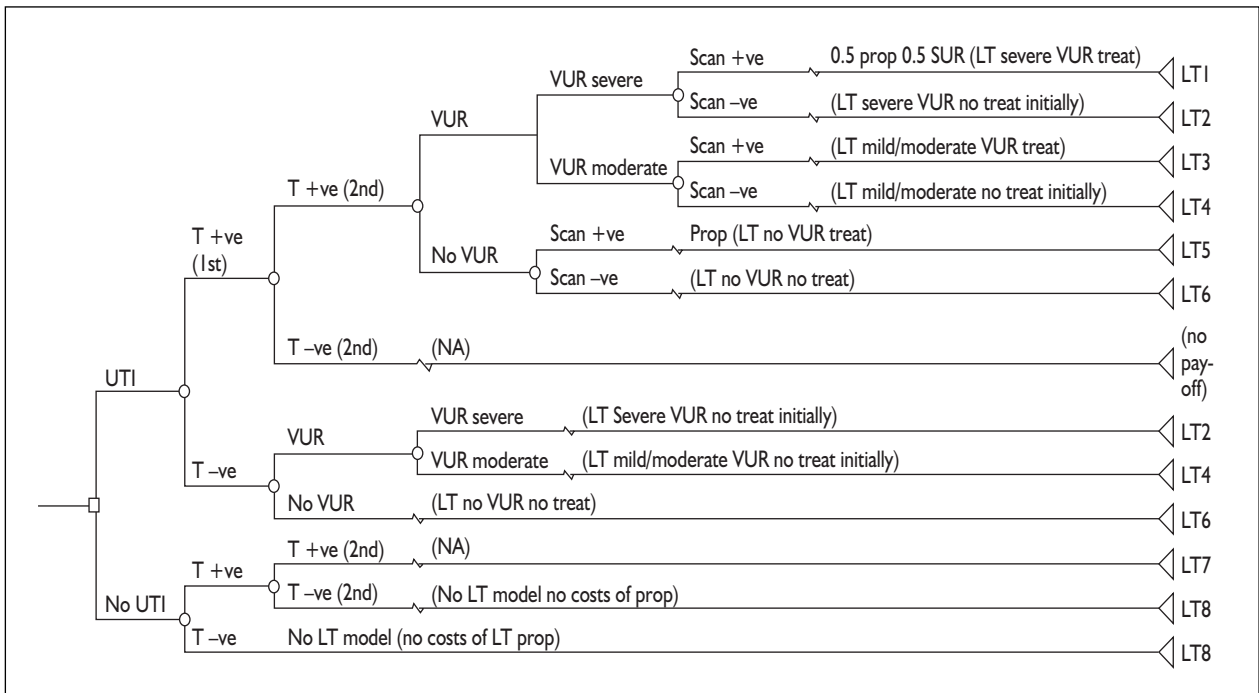


FIGURE 38 Diagnostic tree for multiple diagnostic tests (confirmatory culture)

Appendix 10

Full results of the cost-effectiveness and
Monte Carlo simulation for all strategies

TABLE 48 Simulation results (girl, < 1 year) excluding glucose

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-63.34	0.323	2	-800.74	0	39	-1169.43	0	58	-1538.13	0	64
2	-61.58	0.677	1	-785.03	0.139	33	-1146.75	0.083	50	-1508.47	0.059	53
3	-282.74	0	43	-867.25	0	52	-1159.51	0.001	53	-1451.77	0.004	51
4	-476.92	0	72	-937.39	0	64	-1167.63	0	57	-1397.86	0	46
5	-475.96	0	71	-884.48	0	56	-1088.74	0	39	-1293.00	0	26
6	-71.17	0	6	-801.48	0	40	-1166.64	0	56	-1531.79	0	59
7	-67.78	0	4	-794.14	0	35	-1157.32	0	52	-1520.51	0	56
9	-68.89	0	5	-796.51	0	36	-1160.32	0	54	-1524.13	0	57
10	-67.25	0	3	-792.95	0	34	-1155.80	0	51	-1518.64	0	55
11	-242.22	0	40	-968.86	0	70	-1332.18	0	70	-1695.51	0	70
12	-241.40	0	39	-967.10	0	69	-1329.95	0	69	-1692.80	0	69
13	-257.96	0	42	-984.31	0	72	-1347.49	0	72	-1710.66	0	72
14	-256.44	0	41	-981.02	0	71	-1343.32	0	71	-1705.61	0	71
15	-72.88	0	7	-799.43	0	37	-1162.70	0	55	-1525.98	0	58
16	-228.67	0	38	-952.11	0	68	-1313.83	0	68	-1675.55	0	68
17	-99.75	0	8	-759.70	0	28	-1089.68	0	40	-1419.66	0	49
18	-129.46	0	14	-746.24	0	25	-1054.62	0.007	30	-1363.01	0.009	37
20	-106.47	0	9	-736.45	0.056	22	-1051.44	0.042	28	-1366.43	0.024	39
21	-131.82	0	15	-740.92	0.028	24	-1045.47	0.029	26	-1350.02	0.037	34
22	-124.69	0	13	-721.70	0	19	-1020.20	0	21	-1318.71	0	29
23	-182.53	0	35	-701.30	0	16	-960.69	0	15	-1220.08	0	14
25	-139.33	0	20	-682.25	0	9	-953.71	0	14	-1225.16	0	16
26	-187.38	0	37	-692.38	0	12	-944.87	0.001	11	-1197.37	0.001	11
27	-124.21	0	12	-694.83	0	14	-980.15	0	18	-1265.46	0	21
28	-181.77	0	34	-659.54	0.013	7	-898.42	0.011	6	-1137.31	0.013	6
30	-138.66	0	18	-645.06	0.254	3	-898.25	0.187	5	-1151.45	0.135	8
31	-186.58	0	36	-647.97	0.084	4	-878.67	0.097	2	-1109.37	0.086	2
32	-301.50	0	46	-921.16	0	61	-1230.99	0	62	-1540.82	0	65
33	-294.07	0	44	-903.29	0	58	-1207.90	0	60	-1512.51	0	54
34	-300.91	0	45	-917.31	0	60	-1225.51	0	61	-1533.70	0	60
35	-341.57	0	58	-938.50	0	66	-1236.97	0	65	-1535.43	0	62
36	-120.06	0	11	-738.92	0.025	23	-1048.35	0.024	27	-1357.78	0.024	35
37	-323.31	0	51	-907.82	0	59	-1200.08	0	59	-1492.33	0	52
38	-352.55	0	62	-876.74	0	54	-1138.84	0	48	-1400.93	0	48
39	-339.75	0	56	-844.87	0	47	-1097.44	0	41	-1350.00	0	33
40	-338.43	0	54	-856.74	0	49	-1115.89	0	45	-1375.04	0	41

continued

TABLE 48 Simulation results (girl, <1 year) excluding glucose (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-414.38	0	70	-897.21	0	57	-1138.63	0	47	-1380.04	0	42
42	-161.05	0	25	-683.61	0.001	10	-944.88	0	12	-1206.16	0	12
43	-321.07	0	50	-781.54	0	32	-1011.77	0	20	-1242.01	0	17
44	-351.81	0	61	-836.01	0	45	-1078.12	0	37	-1320.22	0	30
45	-338.94	0	55	-800.50	0	38	-1031.27	0	22	-1262.05	0	20
46	-337.67	0	53	-814.83	0	42	-1053.40	0	29	-1291.98	0	25
47	-413.50	0	69	-848.58	0	48	-1066.13	0	33	-1283.67	0	24
48	-160.31	0	24	-642.58	0.184	2	-883.72	0.182	3	-1124.86	0.15	3
49	-320.11	0	49	-728.63	0.02	20	-932.89	0.106	9	-1137.15	0.22	5
50	-117.12	0	10	-777.07	0	31	-1107.05	0	43	-1437.03	0	50
51	-140.33	0	21	-757.10	0	27	-1065.49	0	32	-1373.88	0	40
53	-132.25	0	16	-762.23	0	29	-1077.23	0	36	-1392.22	0	44
54	-144.14	0	23	-753.24	0.001	26	-1057.79	0.001	31	-1362.34	0.002	36
55	-139.05	0	19	-736.06	0	21	-1034.56	0	24	-1333.07	0	31
56	-174.48	0	29	-693.26	0	13	-952.65	0	13	-1212.03	0	13
58	-162.53	0	27	-705.45	0	17	-976.91	0	17	-1248.36	0	18
59	-180.35	0	33	-685.34	0	11	-937.84	0	10	-1190.34	0.001	10
60	-138.56	0	17	-709.19	0	18	-994.51	0	19	-1279.82	0	23
61	-173.72	0	28	-651.49	0.021	5	-890.38	0.021	4	-1129.26	0.019	4
63	-161.86	0	26	-668.26	0	8	-921.45	0	8	-1174.65	0	9
64	-179.54	0	32	-640.94	0.169	1	-871.63	0.207	1	-1102.33	0.214	1
65	-312.86	0	47	-932.52	0	63	-1242.35	0	66	-1552.18	0	66
66	-317.65	0	48	-926.87	0	62	-1231.48	0	63	-1536.09	0	63
67	-329.56	0	52	-945.95	0	67	-1254.15	0	67	-1562.35	0	67
68	-341.48	0	57	-938.41	0	65	-1236.87	0	64	-1535.34	0	61
69	-143.46	0	22	-762.32	0	30	-1071.75	0	35	-1381.18	0	43
70	-346.14	0	60	-870.34	0	53	-1132.44	0	46	-1394.53	0	45
71	-353.85	0	64	-858.98	0	50	-1111.54	0	44	-1364.10	0	38
72	-363.74	0	66	-882.04	0	55	-1141.20	0	49	-1400.35	0	47
73	-381.15	0	68	-863.98	0	51	-1105.40	0	42	-1346.81	0	32
74	-177.03	0	31	-699.58	0	15	-960.85	0	16	-1222.13	0	15
75	-345.40	0	59	-829.61	0	44	-1071.72	0	34	-1313.82	0	27
76	-353.05	0	63	-814.60	0	41	-1045.38	0	25	-1276.16	0	22
77	-362.98	0	65	-840.13	0	46	-1078.71	0	38	-1317.29	0	28
78	-380.26	0	67	-815.35	0	43	-1032.90	0	23	-1250.44	0	19
79	-176.28	0	30	-658.56	0.001	6	-899.70	0.001	7	-1140.84	0.002	7

TABLE 49 Simulation results (girl, 1–2 years)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-38.35	0.318	2	-518.50	0	36	-758.58	0	42	-998.66	0	57
2	-36.63	0.682	1	-502.87	0.26	26	-735.99	0.151	33	-969.11	0.099	46
3	-260.65	0	43	-629.18	0	45	-813.45	0	52	-997.72	0	56
4	-457.59	0	71	-739.38	0	72	-880.28	0	63	-1021.18	0	58
5	-457.77	0	72	-702.19	0	70	-824.40	0	55	-946.60	0	41
6	-46.21	0	6	-519.31	0	37	-755.86	0	39	-992.41	0	55
7	-42.83	0	4	-511.99	0	31	-746.58	0	36	-981.16	0	49
9	-43.92	0	5	-514.30	0	33	-749.50	0	37	-984.69	0	50
10	-42.27	0	3	-510.74	0	30	-744.98	0	35	-979.21	0	47
11	-217.27	0	40	-686.73	0	64	-921.46	0	70	-1156.19	0	70
12	-216.45	0	39	-684.95	0	63	-919.20	0	69	-1153.45	0	69
13	-233.04	0	42	-702.23	0	71	-936.82	0	72	-1171.42	0	72
14	-231.51	0	41	-698.92	0	69	-932.62	0	71	-1166.32	0	71
15	-47.86	0	7	-517.14	0	35	-751.78	0	38	-986.42	0	52
16	-203.71	0	38	-669.94	0	58	-903.06	0	68	-1136.18	0	68
17	-76.05	0	8	-499.74	0	25	-711.59	0	30	-973.43	0	36
18	-106.51	0	14	-498.59	0.007	24	-694.64	0.008	25	-890.68	0.006	28
20	-83.49	0	9	-484.90	0.049	19	-685.61	0.027	21	-886.31	0.024	26
21	-109.45	0	16	-495.79	0.011	22	-688.97	0.027	24	-882.14	0.033	23
22	-102.25	0	12	-482.05	0	18	-671.94	0	20	-861.84	0	20
23	-161.51	0	34	-485.23	0	20	-647.09	0	16	-808.95	0	16
25	-118.29	0	20	-458.73	0	9	-628.95	0.001	10	-799.18	0.001	13
26	-167.49	0	36	-481.03	0	15	-637.80	0	12	-794.58	0	12
27	-102.34	0	13	-463.21	0	10	-643.65	0.001	14	-824.09	0	18
28	-161.66	0	35	-455.94	0.007	8	-603.08	0.008	7	-750.22	0.007	7
30	-118.41	0	21	-432.52	0.296	1	-589.57	0.254	3	-746.62	0.21	5
31	-167.64	0	37	-449.79	0.052	5	-590.86	0.072	4	-731.93	0.076	2
32	-278.23	0	46	-672.56	0	60	-869.73	0	61	-1066.89	0	62
33	-271.40	0	44	-658.07	0	52	-851.41	0	58	-1044.75	0	60
34	-278.01	0	45	-670.27	0	59	-866.40	0	60	-1062.52	0	61
35	-319.71	0	58	-697.63	0	67	-886.59	0	66	-1075.54	0	65
36	-97.58	0	11	-490.57	0.029	21	-687.06	0.033	23	-883.56	0.035	24
37	-300.89	0	49	-669.42	0	57	-853.69	0	59	-1037.95	0	59
38	-330.92	0	61	-658.44	0	53	-822.21	0	54	-985.97	0	51
39	-319.30	0	56	-633.27	0	47	-790.26	0	47	-947.25	0	42
40	-317.49	0	53	-641.67	0	49	-803.76	0	49	-965.85	0	45

continued

TABLE 49 Simulation results (girl, 1–2 years) (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-395.39	0	69	-693.87	0	65	-843.11	0	57	-992.35	0	54
42	-140.99	0	24	-466.04	0	11	-628.56	0	9	-791.08	0	11
43	-301.41	0	50	-583.20	0	39	-724.10	0	31	-864.99	0	21
44	-331.06	0	62	-629.81	0	46	-779.18	0	45	-928.55	0	38
45	-319.45	0	57	-602.10	0	40	-743.43	0	34	-884.75	0	25
46	-317.63	0	54	-612.48	0	41	-759.90	0	43	-907.32	0	33
47	-395.55	0	70	-659.80	0	55	-791.92	0	48	-924.05	0	37
48	-141.13	0	25	-436.92	0.178	2	-584.81	0.216	2	-732.71	0.212	3
49	-301.59	0	51	-546.00	0.004	38	-668.21	0.028	19	-790.42	0.094	10
50	-93.34	0	10	-517.03	0	34	-728.87	0	32	-940.72	0	40
51	-117.28	0	19	-509.37	0	28	-705.42	0	27	-901.46	0	31
53	-109.15	0	15	-510.56	0	29	-711.26	0	29	-911.97	0	34
54	-121.43	0	23	-507.77	0	27	-700.95	0	26	-894.12	0.001	29
55	-116.54	0	17	-496.33	0	23	-686.23	0	22	-876.13	0	22
56	-153.44	0	28	-477.16	0	13	-639.02	0	13	-800.88	0	14
58	-141.30	0	26	-481.74	0	17	-651.97	0	17	-822.19	0	17
59	-159.88	0	32	-473.42	0	12	-630.19	0	11	-786.97	0	9
60	-116.63	0	18	-477.50	0	14	-657.94	0	18	-838.38	0	19
61	-153.58	0	29	-447.86	0.015	4	-595.00	0.029	5	-742.15	0.031	4
63	-141.43	0	27	-455.53	0	7	-612.58	0	8	-769.64	0	8
64	-160.03	0	33	-442.18	0.091	3	-583.25	0.144	1	-724.32	0.17	1
65	-289.72	0	47	-684.05	0	62	-881.22	0	64	-1078.39	0	66
66	-294.75	0	48	-681.42	0	61	-874.76	0	62	-1068.09	0	63
67	-306.33	0	52	-698.58	0	68	-894.71	0	67	-1090.84	0	67
68	-318.72	0	55	-696.64	0	66	-885.60	0	65	-1074.55	0	64
69	-120.84	0	22	-513.83	0	32	-710.32	0	28	-906.82	0	32
70	-324.93	0	59	-652.46	0	51	-816.22	0	53	-979.99	0	48
71	-333.10	0	63	-647.08	0	50	-804.07	0	50	-961.05	0	44
72	-342.41	0	65	-666.59	0	56	-828.68	0	56	-990.77	0	53
73	-360.67	0	67	-659.15	0	54	-808.39	0	51	-957.63	0	43
74	-156.64	0	30	-481.68	0	16	-644.21	0	15	-806.73	0	15
75	-325.08	0	60	-623.82	0	43	-773.19	0	44	-922.57	0	35
76	-333.25	0	64	-615.91	0	42	-757.23	0	41	-898.56	0	30
77	-342.55	0	66	-637.39	0	48	-784.81	0	46	-932.23	0	39
78	-360.83	0	68	-625.08	0	44	-757.20	0	40	-889.32	0	27
79	-156.78	0	31	-452.57	0.001	6	-600.46	0.001	6	-748.36	0.001	6

TABLE 50 Simulation results (girl, 2–3 years)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-45.47	0.288	2	-688.82	0.001	31	-1010.50	0	38	-1332.17	0	46
2	-43.53	0.712	1	-672.76	0.315	15	-987.37	0.208	31	-1301.99	0.156	35
3	-268.42	0	43	-816.70	0	41	-1090.84	0.006	48	-1364.98	0.013	53
4	-466.53	0	71	-944.97	0	72	-1184.18	0	71	-1423.40	0	61
5	-467.21	0	72	-915.05	0	71	-1138.97	0	59	-1362.89	0	52
6	-53.25	0	6	-689.45	0	32	-1007.55	0	37	-1325.65	0	43
7	-49.84	0	4	-682.07	0	24	-998.18	0	34	-1314.30	0	39
9	-50.88	0	5	-684.27	0	28	-1000.97	0	35	-1317.67	0	40
10	-49.24	0	3	-680.74	0	21	-996.48	0	33	-1312.23	0	38
11	-224.27	0	40	-856.77	0	57	-1173.03	0	69	-1489.28	0	70
12	-223.42	0	39	-854.93	0	55	-1170.69	0	68	-1486.44	0	69
13	-239.99	0	42	-872.18	0	66	-1188.27	0	72	-1504.36	0	72
14	-238.43	0	41	-868.79	0	64	-1183.97	0	70	-1499.16	0	71
15	-54.85	0	7	-887.17	0	30	-1003.33	0	36	-1319.49	0	41
16	-210.65	0	38	-839.88	0	48	-1154.49	0	64	-1469.11	0	68
17	-83.93	0	8	-679.49	0	20	-977.27	0	26	-1275.05	0	31
18	-114.31	0	14	-682.72	0.016	26	-966.93	0.019	24	-1251.13	0.018	25
20	-91.84	0	9	-668.15	0.105	12	-956.31	0.068	20	-1244.47	0.055	22
21	-117.77	0	15	-681.24	0.058	22	-962.97	0.078	22	-1244.70	0.09	23
22	-111.07	0	12	-671.19	0	14	-951.26	0	19	-1231.32	0	20
23	-170.38	0	33	-684.09	0	27	-940.95	0	16	-1197.81	0.001	15
25	-128.13	0	20	-655.25	0	5	-918.82	0.002	9	-1182.38	0.002	11
26	-177.35	0	36	-682.27	0.002	25	-934.73	0.001	14	-1187.19	0	12
27	-111.41	0	13	-656.04	0	6	-928.35	0	12	-1200.66	0	16
28	-170.91	0	35	-660.64	0.009	10	-905.50	0.012	7	-1150.37	0.011	7
30	-128.60	0	21	-634.15	0.213	1	-886.92	0.191	2	-1139.69	0.166	4
31	-177.92	0	37	-657.16	0.038	7	-896.78	0.053	4	-1136.40	0.062	3
32	-286.57	0	46	-856.72	0	56	-1141.79	0	60	-1426.86	0	62
33	-279.83	0	44	-843.27	0	51	-1124.98	0	56	-1406.70	0	58
34	-286.26	0	45	-854.54	0	54	-1138.68	0	58	-1422.82	0	60
35	-328.15	0	56	-884.14	0	69	-1162.14	0	66	-1440.13	0	66
36	-105.65	0	11	-674.74	0.067	16	-959.29	0.069	21	-1243.83	0.059	21
37	-310.16	0	49	-858.44	0	59	-1132.59	0	57	-1406.73	0	59
38	-340.75	0	61	-857.15	0	58	-1115.34	0	54	-1373.54	0	55
39	-329.37	0	57	-834.36	0	46	-1086.85	0	46	-1339.35	0	48
40	-327.22	0	53	-840.45	0	49	-1097.06	0	49	-1353.68	0	50

continued

TABLE 50 Simulation results (girl, 2–3 years) (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-405.54	0	69	-897.27	0	70	-1143.14	0	61	-1389.00	0	57
42	-150.35	0	24	-664.88	0	11	-922.14	0.003	10	-1179.41	0.004	9
43	-311.86	0	50	-790.29	0	39	-1029.50	0	39	-1268.72	0	29
44	-341.27	0	62	-834.13	0	45	-1080.56	0	45	-1326.99	0	44
45	-329.93	0	58	-809.29	0	40	-1048.97	0	40	-1288.66	0	32
46	-327.75	0	54	-816.80	0	42	-1061.33	0	41	-1305.86	0	37
47	-406.16	0	70	-869.75	0	65	-1101.54	0	51	-1333.33	0	47
48	-150.87	0	25	-641.46	0.114	2	-886.76	0.14	1	-1132.05	0.144	2
49	-312.53	0	51	-760.37	0.001	38	-984.29	0.015	30	-1208.21	0.051	18
50	-101.48	0	10	-697.04	0	36	-994.83	0	32	-1292.61	0	33
51	-125.85	0	18	-694.26	0	34	-978.46	0	27	-1262.67	0	27
53	-117.99	0	16	-694.31	0	35	-982.46	0	28	-1270.62	0	30
54	-130.30	0	23	-693.76	0.001	33	-975.50	0.005	25	-1257.23	0.008	26
55	-125.61	0	17	-685.74	0	29	-965.81	0	23	-1245.87	0	24
56	-163.38	0	28	-677.09	0	18	-933.95	0	13	-1190.81	0	13
58	-151.59	0	26	-678.72	0	19	-942.28	0	17	-1205.84	0	17
59	-170.31	0	32	-675.23	0	17	-927.69	0	11	-180.15	0	10
60	-125.96	0	19	-670.58	0	13	-942.90	0	18	-1215.21	0	19
61	-163.91	0	29	-653.64	0.005	4	-898.51	0.008	5	-1143.37	0.011	5
63	-152.07	0	27	-657.61	0	8	-910.38	0	8	-1163.16	0	8
64	-170.87	0	34	-650.11	0.055	3	-889.73	0.121	3	-1129.36	0.148	1
65	-298.38	0	47	-868.52	0	63	-1153.59	0	63	-1438.67	0	64
66	-303.67	0	48	-867.10	0	62	-1148.82	0	62	-1430.54	0	63
67	-315.30	0	52	-883.58	0	67	-1167.72	0	67	-1451.85	0	67
68	-328.05	0	55	-884.04	0	68	-1162.04	0	65	-1440.03	0	65
69	-129.54	0	22	-698.63	0	37	-983.17	0	29	-1267.72	0	28
70	-334.97	0	59	-851.36	0	53	-1109.56	0	53	-1367.76	0	54
71	-343.61	0	63	-848.60	0	52	-1101.10	0	50	-1353.59	0	49
72	-352.97	0	65	-866.20	0	61	-1122.81	0	55	-1379.43	0	56
73	-371.87	0	67	-863.60	0	60	-1109.46	0	52	-1355.33	0	51
74	-166.77	0	30	-681.30	0	23	-938.56	0	15	-1195.83	0	14
75	-335.48	0	60	-828.35	0	44	-1074.78	0	44	-1321.21	0	42
76	-344.17	0	64	-823.54	0	43	-1063.22	0	42	-1302.90	0	36
77	-353.50	0	66	-842.55	0	50	-1087.08	0	47	-1331.61	0	45
78	-372.49	0	68	-836.08	0	47	-1067.87	0	43	-1299.66	0	34
79	-167.29	0	31	-657.88	0	9	-903.18	0.001	6	-1148.47	0.001	6

TABLE 51 Simulation results (girl, >3 years)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-27.46	0.335	2	-219.67	0.001	6	-315.78	0	23	-411.89	0	36
2	-25.71	0.665	1	-203.99	0.722	1	-293.13	0.599	2	-382.27	0.52	14
3	-252.71	0	43	-389.54	0	42	-457.95	0	40	-526.36	0	42
4	-452.29	0	71	-551.25	0	72	-600.73	0	72	-650.21	0	72
5	-453.62	0	72	-536.53	0	71	-577.99	0	71	-619.45	0	71
6	-35.29	0	6	-220.43	0	7	-312.99	0	18	-405.56	0	33
7	-31.92	0	4	-213.13	0	3	-303.74	0	7	-394.35	0	25
9	-33.06	0	5	-215.56	0	4	-306.81	0	11	-398.06	0	28
10	-31.36	0	3	-211.88	0	2	-302.13	0	5	-392.39	0	23
11	-206.36	0	40	-387.87	0	41	-478.62	0	45	-569.37	0	60
12	-205.53	0	39	-386.06	0	40	-476.32	0	44	-566.59	0	56
13	-222.08	0	42	-403.26	0	45	-493.85	0	54	-584.44	0	66
14	-220.57	0	41	-399.98	0	44	-489.68	0	52	-579.39	0	64
15	-37.00	0	7	-218.37	0	5	-309.06	0	12	-399.74	0	30
16	-192.79	0	38	-371.07	0	38	-460.21	0	41	-549.35	0	50
17	-66.77	0	8	-230.90	0	10	-312.96	0	17	-395.03	0	27
18	-97.71	0	14	-246.09	0	16	-320.28	0	25	-394.47	0	26
20	-74.51	0	9	-228.26	0.008	9	-305.13	0.014	8	-382.01	0.019	13
21	-101.03	0	16	-246.76	0	18	-319.63	0.001	24	-392.50	0.001	24
22	-94.41	0	12	-239.33	0	14	-311.79	0	15	-384.25	0	17
23	-154.60	0	33	-273.00	0	36	-332.20	0	36	-391.41	0	22
25	-110.95	0	20	-238.41	0	13	-302.14	0	6	-365.87	0	7
26	-161.28	0	36	-275.22	0	37	-332.20	0	35	-389.17	0	21
27	-95.08	0	13	-231.87	0	11	-300.26	0.001	4	-368.65	0	10
28	-155.65	0	35	-261.33	0.006	31	-314.18	0.009	21	-367.02	0.007	8
30	-111.88	0	21	-228.17	0.145	8	-286.32	0.167	1	-344.47	0.16	1
31	-162.39	0	37	-262.87	0.012	32	-313.11	0.022	19	-363.35	0.038	5
32	-269.69	0	46	-419.34	0	50	-494.17	0	55	-568.99	0	59
33	-263.04	0	44	-408.80	0	46	-481.68	0	47	-554.56	0	51
34	-269.66	0	45	-417.94	0	48	-492.09	0	53	-566.23	0	55
35	-310.58	0	53	-451.92	0	65	-522.59	0	67	-593.26	0	67
36	-88.66	0	11	-237.71	0.001	12	-312.23	0.002	16	-386.75	0.011	18
37	-293.05	0	49	-429.87	0	55	-498.29	0	59	-566.70	0	57
38	-324.48	0	61	-445.08	0	62	-505.38	0	62	-565.67	0	54
39	-313.20	0	57	-427.21	0	51	-484.21	0	49	-541.22	0	46
40	-311.45	0	55	-429.76	0	54	-488.92	0	51	-548.08	0	48

continued

TABLE 51 Simulation results (girl, > 3 years) (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-387.93	0	69	-494.53	0	70	-547.84	0	70	-601.14	0	70
42	-133.81	0	25	-253.36	0	21	-313.13	0	20	-372.90	0	11
43	-296.21	0	50	-395.17	0	43	-444.65	0	39	-494.12	0	39
44	-325.51	0	62	-433.78	0	58	-487.92	0	50	-542.06	0	47
45	-314.31	0	58	-414.87	0	47	-465.15	0	42	-515.43	0	40
46	-312.50	0	56	-418.13	0	49	-470.94	0	43	-523.75	0	41
47	-389.15	0	70	-481.02	0	69	-526.95	0	69	-572.89	0	61
48	-134.85	0	27	-241.88	0.067	15	-295.40	0.113	3	-348.92	0.127	2
49	-297.54	0	51	-380.45	0	39	-421.91	0.009	38	-463.36	0.021	38
50	-84.11	0	10	-248.24	0	19	-330.30	0	31	-412.37	0	37
51	-108.77	0	18	-257.16	0	26	-331.35	0	32	-405.54	0	32
53	-99.99	0	15	-253.73	0	23	-330.61	0	33	-407.48	0	34
54	-113.08	0	23	-258.81	0	28	-331.68	0	34	-404.55	0	31
55	-108.73	0	17	-253.65	0	22	-326.11	0	29	-398.57	0	29
56	-147.04	0	28	-265.44	0	33	-324.64	0	26	-383.85	0	16
58	-133.77	0	24	-261.22	0	30	-324.95	0	28	-388.68	0	19
59	-153.77	0	32	-267.72	0	34	-324.70	0	27	-381.67	0	12
60	-109.41	0	19	-246.19	0	17	-314.58	0	22	-382.98	0	15
61	-148.09	0	29	-253.77	0.005	24	-306.62	0.005	10	-359.46	0.01	4
63	-134.69	0	26	-250.99	0	20	-309.13	0	13	-367.28	0	9
64	-154.89	0	34	-255.37	0.033	25	-305.61	0.058	9	-355.85	0.086	3
65	-281.17	0	47	-430.82	0	56	-505.64	0	63	-580.47	0	65
66	-286.48	0	48	-432.25	0	57	-505.13	0	61	-578.01	0	63
67	-298.19	0	52	-446.47	0	63	-520.61	0	66	-594.76	0	69
68	-310.78	0	54	-452.12	0	66	-522.79	0	68	-593.46	0	68
69	-111.99	0	22	-261.03	0	29	-335.55	0	37	-410.08	0	35
70	-318.44	0	59	-439.04	0	59	-499.33	0	60	-559.63	0	53
71	-327.13	0	63	-441.14	0	60	-498.14	0	58	-555.15	0	52
72	-336.59	0	65	-454.90	0	67	-514.06	0	64	-573.22	0	62
73	-355.32	0	67	-461.93	0	68	-515.23	0	65	-568.53	0	58
74	-149.70	0	30	-269.24	0	35	-329.01	0	30	-388.78	0	20
75	-319.47	0	60	-427.74	0	52	-481.88	0	48	-536.02	0	44
76	-328.24	0	64	-428.80	0	53	-479.08	0	46	-529.36	0	43
77	-337.64	0	66	-443.27	0	61	-496.08	0	57	-548.89	0	49
78	-356.54	0	68	-448.41	0	64	-494.35	0	56	-540.28	0	45
79	-150.73	0	31	-257.76	0	27	-311.28	0	14	-364.80	0	6

TABLE 52 Simulation results (boy, < 1 year)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-40.80	0.317	2	-407.38	0	27	-590.68	0	38	-773.97	0	43
2	-38.95	0.683	1	-391.49	0.398	12	-567.75	0.252	27	-744.02	0.169	33
3	-263.82	0	43	-543.76	0	42	-683.73	0	48	-823.70	0	53
4	-461.30	0	71	-672.52	0	72	-778.12	0	72	-883.73	0	62
5	-461.74	0	72	-644.16	0	71	-735.37	0	62	-826.58	0	54
6	-48.59	0	6	-408.05	0	30	-587.77	0	37	-767.50	0	42
7	-45.18	0	4	-400.66	0	18	-578.40	0	32	-756.14	0	36
9	-46.21	0	5	-402.83	0	22	-581.14	0	35	-759.45	0	37
10	-44.65	0	3	-399.46	0	16	-576.87	0	31	-754.27	0	35
11	-219.61	0	40	-575.35	0	54	-753.22	0	67	-931.09	0	70
12	-218.81	0	39	-573.62	0	52	-751.03	0	65	-928.43	0	69
13	-235.40	0	42	-590.91	0	62	-768.66	0	71	-946.42	0	72
14	-233.86	0	41	-587.57	0	60	-764.43	0	70	-941.28	0	71
15	-50.23	0	7	-405.83	0	24	-583.63	0	36	-761.43	0	39
16	-206.06	0	38	-558.59	0	45	-734.86	0	61	-911.13	0	68
17	-79.25	0	8	-401.89	0	20	-563.21	0	25	-724.54	0	30
18	-109.86	0	14	-407.92	0.003	29	-556.96	0.003	24	-705.99	0.009	25
20	-87.26	0	9	-392.72	0.049	13	-545.45	0.053	20	-698.18	0.043	22
21	-112.81	0	15	-406.72	0.006	26	-553.68	0.016	23	-700.63	0.023	24
22	-106.04	0	12	-393.74	0	14	-537.59	0	19	-681.44	0	20
23	-165.58	0	34	-409.31	0	32	-531.18	0	17	-653.05	0	16
25	-123.17	0	20	-380.18	0	6	-508.68	0	9	-637.19	0.001	11
26	-171.54	0	36	-407.88	0	28	-526.05	0	14	-644.22	0	12
27	-106.26	0	13	-379.33	0	4	-515.86	0	11	-652.39	0	15
28	-165.92	0	35	-386.91	0.011	10	-497.40	0.014	7	-607.89	0.012	7
30	-123.47	0	21	-360.17	0.274	1	-478.52	0.258	1	-596.87	0.239	4
31	-171.90	0	37	-384.13	0.032	8	-490.24	0.06	5	-596.35	0.07	3
32	-282.10	0	46	-581.80	0	57	-731.65	0	59	-881.50	0	61
33	-274.95	0	44	-568.96	0	51	-715.97	0	56	-862.97	0	58
34	-281.44	0	45	-579.79	0	56	-728.96	0	58	-878.13	0	60
35	-321.94	0	54	-609.13	0	68	-752.72	0	66	-896.32	0	65
36	-101.01	0	11	-399.97	0.02	17	-549.44	0.032	21	-698.92	0.03	23
37	-305.17	0	49	-585.10	0	59	-725.07	0	57	-865.04	0	59
38	-335.97	0	61	-582.64	0	58	-705.98	0	54	-829.31	0	55
39	-323.72	0	57	-560.15	0	46	-678.36	0	46	-796.58	0	48
40	-321.71	0	53	-565.83	0	49	-687.89	0	49	-809.94	0	49

continued

TABLE 52 Simulation results (boy, < 1 year) (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-397.39	0	69	-621.48	0	70	-733.53	0	60	-845.58	0	57
42	-145.21	0	24	-390.46	0	11	-513.08	0	10	-635.70	0	9
43	-306.22	0	50	-517.44	0	39	-623.04	0	39	-728.65	0	31
44	-336.30	0	62	-560.77	0	47	-673.00	0	45	-785.23	0	45
45	-324.08	0	58	-536.40	0	40	-642.55	0	40	-748.71	0	34
46	-322.05	0	55	-543.48	0	41	-654.19	0	41	-764.90	0	41
47	-397.79	0	70	-595.45	0	66	-694.28	0	51	-793.11	0	47
48	-145.55	0	25	-368.31	0.145	2	-479.69	0.177	2	-591.08	0.193	2
49	-306.66	0	51	-489.08	0	38	-580.29	0.015	33	-671.50	0.053	19
50	-96.78	0	10	-419.42	0	36	-580.74	0	34	-742.07	0	32
51	-121.23	0	19	-419.30	0.001	35	-568.33	0.001	28	-717.36	0.001	27
53	-113.47	0	16	-418.93	0	33	-571.66	0	29	-724.39	0	29
54	-125.25	0	23	-419.16	0	34	-566.11	0	26	-713.07	0	26
55	-120.55	0	17	-408.25	0	31	-552.10	0	22	-695.95	0	21
56	-158.26	0	28	-402.00	0	21	-523.87	0	13	-645.74	0	13
58	-146.66	0	26	-403.67	0	23	-532.18	0	18	-660.69	0	17
59	-164.51	0	32	-400.85	0	19	-519.01	0	12	-637.18	0	10
60	-120.77	0	18	-393.84	0	15	-530.37	0	16	-666.91	0	18
61	-158.61	0	29	-379.59	0.009	5	-490.08	0.015	4	-600.57	0.019	5
63	-146.96	0	27	-383.66	0	7	-502.01	0	8	-620.37	0	8
64	-164.87	0	33	-377.09	0.052	3	-483.20	0.102	3	-589.31	0.136	1
65	-293.81	0	47	-593.51	0	65	-743.36	0	64	-893.21	0	64
66	-298.73	0	48	-592.74	0	64	-739.74	0	63	-886.75	0	63
67	-310.23	0	52	-608.57	0	67	-757.74	0	69	-906.91	0	67
68	-322.77	0	56	-609.96	0	69	-753.55	0	68	-897.14	0	66
69	-124.70	0	22	-423.66	0	37	-573.14	0	30	-722.61	0	28
70	-329.97	0	59	-576.64	0	55	-699.97	0	52	-823.31	0	52
71	-337.98	0	63	-574.41	0	53	-692.63	0	50	-810.84	0	50
72	-347.17	0	65	-591.29	0	63	-713.35	0	55	-835.40	0	56
73	-365.70	0	67	-589.79	0	61	-701.84	0	53	-813.89	0	51
74	-161.34	0	30	-406.59	0	25	-529.21	0	15	-651.84	0	14
75	-330.30	0	60	-554.76	0	44	-667.00	0	44	-779.23	0	44
76	-338.34	0	64	-550.66	0	43	-656.81	0	42	-762.97	0	40
77	-347.51	0	66	-568.94	0	50	-679.65	0	47	-790.36	0	46
78	-366.10	0	68	-563.76	0	48	-662.59	0	43	-761.42	0	38
79	-161.69	0	31	-384.45	0	9	-495.83	0.002	6	-607.21	0.002	6

TABLE 53 Simulation results (boy, 1–2 years)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-36.38	0.3	2	-354.17	0	12	-513.07	0	36	-671.97	0	38
2	-34.39	0.7	1	-338.01	0.532	2	-489.82	0.349	22	-641.63	0.26	30
3	-260.43	0	43	-504.25	0	40	-626.16	0	43	-748.07	0	48
4	-458.82	0	71	-647.52	0	72	-741.88	0	72	-836.23	0	70
5	-459.65	0	72	-625.53	0	71	-708.47	0	71	-791.40	0	58
6	-44.14	0	6	-354.77	0	14	-510.09	0	34	-665.40	0	37
7	-40.68	0	4	-347.29	0	6	-500.59	0	28	-653.89	0	34
9	-41.82	0	5	-349.70	0	8	-503.64	0	29	-657.58	0	35
10	-40.13	0	3	-346.05	0	5	-499.00	0	27	-651.96	0	32
11	-215.10	0	40	-521.96	0	45	-675.39	0	62	-828.82	0	69
12	-214.30	0	39	-520.24	0	44	-673.21	0	61	-826.18	0	68
13	-230.86	0	42	-537.44	0	53	-690.73	0	66	-844.02	0	72
14	-229.30	0	41	-534.07	0	50	-686.45	0	65	-838.84	0	71
15	-45.72	0	7	-352.44	0	10	-505.80	0	30	-659.17	0	36
16	-201.52	0	38	-505.14	0	41	-656.95	0	53	-808.76	0	63
17	-75.41	0	8	-355.73	0	17	-495.89	0	25	-636.05	0	26
18	-106.69	0	14	-366.13	0.001	25	-495.85	0.004	24	-625.57	0.004	24
20	-83.37	0	9	-349.73	0.05	9	-482.92	0.054	16	-616.10	0.057	20
21	-109.56	0	16	-365.45	0.006	24	-493.40	0.013	23	-621.35	0.016	22
22	-102.69	0	12	-355.09	0	15	-481.29	0	15	-607.49	0	19
23	-163.49	0	34	-379.40	0	36	-487.36	0	20	-595.32	0	16
25	-119.64	0	20	-347.55	0.001	7	-461.51	0.001	8	-575.47	0	9
26	-169.31	0	36	-379.00	0	35	-483.85	0	17	-588.70	0.002	14
27	-103.11	0	13	-343.94	0	4	-464.36	0	10	-584.78	0	12
28	-164.14	0	35	-362.04	0.004	23	-460.99	0.008	7	-559.94	0.01	7
30	-120.22	0	21	-332.22	0.22	1	-438.22	0.229	1	-544.22	0.198	2
31	-170.00	0	37	-360.58	0.024	22	-455.87	0.048	5	-551.16	0.056	4
32	-278.47	0	46	-539.09	0	55	-669.39	0	60	-799.70	0	61
33	-271.68	0	44	-527.65	0	47	-655.63	0	52	-783.62	0	56
34	-278.22	0	45	-537.46	0	54	-667.08	0	57	-796.70	0	60
35	-319.73	0	54	-569.58	0	67	-694.50	0	67	-819.43	0	65
36	-97.78	0	11	-357.75	0.014	18	-487.73	0.026	21	-617.72	0.035	21
37	-302.57	0	49	-546.39	0	60	-668.30	0	58	-790.21	0	57
38	-333.02	0	61	-551.13	0	61	-660.18	0	54	-769.23	0	54
39	-321.42	0	57	-531.28	0	49	-636.21	0	47	-741.14	0	46
40	-319.55	0	53	-535.22	0	52	-643.05	0	49	-750.89	0	49

continued

TABLE 53 Simulation results (boy, 1–2 years) (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-397.09	0	69	-596.33	0	70	-695.95	0	69	-795.57	0	59
42	-142.98	0	24	-359.88	0.001	21	-468.33	0.001	11	-576.78	0.002	10
43	-304.53	0	50	-493.24	0	39	-587.59	0	39	-681.94	0	39
44	-333.66	0	62	-534.15	0	51	-634.39	0	46	-734.64	0	45
45	-322.11	0	58	-512.88	0	42	-608.27	0	40	-703.65	0	40
46	-320.21	0	55	-517.80	0	43	-616.60	0	41	-715.39	0	41
47	-397.85	0	70	-576.13	0	69	-665.28	0	56	-754.42	0	50
48	-143.63	0	26	-342.68	0.103	3	-442.21	0.163	2	-541.73	0.187	1
49	-305.36	0	51	-471.24	0.001	38	-554.18	0.009	38	-637.12	0.025	28
50	-93.07	0	10	-373.39	0	30	-513.55	0	37	-653.71	0	33
51	-118.14	0	19	-377.57	0	33	-507.29	0.001	32	-637.01	0.001	27
53	-109.49	0	15	-375.86	0	31	-509.04	0	33	-642.22	0	31
54	-122.32	0	23	-378.22	0	34	-506.17	0	31	-634.12	0	25
55	-117.39	0	17	-369.79	0	26	-495.99	0	26	-622.19	0	23
56	-156.12	0	28	-372.03	0	28	-479.99	0	14	-587.95	0	13
58	-143.14	0	25	-371.05	0	27	-485.00	0	19	-598.96	0	17
59	-162.65	0	32	-372.35	0	29	-477.20	0	12	-582.05	0	11
60	-117.80	0	18	-358.64	0	19	-479.06	0	13	-599.47	0	18
61	-156.77	0	29	-354.67	0.004	13	-453.62	0.012	4	-552.57	0.016	5
63	-143.71	0	27	-355.71	0	16	-461.71	0	9	-567.72	0	8
64	-163.34	0	33	-353.92	0.038	11	-449.21	0.081	3	-544.50	0.13	3
65	-290.71	0	47	-551.33	0	62	-681.64	0	64	-811.95	0	64
66	-295.70	0	48	-551.68	0	63	-679.66	0	63	-807.65	0	62
67	-307.44	0	52	-566.68	0	66	-696.31	0	70	-825.93	0	67
68	-320.25	0	56	-570.10	0	68	-695.02	0	68	-819.95	0	66
69	-121.63	0	22	-381.60	0	37	-511.59	0	35	-641.57	0	29
70	-327.83	0	59	-545.93	0	59	-654.99	0	51	-764.04	0	53
71	-335.96	0	63	-545.82	0	58	-650.75	0	50	-755.68	0	51
72	-345.51	0	65	-561.18	0	64	-669.01	0	59	-776.85	0	55
73	-364.47	0	67	-563.71	0	65	-663.33	0	55	-762.95	0	52
74	-159.22	0	30	-376.12	0	32	-484.57	0	18	-593.02	0	15
75	-328.47	0	60	-528.95	0	48	-629.20	0	44	-729.44	0	44
76	-336.65	0	64	-527.42	0	46	-622.81	0	42	-718.19	0	42
77	-346.17	0	66	-543.76	0	57	-642.56	0	48	-741.36	0	47
78	-365.23	0	68	-543.51	0	56	-632.66	0	45	-721.80	0	43
79	-159.87	0	31	-358.92	0.001	20	-458.44	0.001	6	-557.97	0.001	6

Table 54 Simulation results (boy, 2–3 years)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-29.14	0.316	2	-219.81	0.002	6	-315.14	0	10	-410.48	0	28
2	-27.43	0.684	1	-204.17	0.755	1	-292.55	0.618	1	-380.92	0.519	3
3	-254.57	0	43	-397.58	0	41	-469.09	0	41	-540.59	0	40
4	-454.15	0	71	-566.91	0	72	-623.30	0	72	-679.68	0	72
5	-455.46	0	72	-555.28	0	71	-605.19	0	71	-655.10	0	71
6	-36.98	0	6	-220.57	0	7	-312.36	0	8	-404.15	0	23
7	-33.61	0	4	-213.27	0	3	-303.10	0	3	-392.93	0	12
9	-34.74	0	5	-215.67	0	4	-306.13	0	5	-396.60	0	16
10	-33.08	0	3	-212.08	0	2	-301.57	0	2	-391.07	0	9
11	-208.06	0	40	-388.01	0	40	-477.99	0	43	-567.97	0	49
12	-207.24	0	39	-386.24	0	39	-475.74	0	42	-565.24	0	46
13	-223.83	0	42	-403.52	0	44	-493.37	0	48	-583.22	0	60
14	-222.27	0	41	-400.14	0	43	-489.08	0	45	-578.01	0	55
15	-38.64	0	7	-218.40	0	5	-308.29	0	6	-398.17	0	18
16	-194.50	0	38	-371.25	0	38	-459.63	0	39	-548.00	0	42
17	-68.58	0	8	-234.93	0	9	-318.11	0	13	-401.28	0	21
18	-99.82	0	14	-252.79	0.001	16	-329.28	0	21	-405.76	0.001	24
20	-76.38	0	9	-233.67	0.03	8	-312.32	0.036	7	-390.96	0.04	8
21	-102.76	0	16	-253.45	0.005	17	-328.80	0.011	20	-404.15	0.014	22
22	-96.26	0	12	-247.22	0	13	-322.70	0	15	-398.18	0	19
23	-156.98	0	34	-286.04	0	36	-350.58	0	36	-415.11	0	32
25	-112.93	0	20	-249.12	0	14	-317.22	0	12	-385.32	0	4
26	-162.92	0	36	-288.30	0	37	-351.00	0	37	-413.69	0	31
27	-96.93	0	13	-241.29	0	11	-313.47	0	9	-385.65	0	5
28	-158.01	0	35	-276.86	0.001	31	-336.28	0.003	26	-395.70	0.007	15
30	-113.85	0	21	-241.00	0.12	10	-304.57	0.147	4	-368.14	0.172	1
31	-164.02	0	37	-278.57	0.006	33	-335.84	0.02	25	-393.11	0.025	13
32	-271.65	0	46	-425.63	0	48	-502.61	0	52	-579.60	0	56
33	-264.88	0	44	-415.59	0	46	-490.94	0	46	-566.30	0	48
34	-271.30	0	45	-424.56	0	47	-501.19	0	49	-577.82	0	54
35	-312.61	0	53	-459.40	0	64	-532.80	0	66	-606.20	0	67
36	-90.61	0	11	-243.93	0.013	12	-320.59	0.02	14	-397.25	0.028	17
37	-294.95	0	49	-437.96	0	52	-509.46	0	56	-580.97	0	58
38	-326.56	0	61	-457.20	0	62	-522.52	0	63	-587.84	0	61
39	-315.05	0	57	-440.42	0	54	-503.10	0	53	-565.78	0	47
40	-312.92	0	55	-442.32	0	56	-507.02	0	54	-571.72	0	51

continued

Table 54 Simulation results (boy, 2–3 years) (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-390.11	0	69	-509.07	0	70	-568.55	0	70	-628.03	0	70
42	-135.93	0	24	-265.52	0	24	-330.32	0	22	-395.12	0	14
43	-298.10	0	50	-410.87	0	45	-467.26	0	40	-523.64	0	39
44	-327.57	0	62	-448.23	0	58	-508.56	0	55	-568.88	0	50
45	-316.16	0	58	-430.67	0	49	-487.92	0	44	-545.18	0	41
46	-313.96	0	56	-433.16	0	50	-492.76	0	47	-552.36	0	43
47	-391.32	0	70	-498.37	0	69	-551.89	0	69	-605.41	0	66
48	-136.96	0	27	-256.41	0.055	19	-316.13	0	11	-375.85	0.116	2
49	-299.42	0	51	-399.24	0	42	-449.15	0.002	38	-499.06	0.005	38
50	-85.96	0	10	-252.31	0	15	-335.48	0	24	-418.65	0	36
51	-110.76	0	18	-263.73	0	22	-340.21	0	29	-416.70	0	35
53	-102.02	0	15	-259.31	0	20	-337.96	0	28	-416.60	0	34
54	-114.86	0	23	-265.55	0	25	-340.90	0	31	-416.25	0	33
55	-110.63	0	17	-261.58	0	21	-337.06	0	27	-412.54	0	30
56	-149.11	0	28	-278.18	0	32	-342.71	0	32	-407.24	0	26
58	-135.96	0	25	-272.15	0	29	-340.25	0	30	-408.35	0	27
59	-155.53	0	32	-280.91	0	34	-343.61	0	33	-406.30	0	25
60	-111.29	0	19	-255.65	0	18	-327.83	0	17	-400.01	0	20
61	-150.15	0	29	-268.99	0.005	27	-328.41	0.007	18	-387.83	0.011	7
63	-136.88	0	26	-264.03	0	23	-327.60	0	16	-391.17	0	10
64	-156.63	0	33	-271.17	0.007	28	-328.44	0.04	19	-385.72	0.06	6
65	-283.09	0	47	-437.07	0	51	-514.06	0	57	-591.05	0	63
66	-288.32	0	48	-439.03	0	53	-514.39	0	58	-589.74	0	62
67	-299.72	0	52	-452.97	0	60	-529.60	0	64	-606.23	0	68
68	-312.72	0	54	-459.52	0	65	-532.92	0	67	-606.32	0	69
69	-114.28	0	22	-267.60	0	26	-344.26	0	34	-420.92	0	37
70	-320.41	0	59	-451.05	0	59	-516.37	0	59	-581.69	0	59
71	-328.98	0	63	-454.34	0	61	-517.02	0	60	-579.71	0	57
72	-337.96	0	65	-467.36	0	67	-532.07	0	65	-596.77	0	65
73	-357.31	0	67	-476.27	0	68	-535.75	0	68	-595.23	0	64
74	-152.26	0	30	-281.85	0	35	-346.65	0	35	-411.45	0	29
75	-321.42	0	60	-442.08	0	55	-502.41	0	51	-562.73	0	45
76	-330.08	0	64	-444.59	0	57	-501.85	0	50	-559.10	0	44
77	-339.00	0	66	-458.20	0	63	-517.80	0	61	-577.40	0	53
78	-358.52	0	68	-465.56	0	66	-519.09	0	62	-572.61	0	52
79	-153.29	0	31	-272.74	0	30	-332.46	0.001	23	-392.18	0.002	11

TABLE 55 Simulation results (boy, > 3 years)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
1	-25.31	0.315	2	-198.27	0.001	6	-284.75	0	7	-371.22	0	18
2	-23.60	0.685	1	-182.64	0.79	1	-262.16	0.709	1	-341.68	0.643	1
3	-251.43	0	43	-382.06	0	43	-447.37	0	41	-512.69	0	41
4	-451.48	0	71	-557.23	0	72	-610.10	0	72	-662.98	0	72
5	-453.00	0	72	-548.23	0	71	-595.85	0	71	-643.47	0	71
6	-33.15	0	6	-199.04	0	7	-281.98	0	6	-364.93	0	11
7	-29.81	0	4	-191.81	0	3	-272.81	0	3	-353.81	0	4
9	-30.82	0	5	-193.94	0	4	-275.50	0	4	-357.06	0	5
10	-29.25	0	3	-190.54	0	2	-271.18	0	2	-351.83	0	3
11	-204.22	0	40	-366.46	0	40	-447.59	0	42	-528.71	0	43
12	-203.41	0	39	-364.72	0	39	-445.37	0	40	-526.02	0	42
13	-219.97	0	42	-381.91	0	42	-462.89	0	45	-543.86	0	49
14	-218.46	0	41	-378.64	0	41	-458.74	0	44	-538.83	0	47
15	-34.85	0	7	-196.96	0	5	-278.01	0	5	-359.06	0	7
16	-190.67	0	38	-349.71	0	38	-429.23	0	38	-508.75	0	40
17	-65.00	0	8	-216.58	0	9	-292.38	0	10	-368.17	0	13
18	-95.93	0	14	-235.52	0.002	15	-305.32	0.002	16	-375.11	0.002	22
20	-73.42	0	9	-216.47	0.032	8	-288.00	0.043	8	-359.53	0.041	8
21	-99.69	0	16	-237.17	0.001	17	-305.91	0.009	17	-374.65	0.016	20
22	-92.85	0	12	-231.80	0	13	-301.28	0	14	-370.76	0	17
23	-152.91	0	33	-272.94	0	36	-332.95	0	36	-392.96	0	36
25	-110.65	0	21	-236.06	0	16	-298.77	0	13	-361.48	0	10
26	-160.39	0	36	-277.06	0	37	-335.39	0	37	-393.73	0	37
27	-93.62	0	13	-227.25	0	10	-294.06	0	11	-360.87	0	9
28	-154.11	0	35	-265.86	0.002	32	-321.74	0.003	30	-377.61	0.004	25
30	-111.73	0	23	-229.66	0.091	12	-288.63	0.105	9	-347.60	0.106	2
31	-161.67	0	37	-269.53	0.004	35	-323.46	0.015	32	-377.39	0.017	24
32	-268.24	0	46	-408.60	0	48	-478.78	0	50	-548.96	0	53
33	-261.62	0	44	-399.15	0	45	-467.91	0	46	-536.67	0	46
34	-267.99	0	45	-407.38	0	47	-477.07	0	48	-546.76	0	52
35	-309.15	0	53	-443.23	0	62	-510.26	0	65	-577.30	0	67
36	-87.41	0	11	-227.31	0.013	11	-297.27	0.018	12	-367.22	0.028	12
37	-291.69	0	49	-422.32	0	51	-487.63	0	52	-552.95	0	56
38	-323.43	0	61	-444.66	0	64	-505.28	0	62	-565.89	0	63
39	-312.17	0	57	-428.84	0	54	-487.18	0	51	-545.52	0	51
40	-309.98	0	55	-429.56	0	55	-489.35	0	55	-549.14	0	54

continued

TABLE 55 Simulation results (boy, > 3 years) (cont'd)

Strategy no.	Threshold value											
	£0			£20,000			£30,000			£40,000		
	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank	Exp. NMB	Prob. ranked 1st	Rank
41	-386.84	0	69	-498.03	0	70	-553.63	0	70	-609.23	0	70
42	-133.11	0	24	-253.68	0	26	-313.97	0	22	-374.25	0	19
43	-295.32	0	50	-401.07	0	46	-453.94	0	43	-506.82	0	39
44	-324.60	0	62	-437.74	0	59	-494.31	0	58	-550.88	0	55
45	-313.44	0	58	-421.31	0	50	-475.24	0	47	-529.17	0	44
46	-311.19	0	56	-422.41	0	52	-478.03	0	49	-533.64	0	45
47	-388.23	0	70	-489.76	0	69	-540.53	0	69	-591.29	0	69
48	-134.30	0	26	-246.67	0.048	21	-302.86	0.061	15	-359.05	0.076	6
49	-296.84	0	52	-392.07	0.002	44	-439.69	0.004	39	-487.31	0.013	38
50	-82.35	0	10	-233.94	0	14	-309.73	0	19	-385.52	0	29
51	-107.09	0	17	-246.69	0	22	-316.48	0	26	-386.28	0	32
53	-99.24	0	15	-242.30	0	19	-313.83	0	21	-385.36	0	28
54	-111.51	0	22	-249.00	0	23	-317.74	0	27	-386.48	0	33
55	-107.21	0	18	-246.17	0	20	-315.64	0	25	-385.12	0	27
56	-145.62	0	28	-265.65	0	31	-325.66	0	33	-385.67	0	30
58	-133.79	0	25	-259.21	0	28	-321.91	0	31	-384.62	0	26
59	-152.52	0	32	-269.19	0	33	-327.52	0	34	-385.86	0	31
60	-107.98	0	19	-241.61	0	18	-308.42	0	18	-375.23	0	23
61	-146.82	0	29	-258.57	0	27	-314.45	0.002	23	-370.32	0.005	15
63	-134.87	0	27	-252.80	0	25	-311.77	0	20	-370.74	0	16
64	-153.80	0	34	-261.66	0.014	29	-315.59	0.029	24	-369.53	0.049	14
65	-279.73	0	47	-420.09	0	49	-490.27	0	56	-560.44	0	62
66	-284.92	0	48	-422.44	0	53	-491.20	0	57	-559.96	0	61
67	-296.56	0	51	-435.95	0	58	-505.64	0	63	-575.34	0	65
68	-309.28	0	54	-443.35	0	63	-510.38	0	66	-577.42	0	68
69	-110.63	0	20	-250.54	0	24	-320.49	0	29	-390.44	0	35
70	-317.38	0	59	-438.61	0	60	-499.23	0	59	-559.84	0	60
71	-325.88	0	63	-442.55	0	61	-500.89	0	60	-559.23	0	59
72	-335.26	0	65	-454.84	0	66	-514.63	0	67	-574.42	0	64
73	-354.15	0	67	-465.34	0	68	-520.94	0	68	-576.54	0	66
74	-148.76	0	30	-269.33	0	34	-329.61	0	35	-389.90	0	34
75	-318.55	0	60	-431.69	0	56	-488.26	0	53	-544.83	0	50
76	-327.15	0	64	-435.02	0	57	-488.95	0	54	-542.88	0	48
77	-336.47	0	66	-447.69	0	65	-503.31	0	61	-558.92	0	58
78	-355.54	0	68	-457.07	0	67	-507.84	0	64	-558.60	0	57
79	-149.94	0	31	-262.32	0	30	-318.51	0	28	-374.69	0	21

Appendix II

Protocol changes

Inclusion criteria

Population

Very few studies were identified that included only children aged less than 5 years, and those that did were generally restricted to children aged less than 2. It was therefore decided to expand the inclusion criteria so that studies that included children aged less than 18 years were eligible for inclusion, provided that they also included children aged less than 5.

Study design

The large majority of studies identified were diagnostic cohort studies, with only a small number of diagnostic case-control studies meeting inclusion criteria. As diagnostic case-control

studies are subject to considerably more bias than diagnostic cohort studies, it was decided to limit the diagnostic accuracy studies included in the review to diagnostic cohort studies.

Existing systematic reviews

It was decided not to include existing systematic reviews in the current review, but instead to use these as sources of potentially relevant studies. This was due to differences between the identified systematic reviews and the proposed inclusion criteria and methods of analysis for the current review.

Feedback

The HTA Programme and the authors would like to know your views about this report.

The Correspondence Page on the HTA website (<http://www.hta.ac.uk>) is a convenient way to publish your comments. If you prefer, you can send your comments to the address below, telling us whether you would like us to transfer them to the website.

We look forward to hearing from you.