PHIMOSIS

Background
At the end of the first year of life, retraction of the foreskin behind the glanular sulcus is possible in only about 50% of boys. The phimosis is either primary (physiological), with no sign of scarring, or secondary (pathological), resulting from scarring due to conditions such as balanitis xerotica obliterans.

Phimosis must be distinguished from normal agglutination of the foreskin to the glans, which is a physiological phenomenon. If the tip remains narrow and glanular adhesions are separated, then the space is filled with urine during voiding, causing the foreskin to balloon outward.
**Treatment**

**Conservative treatment**
Administration of a corticoid ointment or cream is an option for primary phimosis with a success rate of > 90%, but with a recurrence rate of 17%. Agglutination of the foreskin does not respond to steroid treatment.

**Circumcision: indication and contraindication**
Childhood circumcision should not be recommended without a medical reason. An absolute indication for circumcision is secondary phimosis. Contraindications for circumcision are acute local infection and congenital anomalies of the penis, particularly hypospadias or buried penis, as the foreskin may be required for a reconstructive procedure.

Plastic circumcision (dorsal incision, partial circumcision) carries the potential for recurrence of phimosis. Associated frenulum breve is corrected by frenulotomy. Meatoplasty is added if necessary.

**Paraphimosis**
It is characterised by retracted foreskin with the constrictive ring localised at the level of the sulcus. A dorsal incision of the constrictive ring may be required, or circumcision is carried out immediately or in a second session.

**MANAGEMENT OF UNDESCENDED TESTES**

**Background**
Cryptorchidism or undescended testis is one of the most common congenital malformations of male neonates. In newborn cases with non-palpable or undescended testes on both sides and any sign of disorders of sex development (DSDs) like concomitant hypospadias, urgent endocrinological and genetic evaluation is required.
Classification
The term cryptorchidism is most often used synonymously for undescended testes. The most useful classification of undescended testes is into palpable and non-palpable testes, and clinical management is decided by the location and presence of the testes. Approximately 80% of all undescended testes are palpable.

Palpable testes include true undescended testes and ectopic testes. Non-palpable testes include intra-abdominal, inguinal, absent, and sometimes also some ectopic testes.

Most importantly, the diagnosis of a palpable or non-palpable testis needs to be confirmed once the child is under general anaesthesia, as the first step of any surgical procedure for undescended testes. See figure 1.

Diagnostic evaluation
History taking and physical examination are key in evaluating boys with undescended testes. Localisation studies using different imaging modalities are usually without any additional benefit.

Management
Treatment should be started at the age of 6 months. After that age, undescended testes rarely descend. Any kind of treatment leading to a scrotally positioned testis should be finished by 12 months, or 18 months at the latest, because histological examination of undescended testes at that age has already revealed a progressive loss of germ cells and Leydig cells. The early timing of treatment is also driven by the final adult results on spermatogenesis and hormone production, as well as on the risk of tumour development. See figure 2.

Medical therapy for testicular descent
Unfortunately, most of the studies on hormonal treatment
have been of poor quality, with heterogeneous and mixed patient populations, testis location, schedules and dosages of hormonal administration. Additionally, long-term data are almost completely lacking.

Hormonal therapy using human chorionic gonadotropin or gonadotrophin-releasing hormone (GnRH) is based on the hormonal dependence of testicular descent, but has a maximum success rate of only 20%. In general, success rates depend on testicular location. The higher the testis is located prior to therapy, the lower the success rate. The Panel consensus is that endocrine treatment to achieve testicular descent is not recommended (LE: 4, GR: C).

**Medical therapy for fertility potential**

Hormonal treatment may improve fertility indices and therefore serve as an additional tool to orchidopexy. It is still unknown whether this effect on testicular histology persists into adulthood but it has been shown that men who were treated in childhood with buserelin had better semen analyses compared with men who had childhood orchidopexy alone or placebo treatment.

Identification of specific subgroups of boys with undescended testes who would benefit from such an approach using hormones is difficult. The Panel consensus recommends endocrine treatment with GnRH analogues for boys with bilateral undescended testes to preserve the fertility potential (LE: 4, GR: C).

**Surgical therapy**

If a testis has not concluded its descent at the age of six months (corrected for gestational age), and since spontaneous testicular descent is unlikely to occur after that age, surgery should be performed within the subsequent year, at age 18 months at the latest.
Palpable testes
Surgery for palpable testes includes orchidofunicolysis and orchidopexy, either via an inguinal or scrotal approach.

Non-palpable testes
For non-palpable testes, surgery must clearly determine whether a testis is present or not. If a testis is found, the decision has to be made to remove it or bring it down to the scrotum. An important step in surgery is a thorough re-examination once the boy is under general anaesthesia, since a previously non-palpable testis might be identifiable and subsequently change the surgical approach to standard inguinal orchidopexy, as described above. Otherwise, the easiest and most accurate way to locate an intra-abdominal testis is diagnostic laparoscopy. Subsequent removal or orchidolysis and orchidopexy can be carried out using the same approach to achieve the therapeutic aims.

In case of a vanishing testis, the procedure is finished once blind-ending spermatic vessels are clearly identified. If the vessels enter the inguinal canal, one may find an atrophic testis upon inguinal exploration or a healthy testis that needs to undergo standard orchidopexy. A peeping testis can be placed down in the scrotum laparoscopically or via an inguinal incision. Placement of an intra-abdominal testis can sometimes be a surgical challenge. Usually, testes lying > 2 cm above the internal inguinal ring may not reach the scrotum without division of the testicular vessels. Under such circumstances, a Fowler–Stephens orchidopexy might be an option.

Undescended testes and fertility
The association of undescended testes with compromised fertility is extensively discussed in the literature and seems to be a result of multiple factors, including germ cell loss,
impaired germ cell maturation, Leydig cell diminution, and testicular fibrosis.

Although boys with one undescended testis have a lower fertility rate, they have the same paternity rate as those with bilateral descended testes. Boys with bilateral undescended testes suffer both, lower fertility and paternity rates.

Regarding preservation of fertility potential, early surgical correction of undescended testes is highly recommended before 12 months of age, and 18 months at the latest.

**Undescended testes and malignancy**

Boys who are treated for an undescended testis have an increased risk of developing testicular malignancy. Screening and self-examination both during and after puberty is therefore recommended.

A systematic review and meta-analysis of the literature concluded that pre-pubertal orchidopexy may reduce the risk of testicular cancer and that early surgical intervention is indicated in boys with undescended testes.
Boys with retractile testes do not need medical or surgical treatment, but close follow-up until puberty is recommended.

Surgical orchidolysis and orchidopexy are strongly recommended before the age of 12 months, and by 18 months at the latest.

Male neonates with bilateral non-palpable testes should be evaluated for possible DSDs.

In case of non-palpable testes and no evidence of DSDs, laparoscopy is recommended because of its excellent sensitivity and specificity in identifying an intra-abdominal testis, as well as the possibility for subsequent treatment in the same session.

Hormonal therapy, either in an adjuvant or neo-adjuvant setting, is not routinely recommended. Patients have to be evaluated on an individual basis.

In case of bilateral undescended testes, endocrine treatment is recommended.

For an undescended testis in a post-pubertal boy or older, with a normal contralateral testis, removal should be discussed with the patient/parents because of the theoretical risk of a later malignancy.

DSD = disorders of sex development.
Figure 1: Classification of undescended testes

Undescended testis

- Palpable
  - Inguinal
  - Ectopic
  - Retractile

- Non-palpable
  - Inguinal
  - Ectopic (Intra-abdominal)
  - Absent
    - Agenesis
    - Vanishing testis
HYDROCELE

**Background**

Incomplete obliteration of the processus vaginalis peritonei results in formation of various types of communicating hydrocele, alone or connected with other intrascrotal pathology (hernia).

*Non-communicating hydroceles* are found secondary to minor
trauma, testicular torsion, epididymitis, or varicocele operation, or may appear as a recurrence after primary repair of a communicating hydrocele.

A *communicating hydrocele* vacillates in size, usually relative to activity. It is diagnosed by medical history and physical investigation, the swelling is translucent, and transillumination of the scrotum confirms the diagnosis. If there are any doubts about the intrascrotal mass, ultrasound (US) should be performed. Contralateral disease should be excluded.

**Surgical treatment**

Surgical treatment of hydrocele is not indicated within the first 12-24 months because of the tendency for spontaneous resolution.

Early surgery is indicated if there is suspicion of a concomitant inguinal hernia or underlying testicular pathology. There is no evidence that this type of hydrocele risks testicular damage.

In the paediatric age group, the operation consists of ligation of the patent processus vaginalis via an inguinal incision, leaving the distal stump open, whereas in hydrocele of the cord, the cystic mass is excised or unroofed. Sclerosing agents should not be used because of the risk of chemical peritonitis in the communicating processus vaginalis peritonei.

The scrotal approach (Lord or Jaboulay technique) is used in the treatment of a secondary non-communicating hydrocele.
In the majority of infants, observe hydrocele for 12 months prior to considering surgical treatment.  

Perform early surgery if there is suspicion of a concomitant inguinal hernia or underlying testicular pathology.  

Perform a scrotal US in case of doubt about the character of an intrascrotal mass.  

Do not use sclerosing agents because of the risk for chemical peritonitis.

**US** = *ultrasound.*

**HYPOSPADIAS**

**Background**

Hypospadias are usually classified according to the anatomical location of the proximally displaced urethral orifice:  
- distal - anterior hypospadias  
  (glanular, coronal or distal penile);  
- intermediate - middle (penile);  
- proximal - posterior (penoscrotal, scrotal, perineal).  
The pathology may be much more severe after skin release.

**Assessment**

Patients with hypospadias should be diagnosed at birth. The diagnostic evaluation also includes an assessment of associated anomalies, which are cryptorchidism and open processus vaginalis or inguinal hernia. Severe hypospadias with unilaterally or bilaterally impalpable testis, or with ambiguous genitalia, require a complete genetic and endocrine work-up immediately after birth to exclude disorders of sex development, especially congenital adrenal hyperplasia.
Trickling urine and ballooning of the urethra require exclusion of meatal stenosis.

The length of the hypospadiac penis may be distorted by penile curvature, by penoscrotal transposition, or may be smaller due to hypogonadism.

Differentiation between functionally necessary and aesthetically feasible operative procedures is important for therapeutic decision-making. As all surgical procedures carry the risk of complications, thorough pre-operative counselling of the parents is crucial. The therapeutic objectives are to correct the penile curvature, to form a neo-urethra of an adequate size, to bring the neomeatus to the tip of the glans, if possible, and to achieve an overall acceptable cosmetic appearance. This goal is achieved by using different surgical techniques according to the individual findings.

**Surgery**
For repeat hypospadias repairs, no definitive guidelines can be given.

**Outcome**
Excellent long-term functional and cosmetic results can be achieved after repair of anterior penile hypospadias. The complication rate in proximal hypospadias repair is higher. Figure 3 provides an algorithm for the management of hypospadias.
Figure 3: Algorithm for the management of hypospadias

DSD = disorders of sex development; GAP = glans approximation procedure; TIP = tubularised incised plate urethroplasty; MAGPI = meatal advancement and glanuloplasty incorporated.
MICROPENIS
Micropenis is defined as a small but otherwise normally formed penis with a stretched length of less than 2.5 cm ± SD below the mean (Table 1).

<table>
<thead>
<tr>
<th>Age</th>
<th>Mean ± SD (cm)</th>
</tr>
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<tbody>
<tr>
<td>Newborns</td>
<td>3.5 ± 0.4</td>
</tr>
<tr>
<td>0-5 months</td>
<td>3.9 ± 0.8</td>
</tr>
<tr>
<td>6-12 months</td>
<td>4.3 ± 0.8</td>
</tr>
<tr>
<td>1-2 y</td>
<td>4.7 ± 0.8</td>
</tr>
<tr>
<td>2-3 y</td>
<td>5.1 ± 0.9</td>
</tr>
<tr>
<td>3-4 y</td>
<td>5.5 ± 0.9</td>
</tr>
<tr>
<td>4-5 y</td>
<td>5.7 ± 0.9</td>
</tr>
<tr>
<td>5-6 y</td>
<td>6.0 ± 0.9</td>
</tr>
<tr>
<td>6-7 y</td>
<td>6.1 ± 0.9</td>
</tr>
<tr>
<td>7-8 y</td>
<td>6.2 ± 1.0</td>
</tr>
<tr>
<td>8-9 y</td>
<td>6.3 ± 1.0</td>
</tr>
<tr>
<td>9-10 y</td>
<td>6.3 ± 1.0</td>
</tr>
<tr>
<td>10-11 y</td>
<td>6.4 ± 1.1</td>
</tr>
<tr>
<td>Adults</td>
<td>13.3 ± 1.6</td>
</tr>
</tbody>
</table>

VARICOCELE IN CHILDREN AND ADOLESCENTS
Background
Varicocele is unusual in boys under 10 years of age, but becomes more frequent at the beginning of puberty. Fertility problems will arise in about 20% of adolescents with varicocele. The adverse influence of varicocele increases with time.

Testicular catch-up growth and improvement in sperm parameters after varicocelectomy has been reported in adolescents. Varicocele is mostly asymptomatic, rarely
causing pain at this age. It may be noticed by the patient or parents, or discovered by the paediatrician at a routine visit. Diagnosis and classification depends upon the clinical finding and US investigation.

**Surgical treatment**
Surgical intervention is based on ligation or occlusion of the internal spermatic veins. Microsurgical lymphatic-sparing repairs (microscopic or laparoscopic) are associated with the lowest recurrence and complication rates. There is no evidence that treatment of varicocele at paediatric age will offer a better andrological outcome than an operation performed later.

**Conservative treatment - Follow-up**
During adolescence, testicular size should be checked annually. After adolescence, repeated sperm analysis is to be recommended. Figure 4 shows an algorithm for the diagnosis of varicocele in children and adolescents, and Figure 5 shows an algorithm for its treatment.
Figure 4: Algorithm for the diagnosis of varicocele in children and adolescents

Varicocele in children and adolescents

Physical examination in the upright position

Grade I - Valsalva positive
Grade II - Palpable
Grade III - Visible

Ultrasound investigation

Venous reflux detected on Doppler ultrasound

Size of the testes
Figure 5: Algorithm for the management of varicocele in children and adolescents

**Varicocele in children and adolescents**

- Surgery
- Conservative treatment

**Indication**
- Small testis (growth arrest)
- Additional testicular pathology
- Bilateral palpable varicocele
- Pathological spermogram
- Symptomatic varicocele

**Type**
- Microsurgical lymphaticsparing repair (microscopic or laparoscopic)

**Indication**
- Symmetrical testes
- Normal spermogram (in older adolescents)

**Type**
- Measurement of testicular size (during adolescence)
- Repeated sperm analysis (after adolescence)

**URINARY TRACT INFECTIONS IN CHILDREN**

**Epidemiology, aetiology and pathophysiology**

Urinary tract infections (UTIs) represent the most common bacterial infection in children. In neonates, the symptoms differ in many aspects from those in infants and children. The prevalence is higher; there is a male predominance; infections not caused by *Escherichia coli* are more frequent, and there is a higher risk of urosepsis.

Classification according to:
- **Site**: Lower urinary tract (cystitis) versus upper urinary tract (pyelonephritis);
• **Episode:** first UTI versus unresolved infection, persistent infection and reinfection;
• **Severity:** simple UTI versus severe UTI;
• **Symptoms:** asymptomatic bacteriuria versus symptomatic UTI;
• **Complicating factors** uncomplicated versus complicated UTI.

**Diagnostic evaluation**
Diagnosis includes a medical history, searching for clinical signs and symptoms and a complete physical examination.

**Urine sampling, analysis and culture**
Urine sampling has to be performed before any antimicrobial agent is administered. The technique for obtaining urine is important to confirm or exclude UTI. Sampling in neonates, infants and non-toilet-trained children:
• **Plastic bag:** (high incidence of false positive results [85-99%]). Only helpful to exclude a UTI if the dipstick is negative for leukocyte esterase and the culture results are negative, otherwise the UTI has to be confirmed by a more specific method.
• **Clean-catch urine collection:** has a false-positive rate of 5% and false-negative rate of 12% and the contamination rate is higher compared to SPA.
• **Bladder catheterisation:** In female infants and in neonates, this technique may be an alternative to SPA, however with a higher contamination rate.
• **Supra-pubic bladder aspiration (SPA):** This is the most sensitive method to obtain an uncontaminated urine sample in non-toilet trained children.
• **Midstream urine** in toilet-trained, children who can void on command, could be an acceptable technique for obtaining urine after cleaning the urethral meatus and perineum.
Urinalysis:
- **Dipsticks:** are ready to use and helpful when the result is positive, because it is highly specific.
- **Microscopy:** can be used after centrifugation as well as in uncentrifuged urine and has been demonstrated to be sensitive for UTI. This is rarely done in an outpatient setting.
- **Flow imaging analysis technology:** is being increasingly used to classify particles in uncentrifuged urine specimens and correlates well with manual methods.
- **Urine culture** is generally not necessary after negative results for dipstick, microscopic or automated urinalysis. If the dipstick result is positive, confirmation by urine culture is strongly recommended.

Pyuria without bacteriuria (sterile pyuria) may be due to incomplete antibiotic treatment, urolithiasis, or foreign bodies in the urinary tract, and infections caused by *Mycobacterium tuberculosis* or *Chlamydia trachomatis*.

**Table 2: Criteria for UTI in children (adapted from the EAU Guidelines on Urological Infections 2015)**

<table>
<thead>
<tr>
<th>Urine specimen from suprapubic bladder puncture</th>
<th>Urine specimen from bladder catheterisation</th>
<th>Urine specimen from midstream void</th>
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</thead>
<tbody>
<tr>
<td>Any number of cfu/mL (at least 10 identical colonies)</td>
<td>$&gt; 10^3 - 10^5$ cfu/mL</td>
<td>$&gt; 10^4$ cfu/mL with symptoms $&gt; 10^5$ cfu/mL without symptoms</td>
</tr>
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</table>

**Imaging**

**Ultrasound** of the kidneys and bladder US as soon as possible is advised in infants with febrile UTI to exclude obstruction of the upper and lower urinary tract and post-void residual urine should be measured in toilet-trained children to exclude
voiding abnormalities as a cause of UTI. **Radionuclide scanning:** changes in dimercaptosuccinic acid (DMSA) clearance during acute UTI (up to 4-6 weeks) indicating pyelonephritis and renal scars can be detected after 3-6 months. This correlates well with the presence of dilating reflux and the risk of further pyelonephritis episodes, breakthrough infections and future renal scarring. **Voiding cystourethrography:** is the gold standard to exclude or confirm vesicoureteral reflux, due to the risk of renal scarring. VCUG is recommended after the first episode of febrile UTI in boys and girls depending on sex, age and clinical presentation (see Figure 6). Another option is doing DMSA first, followed by VCUG if there is renal cortical uptake deficiency after UTI.

**Bladder and bowel dysfunction (BBD)** are risk factors for which each child with UTI should be screened upon presentation. If there are signs of BBD at infection-free intervals, further diagnosis and effective treatment are strongly recommended.

Status of circumcision should be checked in boys and treatment of the phimosis considered in those with pyelonephritis.

**Management**

**Administration route** the choice between oral and parenteral therapy should be based on patient age; clinical suspicion of urosepsis; illness severity; refusal of fluids, food and/or oral medication; vomiting; diarrhoea; non-compliance; and complicated pyelonephritis (e.g. urinary obstruction). Febrile UTI in early infancy should be treated by i.v. fluids and antibiotics and under close monitoring within the hospital.

**Duration of therapy:** outcomes of short courses (1-3 days) are inferior to those of 7-14 day courses. In late infancy, oral therapy with a third-generation cephalosporin (e.g. cefixime
or ceftibuten) is equivalent to the usual 2-4 days intravenous therapy followed by oral treatment in uncomplicated UTI’s. In complicated UTI parenteral treatment with broad-spectrum antibiotics is preferred.

**Figure 6: Algorithm for the management of a first febrile UTI**

- **First febrile UTI**
  - Ultrasound upper and lower urinary tract
  - **No pathological findings**
    - Boys > 12 months
    - Imaging after recurrent infections
    - Toilet trained children: exclusion of BBD
  - Infant/girl
    - Exclusion of reflux/VCUG/DMSA
  - **Suspicion of VUR and/or pyelonephritis**
  - **Upper tract dilatation/hydronephrosis**
    - Complicated UTI/close monitoring i.v. antibiotic treatment
    - If good response:
      - Further evaluation of upper tract function (renal scan/MRI)
      - Exclusion of VUR (VCUG)
    - If critical clinical status or no response:
      - Consider transient urinary diversion

*BBD = Bladder Bowel Dysfunction; DMSA = technetium⁹⁹⁻ labelled dimercaptosuccinic acid; MRI = magnetic resonance imaging; UTI = urinary tract infection; VCUG = voiding cystourethrography; VUR = vesicoureteral reflux.*
Monitoring of UTI
Urine usually becomes sterile after 24 h, and leukocyturia normally disappears within 3-4 days. Normalisation of body temperature can be expected within 24-48 h after the start of therapy in 90% of cases. In patients with prolonged fever and failing recovery, treatment-resistant uropathogens or the presence of congenital uropathy or acute urinary obstruction should be considered. Immediate US examination is recommended.

Procalcitonin (among other laboratory inflammatory parameters such as C-reactive protein and leukocyte count) is a reliable serum marker for early prediction of renal parenchymal inflammation. In patients with febrile UTI, serum electrolytes and blood cell counts should be obtained.
<table>
<thead>
<tr>
<th>Recommendations</th>
<th>LE</th>
<th>GR</th>
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<tr>
<td>Take a medical history, assess clinical signs and symptoms and perform a physical examination to diagnose children suspected of having a UTI.</td>
<td>3</td>
<td>A</td>
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<tr>
<td>Exclude bladder- and bowel dysfunction and obstruction in any child with febrile and/or recurrent UTI.</td>
<td>3</td>
<td>A</td>
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<td>Do not delay diagnosis and treatment of bladder-bowel-dysfunction.</td>
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<td>A</td>
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<tr>
<td>Collect an uncontaminated urine sample in an infant through suprapubic bladder aspiration.</td>
<td>2a</td>
<td>B</td>
</tr>
<tr>
<td>Bladder catheterisation is an alternative (traumatic especially in boys).</td>
<td>2a</td>
<td>B</td>
</tr>
<tr>
<td>Do not use plastic bags to for urine sampling in non-toilet-trained children since it has a high risk of false-positive results.</td>
<td>2a</td>
<td>B</td>
</tr>
<tr>
<td>Clean catch urine is an acceptable technique for toilet-trained children.</td>
<td>2a</td>
<td>B</td>
</tr>
<tr>
<td>Urinalysis by dipstick yields rapid results, but it should be used with caution. Microscopic investigation is the standard method of assessing pyuria after centrifugation. Using flow imaging analysis, the numbers of WBCs, squamous epithelial cells and red cells correlate well with manual methods.</td>
<td>2a</td>
<td>B</td>
</tr>
<tr>
<td>The choice between oral and parenteral therapy should be based on patient age; clinical suspicion of urosepsis; illness severity; refusal of fluids, food and/or oral medication; vomiting; diarrhoea; non-compliance; complicated pyelonephritis.</td>
<td>2a</td>
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Treat UTIs with 4-7 day courses of oral or parenteral therapy. Do not use of short courses (1-3 days) since outcomes are inferior.

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Offer long-term antibacterial prophylaxis in case of high susceptibility to UTI and risk of acquired renal damage and LUTS.

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Treat complicated UTI, with broad-spectrum antibiotics (parenteral).

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In infants with febrile UTI, use renal and bladder ultrasonography to exclude obstruction of the upper and lower urinary tract.

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<td>3</td>
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In all infants, exclude VUR after the first episode of febrile UTI, using VCUG or a DMSA-scan first (in case of a positive DMSA-scan, follow-up with VCUG). In boys > 1 year of age, exclude VUR after the second febrile UTI.

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<tbody>
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</table>

DMSA = dimercaptosuccinic acid; LUTS = lower urinary tract symptoms; UTI = urinary tract infections; VCUG = voiding cystourethography; VUR = vesicoureteral reflux; WBC = white blood cell.

**MONOSYMPATOMATC NOCTURNAL ENURESIS**

**Background**

Enuresis is incontinence during the night. Any wetting during sleep above the age of five years is enuresis. It is important to note that there is a single symptom only. Due to an imbalance between night-time urine output and night-time bladder capacity, the bladder can easily become full at night, and the child will either wake-up to empty the bladder or will void during sleep.
Assessment
A voiding diary, registering the day-time bladder function and the night-time urine output will help guide the treatment. Measuring the day-time bladder capacity gives an estimate of bladder capacity to compare with normal values for age. Figure 7 presents an algorithm for the diagnosis and treatment of monosymptomatic nocturnal enuresis.
Fig. 7: Algorithm for the diagnosis and management of monosymptomatic nocturnal enuresis

Ab = antibody; Ach = acetylcholine.
**Vesicoureteric reflux (VUR) in children**
VUR presents with a wide range of severities, and the majority of reflux patients will not develop renal scars and probably will not need any intervention. The main goal in management is the preservation of kidney function.

**Diagnosis**
The diagnostic work-up should evaluate the overall health and development of the child. A basic diagnostic work-up includes a detailed medical history (including family history, and screening for lower urinary tract dysfunction [LUTD]), physical examination including blood pressure measurement, urinalysis (assessing proteinuria), urine culture, and serum creatinine in patients with bilateral renal parenchymal abnormalities.

**Prenatally diagnosed hydronephrosis**
Ultrasound of the kidney and bladder is the first standard evaluation tool for children with prenatally diagnosed hydronephrosis. It should be delayed until the end of first week after birth because of early oliguria in the neonate. It is essential to evaluate the bladder, as well as the kidneys.

<table>
<thead>
<tr>
<th><strong>Recommendations for paediatric screening of VUR</strong></th>
</tr>
</thead>
<tbody>
<tr>
<td>Inform parents of children with VUR that siblings and offspring have a high prevalence of VUR.</td>
</tr>
<tr>
<td>Use renal US for screening of sibling(s).</td>
</tr>
<tr>
<td>Use VCUG if there is evidence of renal scarring on US or a history of UTI.</td>
</tr>
<tr>
<td>Do not screen older toilet-trained children since there is no added value in screening for VUR.</td>
</tr>
</tbody>
</table>

*US = ultrasound; UTI = urinary tract infection; VCU = voiding cystourethrography; VUR = vesicoureteral reflux.*
Conservative therapy
The objective of conservative therapy is prevention of febrile UTI. It is based on the understanding that:

- VUR resolves spontaneously, mostly in young patients with low-grade reflux. However, spontaneous resolution is low for bilateral high-grade reflux.
- VUR does not damage the kidney when patients are free of infection and have normal LUT function.
- There is no evidence that small scars can cause hypertension, renal insufficiency or problems during pregnancy.
- The conservative approach includes watchful waiting, intermittent or continuous antibiotic prophylaxis, and bladder rehabilitation in those with LUTD.
- Circumcision during early infancy may be considered as part of the conservative approach, because it is effective in reducing the risk of infection in normal children.

Surgical treatment
Surgical treatment comprises endoscopic injection of bulking agents or ureteral reimplantation.

Subureteric infection of bulking agents: Due to the availability of biodegradable substances, endoscopic subureteric injection of bulking agents has become an alternative to long-term antibiotic prophylaxis and surgical intervention.

Open surgical techniques: Overall, all surgical procedures offer very high and similar success rates for correcting VUR.

Laparoscopy: A laparoscopic approach cannot be recommended as a routine procedure. It can be offered as an alternative to the parents in centres where there is enough experience.
<table>
<thead>
<tr>
<th>Recommendations for the management of vesicoureteric reflux in childhood</th>
<th>GR</th>
</tr>
</thead>
<tbody>
<tr>
<td>Initially treat all patients diagnosed within the first year of life with continuous antibiotic prophylaxis, regardless of the grade of reflux or presence of renal scars.</td>
<td>C</td>
</tr>
<tr>
<td>Offer immediate, parenteral antibiotic treatment for febrile breakthrough infections.</td>
<td>A</td>
</tr>
<tr>
<td>Offer definitive surgical or endoscopic correction to patients with frequent breakthrough infections.</td>
<td>A</td>
</tr>
<tr>
<td>Offer surgical correction to patients with persistent high-grade reflux (grades IV/V) if intervention is needed; the outcome of open surgical correction is better than endoscopic correction for higher grades of reflux, whereas satisfactory results can be achieved by endoscopic injection for lower grades.</td>
<td>B</td>
</tr>
<tr>
<td>Initially manage all children presenting at age 1-5 years conservatively.</td>
<td>B</td>
</tr>
<tr>
<td>Offer surgical repair to children presenting with high-grade reflux or abnormal renal parenchyma.</td>
<td>B</td>
</tr>
<tr>
<td>Offer close surveillance without antibiotic prophylaxis to children presenting with lower grades of reflux and without symptoms.</td>
<td>B</td>
</tr>
<tr>
<td>Ensure that a detailed investigation for the presence of LUTD is done in all children after toilet-training. If LUTD is found, the initial treatment should always be for LUTD.</td>
<td>A</td>
</tr>
<tr>
<td>Consider surgical correction, if parents prefer definitive therapy to conservative management. Endoscopic treatment is an option for all children with low grades of reflux.</td>
<td>B</td>
</tr>
</tbody>
</table>
Select the most appropriate management option based on:
- the presence of renal scars;
- clinical course;
- the grade of reflux;
- ipsilateral renal function;
- bilaterality;
- bladder function;
- associated anomalies of the urinary tract;
- age and gender;
- compliance;
- parental preference.

In high-risk patients who already have renal impairment, a more aggressive, multidisciplinary approach is needed.

*LUTD = lower urinary tract dysfunction.*

### Table 3: Management and follow-up according to different risk groups

<table>
<thead>
<tr>
<th>Risk Groups</th>
<th>Presentation</th>
<th>Initial treatment</th>
</tr>
</thead>
<tbody>
<tr>
<td>High</td>
<td>Symptomatic male or female patients after toilet-training with high-grade reflux (grades IV/V), abnormal kidneys and LUTD</td>
<td>Initial treatment is always for LUTD with CAP; intervention may be considered in cases of breakthrough infections or persistent reflux</td>
</tr>
<tr>
<td>High</td>
<td>Symptomatic male or female patients after toilet-training with high-grade reflux (grade IV/V), abnormal kidneys and no LUTD</td>
<td>Intervention should be considered</td>
</tr>
<tr>
<td>Moderate</td>
<td>Symptomatic male or female patients before toilet-training, with high-grade reflux and abnormal kidneys</td>
<td>CAP is the initial treatment. Intervention may be considered in cases of breakthrough infections or persistent reflux</td>
</tr>
<tr>
<td>Moderate</td>
<td>Asymptomatic patients (PNH or sibling) with high-grade reflux and abnormal kidneys</td>
<td>CAP is the initial treatment. Intervention may be considered in cases of breakthrough infections or persistent reflux</td>
</tr>
<tr>
<td>Risk Groups</td>
<td>Presentation</td>
<td>Initial treatment</td>
</tr>
<tr>
<td>-------------</td>
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</tr>
<tr>
<td>Category</td>
<td>Description</td>
<td>Treatment Options</td>
</tr>
<tr>
<td>----------</td>
<td>-------------</td>
<td>-------------------</td>
</tr>
<tr>
<td>Moderate</td>
<td>Symptomatic male or female patients after toilet-training, with high-grade reflux and normal kidneys with LUTD</td>
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</tr>
<tr>
<td>Moderate</td>
<td>Symptomatic male or female patients after toilet-training, with high-grade reflux and normal kidneys with LUTD</td>
<td>Choice of treatment is controversial. Endoscopic treatment may be an option. LUTD treatment should be given if needed</td>
</tr>
<tr>
<td>Moderate</td>
<td>All symptomatic patients with normal kidneys, with low-grade reflux, with LUTD</td>
<td>Initial treatment is always for LUTD with or without CAP</td>
</tr>
<tr>
<td>Low</td>
<td>All symptomatic patients with normal kidneys, with low-grade reflux, with no LUTD</td>
<td>No treatment or CAP</td>
</tr>
<tr>
<td>Low</td>
<td>All asymptomatic patients with normal kidneys with low-grade reflux</td>
<td>No treatment or CAP in infants</td>
</tr>
</tbody>
</table>

CAP = continuous antibiotic prophylaxis; LUTD = lower urinary tract dysfunction; PNH = prenatal diagnosed hydronephrosis; UTI = urinary tract infection; VCUG = voiding cystourethrography.
<table>
<thead>
<tr>
<th>Symptomatology</th>
<th>Treatment</th>
<th>Follow-up for UTI, LUTD, and kidney status until after puberty</th>
</tr>
</thead>
<tbody>
<tr>
<td>Moderate Symptomatic</td>
<td>Initial treatment is always for LUTD with or without CAP. Intervention may be considered in cases of breakthrough infections or persistent reflux.</td>
<td></td>
</tr>
<tr>
<td>All symptomatic patients with normal kidneys and low-grade reflux, with LUTD</td>
<td>Endoscopic treatment may be an option. LUTD treatment should be given if needed.</td>
<td></td>
</tr>
<tr>
<td>Low Symptomatic</td>
<td>No treatment or CAP in infants. If no treatment is given, parents should be informed about risk of infection.</td>
<td></td>
</tr>
<tr>
<td>All asymptomatic</td>
<td>Follow-up for UTI and LUTD</td>
<td></td>
</tr>
<tr>
<td>patients with normal kidneys and low-grade reflux</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

In case of persistent LUTD, despite urotherapy, intervention should be considered. The choice of intervention is controversial. Follow-up for UTI and LUTD, kidney status; full re-evaluation after successful urotherapy.