

Guidelines on Neuro-Urology

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1. INTRODUCTION

1.1 Aim

The European Association of Urology (EAU) Neuro-Urology Guidelines aim to provide information for clinical practitioners on the incidence, definitions, diagnosis, therapy, and follow-up of neuro-urological disorders. These Guidelines reflect the current opinion of experts in this specific pathology and thus represent a state-of-the-art reference for all clinicians, as of the publication date.

The terminology used and the diagnostic procedures advised throughout these Guidelines follow the recommendations for investigations on the lower urinary tract (LUT) as published by the International Continence Society (ICS) [1-4]. Readers are advised to consult the other EAU Guidelines which may address different aspects of the topics discussed in this document.

1.2 Publication history

The EAU published the first Neuro-Urology Guidelines in 2003 with updates in 2008, 2014 and 2015. In 2009, a review paper was published in European Urology [5].

Standard procedure for EAU Guidelines includes an annual assessment of newly published literature in the field to guide future updates. A shorter reference document, the Pocket Guidelines, is also available, both in print and as a mobile application, presenting the main findings of the Neuro-Urology Guidelines. These versions are abridged and therefore may require consultation with the full text version. All are available through the EAU website: <http://www.uroweb.org/guidelines/>.

For this 2015 print updates were made to:

- Chapter 3A: A new table summarising epidemiology of neuro-urological disorders has been added (Table 1) and text in this chapter has consequently been replaced.
- Chapter 3D: The sections on botulinum toxin sphincter injection (3D.2.5.4) and surgical treatment (3D.2.6) have been revised and updated.
- Chapter 3F: Sexual (dys)function and fertility has been revised and updated.

Additionally, the text has been significantly reduced so that only key information is included and re-formatted according to the EAU template for non-oncology Guidelines so that all Guidelines follow a similar format.

This document was peer-reviewed prior to publication.

1.3 Panel composition

The EAU Neuro-Urology Guidelines panel consists of an international multidisciplinary group of experts, including urologists specialised in the care of spinal cord injured (SCI) patients and a specialist in the field of urodynamic technologies.

1.4 Background

The function of the LUT is mainly storage and voiding of urine, which is regulated by the nervous system that coordinates the activity of the urinary bladder and bladder outlet. Any disturbance of the nervous system involved, including the peripheral nerves in the pelvis, can result in neuro-urological symptoms. Depending on the extent and location of the disturbance, a variety of different LUT changes might occur, which can be symptomatic or asymptomatic. Moreover, neuro-urological symptoms can cause a variety of long-term complications; the most dangerous being deterioration in renal function. Since symptoms and long-term complications do not correlate [6], it is important to identify patients with neuro-urological symptoms, and establish if they have a low or high-risk of subsequent complications.

According to current knowledge, elevated storage pressure in the bladder, either alone or combined with vesicoureteric reflux (VUR), is the most important risk factor for renal damage [7]. Sustained elevated storage pressure in the bladder is mainly due to a combination of increased detrusor activity during the storage phase (detrusor overactivity [DO] or low compliance), combined with detrusor-sphincter dyssynergia (DSD). The combination of these findings is usually caused by suprasacral infrapontine spinal lesions. Furthermore, elevated detrusor leak point pressure has been demonstrated to be a risk factor for renal deterioration in patients with meningomyelocele [8]. Therefore, renal failure has been the leading cause of death in patients with SCI for a long time [9]. Even today, 26% of patients with meningomyelocele who do not undergo urological treatment develop renal damage. Detrusor leak point pressure > 40 cm H₂O and low bladder compliance are the main risk factors for renal damage [10].

In recent years, adequate diagnosis and treatment of neuro-urological symptoms in patients with spinal cord lesions have improved the situation of these patients. Nowadays, respiratory diseases are the most frequent (21%) cause of death in patients with SCI [11].

In all other patients with neuro-urological symptoms, the risk of renal damage is significantly lower. However, in multiple sclerosis (MS), urodynamics and clinical symptoms may not correlate, which means that asymptomatic patients can present with abnormal urodynamic findings [12]. LUT symptoms do not always lead to urological evaluation in MS patients, even if the symptoms are troublesome [13]. Therefore, urological assessment is important [14]; although respiratory diseases are currently the leading cause of death in MS patients [15].

In Parkinson's disease (PD), neuro-urological disorders have not been reported as a significant cause of death. Moreover, patients with PD commonly suffer from overactive bladder without DSD [16], which does not seem to be as threatening to the upper urinary tract (UUT) as DO with DSD. In PD, urodynamic diagnosis of DO correlates well with diagnosis made by questionnaires [17]. Therefore, regular urodynamic follow-up might be less important in patients with PD compared with MS or SCI. The same is true for diabetes mellitus, which frequently leads to neuro-urological symptoms [18], but cardiovascular diseases are the main cause of death in these patients [19].

In summary, treatment and intensity of follow-up examinations are based on the type of neuro-urological disorder and the underlying cause.

2. METHODS

There is a need for ongoing re-evaluation of the information presented in the current Guidelines by an expert Panel. It must be emphasised that clinical guidelines present the best evidence available to the experts but following guideline recommendations will not necessarily result in the best outcome. Guidelines can never replace clinical expertise when making treatment decisions for individual patients, but rather help to focus decisions - also taking personal values and preferences/individual circumstances of patients into account.

Literature searches were carried out for all sections of the Neuro-Urology Guidelines. Focus of all searches was identification of all level 1 scientific papers (systematic reviews and meta-analyses of randomised controlled trials) in accordance with EAU methodology. If sufficient data was identified to answer the clinical question, the search was not expanded to include lower level literature. Searches were carried out in Medline and Embase on the Ovid platform. The searches used the controlled terminology of the respective databases.

References used in this text are graded according to their level of evidence (LE) and Guidelines are given a grade of recommendation (GR), according to a classification system modified from the Oxford Centre for Evidence-Based Medicine Levels of Evidence [20]. The aim of grading recommendations is to provide transparency between the underlying evidence and the recommendation given. In this 2015 EAU Guidelines compilation, all standard information on LE and GR has been taken out of the individual Guidelines topics for the sake of brevity. The methodology section (see the introduction chapter of the complete book) outlines the LE and GR criteria which are used throughout the Guidelines.

3. THE GUIDELINE

3A EPIDEMIOLOGY, AETIOLOGY AND PATHOPHYSIOLOGY

3A.1 Introduction

Neuro-urological symptoms may be caused by various diseases and events affecting the nervous systems controlling the LUT. The resulting neuro-urological symptoms depend grossly on the location and the extent of the neurological lesion. There are no exact figures on the overall prevalence of neuro-urological disorders in the general population, but data are available on the prevalence of the underlying conditions and the relative risk of those for the development of neuro-urological symptoms. It is important to note that the majority of the data shows a very wide range of prevalence/incidence figures. This reflects the variability in the cohort (e.g. early or late stage disease) and the frequently smaller sample sizes, resulting in low level of evidence in most published data (summarised in Table 1).

Table 1: Epidemiology of Neuro-Urological Disorders

Suprapontine and pontine lesions and diseases		
Neurological Disease	Frequency in General Population	Type and Frequency of Neuro-Urological Symptoms
Cerebrovascular accident (Strokes)	450 cases/100,000/yr (Europe) [21] (10% of cardiovascular mortality)	Nocturia - OAB - UUI - DO (other patterns less frequent) [22]. 57-83% of neuro-urological symptoms at 1 month post stroke, 80% of spontaneous recovery at 6 months [23]. Persistence of UI correlates with poor prognosis [24].
Dementias: Alzheimer's disease (80%), Vascular (10%), Other (10%)	6.4% of adults > 65 yrs [25, 26]	OAB - UUI - DO 25% of incontinence in Alzheimer's disease, ≥ 25% in other dementias: Lewy body, NPH, Binswanger, Nasu-Hakola, Pick Disease [27]. Incontinence 3 times more frequent in geriatric patients with dementia than without (42.3/1000 women and 33.5/1000 men vs 19.6/1000 women, 18.6/1000 men) [28].
Parkinsonian syndrome Idiopathic Parkinson's disease (IPD): 75-80% of Parkinsonian syndromes Non-IPD: Parkinson's-plus (18%): Multi system atrophy (MSA), - Progressive supranuclear palsy, - Corticobasal degeneration, - Dementia with Lewy bodies. Secondary Parkinson's (2%)	1.5% in > 65 yrs [29] 2nd neurodegenerative disease after Alzheimer's disease Prevalence: 150/100,000/yr Incidence: 20/100,000/yr MSA is the most frequent non-IPD.	LUTS frequency 30% at onset, 70% after 5 yrs. Storage phase symptoms: Nocturia (60%) OAB - UUI - DO [30]. OAB and DO at the initial phase, intrinsic sphincter deficiency and impaired contractility appear as the disease progress. Complications of neuro-urological symptoms (infections) account for a major cause of mortality in MSA [31].
Brain tumors	26.8/100,000/yr in adult (> 19 yrs) (17.9 benign, 8.9 malignant) [32].	Neuro-urological symptoms vary according to tumour location. Incontinence occurs mainly in frontal location (part of frontal syndrome or isolated in frontal location) [33]. Voiding dysfunction may occur in other location.
Mental retardation and cerebral palsy <i>Intellectual disability in children is a very heterogenous group: including perinatal injury, materno-foetal infections, metabolic disease, genetic disorders and cerebral palsy</i>	Mental retardation other than cerebral palsy Cerebral Palsy: 3.1-3.6/1,000 in children aged 8 yrs [34].	Incontinence: In 65% of severe and profoundly retarded adult patients [35, 36]. DO and impaired contractility also reported. 89% incontinence, 70% uninhibited detrusor contraction at urodynamic examination. Recurrent urinary tract infection and radiologic abnormalities in > 10% of cases.

Lesions and diseases between caudal brainstem and sacral spinal cord		
Spinal cord injury	Non-congenital SCI cases exceed 200,000 in the US with new cases 8/10,000/yr.	Suprasacral lesion leads to DO and DSD (95%). Lower lesions (sacral conus) lead to detrusor hypocontractility (83%) and to complete EUS denervation (60%) [37-39].
Myelomeningocele and nerve tube defects	Spina bifida and congenital nerve tube defects in G8 = 3-4/10,000 live birth/stillbirths with/without pregnancy termination [40]. Lumbar and lumbosacral form are the most common (60%).	Urethrovessical dysfunction in myelomeningocele is very high (90-97%). 50% of these children demonstrate DO. Low compliance is also frequent (alone/associated with can develop with time). Urethral behaviour varies from dyssynergia (50%), normal reflexes (25%) and denervation (25%) [41].
Lesions and diseases of the peripheral nervous system		
Lumbar spine Degenerative disease Disk prolapse	Male (5%) and female (3%) > 35 yrs have had a lumbosacral episode related to disc prolapse.	26% difficulty to void and acontractile detrusor at urodynamic testing. 14% frequent voiding while normal urodynamics testing [42]. Cauda equina lesions lead to detrusor hypocontractility (83%) and complete EUS denervation (60%) [37-39].
Lumbar canal stenosis	Incidence: approx. 5/100,000/yr More common: > 45 yrs, females.	27% significant LUTS (mainly difficulty to void) [42].
Iatrogenic pelvic nerve lesions	Rectum cancer Cervical cancer (multimodal therapy, radiotherapy and surgery) Endometriosis surgery	After APR: 50% urinary retention. After TME: 10-30% voiding dysfunction. Voiding phase dysfunction described. Voiding phase dysfunction if no nerve sparing approach.
Peripheral neuropathy Diabetes Other causes of peripheral neuropathy can cause neuro-urological symptoms: alcohol abuse, lumbosacral zona and genital herpes, Guillain Barré syndrome, porphyria, sarcoidosis.	In Europe, prevalence of pharmacologically treated diabetes ranges from 2.8-3.8%. 50% of patients will develop neuropathy, with 75-100% of these developing neuro-urological symptoms.	"Diabetic Cystopathy"[18, 43]. OAB and DO initially. Hyposensitive and hypocontractile detrusor at later phase.

Disseminated central diseases		
Multiple Sclerosis	Prevalence: 1/1,000 adult in developed country, geographic variation (north > south). First neurological disorder in young adults [44].	80% of patients present neuro-urological symptoms after 10 yrs. 10% of MS patients present voiding dysfunction at disease onset. DO due to suprapontine lesions most frequent dysfunction (> 60%). DSD due to spinal cord lesions in 25%. Hypocontractility in 20%. Dysfunction may change during the course of the disease [45].

APR = abdominoperineal resection; DO = detrusor overactivity; DSD = detrusor sphincter dyssynergia; G8 = 8 most developed countries; IPD = idiopathic Parkinson's disease; LUTS = lower urinary tract symptoms; MSA = multi system atrophy; NPH = normal pressure hydrocephalus; OAB = overactive bladder; SCI = spinal cord injury; TME = total mesorectal excision; UUI = urinary urge incontinence.

3B CLASSIFICATION SYSTEMS

3B.1 Introduction

Several national and international guidelines have already been published for the care of patients with neuro-urological disorders [1, 46-48]. The ICS neuro-urological standardisation report [1] deals specifically with the standardisation of terminology and urodynamic investigation in neuro-urological patients. Other relevant definitions are found in the general ICS standardisation report [49].

Section 3B.2 lists the definitions from these references, partly adapted, and other definitions considered useful for clinical practice (Tables 2 and 3). For specific definitions relating to urodynamic investigation, the reader is referred to the appropriate ICS report [1].

3B.2 Definitions

Table 2: Definitions useful in clinical practice

Acontractility, detrusor	See below under voiding phase (Table 3)
Acontractility, urethral sphincter	See below under storage phase (Table 3)
Autonomic dysreflexia	Increase of sympathetic reflex due to noxious stimuli with symptoms or signs of headache, hypertension, flushing face, and perspiration
Capacity	See below under storage phase
Catheterisation, indwelling	Emptying of the bladder by a catheter that is introduced (semi-) permanently
Catheterisation, intermittent (IC)	Emptying of the bladder by a catheter that is removed after the procedure, mostly at regular intervals
• Aseptic IC	The catheters remain sterile, the genitals are disinfected or washed, and disinfecting lubricant might be used
• Clean IC	Disposable or cleansed re-usable catheters, genitals washed
• Sterile IC	Complete sterile setting, including sterile gloves, forceps, gown and mask
• Intermittent self-catheterisation	IC performed by the patient
Compliance, bladder	See below under storage phase
Condition	Evidence of relevant pathological processes
Diary, bladder	Record of times of micturitions and voided volumes, incontinence episodes, pad usage, and other relevant information

• Frequency volume chart (FVC)	Times of micturitions and voided volumes only
• Micturition time chart	Times of micturitions only
Filling rate, physiological	Below the predicted maximum: body weight (kg) /4 in mL/s [2, 50]
Hesitancy	Difficulty in initiating micturition; delay in the onset of micturition after the individual is ready to pass urine
Intermittency	Urine flow stops and starts on one or more occasions during voiding
Leak point pressure	See below under storage phase
Lower motor neuron lesion (LMNL)	Lesion at or below the S1-S2 spinal cord level
NLUTD	LUTD secondary to confirmed pathology of the nervous supply
Observation, specific	Observation made during specific diagnostic procedure
Overactivity, bladder	See below under symptom syndrome (Table 3)
Overactivity, detrusor	See below under storage phase
Rehabilitation, LUT	Non-surgical non-pharmacological treatment for LUTD
Sign	To verify symptoms and classify them
Sphincter, urethral, non-relaxing	See below under voiding phase
Symptom	Subjective indicator of a disease or change in condition, as perceived by the patient, carer, or partner that may lead the patient to seek help from healthcare professionals
Upper motor neuron lesion (UMNL)	Lesion above the S1-S2 spinal cord level
Voiding, balanced: In patients with neurourological disorders	Voiding with physiological detrusor pressure and low residual (< 80 mL or < 20% of bladder volume)
Voiding, triggered	Voiding initiated by manoeuvres to elicit reflex detrusor contraction by exteroceptive stimuli
Volume, overactivity	See below under storage phase

Table 3: Further definitions useful in clinical practice

Storage phase	
Maximum anaesthetic bladder capacity	Maximum bladder filling volume under deep general or spinal anaesthesia
Increased daytime frequency	Self-explanatory; the normal frequency can be estimated at about 8 times per day [51]
Nocturia	Waking at night one or more times to void
Urgency	The symptom of a sudden compelling desire to pass urine that is difficult to defer
Urinary incontinence	Any involuntary leakage of urine
• Stress urinary incontinence	On effort or exertion, or on sneezing or coughing
• Urgency urinary incontinence	Accompanied by or immediately preceded by urgency
• Mixed urinary incontinence	Associated with urgency but also exertion, effort, sneezing, or coughing
• Continuous urinary incontinence	
Bladder sensation	
<i>Normal</i>	
• Symptom and history	Awareness of bladder filling and increasing sensation up to a strong desire to void
• Urodynamics	First sensation of bladder filling, first desire to void, and strong desire to void at realistic bladder volumes
<i>Increased</i>	
• Symptom and history	An early and persistent desire to void
• Urodynamics	Any of the three urodynamic parameters mentioned under 'normal' persistently at low bladder volume
<i>Reduced</i>	
• Symptom and history	Awareness of bladder filling but no definite desire to void
• Urodynamics	Diminished sensation throughout bladder filling
Absent	
No sensation of bladder filling or desire to void	
Non-specific	Perception of bladder filling as abdominal fullness, vegetative symptoms, or spasticity

<i>Definitions valid after urodynamic confirmation only</i>	
Cystometric capacity	Bladder volume at the end of the filling cystometry
• Maximum cystometric capacity	Bladder volume at strong desire to void
• High-capacity bladder	Bladder volume at cystometric capacity far over the mean voided volume, estimated from the bladder diary, with no significant increase in detrusor pressure under non-anaesthetised condition
Normal detrusor function	Little or no pressure increase during filling; no involuntary phasic contractions despite provocation
Detrusor overactivity	Involuntary detrusor contractions during filling; spontaneous or provoked
• Phasic DO	Characteristic phasic contraction
• Terminal DO	A single contraction at cystometric capacity
• High pressure DO	Maximal detrusor pressure > 40 cm H ₂ O [1, 52]
• Overactivity volume	Bladder volume at first occurrence of DO
• Detrusor overactivity incontinence	Self-explanatory
Leak point pressure	
• Detrusor leak point pressure (DLPP)	Lowest value of detrusor pressure at which leakage is observed in the absence of abdominal strain or detrusor contraction
• Abdominal leak point pressure	Lowest value of intentionally increased intravesical pressure that provokes leakage in the absence of a detrusor contraction
Bladder compliance	Relationship between change in bladder volume (ΔV) and change in detrusor pressure (Δp_{det}): $C = \Delta V / \Delta p_{det}$ (mL/cm H ₂ O)
• Low bladder compliance	Compliance $C = \Delta V / \Delta p_{det} < 20$ mL/cm H ₂ O [53]
Break volume	Bladder volume after which a sudden significant decrease in bladder compliance is observed
Urethral sphincter acontractility	No evidence of sphincter contraction during filling, particularly at higher bladder volumes, or during abdominal pressure increase
Voiding phase	
• Slow stream	Reduced urine flow rate
• Intermittent stream (intermittency)	Stopping and starting of urine flow during micturition
• Hesitancy	Difficulty in initiating micturition
• Straining	Muscular effort to initiate, maintain, or improve urinary stream
• Terminal dribble	Prolonged final part of micturition when the flow has slowed to a trickle/dribble
<i>Definitions valid after urodynamic confirmation only</i>	
Normal detrusor function	Voluntarily initiated detrusor contraction that causes complete bladder emptying within a normal time span
Detrusor underactivity	Contraction of reduced strength/duration
Acontractile detrusor	Absent contraction
Non-relaxing urethral sphincter	Self-explanatory
Detrusor sphincter dyssynergia (DSD)	Detrusor contraction concurrent with an involuntary contraction of the urethra and/or periurethral striated musculature
Post-micturition phase	
Feeling of incomplete emptying (symptom only).	
Post-micturition dribble: involuntary leakage of urine shortly after finishing the micturition.	
Pain, discomfort or pressure sensation in the LUT and genitalia that may be related to bladder filling or voiding, may be felt after micturition, or be continuous.	
Symptom syndrome: combination of symptoms	
Overactive bladder syndrome: urgency with or without urgency incontinence, usually with frequency and nocturia.	
Synonyms: urgency syndrome, urgency-frequency syndrome.	

3C DIAGNOSTIC EVALUATION

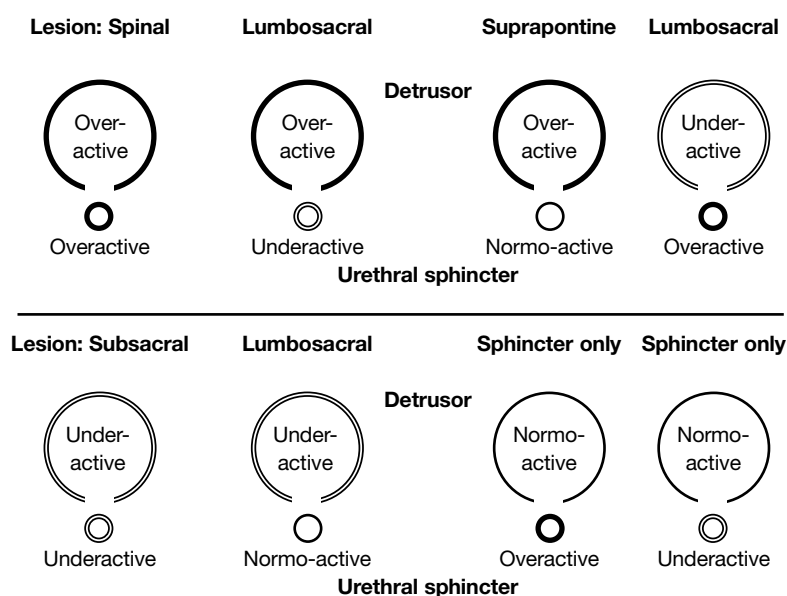
3C.1 Introduction

The normal physiological function of the LUT depends on an intricate interplay between the sensory and motor nervous systems. When diagnosing neuro-urological symptoms, the aim is to describe the type of dysfunction involved. A thorough medical history, physical examination and bladder diary are mandatory before any additional diagnostic investigations can be planned. Results of the initial evaluation are used to decide the patient's long-term treatment and follow-up.

3C.2 Classification systems

Several classification systems for neuro-urological symptoms have been proposed. The Madersbacher [54] (LE: 4) classification describes neuro-urological function in terms of the contraction state of the bladder and external urethral sphincter during filling and voiding phases, which can then be used to decide on the appropriate therapeutic approach [54] (Figure 1).

Figure 1: Madersbacher classification system [54] showing typical neurogenic lesions*



*Adapted from Madersbacher et al.

3C.3 The timing of diagnosis and treatment

Early diagnosis and treatment are essential in both congenital and acquired neuro-urological disorders [55]. This helps to prevent irreversible changes within the LUT, even in the presence of normal reflexes [56, 57] (LE: 3). Furthermore, urological symptoms can be the presenting feature of neurological pathology [58, 59] (LE: 3). Early intervention can prevent irreversible deterioration of the LUT and UUT [60] (LE: 3).

3C.4 Patient history

History taking should include past and present symptoms and disorders (Table 4). It is the cornerstone of evaluation, as the answers will aid in diagnostic investigations and treatment options.

- In non-traumatic neuro-urological patients with a slow insidious onset, history may find that the condition started in childhood or adolescence [61] (LE: 4).
- Urinary history consists of symptoms associated with both urine storage and evacuation.
- Bowel history is important because patients with neuro-urological symptoms may also have a related neuropathic lower gastrointestinal tract [62] (LE: 4).
- Sexual function may be impaired because of the neurological condition.
- Special attention should be paid to possible warning signs and symptoms (e.g. pain, infection, haematuria and fever) requiring further investigation.
- Patients with SCI usually find it difficult to report UTI-related symptoms accurately [1, 63, 64] (LE: 3).

Table 4: History taking in patients with suspected neuro-urological disorders*

Past history
Childhood through to adolescence and in adulthood
Hereditary or familial risk factors
Menarche (age); this may suggest a metabolic disorder
Obstetric history
History of diabetes; in some cases, correction will resolve the neurological problem
Diseases, e.g. syphilis, parkinsonism, multiple sclerosis, encephalitis
Accidents and operations, especially those involving the spine and central nervous system
Present history
Present medication
Lifestyle (smoking, alcohol and drugs); may influence urinary, sexual and bowel function
Quality of life
Specific urinary history
Onset of urological history
Relief after voiding; to detect the extent of a neurological lesion in the absence of obstructive uropathy
Bladder sensation
Initiation of micturition (normal, precipitate, reflex, strain, Credé)
Interruption of micturition (normal, paradoxical, passive)
Enuresis
Mode and type of voiding (catheterisation)
Frequency, volumes voided, incontinence, urge episodes
Bowel history
Frequency and faecal incontinence
Desire to defecate
Defecation pattern
Rectal sensation
Initiation of defecation (digitation)
Sexual history
Genital or sexual dysfunction symptoms
Sensation in genital area
Specific male: erection, (lack of) orgasm, ejaculation
Specific female: dyspareunia, (lack of) orgasm
Neurological history
Acquired or congenital neurological condition
Mental status and comprehension
Neurological symptoms (somatic and sensory), with onset, evolution and any treatment
Spasticity or autonomic dysreflexia (especially in lesions at or above level Th 6)
Mobility and hand function

* Adapted from Bors and Turner [61] (LE: 4; GR: C) and Stöhrer et al. [1] (LE: 4; GR: C).

3C.4.1 Bladder diaries

Bladder diaries provide data on the number of voids, volume voided, pad weight, incontinence and urge episodes. Although a 24-hour bladder diary (recording should be done for three consecutive days) is reliable in women with UI [65, 66] (LE: 3) and helpful in IC [1] (LE: 4), no research has been done on bladder diaries in neuro-urological patients. Nevertheless, bladder diaries are considered a valuable diagnostic tool.

3C.5 Quality of life

An assessment of the patient's present and expected future quality of life (QoL) is important to evaluate the effect of any therapy (or refrained from using) on this parameter. Despite the limitations associated with neurological diseases, adequate treatment with social independence is possible in most patients.

QoL is a very important aspect of the overall management of neuro-urological patients, e.g. to evaluate treatment related changes on a patient's QoL [67] (LE: 2a). The type of bladder management has been shown to affect health-related QoL (HRQoL) in patients with SCI [68]. Other research has also highlighted the importance of urological treatment and its impact on the urodynamic functionality of the neuro-urological patient in determining patient QoL [69].

QoL is related to an individual's ability to cope with a new life situation [70]. QoL can be influenced by several factors, including family support, coping ability, productivity, self-esteem, financial stability, education, and the physical and social environment [71] (LE: 3). Age, sex, ethnicity and the patient's acceptance of the condition also need to be considered when assessing QoL [72] (LE: 3).

Although several questionnaires have been developed to assess QoL, there are no specific QoL questionnaires for the neuro-urological patient. However, a validated specific tool for QoL in SCI and MS patients (Qualiveen®) appears to be a discriminative evaluation instrument [69, 73, 74]. A short-form is available [75] and various validated translations [76-79].

A patient's QoL can be assessed secondarily by generic HRQoL questionnaires, including the Incontinence Quality of Life Instrument (I-QOL), King's Health Questionnaire (KHQ), Short Form 36 Health Survey Questionnaire (SF-36), Euro Quality of Life-5 Domains (EQ-5D), Short Form 6D Health Survey Questionnaire (SF-6D), or the Health Utilities Index (HUI). In addition, the quality-adjusted life year (QALY) quantifies outcomes by weighing years of life spent in a specified health state by a factor representing the value placed by society or patients on the specific health state [80] (LE: 3).

3C.5.1 Recommendations

Recommendations	GR
Quality of life should be assessed when evaluating and treating the neuro-urological patient.	B
The available validated tools are Qualiveen®, a specific long-form and short-form tool for spinal cord lesion and multiple sclerosis patients. In addition, generic (SF-36) or specific tools for incontinence (I-QOL) questionnaires can be used.	B

I-QOL = incontinence quality of life instrument.

3C.6 Physical examination

In addition to a detailed patient history, attention should be paid to possible physical and mental handicaps with respect to the planned investigation. Neurological status should be described as completely as possible (Figure 1). Patients with very high neurological lesions may suffer from a significant drop in blood pressure when moved into a sitting or standing position. All sensations and reflexes in the urogenital area must be tested. Furthermore, detailed testing of the anal sphincter and pelvic floor functions must be performed (Figure 2). It is essential to have this clinical information to reliably interpret later diagnostic investigations.

3C.6.1 Autonomic dysreflexia

Autonomic dysreflexia (AD) is a sudden and exaggerated autonomic response to various stimuli in patients with SCI or spinal dysfunction. It can present in any type of suprasacral lesion but generally manifests above level Th 5-Th 6. The stimulus can be distended bladder or bowel. It can also be secondary to a noxious stimulus, e.g. infected toe nail or pressure sore. Hypertension is a relatively common manifestation of AD and can have life-threatening results if not properly managed [81-83] (LE: 3; GR: C).

Figure 2: The neurological status of a patient with neuro-urological symptoms must be described as completely as possible: (a) dermatomes of spinal cord levels L2-S4; (b) urogenital and other reflexes in the lower spinal cord.

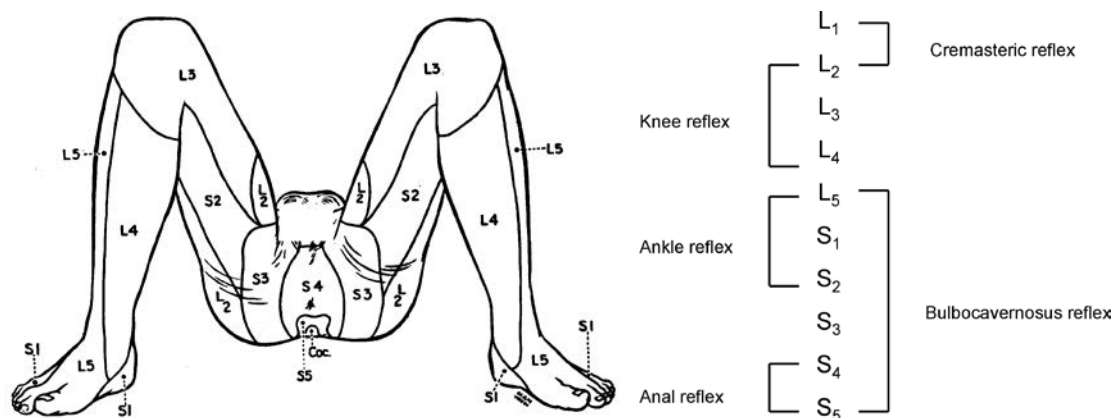


Table 5: Neurological items to be specified*

Sensations S2-S5 (both sides)
Presence (increased/normal/reduced/absent)
Type (light touch/pin prick)
Affected dermatomes
Reflexes (increased/normal/reduced/absent)
Bulbocavernosus reflex
Perianal/anal reflex
Knee and ankle reflexes
Plantar responses (Babinski)
Anal sphincter tone
Presence (increased/normal/reduced/absent)
Voluntary contractions of anal sphincter and pelvic muscles (increased/normal/reduced/absent)
Prostate palpation
Descensus (prolapse) of pelvic organs

*Adapted from Stöhrer et al. [1] (LE: 4; GR: C).

3C.6.2 Recommendations for history taking and physical examination*

History taking	GR
An extensive general history is mandatory, concentrating on past and present symptoms including urinary, sexual, bowel, and neurological functions.	A
Special attention should be paid to the possible existence of alarm signs, e.g. pain, infection, haematuria, fever, that warrant further specific diagnosis.	A
A specific history should be taken for each of the four mentioned functions.	A
Physical examination	
Individual patient handicaps should be acknowledged in planning further investigations.	A
The neurological status should be described as completely as possible. Sensations and reflexes in the urogenital area must all be tested.	A
The anal sphincter and pelvic floor functions must be tested.	A
Urinalysis, blood chemistry, bladder diary, residual and free flowmetry, incontinence quantification and urinary tract imaging should be performed.	A

* All grade A recommendations are based on panel consensus.

3C.7 Urodynamics

3C.7.1 Introduction

Urodynamic investigation is the only method that can objectively assess the (dys-) function of the LUT. In these patients, the invasive urodynamic investigation is even more provocative than in general patients. Any technical source of artifacts must be critically considered. It is essential to maintain the quality of the urodynamic

recording and its interpretation [2]. Same session repeat urodynamic investigations can be helpful in clinical decision making, since repeat measurements may yield completely different results [84].

In patients at risk for AD, it is advisable to measure blood pressure during the urodynamic study. The rectal ampulla should be empty of stool before the start of the investigation. All urodynamic findings must be reported in detail and performed, according to ICS technical recommendations and standards [1, 2, 85].

3C.7.2 **Urodynamic tests**

Free uroflowmetry and assessment of residual urine: This provides a first impression of the voiding function and is compulsory prior to planning any invasive urodynamics. For reliable information, it should be repeated at least 2-3 times [1, 2]. Possible pathological findings include a low flow rate, low voided volume, intermittent flow, hesitancy and residual urine. Care must be taken when assessing the results in patients unable to void in a normal position, as both flow pattern and rate may be modified by inappropriate positions.

Filling cystometry: This is the only method for quantifying the filling function (undertaken at a very slow rate ~20 mL/min). The status of LUT function must be documented during the filling phase. However, this technique has limited use as a solitary procedure. It is much more effective combined with bladder pressure measurement during micturition and even more effective in video-urodynamics.

The bladder should be empty at the start of filling. A physiological filling rate should be used with body-warm saline, as fast filling and room-temperature saline are provocative. Possible pathological findings include DO, low bladder compliance, abnormal bladder sensations, incontinence, and an incompetent or relaxing urethra.

Detrusor leak point pressure (DLPP) [52]: This appears to have no use as a diagnostic tool. Some positive findings have been reported [86, 87], but sensitivity is too low to estimate the risk to the UUT or for secondary bladder damage [88].

Pressure flow study: This reflects the co-ordination between detrusor and urethra or pelvic floor during the voiding phase. It is even more powerful if combined with filling cystometry and with video-urodynamics. LUT function must be recorded during the voiding phase. Possible pathological findings include detrusor hypocontractility, DSD, a high urethral resistance, and residual urine.

Most types of obstruction caused by neuro-urological disorders are due to DSD [89, 90], non-relaxing urethra, or non-relaxing bladder neck [1, 91, 92]. Pressure-flow analysis mostly assesses the amount of mechanical obstruction caused by the urethra's inherent mechanical and anatomical properties and has limited value in patients with neuro-urological disorders.

Electromyography (EMG): This reflects the activity of the external urethral sphincter, the peri-urethral striated musculature, the anal sphincter, and the striated pelvic floor muscles. Correct interpretation may be difficult due to artefacts introduced by other equipment. In the urodynamic setting, an EMG is useful as a gross indication of the patient's ability to control the pelvic floor. Possible pathological findings include inadequate recruitment upon specific stimuli (e.g. bladder filling, hyper-reflexive contractions, onset of voiding, coughing, Valsalva manoeuvre) suggesting a diagnosis of DSD.

Urethral pressure measurement: This has a very limited role in neuro-urological disorders. There is no consensus on parameters indicating pathological findings [93].

Video-urodynamics: This is the combination of filling cystometry and pressure flow study with imaging. It is the gold standard for urodynamic investigation in neuro-urological disorders [1]. Possible pathological findings include all those described in the cystometry and the pressure flow study sections, and any morphological pathology of the LUT and UUT.

Ambulatory urodynamics: This is the functional investigation of the urinary tract, which uses the predominantly natural filling of the urinary tract to reproduce the patient's normal activity [94]. Although this type of study might be considered when conventional urodynamics do not reproduce the patient's symptoms, the role in the neuro-urological patient needs to be determined.

Provocative tests during urodynamics: LUT function can be provoked by coughing, triggered voiding, or anal stretch. Fast-filling cystometry with cooled saline (the 'ice water test') will discriminate between upper and lower motor neuron lesions (UMNL/LMNL) [95, 96]. Patients with UMNL develop a detrusor contraction if the detrusor muscle is intact, while patients with LMNL do not. However, the test gives false-positive results in young children [97] and does not seem to fully discriminative in other types of patient [98].

Previously, a positive bethanechol test [99] (detrusor contraction > 25 cm H₂O) was thought to indicate detrusor denervation hypersensitivity and the muscular integrity of an acontractile detrusor. However, in practice, the test has given equivocal results. A variation of this method was reported using intravesical electromotive administration of the bethanechol [100], but there was no published follow-up.

3C.7.3 **Specialist uro-neurophysiological tests**

The following tests are advised as part of the neurological work-up:

- Electromyography (in a neurophysiological setting) of pelvic floor muscles, urethral sphincter and/or anal sphincter;
- Nerve conduction studies of pudendal nerve;
- Reflex latency measurements of bulbocavernosus and anal reflex arcs;
- Evoked responses from clitoris or glans penis;
- Sensory testing on bladder and urethra.

Other elective tests for specific conditions may become obvious during the work-up and urodynamic investigations.

3C.7.4 **Recommendations for urodynamics and uro-neurophysiology**

Recommendations	GR
The recording of a bladder diary is advisable.	A
Non-invasive testing is mandatory before invasive urodynamics is planned.	A
Urodynamic investigation is necessary to detect and specify lower urinary tract (dys-) function and help with formulating a management plan.	A
Same session repeat measurement can be helpful in clinical decision making.	C
Video-urodynamics is the gold standard for invasive urodynamics in neuro-urological patients. If this is not available, then a filling cystometry continuing into a pressure flow study should be performed.	A
A physiological filling rate and body-warm saline should be used.	A
Specific uro-neurophysiological tests are elective procedures.	C

3C.7.5 **Typical manifestations of neuro-urological disorders**

Table 6 lists typical signs indicating further neurological evaluation, as neuro-urological symptoms may be the presenting symptom [59].

Table 6: Typical findings in neuro-urological disorders

Filling phase
Hyposensitivity or hypersensitivity
Vegetative sensations
Low compliance
High-capacity bladder
Detrusor overactivity, spontaneous or provoked
Sphincter underactivity
Voiding phase
Detrusor underactivity or acontractility
Detrusor sphincter dyssynergia
Non-relaxing urethra
Non-relaxing bladder neck

3C.8 **Renal function**

In many patients with neuro-urological disorders, the UUT is at risk, particularly in patients who develop high detrusor pressure during the filling phase. Although effective treatment can reduce this risk, there is still a relatively high incidence of renal morbidity [56].

Caregivers must be informed of this condition and instructed to watch carefully for any signs or symptoms of a possible deterioration in the patient's renal function. If necessary, the renal function should be checked regularly.

3D DISEASE MANAGEMENT

3D.1 Introduction

The primary aims for treatment of neuro-urological symptoms and their priorities are [101, 102]:

- protection of the UUT;
- achievement of urinary continence;
- restoration of (parts of) the LUT function;
- improvement of the patient's QoL.

Further considerations are the patient's disability, cost-effectiveness, technical complexity, and possible complications [102].

Renal failure is the main mortality factor in SCI patients who survive the trauma [9, 103, 104].

Keeping the detrusor pressure during both the filling and voiding phases within safe limits significantly reduces the mortality from urological causes in these patients [105, 106] and has consequently become the golden rule in the treatment of patients with neuro-urological symptoms [101, 102].

In patients with high detrusor pressure during the filling phase (DO, low bladder compliance), treatment is aimed primarily at "conversion of an active, aggressive high-pressure bladder into a passive low-pressure reservoir" despite the resulting residual urine [101]. Reduction of the detrusor pressure contributes to urinary continence, and consequently to social rehabilitation and QoL. It is also pivotal in preventing UTI [9, 104]. Complete continence can however not always be obtained.

3D.2 Non-invasive conservative treatment

3D.2.1 Assisted bladder emptying - Credé manoeuvre, Valsalva manoeuvre, triggered reflex voiding

Incomplete bladder emptying is a serious risk factor for UTI, high intravesical pressure during the filling phase, and incontinence. Methods to improve the voiding process are therefore practiced.

Bladder expression (Credé manoeuvre) and voiding by abdominal straining (Valsalva manoeuvre): The downwards movement of the lower abdomen by suprapubic compression (Credé) or by abdominal straining (Valsalva) leads to an increase in intravesical pressure, and generally also causes a reflex sphincter contraction [107, 108]. The latter may increase bladder outlet resistance and lead to inefficient emptying. The high pressures created during these procedures are hazardous for the urinary tract [109, 110]. Their use should therefore be discouraged unless urodynamics show that the intravesical pressure remains within safe limits [107, 110-113].

Long-term complications are unavoidable for both methods of bladder emptying [108]. The already weak pelvic floor function may be further impaired, thus introducing or exacerbating already existing stress urinary incontinence (SUI) [110].

Triggered reflex voiding: Stimulation of the sacral or lumbar dermatomes in patients with UMNL can elicit a reflex detrusor contraction [110]. The risk of high pressure voiding is present and interventions to decrease outlet resistance may be necessary [114]. Triggering can induce AD in patients with high level SCI (above Th 6) [115]. All assisted bladder emptying techniques require low outlet resistance. Even then, high detrusor pressures may still be present. Patients hence need dedicated education and close urodynamic and urological surveillance [110, 111, 113, 116].

Note: In the literature, including some of the references cited here, the concept "reflex voiding" is sometimes used to cover all three assisted voiding techniques described in this section.

External appliances: Social continence may be achieved by collecting urine during incontinence, for instance using pads [101, 117]. Condom catheters with urine collection devices are a practical method for men [117]. The infection risk must be closely observed [117]. The penile clamp is absolutely contraindicated in case of DO or low bladder compliance because of the risk of developing high intravesical pressure, and in case of significant reflux.

3D.2.2 Lower urinary tract rehabilitation

3D.2.2.1 Bladder rehabilitation including electrical stimulation

The term bladder rehabilitation summarises treatment options that aim to re-establish bladder function in patients with neuro-urological symptoms. Strong contraction of the urethral sphincter and/or pelvic floor, as well as anal dilatation, manipulation of the genital region, and physical activity inhibit micturition in a reflex manner [117, 118]. The first mechanism is affected by activation of efferent nerve fibres, and the latter ones are produced by activation of afferent fibres [88]. Electrical stimulation of the pudendal nerve afferents strongly inhibits the micturition reflex and detrusor contraction [119]. This stimulation might then support the restoration

of the balance between excitatory and inhibitory inputs at the spinal or supraspinal level [117, 120, 121]. Evidence for bladder rehabilitation using electrical stimulation in neurological patients is mainly based on pilot studies with small patient numbers.

Peripheral temporary electrostimulation: Percutaneous tibial nerve stimulation and external (e.g. penile/clitoral or intracavitary) temporary electrical stimulation suppress neurogenic DO during acute stimulation [122, 123]. Both techniques have also demonstrated sustained effects in patients with MS [124-126]. LUT function remained improved 2 years after transcutaneous electrical stimulation of the bladder in patients with SCI [127]. Electrostimulation also improved continence in children with MMC [128].

In MS patients, combining active neuromuscular electrical stimulation with pelvic floor muscle training and EMG biofeedback can achieve a substantial reduction of neuro-urological symptoms [129]. Furthermore, this treatment combination is significantly superior to electrostimulation alone. Biofeedback can be used for supporting the alleviation of neuro-urological symptoms [130].

Intravesical electrostimulation: Intravesical electrostimulation can increase bladder capacity and improve bladder compliance and bladder filling sensation in patients with incomplete SCI or MMC [131]. In patients with neurogenic detrusor underactivity, intravesical electrostimulation may also improve voiding and reduce residual volume [132, 133].

Chronic peripheral pudendal stimulation: A pilot study in patients with incomplete SCI showed that chronic peripheral pudendal stimulation (defined as 15 min, twice daily, during two weeks) may produce neuromodulatory effects in the brain. These effects are correlated with clinical improvement [134]. Semiconditional electrical stimulation of the dorsal penile nerve during 14-28 days improved bladder storage function in patients with SCI [135].

Repetitive transcranial magnetic stimulation: Although improvement of neuro-urological symptoms has been described in PD and MS patients, this technique is still under investigation [136, 137].

Summary: To date, bladder rehabilitation techniques are mainly based on electrical or magnetic stimulation. However, there is a lack of well-designed studies.

3D.2.3 **Drug treatment**

A single, optimal, medical therapy for neuro-urological symptoms is not yet available. Commonly, a combination of different therapies (e.g. intermittent catheterisation and antimuscarinic drugs) is advised to prevent urinary tract damage and improve long-term outcomes, particularly in patients with SCI with a suprasacral lesion or MS [110, 138-142].

3D.2.3.1 *Drugs for treatment of storage neuro-urological symptoms*

Antimuscarinic drugs: They are the first-line choice for treating neurogenic detrusor overactivity (NDO), increasing bladder capacity, reducing episodes of urinary incontinence secondary to NDO by the inhibition of parasympathetic pathways [5, 143-149].

Although antimuscarinic drugs have been used for many years to treat patients with NDO, the evidence is still limited [145, 146, 150], and the responses of individual patients to antimuscarinic treatment are variable. Only a recent meta-analysis has confirmed the clinical and urodynamic efficacy of antimuscarinic therapy compared to placebo in adult NDO [146]. In children, only oxybutynin is approved, despite prospective trials supporting the efficacy and tolerability of tolterodine, propiverine and solifenacin [151-153]. A prospective randomised study using fesoterodine in children with NDO is ongoing [154].

Higher doses or a combination of antimuscarinic agents may be an option to maximise outcomes in neurological patients [138, 140, 155-158] (LE: 3). However, these drugs have a high incidence of adverse events, which may lead to early discontinuation of therapy [146, 155, 157]. Dry mouth is the most frequent side effect.

Choice of antimuscarinic agent: Oxybutynin [5, 138, 140, 144-146, 149, 155, 156, 159-161], trospium [146, 157, 162], tolterodine [151, 163, 164] and propiverine [5, 146, 160, 165-168] are established, effective and well tolerated treatments even in long-term use (LE: 1a).

Darifenacin and solifenacin have been evaluated recently in NDO secondary to SCI and MS [146, 169-172] with results similar to other antimuscarinic drugs. A study using solifenacin in NDO due to Parkinson's disease is currently suspended [173]. The relatively new fesoterodine, an active metabolite of tolterodine, has also been introduced, even though to date there has been no published clinical evidence of its use in the treatment of neuro-urological disorders.

Side effects: Controlled release antimuscarinics have some minor side effects, e.g. dry mouth. It has been suggested that different ways of administration may help to reduce side effects. In a selected group of patients, transdermal oxybutynin was found to be well tolerated and effective [174-176]. Instead, although there are several studies reporting the efficacy and safety of intravesical oxybutynin, there are no standard protocols yet for its use [177-179]. Therefore, further research is needed into the use of alternative methods of administration, particularly long-term results (LE: 1b).

Other agents

Phosphodiesterase inhibitors (PDE5Is): In vivo and pilot studies seem to support that PDE5Is may become an alternative or adjunct to antimuscarinic treatment for NDO [180-182].

Beta₃-adrenergic receptor agonist: They have recently been introduced and evaluated in OAB, but clinical experience in neuro-urological patients is limited. Studies on safety and effectiveness in NDO are ongoing. In the future, combined therapy with antimuscarinics may be an attractive option [183-185].

3D.2.3.2 Drugs for voiding neuro-urological symptoms

Detrusor underactivity: Cholinergic drugs, such as bethanechol and distigmine, have been considered to enhance detrusor contractility and promote bladder emptying, but are not routinely used in clinical practice [186]. Only preclinical studies have documented the potential benefits of cannabinoid agonists on improving detrusor contractility administered intravesically [187, 188]. Conversely, a randomised controlled study on the use of oromucosal nabixinols (an endocannabinoid modulator), did not report any significant reduction of incontinence episodes in MS patients, although a statistically significant improvement in frequency, urgency and nocturia was documented [189].

Decreasing bladder outlet resistance: α -blockers (e.g. tamsulosin and naftopidil) seem to be effective for decreasing bladder outlet resistance, postvoid residual and autonomic dysreflexia [49]. Combination therapy with a cholinergic drug and an α -blocker appears to be more useful than monotherapy with either agent [190, 191].

Increasing bladder outlet resistance: Several drugs have shown efficacy in selected cases of mild stress urinary incontinence, but there are no high level evidence studies in neurological patients [149].

3D.2.4 Recommendations for drug treatments

Recommendations	LE	GR
For NDO, antimuscarinic therapy is the recommended first-line medical treatment.	1a	A
Alternative routes of administration (i.e., transdermal or intravesical) of antimuscarinic agents may be used.	2	A
Outcomes for NDO may be maximised by considering a combination of antimuscarinic agents.	3	B
To decrease bladder outlet resistance, alpha-blockers could be prescribed.	1b	A
For underactive detrusor, no parasympathomimetics should be prescribed.	1a	A
In neurogenic stress urinary incontinence, drug treatment should not be prescribed.	4	A

NDO = neurogenic detrusor overactivity.

3D.2.5 Minimal invasive treatment

3D.2.5.1 Catheterisation

Intermittent self- or third-party catheterisation [192, 193] is the preferred management for neuro-urological patients who cannot effectively empty their bladders [101, 117].

Sterile IC, as originally proposed by Guttmann and Frankel [192], significantly reduces the risk of UTI and/or bacteriuria [117, 159, 194, 195] compared with clean IC introduced by Lapidis et al. [193]. However, it cannot be considered a routine procedure [117, 195].

Aseptic IC is an alternative [101, 196] that provides a significant benefit by reducing external contamination of the catheter [197-199]. Contributing factors to contamination are insufficient patient education and the inherently greater risk of UTI in neuro-urological patients [117, 198, 200-202]. The average frequency of catheterisations per day is 4-6 times [203] and the catheter size most often used are between 12-16 Fr. In aseptic IC, an optimum frequency of 5 times showed a reduction of UTI [203]. Ideally, bladder volume at catheterisation should, as a rule, not exceed 400-500 mL.

Indwelling transurethral catheterisation and, to a lesser extent, suprapubic cystostomy are associated with a range of complications as well as an enhanced risk for UTI [112, 117, 204-211]. Both

procedures should therefore be avoided when possible.

Silicone catheters are preferred because they are less susceptible to encrustation and because of the high incidence of latex allergy in the neuro-urological patient population [212].

Recommendations for catheterisation

Recommendations	LE	GR
Intermittent catheterisation - whenever possible aseptic technique - should be used as a standard treatment for patients who are unable to empty their bladder.	3	A
Patients must be well instructed in the technique and risks of IC.	3	A
The catheter size should be 12-16 Fr.	4	B
Whenever possible, indwelling transurethral and suprapubic catheterisation should be avoided.	3	A

IC = intermittent catheterisation.

3D.2.5.2 Intravesical drug treatment

To reduce DO, anticholinergics can also be applied intravesically [213-216]. This approach may reduce adverse effects because the anticholinergic drug is metabolised differently [214] and a greater amount is sequestered in the bladder, even more than with electromotive administration [213].

The vanilloids, capsaicin and resiniferatoxin, desensitise the C-fibres and thereby decrease DO for a period of a few months until the sensation of these fibres has been restored [217-219]. The dosage is 1-2 mMol capsaicin in 100 mL 30% alcohol, or 10-100 nMol resiniferatoxin in 100 mL 10% alcohol for 30 minutes. Resiniferatoxin has about a 1,000-fold potency compared to capsaicin, with less pain during the instillation, and is effective in patients refractory to capsaicin. Clinical studies have shown that resiniferatoxin has limited clinical efficacy compared to botulinum toxin A (BTX-A) injections in the detrusor [218].

3D.2.5.3 Intravesical electrostimulation

Intravesical electrostimulation [220] enhances the sensation for bladder filling and urge to void and may restore the volitional control of the detrusor [221, 222]. Daily stimulation sessions of 90 minutes with 10 mA pulses of 2 ms duration at a frequency of 20 Hz [132, 222] are used for at least 1 week [132]. It appears that patients with peripheral lesions are the best candidates, that the muscle must be intact, and that at least some afferent connection between the detrusor and the brain must still be present [132, 222]. Also, the positioning of the stimulating electrodes and bladder filling are important parameters [223]. With these precautions, the results in the literature are still not unequivocal: both positive [131, 132, 221, 224] and negative [225, 226] (LE: 3) results have been reported.

3D.2.5.4 Botulinum toxin injections in the bladder

BTX-A causes a long-lasting but reversible chemical denervation that lasts for about 9 months [12, 227]. The toxin injections are mapped over the detrusor in a dosage that depends on the preparation used. BTX-A has been proven effective in patients with neuro-urological disorders in phase III RCTs [228-230]. Repeated injections seem to be possible without loss of efficacy [12, 230, 231]. Generalised muscular weakness is an occasional adverse effect [12, 229, 231]. Histological studies have not found ultrastructural changes after injection [232].

3D.2.5.5 Bladder neck and urethral procedures

Reduction of the bladder outlet resistance may be necessary to protect the UUT. This can be achieved by chemical denervation of the sphincter or by surgical interventions (bladder neck or sphincter incision or urethral stent). Incontinence may result and can be managed by external devices (see Section 3D.2.1).

BTX-A: This can be used to treat detrusor sphincter dyssynergia effectively by injection at a dose that depends on the preparation used. The dyssynergia is abolished for a few months, necessitating repeat injections. The efficacy of this treatment has been reported to be high and with few adverse effects [233-235]. However, a recent Cochrane report concluded that because of limited evidence future RCTs assessing the effectiveness of BTX injections also need to address the uncertainty about the optimal dose and mode of injection [236]. In addition, this therapy is not registered.

Balloon dilatation: Favourable immediate results were reported [237], but there are no further reports since 1994 so this method is no longer recommended.

Sphincterotomy: By staged incision, bladder outlet resistance can be reduced without completely losing the

closure function of the urethra [101, 117, 228]. Different techniques are used, and laser treatment appears to be advantageous [238, 239]. Sphincterotomy needs to be repeated at regular intervals in many patients [240], but it is efficient and does not cause severe adverse effects [101, 237]. Secondary narrowing of the bladder neck may occur, for which combined bladder neck incision might be considered [241].

Bladder neck incision: This is indicated only for secondary changes at the bladder neck (fibrosis) [101, 238]. This procedure is not recommended in patients with detrusor hypertrophy, which causes thickening of the bladder neck [101].

Stents: Implantation of urethral stents results in continence being dependent on adequate closure of the bladder neck [102]. The results are comparable with sphincterotomy and the stenting procedure has a shorter duration of surgery and hospital stay [242, 243]. However, the costs [101], possible complications and re-interventions [244, 245] are limiting factors in its use [246-249].

Increasing bladder outlet resistance: This can improve the continence condition. Despite early positive results with urethral bulking agents, a relative early loss of continence is reported in patients with neuro-urological disorders [102, 250, 251].

Urethral inserts: Urethral plugs or valves for management of (female) stress incontinence have not been applied in neuro-urological patients. The experience with active pumping urethral prosthesis for treatment of the underactive or acontractile detrusor was disappointing [252].

3D.2.5.6 Recommendations for minimal invasive treatment*

Recommendations	GR
Botulinum toxin injection in the detrusor is the most effective minimally invasive treatment to reduce neurogenic detrusor overactivity.	A
Bladder neck incision is effective in a fibrotic bladder neck.	B

*Recommendations for catheterisation are listed separately under Section 3D.2.5.1

3D.2.6 Surgical treatment

3D.2.6.1 Urethral and bladder neck procedures

Increasing the bladder outlet resistance has the inherent risk of causing high intravesical pressure during filling, which may become even higher during the voiding phase. Procedures to treat sphincteric incontinence are suitable only when the detrusor activity is, or can be, controlled, when no significant reflux is present. Moreover, these procedures require the urethra and bladder neck to be in good condition and mostly result in IC being performed after the procedure [102].

Urethral sling: Various materials have been used for this procedure with enduring positive results. The procedure is established in women with the ability to self-catheterise [102, 253-258]. In men there are a growing number of reports suggesting that both autologous and synthetic slings may also be an alternative [259-261].

Artificial urinary sphincter: This device has stood the test of time in patients with neuro-urological disorders [102]. It was introduced by Light and Scott [262] for this patient group, and the need for revisions [263] has decreased significantly with new generations of devices allowing one to obtain an acceptable long-term outcome [264-270].

Functional sphincter augmentation: By transposing the gracilis muscle to the bladder neck [271] or proximal urethra [272], there is a possibility of creating a functional autologous sphincter by electrical stimulation [271-273]. This opens the possibility of restoring control over the urethral closure.

Bladder neck and urethra reconstruction: The classical Young-Dees-Leadbetter [274] procedure for bladder neck reconstruction in children with bladder exstrophy, and Kropp urethra lengthening [275] improved by Salle [276], are established methods to restore continence provided that IC is practiced and/or bladder augmentation is performed [277].

Urethral inserts: See section 3D.2.5.5.

3D.2.6.2 Denervation, deafferentation, sacral neuromodulation

Sacral rhizotomy, also known as sacral deafferentation, has achieved some success in reducing detrusor overactivity [278-280], but nowadays, it is used mostly as an adjuvant to sacral anterior root stimulation (SARS) [281-285]. Alternatives to rhizotomy are sought in this treatment combination [286-288].

SARS is aimed at producing detrusor contraction. The technique was developed by Brindley [289] and is only applicable to complete lesions above the implant location, because its stimulation amplitude is over the pain threshold. The urethral sphincter efferents are also stimulated, but because the striated muscle relaxes faster than the smooth muscle of the detrusor, so-called "post-stimulus voiding" occurs. This approach has been successful in highly selected patients [282, 290, 291]. By changing the stimulation parameters, this method can also induce defecation or erection.

Sacral neuromodulation (SNM) [292] might be effective and safe for treating neuro-urological symptoms but there is a lack of RCTs and it is unclear which neurological patient is most suitable [293].

3D.2.6.3 Bladder covering by striated muscle

When the bladder is covered by striated muscle that can be stimulated electrically, or ideally that can be contracted voluntarily, voiding function can be restored to an acontractile bladder. The rectus abdominis [294] and latissimus dorsi [295] have been used successfully in patients with neuro-urological symptoms [296, 297].

3D.2.6.4 Bladder augmentation

The aim of auto-augmentation (detrusor myectomy) is to reduce detrusor overactivity or improve low bladder compliance. The advantages are: low surgical burden, low rate of long-term adverse effects, positive effect on patient QoL, and it does not preclude further interventions [101, 102, 298-304].

Replacing or expanding the bladder by intestine or other passive expandable coverage will reduce bladder compliance and at least reduce the pressure effect of detrusor overactivity [305]. Inherent complications associated with these procedures are: recurrent infection, stone formation, perforation or diverticula, possible malignant changes, and for intestine metabolic abnormality, mucus production and impaired bowel function [102, 306-308]. The procedure should be used with caution in patients with neuro-urological symptoms, but may become necessary if all less-invasive treatment methods have failed.

Bladder augmentation is a valid option to decrease detrusor pressure and increase bladder capacity, whenever more conservative approaches have failed. Several different techniques have been published, with comparable and satisfactory results [300, 309-317]. Bladder substitution to create a low-pressure reservoir is indicated in patients with a severely thick and fibrotic bladder wall [318, 319].

3D.2.6.5 Urinary diversion

When no other therapy is successful, urinary diversion must be considered for the protection of the UUT and for the patient's QoL [102, 320].

Continent diversion: This should be the first choice for urinary diversion. Patients with limited dexterity may prefer a stoma instead of using the urethra for catheterisation [102]. A continent stoma is created using various techniques. However, all of them have frequent complications, including leakage or stenosis [102, 321]. The short-term continence rates are > 80% and good protection of the UUT is achieved [102, 322-330]. For cosmetic reasons, the umbilicus is often used for the stoma site [326, 329-336].

Incontinent diversion: If catheterisation is impossible, incontinent diversion with a urine-collecting device is indicated. Ultimately, it could be considered in patients who are wheelchair bound or bed-ridden with intractable and untreatable incontinence, in patients with LUT destruction, when the UUT is severely compromised, and in patients who refuse other therapy [102]. An ileal segment is used for the deviation in most cases [102, 337-341].

Undiversion: Long-standing diversions may be successfully undiverted or an incontinent diversion changed to a continent one with the emergence of new and better techniques for control of detrusor pressure and incontinence [102]. The patient must be carefully counselled and must comply meticulously with the instructions [102]. Successful undiversion can then be performed [342].

3D.2.6.6 Recommendations for surgical treatment

Recommendations	LE	GR
In order to treat refractory detrusor overactivity, bladder augmentation is recommended. Detrusor myectomy is an acceptable alternative in highly selected cases.	3	A
In female patients with neurogenic stress urinary incontinence who are able to self-catheterise, placement of an autologous urethral sling should be used.	4	B
In male patients with neurogenic stress urinary incontinence, artificial urinary sphincter should be used.	3	A

3E URINARY TRACT INFECTION IN NEURO-UROLOGICAL PATIENTS

3E.1 Epidemiology, aetiology and pathophysiology

Urinary tract infection (UTI) is the onset of signs and/or symptoms accompanied by laboratory findings of a UTI (bacteriuria, leukocyturia and positive urine culture) [343]. There are no evidence-based cutoff values for the quantification of these findings. The published consensus is that a significant bacteriuria in persons performing IC is present with $> 10^2$ colony-forming units (cfu)/mL, $> 10^4$ cfu/mL in clean-void specimens and any detectable concentration in suprapubic aspirates. Regarding leukocyturia, 10 or more leukocytes in centrifuged urine samples per microscopic field (400x) are regarded as significant [343].

Individuals with neuro-urological symptoms, especially those with SCI, may have other signs and symptoms in addition to or instead of traditional signs and symptoms of a UTI in able-bodied individuals. Other problems, such as autonomic dysreflexia, may develop or worsen due to a UTI [344]. The most common signs and symptoms suspicious of a UTI in those with neuro-urological disorders are fever, new onset or increase in incontinence, including leaking around an indwelling catheter, increased spasticity, malaise, lethargy or sense of unease, cloudy urine with increased urine odour, discomfort or pain over the kidney or bladder, dysuria, or autonomic dysreflexia [344, 345].

3E.2 Diagnostic evaluation

The gold standard for diagnosis is urine culture and urinalysis. A dipstick test may be more useful to exclude than to prove UTI [346, 347]. As bacterial strains and resistance patterns in persons with neuro-urological disorders may differ from those of able-bodied patients, microbiologic testing is mandatory [348].

3E.3 Disease management

Bacteriuria in patients with neuro-urological disorders should not be treated. Treatment of asymptomatic bacteriuria results in significantly more resistant bacterial strains without improving the outcome [349]. UTI in persons with neuro-urological disorders are by definition complicated UTI. Therefore, single-dose treatment is not advised. There is no consensus in the literature about the duration of treatment. It depends on the severity of the UTI and the involvement of kidneys and the prostate. Generally, a 5-7 day course of antibiotic treatment is advised, that can be extended up to 14 days according to the extent of the infection [349]. The choice of the antibiotic therapy should be based on the results of the microbiologic testing. If immediate treatment is mandatory (e.g. fever, septicaemia, intolerable clinical symptoms, extensive autonomic dysreflexia), the choice of treatment should be based on local and individual resistance profiles [350].

3E.3.1 Recurrent UTI

Recurrent UTI in patients with neuro-urological disorders may indicate a suboptimal management of the underlying functional problem, e.g. high bladder pressure during storage and voiding, incomplete voiding or bladder stones. The improvement of bladder function, e.g. by treating detrusor overactivity by BTX-A injection in the detrusor [351], and the removal of bladder stones or other direct supporting factors, especially indwelling catheters, as early as possible, are mandatory [348].

3E.3.2 Prevention

If the improvement of bladder function and removal of foreign bodies/stones is not successful, additional UTI prevention strategies should be utilised. In men performing IC, the use of hydrophilic catheters is associated with a lower rate of UTI; in women this effect is not demonstrated [352]. Bladder irrigation has not been proven effective [353].

Various medical approaches have been tested as UTI prophylaxis in patients with neuro-urological disorders. The benefit of cranberry juice for the prevention of UTI could not be demonstrated in RCTs [354]. Methenamine hippurate is not effective in individuals with neuro-urological symptoms [355]. There is not sufficient evidence to support the use of L-methionine for urine acidification to prevent recurrent UTI [356]. There is only weak evidence that oral immunotherapy reduces bacteriuria in patients with SCI, and no evidence that recurrent UTI are reduced [357]. Low-dose, long-term, antibiotic prophylaxis cannot reduce UTI frequency, but increases bacterial resistance and is therefore not recommended [349].

A newly proposed application scheme of antibiotic substances for antibiotic prophylaxis provided positive results, but the results of this trial need to be confirmed in further studies [358]. Another possible future option, the inoculation of apathogenic *E. coli* strains into the bladder, has provided positive results in initial studies, but because of the paucity of data [359], cannot be recommended as a treatment option.

In summary, based on the criteria of evidence-based medicine, there is currently no preventive measure for recurrent UTI in patients with neuro-urological disorders that can be recommended without limitations. Therefore, individualised concepts should be taken into consideration, including immunostimulation, phytotherapy and complementary medicine [360]. Prophylaxis in patients with neuro-urological disorders is important to pursue, but since there are no data favouring one approach over another, prophylaxis is essentially a trial and error approach.

3E.4 Recommendations for the treatment of UTI

Recommendations	LE	GR
Asymptomatic bacteriuria in patients with neuro-urological disorders should not be treated.	4	A
The use of long-term antibiotics in recurrent UTIs should be avoided.	2a	A
In patients with recurrent UTI, treatment of neuro-urological symptoms should be optimised and foreign bodies (e.g. stones, indwelling catheters) should be removed from the urinary tract.	3	A
In patients with neuro-urological disorders, UTI prophylaxis must be individualised since there is no optimal prophylactic measure available.	4	C

UTI = urinary tract infection.

3F SEXUAL (DYS)FUNCTION AND FERTILITY

These Guidelines specifically focus on sexual dysfunction and infertility in patients with a neurological disease [361]. Non-neurogenic, male sexual dysfunction and infertility are covered in separate EAU Guidelines [362, 363]. Adopting a systematic approach, such as the PLISSIT model (Permission, Limited Information, Specific Suggestions and Intensive Therapy) [364], provides a framework for counselling and treatment involving a stepwise approach to the management of neurogenic sexual dysfunction.

3F.1 Erectile dysfunction

3F.1.1 Phosphodiesterase type 5 inhibitors

Phosphodiesterase type 5 inhibitors (PDE5Is) are recommended as first-line treatment in neurogenic erectile dysfunction (ED) [361]. All currently available PDE5Is appear to be effective and safe, although there are no high-evidence level studies in neuro-urological patients investigating efficacy and side effects across different PDE5Is, dosages and formulations. A recent network meta-analysis on a mixed ED population has suggested that tadalafil is the most effective agent [365]. Most common side effects of PDE5Is are headache, flushing, dyspepsia and nasal congestion, while PDE5Is may induce relevant hypotension in patients with tetraplegia/high-level paraplegia and multiple system atrophy [366, 367].

Several studies, including RCTs, show the efficacy and safety of PDE5Is for treating ED in patients with SCI [366, 368-371], MS [372-374], PD [375-377], diabetes mellitus [377-380], spina bifida [379] and after radical prostatectomy [381].

Most neuro-urological patients require long-term therapy for ED but some have a low compliance rate or stop therapy because of side effects [366, 367]. As a prerequisite for successful PDE5I-therapy, some residual nerve function is required to induce erection.

Since many patients with SCI use on-demand nitrates for the treatment of autonomic dysreflexia, they must be counselled that PDE5Is are contraindicated when using nitrate medication.

3F.1.2 **Mechanical devices**

Mechanical devices (vacuum tumescence devices and penile rings) may be effective but are less popular [382-386].

3F.1.3 **Intracavernous injections and intraurethral application**

Patients not responding to oral drugs may be offered intracavernous injections (alprostadil, papaverine and phentolamine) that have been shown to be effective in a number of neurological conditions, including SCI, MS, and diabetes mellitus [387-392], but their use requires careful dose titration and some precautions. Complications of intracavernous drugs include pain, priapism and corpora cavernosa fibrosis.

Intracavernous vasoactive drug injection is the first therapeutic option in patients taking nitrate medications, for whom there are concerns about drug interactions with PDE5Is, or in patients for whom PDE5Is are ineffective. The impact of intracavernous injections on ejaculation and orgasmic function, their early use for increasing the recovery rate of a spontaneous erection, and their effectiveness and tolerability in the long-term are unclear [366].

Intraurethral alprostadil application is an alternative but less effective route of administration [393].

3F.1.4 **Penile prostheses**

Penile prostheses may be considered for treatment of neurogenic ED when all conservative treatments have failed. Serious complications, including infection and prosthesis perforation, may occur in about 10% of patients, depending on implant type [394-396].

3F.1.5 **Recommendations for erectile dysfunction**

Recommendations	LE	GR
In neurogenic ED, oral PDE5Is are the recommended first-line medical treatment.	1b	A
In neurogenic ED, intracavernous injections of vasoactive drugs (alone or in combination) are the recommended second-line medical treatment.	3	A
In neurogenic ED, mechanical devices such as vacuum devices and rings can be effective and may be offered to patients.	3	B
In neurogenic ED, penile prostheses should be reserved for selected patients.	4	B

ED = erectile dysfunction; PDE5Is = phosphodiesterase type 5 inhibitors.

3F.2 **Male fertility**

Among the major conditions contributing to neurogenic infertility are pelvic and retroperitoneal surgery, diabetes mellitus, spina bifida, MS and SCI [397]. ED is managed as described previously. Retrograde ejaculation may be reversed by sympathomimetic agents contracting the bladder neck, including imipramine, ephedrine, pseudoephedrine, and phenylpropanolamine [397]. The use of a balloon catheter to obstruct the bladder neck may be effective in obtaining antegrade ejaculation [398]. If antegrade ejaculation is not achieved, the harvest of semen from the urine may be considered [397]. Prostatic massage is safe and easy to use for obtaining semen in men with lesions above T10 [399]. In several patients, vibrostimulation or transrectal electroejaculation are needed for sperm retrieval [397, 400-403]. Semen retrieval is more likely with vibrostimulation in men with lesions above T10 [404-406]. In men with SCI, especially at or above T6, AD might occur during sexual activity and ejaculation [407, 408]; patients at risk and fertility clinics must be informed and aware of this potentially life-threatening condition.

Surgical procedures, such as microsurgical epididymal sperm aspiration (MESA) or testicular sperm extraction (TESE), may be used if vibrostimulation and electroejaculation are not successful [409, 410]. Pregnancy rates in patients with SCI are lower than in the general population, but since the introduction of intracytoplasmic sperm injection (ICSI), men with SCI now have a good chance of becoming biological fathers [411-413].

3F.2.1 **Sperm quality and motility**

The following has been reported on sperm quality and motility:

- Vibrostimulation produces samples with better sperm motility than electrostimulation [402, 414].
- Electroejaculation with interrupted current produces better sperm motility than continuous current [415].
- Bladder management with clean IC may improve semen quality compared to indwelling catheterisation, reflex voiding or bladder expression [416].
- Sperm quality in men with SCI is enhanced by processing in able-bodied seminal plasma [417].
- Freezing of sperm is unlikely to improve fertility rates in men with SCI [400].

3F.2.2 Recommendations for male fertility

Recommendations	LE	GR
In men with SCI, vibrostimulation and transrectal electroejaculation are effective methods of sperm retrieval.	3	B
In men with SCI; MESA, TESE or ICSI may be used after failed vibrostimulation and/or transrectal electroejaculation.	3	B
In men with SCI, especially at or above T6, it is essential to counsel patients at risk and fertility clinics about the potentially life-threatening condition of autonomic dysreflexia.	3	A

SCI = spinal cord injury.

3F.3 Female sexuality

The most relevant publications on neurogenic female sexual dysfunction are in women with SCI and MS. After SCI, about 65-80% of women continue to be sexually active, but to a much lesser extent than before the injury, and about 25% report a decreased satisfaction with their sexual life [418-420]. Although sexual dysfunction is very common in women with MS, it is still often overlooked by medical professionals [421, 422].

The greatest physical barrier to sexual activity is urinary incontinence. Problems with positioning and spasticity affect mainly tetraplegic patients. Peer support may help to optimise the sexual adjustment of women with SCI in achieving a more positive self-image, self-esteem and feelings of being attractive to themselves and others [418, 423-425].

The use of specific drugs for sexual dysfunction is indicated to treat inadequate lubrication. Sildenafil may partially reverse subjective sexual arousal difficulties, while manual and vibratory clitoral stimulation may increase genital responsiveness [426, 427]. Although good evidence exists that psychological interventions are effective in the treatment of female hypoactive sexual desire disorder and female orgasmic disorder [428], there is a lack of high-evidence level studies in the neurological population.

Neurophysiological studies have shown that women with the ability to perceive T11-L2 pin-prick sensations may have psychogenic genital vasocongestion. Reflex lubrication and orgasm is more prevalent in women with SCI who have preserved the sacral reflex arc (S2-S5), even when it has not been shown in an individual woman that a specific level and degree of lesion is the cause of a particular sexual dysfunction. In SCI women with a complete lesion of the sacral reflex, arousal and orgasm may be evoked through stimulation of other erogenous zones above the level of lesions [429-431].

Women with SCI reported dissatisfaction with the quality and quantity of sexuality-related rehabilitation services and were less likely to receive sexual information than men [429, 432, 433].

3F.3.1 Recommendation for female sexuality

Recommendation	LE	GR
There is no effective medical therapy for the treatment of neurogenic sexual dysfunction in women.	4	A

3F.4 Female fertility

There are few studies on female fertility in neurological patients. More than a third (38%) of women with epilepsy had infertility and the relevant predictors were exposure to multiple (three or more) antiepileptic drugs, older age and lower education [434].

Although it seems that the reproductive capacity of women with SCI is only temporarily affected by SCI with cessation of menstruation for approximately 6 months after SCI [435], there are no high-evidence level studies. About 70% of sexually active women use some form of contraception after injury, but fewer women use the birth control pill compared to before their injury [436].

Women with SCI are more likely to suffer complications during pregnancy, labour and delivery compared to able-bodied women. Complications of labour and delivery include bladder problems, spasticity, pressure sores, anaemia, and AD [437, 438]. Obstetric outcomes include higher rates of Caesarean sections and an increased incidence of low birth-weight babies [436].

Epidural anaesthesia is chosen and effective for most patients with AD during labour and delivery [439, 440].

There is very little published data on women's experience of the menopause following SCI [441].

3F.4.1 Recommendation for female fertility

Recommendation	LE	GR
In women with a neurological disease, the management of fertility, pregnancy and delivery requires a multidisciplinary approach tailored to individual patient's needs and preferences.	4	A

3G FOLLOW-UP

3G.1 Introduction

Neuro-urological disorders are often unstable and the symptoms may vary considerably, even within a relatively short period. Regular follow-up is therefore necessary [46, 101, 305, 311, 338, 442-453].

Depending on the type of the underlying neurological pathology and the current stability of the neuro-urological symptoms, the interval between initial investigations and control diagnostics may vary and in many cases should not exceed 1-2 years. In high-risk neuro-urological patients this interval should be much shorter. Urinalysis should be performed regularly; the frequency to be guided by patient symptoms. The UUT should be checked by ultrasonography at regular intervals in high-risk patients; at least once every 6 months. In these patients, physical examination and urine laboratory should take place every year. Any significant clinical change warrants further, specialised, investigation.

3G.2 Recommendations for follow-up

Recommendations	LE	GR
In high-risk patients, the upper urinary tract should be assessed at least every six months.	4	A
In high-risk patients, physical examination, and urine laboratory should take place every year.	4	A
Any significant clinical changes should instigate further, specialised, investigation.	4	A
Urodynamic investigation is a mandatory baseline diagnostic and in high-risk patients, should be done at regular intervals.	3	A

3H CONCLUSIONS

Neuro-urological disorders have a multi-faceted pathology. They require an extensive and specific diagnosis before one can embark on an individualised therapy, which takes into account the medical and physical condition of the patient and the patient's expectations about his/her future.

The urologist or paediatric urologist can select from a wealth of therapeutical options, each with its own pros and cons. Notwithstanding the success of any therapy embarked upon, a close surveillance is necessary for the patient's entire life.

These Guidelines offer you expert advice on how to define the patient's neuro-urological symptoms as precisely as possible and how to select, together with the patient, the appropriate therapy. This last choice, as always, is governed by the golden rule: as effective as needed, as less invasive as possible.

4. REFERENCES

1. Stohrer M, et al. The standardization of terminology in neurogenic lower urinary tract dysfunction: with suggestions for diagnostic procedures. International Continence Society Standardization Committee. *Neurourol Urodyn*, 1999. 18(2): p. 139-58.
<http://www.ncbi.nlm.nih.gov/pubmed/10081953>
2. Schafer W, et al. Good urodynamic practices: uroflowmetry, filling cystometry, and pressure-flow studies. *Neurourol Urodyn*, 2002. 21(3): p. 261-74.
<http://www.ncbi.nlm.nih.gov/pubmed/11948720>

3. Abrams P, et al. Reviewing the ICS 2002 terminology report: the ongoing debate. *Neurourol Urodyn*, 2009. 28(4): p. 287.
<http://www.ncbi.nlm.nih.gov/pubmed/19350662>
4. Abrams P, et al. The standardisation of terminology of lower urinary tract function: report from the Standardisation Sub-committee of the International Continence Society. *Neurourol Urodyn*, 2002. 21(2): p. 167-78.
<http://www.ncbi.nlm.nih.gov/pubmed/11857671>
5. Stohrer M, et al. EAU guidelines on neurogenic lower urinary tract dysfunction. *Eur Urol*, 2009. 56(1): p. 81-8.
<http://www.ncbi.nlm.nih.gov/pubmed/19403235>
6. Nosseir M, et al. Clinical usefulness of urodynamic assessment for maintenance of bladder function in patients with spinal cord injury. *Neurourol Urodyn*, 2007. 26(2): p. 228-33.
<http://www.ncbi.nlm.nih.gov/pubmed/16998859>
7. Gerridzen RG, et al. Risk factors for upper tract deterioration in chronic spinal cord injury patients. *J Urol*, 1992. 147(2): p. 416-8.
<http://www.ncbi.nlm.nih.gov/pubmed/1732606>
8. McGuire EJ, et al. Prognostic value of urodynamic testing in myelodysplastic patients. *J Urol*, 1981. 126(2): p. 205-9.
<http://www.ncbi.nlm.nih.gov/pubmed/7196460>
9. Hackler RH. A 25-year prospective mortality study in the spinal cord injured patient: comparison with the long-term living paraplegic. *J Urol*, 1977. 117(4): p. 486-8.
<http://www.ncbi.nlm.nih.gov/pubmed/850323>
10. Bruschini H, et al. Upper and lower urinary tract evaluation of 104 patients with myelomeningocele without adequate urological management. *World J Urol*, 2006. 24(2): p. 224-8.
<http://www.ncbi.nlm.nih.gov/pubmed/16758253>
11. Lidal IB, et al. Mortality after spinal cord injury in Norway. *J Rehabil Med*, 2007. 39(2): p. 145-51.
<http://www.ncbi.nlm.nih.gov/pubmed/17351697>
12. Del Popolo G, et al. Neurogenic detrusor overactivity treated with english botulinum toxin a: 8-year experience of one single centre. *Eur Urol*, 2008. 53(5): p. 1013-19.
<http://www.ncbi.nlm.nih.gov/pubmed/17950989>
13. Marrie RA, et al. Disparities in the management of multiple sclerosis-related bladder symptoms. *Neurology*, 2007. 68(23): p. 1971-8.
<http://www.ncbi.nlm.nih.gov/pubmed/17548546>
14. de Seze M, et al. The neurogenic bladder in multiple sclerosis: review of the literature and proposal of management guidelines. *Mult Scler*, 2007. 13(7): p. 915-28.
<http://www.ncbi.nlm.nih.gov/pubmed/17881401>
15. Ragonese P, et al. Mortality in multiple sclerosis: a review. *Eur J Neurol*, 2008. 15(2): p. 123-7.
<http://www.ncbi.nlm.nih.gov/pubmed/18217882>
16. Sakakibara R, et al. Videourodynamic and sphincter motor unit potential analyses in Parkinson's disease and multiple system atrophy. *J Neurol Neurosurg Psychiatry*, 2001. 71(5): p. 600-6.
<http://www.ncbi.nlm.nih.gov/pubmed/11606669>
17. Paleschi G, et al. Correlation between the Overactive Bladder questionnaire (OAB-q) and urodynamic data of Parkinson disease patients affected by neurogenic detrusor overactivity during antimuscarinic treatment. *Clin Neuropharmacol*, 2006. 29(4): p. 220-9.
<http://www.ncbi.nlm.nih.gov/pubmed/16855424>
18. Fridodt-Moller C. Diabetic cystopathy: epidemiology and related disorders. *Ann Intern Med*, 1980. 92(2 Pt 2): p. 318-21.
<http://www.ncbi.nlm.nih.gov/pubmed/7356221>
19. Brown SH, et al. Trials review: cardiovascular outcome with intensive glycemic control and implications for patients with type 2 diabetes. *Postgrad Med*, 2009. 121(5): p. 31-41.
<http://www.ncbi.nlm.nih.gov/pubmed/19820272>
20. Phillips B, et al. Oxford Centre for Evidence-based Medicine Levels of Evidence. Updated by Jeremy Howick March 2009. 1998.
<http://www.cebm.net/oxford-centre-evidence-based-medicine-levels-evidence-march-2009/>
21. Nichols M, et al. Cardiovascular disease in Europe: epidemiological update. *Eur Heart J*, 2013. 34(39): p. 3028-34.
<http://www.ncbi.nlm.nih.gov/pubmed/24014390>
22. Tibaek S, et al. Prevalence of lower urinary tract symptoms (LUTS) in stroke patients: a cross-sectional, clinical survey. *Neurourol Urodyn*, 2008. 27(8): p. 763-71.
<http://www.ncbi.nlm.nih.gov/pubmed/18551565>

23. Marinkovic SP, et al. Voiding and sexual dysfunction after cerebrovascular accidents. *J Urol*, 2001. 165(2): p. 359-70.
<http://www.ncbi.nlm.nih.gov/pubmed/11176374>
24. Rotar M, et al. Stroke patients who regain urinary continence in the first week after acute first-ever stroke have better prognosis than patients with persistent lower urinary tract dysfunction. *Neurourol Urodyn*, 2011. 30(7): p. 1315-8.
<http://www.ncbi.nlm.nih.gov/pubmed/21488096>
25. Berr C, et al. [Epidemiology of dementia]. *Presse Med*, 2007. 36(10 Pt 2): p. 1431-41. [Article in French]
<http://www.ncbi.nlm.nih.gov/pubmed/17560760>
26. Lobo A, et al. Prevalence of dementia and major subtypes in Europe: A collaborative study of population-based cohorts. *Neurologic Diseases in the Elderly Research Group. Neurology*, 2000. 54(11 Suppl 5): p. S4-9.
<http://www.ncbi.nlm.nih.gov/pubmed/10854354>
27. Na HR, et al. Urinary incontinence in Alzheimer's disease is associated with Clinical Dementia Rating-Sum of Boxes and Barthel Activities of Daily Living. *Asia Pac Psychiatry*, 2012. 7(1): p. 113-20.
<http://www.ncbi.nlm.nih.gov/pubmed/23857871>
28. Grant RL, et al. First diagnosis and management of incontinence in older people with and without dementia in primary care: a cohort study using The Health Improvement Network primary care database. *PLoS Med*, 2013. 10(8): p. e1001505.
<http://www.ncbi.nlm.nih.gov/pubmed/24015113>
29. Winge K, et al. Bladder dysfunction in Parkinsonism: mechanisms, prevalence, symptoms, and management. *Mov Disord*, 2006. 21(6): p. 737-45.
<http://www.ncbi.nlm.nih.gov/pubmed/16570299>
30. Ragab MM, et al. Idiopathic Parkinson's disease patients at the urologic clinic. *Neurourol Urodyn*, 2011. 30(7): p. 1258-61.
<http://www.ncbi.nlm.nih.gov/pubmed/21404318>
31. Papatsoris AG, et al. Urinary and erectile dysfunction in multiple system atrophy (MSA). *Neurourol Urodyn*, 2008. 27(1): p. 22-7.
<http://www.ncbi.nlm.nih.gov/pubmed/17563111>
32. Dolecek TA, et al. CBTRUS statistical report: primary brain and central nervous system tumors diagnosed in the United States in 2005-2009. *Neuro Oncol*, 2012. 14 Suppl 5: p. v1-49.
<http://www.ncbi.nlm.nih.gov/pubmed/23095881>
33. Maurice-Williams RS. Micturition symptoms in frontal tumours. *J Neurol Neurosurg Psychiatry*, 1974. 37(4): p. 431-6.
<http://www.ncbi.nlm.nih.gov/pubmed/4365244>
34. Christensen D, et al. Prevalence of cerebral palsy, co-occurring autism spectrum disorders, and motor functioning - Autism and Developmental Disabilities Monitoring Network, USA, 2008. *Dev Med Child Neurol*, 2014. 56(1): p. 59-65.
<http://www.ncbi.nlm.nih.gov/pubmed/24117446>
35. Reid AH, et al. Behavioural syndromes identified by cluster analysis in a sample of 100 severely and profoundly retarded adults. *Psychol Med*, 1978. 8(3): p. 399-412.
<http://www.ncbi.nlm.nih.gov/pubmed/704707>
36. Hellstrom PA, et al. Bladder function in the mentally retarded. *Br J Urol*, 1990. 66(5): p. 475-8.
<http://www.ncbi.nlm.nih.gov/pubmed/2249114>
37. Kaplan SA, et al. Bladder and sphincter behavior in patients with spinal cord lesions. *J Urol*, 1991. 146(1): p. 113-7.
<http://www.ncbi.nlm.nih.gov/pubmed/2056568>
38. Weld KJ, et al. Association of level of injury and bladder behavior in patients with post-traumatic spinal cord injury. *Urology*, 2000. 55(4): p. 490-4.
<http://www.ncbi.nlm.nih.gov/pubmed/10736489>
39. Hagen EM, et al. Traumatic spinal cord injury and concomitant brain injury: a cohort study. *Acta Neurol Scand Suppl*, 2010(190): p. 51-7.
<http://www.ncbi.nlm.nih.gov/pubmed/20586736>
40. Kondo A, et al. Neural tube defects: prevalence, etiology and prevention. *Int J Urol*, 2009. 16(1): p. 49-57.
<http://www.ncbi.nlm.nih.gov/pubmed/19120526>

41. Spindel MR, et al. The changing neurourologic lesion in myelodysplasia. *JAMA*, 1987. 258(12): p. 1630-3.
<http://www.ncbi.nlm.nih.gov/pubmed/3625970>
42. Bartolin Z, et al. Relationship between clinical data and urodynamic findings in patients with lumbar intervertebral disk protrusion. *Urol Res*, 2002. 30(4): p. 219-22.
<http://www.ncbi.nlm.nih.gov/pubmed/12202938>
43. Daneshgari F, et al. Temporal differences in bladder dysfunction caused by diabetes, diuresis, and treated diabetes in mice. *Am J Physiol Regul Integr Comp Physiol*, 2006. 290(6): p. R1728-35.
<http://www.ncbi.nlm.nih.gov/pubmed/16439670>
44. Nortvedt MW, et al. Prevalence of bladder, bowel and sexual problems among multiple sclerosis patients two to five years after diagnosis. *Mult Scler*, 2007. 13(1): p. 106-12.
<http://www.ncbi.nlm.nih.gov/pubmed/17294618>
45. de Seze M, et al. The neurogenic bladder in multiple sclerosis: review of the literature and proposal of management guidelines. *Mult Scler*, 2007. 13(7): p. 915-28.
<http://www.ncbi.nlm.nih.gov/pubmed/17881401>
46. Burgdörfer H, et al. [Guidelines for the urological management of paraplegic patients]. *Urologe A*, 1998. 37: p. 222-8. [Article in German]
47. Wyndaele JJ, et al., Neurologic urinary and faecal incontinence, in *Incontinence*, P. Abrams, et al., Editors. 2005, Health Publications: Plymouth. p. 1061-2.
http://www.ics.org/publications/ICI_3/v2.pdf/chap17.pdf
48. Consortium for Spinal Cord Medicine. Bladder management for adults with spinal cord injury: a clinical practice guideline for health-care providers. *J Spinal Cord Med*, 2006. 29(5): p. 527-73.
<http://www.ncbi.nlm.nih.gov/pubmed/17274492>
49. Abrams P, et al. Tamsulosin: efficacy and safety in patients with neurogenic lower urinary tract dysfunction due to suprasacral spinal cord injury. *J Urol*, 2003. 170(4 Pt 1): p. 1242-51.
<http://www.ncbi.nlm.nih.gov/pubmed/14501734>
50. Klevmark B. Natural pressure-volume curves and conventional cystometry. *Scand J Urol Nephrol Suppl*, 1999. 201: p. 1-4.
<http://www.ncbi.nlm.nih.gov/pubmed/10573769>
51. Homma Y, et al. Voiding and incontinence frequencies: variability of diary data and required diary length. *Neurourol Urodyn*, 2002. 21(3): p. 204-9.
<http://www.ncbi.nlm.nih.gov/pubmed/11948713>
52. McGuire EJ, et al. Leak-point pressures. *Urol Clin North Am*, 1996. 23(2): p. 253-62.
<http://www.ncbi.nlm.nih.gov/pubmed/8659025>
53. Hollabaugh RS, Jr., et al. Neuroanatomy of the pelvis: implications for colonic and rectal resection. *Dis Colon Rectum*, 2000. 43(10): p. 1390-7.
<http://www.ncbi.nlm.nih.gov/pubmed/11052516>
54. Madersbacher H. The various types of neurogenic bladder dysfunction: an update of current therapeutic concepts. *Paraplegia*, 1990. 28(4): p. 217-29.
<http://www.ncbi.nlm.nih.gov/pubmed/2235029>
55. Del Popolo G, et al. Diagnosis and therapy for neurogenic bladder dysfunctions in multiple sclerosis patients. *Neurol Sci*, 2008. 29 Suppl 4: p. S352-5.
<http://www.ncbi.nlm.nih.gov/pubmed/19089675>
56. Satar N, et al. The effects of delayed diagnosis and treatment in patients with an occult spinal dysraphism. *J Urol*, 1995. 154(2 Pt 2): p. 754-8.
<http://www.ncbi.nlm.nih.gov/pubmed/7609171>
57. Watanabe T, et al. High incidence of occult neurogenic bladder dysfunction in neurologically intact patients with thoracolumbar spinal injuries. *J Urol*, 1998. 159(3): p. 965-8.
<http://www.ncbi.nlm.nih.gov/pubmed/9474194>
58. Ahlberg J, et al. Neurological signs are common in patients with urodynamically verified "idiopathic" bladder overactivity. *Neurourol Urodyn*, 2002. 21(1): p. 65-70.
<http://www.ncbi.nlm.nih.gov/pubmed/11835426>
59. Bemelmans BL, et al. Evidence for early lower urinary tract dysfunction in clinically silent multiple sclerosis. *J Urol*, 1991. 145(6): p. 1219-24.
<http://www.ncbi.nlm.nih.gov/pubmed/2033697>
60. Klausner AP, et al. The neurogenic bladder: an update with management strategies for primary care physicians. *Med Clin North Am*, 2011. 95(1): p. 111-20.
<http://www.ncbi.nlm.nih.gov/pubmed/21095415>

61. Bors E, et al. History and physical examination in neurological urology. *J Urol*, 1960. 83: p. 759-67. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/13802958>
62. Jayawardena V, et al. Significance of bacteriuria in neurogenic bladder. *J Spinal Cord Med*, 2004. 27(2): p. 102-5.
<http://www.ncbi.nlm.nih.gov/pubmed/15162878>
63. Linsenmeyer TA, et al. Accuracy of individuals with spinal cord injury at predicting urinary tract infections based on their symptoms. *J Spinal Cord Med*, 2003. 26(4): p. 352-7.
<http://www.ncbi.nlm.nih.gov/pubmed/14992336>
64. Massa LM, et al. Validity, accuracy, and predictive value of urinary tract infection signs and symptoms in individuals with spinal cord injury on intermittent catheterization. *J Spinal Cord Med*, 2009. 32(5): p. 568-73.
<http://www.ncbi.nlm.nih.gov/pubmed/20025153>
65. Honjo H, et al. Impact of convenience void in a bladder diary with urinary perception grade to assess overactive bladder symptoms: a community-based study. *Neurourol Urodyn*, 2010. 29(7): p. 1286-9.
<http://www.ncbi.nlm.nih.gov/pubmed/20878998>
66. Naoemova I, et al. Reliability of the 24-h sensation-related bladder diary in women with urinary incontinence. *Int Urogynecol J Pelvic Floor Dysfunct*, 2008. 19(7): p. 955-9.
<http://www.ncbi.nlm.nih.gov/pubmed/18235981>
67. Henze T. Managing specific symptoms in people with multiple sclerosis. *Int MS J*, 2005. 12(2): p. 60-8.
<http://www.ncbi.nlm.nih.gov/pubmed/16417816>
68. Liu CW, et al. The relationship between bladder management and health-related quality of life in patients with spinal cord injury in the UK. *Spinal Cord*, 2010. 48(4): p. 319-24.
<http://www.ncbi.nlm.nih.gov/pubmed/19841636>
69. Pannek J, et al. Does optimizing bladder management equal optimizing quality of life? Correlation between health-related quality of life and urodynamic parameters in patients with spinal cord lesions. *Urology*, 2009. 74(2): p. 263-6.
<http://www.ncbi.nlm.nih.gov/pubmed/19428089>
70. Ku JH. The management of neurogenic bladder and quality of life in spinal cord injury. *BJU Int*, 2006. 98(4): p. 739-45. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/16978269>
71. Whiteneck G, et al. Environmental factors and their role in participation and life satisfaction after spinal cord injury. *Arch Phys Med Rehabil*, 2004. 85(11): p. 1793-803.
<http://www.ncbi.nlm.nih.gov/pubmed/15520974>
72. Marschall-Kehrel D, et al. Patient-reported outcomes in overactive bladder: the influence of perception of condition and expectation for treatment benefit. *Urology*, 2006. 68(2 Suppl): p. 29-37.
<http://www.ncbi.nlm.nih.gov/pubmed/16908338>
73. Bonniaud V, et al. Qualiveen, a urinary-disorder specific instrument: 0.5 corresponds to the minimal important difference. *J Clin Epidemiol*, 2008. 61(5): p. 505-10.
<http://www.ncbi.nlm.nih.gov/pubmed/18394545>
74. Pappalardo A, et al. [Management of neuropathic bladder in multiple sclerosis]. *Clin Ter*, 2004. 155(5): p. 183-6. [Article in Italian]
<http://www.ncbi.nlm.nih.gov/pubmed/15344566>
75. Bonniaud V, et al. Development and validation of the short form of a urinary quality of life questionnaire: SF-Qualiveen. *J Urol*, 2008. 180(6): p. 2592-8.
<http://www.ncbi.nlm.nih.gov/pubmed/18950816>
76. Bonniaud V, et al. Italian version of Qualiveen-30: cultural adaptation of a neurogenic urinary disorder-specific instrument. *Neurourol Urodyn*, 2011. 30(3): p. 354-9.
<http://www.ncbi.nlm.nih.gov/pubmed/21305589>
77. Ciudin A, et al. Quality of life of multiple sclerosis patients: translation and validation of the Spanish version of Qualiveen. *Neurourol Urodyn*, 2012. 31(4): p. 517-20.
<http://www.ncbi.nlm.nih.gov/pubmed/22396437>
78. D'Ancona CA, et al. Quality of life of neurogenic patients: translation and validation of the Portuguese version of Qualiveen. *Int Urol Nephrol*, 2009. 41(1): p. 29-33.
<http://www.ncbi.nlm.nih.gov/pubmed/18528780>

79. Pannek J, et al. [Quality of life in German-speaking patients with spinal cord injuries and bladder dysfunctions. Validation of the German version of the Qualiveen questionnaire]. *Urologe A*, 2007. 46(10): p. 1416-21. [Article in German]
<http://www.ncbi.nlm.nih.gov/pubmed/17605119>
80. Hollingworth W, et al. Exploring the impact of changes in neurogenic urinary incontinence frequency and condition-specific quality of life on preference-based outcomes. *Qual Life Res*, 2010. 19(3): p. 323-31.
<http://www.ncbi.nlm.nih.gov/pubmed/20094804>
81. Assadi F, et al. Autonomic dysreflexia manifested by severe hypertension. *Med Sci Monit*, 2004. 10(12): p. 77-9.
<http://www.ncbi.nlm.nih.gov/pubmed/15567988>
82. Braddom RL, et al. Autonomic dysreflexia. A survey of current treatment. *Am J Phys Med Rehabil*, 1991. 70(5): p. 234-41.
<http://www.ncbi.nlm.nih.gov/pubmed/1910647>
83. Silver JR. Early autonomic dysreflexia. *Spinal Cord*, 2000. 38(4): p. 229-33.
<http://www.ncbi.nlm.nih.gov/pubmed/10822393>
84. Bellucci CH, et al. Neurogenic lower urinary tract dysfunction--do we need same session repeat urodynamic investigations? *J Urol*, 2012. 187(4): p. 1318-23.
<http://www.ncbi.nlm.nih.gov/pubmed/22341264>
85. Gammie A, et al. International Continence Society guidelines on urodynamic equipment performance. *Neurourol Urodyn*, 2014. 33(4): p. 370-9.
<http://www.ncbi.nlm.nih.gov/pubmed/24390971>
86. Ozkan B, et al. Which factors predict upper urinary tract deterioration in overactive neurogenic bladder dysfunction? *Urology*, 2005. 66(1): p. 99-104.
<http://www.ncbi.nlm.nih.gov/pubmed/15992868>
87. Wang QW, et al. Is it possible to use urodynamic variables to predict upper urinary tract dilatation in children with neurogenic bladder-sphincter dysfunction? *BJU Int*, 2006. 98(6): p. 1295-300.
<http://www.ncbi.nlm.nih.gov/pubmed/17034510>
88. Linsenmeyer TA, et al. The impact of urodynamic parameters on the upper tracts of spinal cord injured men who void reflexly. *J Spinal Cord Med*, 1998. 21(1): p. 15-20.
<http://www.ncbi.nlm.nih.gov/pubmed/9541882>
89. Krongrad A, et al. Bladder neck dysynergia in spinal cord injury. *Am J Phys Med Rehabil*, 1996. 75(3): p. 204-7.
<http://www.ncbi.nlm.nih.gov/pubmed/8663928>
90. Weld KJ, et al. Clinical significance of detrusor sphincter dyssynergia type in patients with posttraumatic spinal cord injury. *Urology*, 2000. 56(4): p. 565-8.
<http://www.ncbi.nlm.nih.gov/pubmed/11018603>
91. Rossier AB, et al. 5-microtransducer catheter in evaluation of neurogenic bladder function. *Urology*, 1986. 27(4): p. 371-8.
<http://www.ncbi.nlm.nih.gov/pubmed/3962062>
92. Al-Ali M, et al. A 10 year review of the endoscopic treatment of 125 spinal cord injured patients with vesical outlet obstruction: does bladder neck dyssynergia exist? *Paraplegia*, 1996. 34(1): p. 34-38.
<http://www.ncbi.nlm.nih.gov/pubmed/8848321>
93. Lose G, et al. Standardisation of urethral pressure measurement: report from the Standardisation Sub-Committee of the International Continence Society. *Neurourol Urodyn*, 2002. 21(3): p. 258-60.
<http://www.ncbi.nlm.nih.gov/pubmed/11948719>
94. van Waalwijk van Doorn E, et al. Standardisation of ambulatory urodynamic monitoring: Report of the Standardisation Sub-Committee of the International Continence Society for Ambulatory Urodynamic Studies. *Neurourol Urodyn*, 2000. 19(2): p. 113-25.
<http://www.ncbi.nlm.nih.gov/pubmed/10679828>
95. Geirsson G, et al. The ice-water test--a simple and valuable supplement to routine cystometry. *Br J Urol*, 1993. 71(6): p. 681-5.
<http://www.ncbi.nlm.nih.gov/pubmed/8343894>
96. Geirsson G, et al. Pressure, volume and infusion speed criteria for the ice-water test. *Br J Urol*, 1994. 73(5): p. 498-503.
<http://www.ncbi.nlm.nih.gov/pubmed/8012770>
97. Geirsson G, et al. Positive bladder cooling test in neurologically normal young children. *J Urol*, 1994. 151(2): p. 446-8.
<http://www.ncbi.nlm.nih.gov/pubmed/8283555>

98. Chancellor MB, et al. Ice-water test in the urodynamic evaluation of spinal cord injured patients. *Tech Urol*, 1998. 4(2): p. 87-91.
<http://www.ncbi.nlm.nih.gov/pubmed/9623622>
99. Lapedes J. Neurogenic bladder. Principles of treatment. *Urol Clin North Am*, 1974. 1(1): p. 81-97. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/4428540>
100. Riedl CR, et al. Electromotive administration of intravesical bethanechol and the clinical impact on acontractile detrusor management: introduction of a new test. *J Urol*, 2000. 164(6): p. 2108-11.
<http://www.ncbi.nlm.nih.gov/pubmed/11061937>
101. Stöhrer M, et al. Diagnosis and treatment of bladder dysfunction in spinal cord injury patients. *Eur Urol Update Series* 1994. 3: p. 170-5.
102. Castro-Diaz D, et al. Surgery for the neuropathic patient. In *Incontinence*, P. Abrams, et al., Editors. 2002, Health Publication: Plymouth. p. 865-891.
103. Donnelly J, et al. Present urologic status of the World War II paraplegic: 25-year followup. Comparison with status of the 20-year Korean War paraplegic and 5-year Vietnam paraplegic. *J Urol*, 1972. 108(4): p. 558-62. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/4651345>
104. Game X, et al. Botulinum toxin A detrusor injections in patients with neurogenic detrusor overactivity significantly decrease the incidence of symptomatic urinary tract infections. *Eur Urol*, 2008. 53(3): p. 613-8.
<http://www.ncbi.nlm.nih.gov/pubmed/17804150>
105. Frankel HL, et al. Long-term survival in spinal cord injury: a fifty year investigation. *Spinal Cord*, 1998. 36(4): p. 266-74.
<http://www.ncbi.nlm.nih.gov/pubmed/9589527>
106. Jamil F. Towards a catheter free status in neurogenic bladder dysfunction: a review of bladder management options in spinal cord injury (SCI). *Spinal Cord*, 2001. 39(7): p. 355-61.
<http://www.ncbi.nlm.nih.gov/pubmed/11464308>
107. Bauer SB. Neurogenic bladder: etiology and assessment. *Pediatr Nephrol*, 2008. 23(4): p. 541-51.
<http://www.ncbi.nlm.nih.gov/pubmed/18270749>
108. Barbalias GA, et al. Critical evaluation of the Crede maneuver: a urodynamic study of 207 patients. *J Urol*, 1983. 130(4): p. 720-3.
<http://www.ncbi.nlm.nih.gov/pubmed/6887405>
109. Reinberg Y, et al. Renal rupture after the Crede maneuver. *J Pediatr*, 1994. 124(2): p. 279-81.
<http://www.ncbi.nlm.nih.gov/pubmed/8301439>
110. Wyndaele JJ, et al. Neurologic urinary incontinence. *Neurourol Urodyn*, 2010. 29(1): p. 159-64.
<http://www.ncbi.nlm.nih.gov/pubmed/20025021>
111. Consortium for Spinal Cord Medicine. Outcomes following traumatic spinal cord injury: clinical practice guidelines for health-care professionals. *J Spinal Cord Med*, 2000. 23(4): p. 289-316.
<http://www.ncbi.nlm.nih.gov/pubmed/17536300>
112. Weld KJ, et al. Effect of bladder management on urological complications in spinal cord injured patients. *J Urol*, 2000. 163(3): p. 768-72.
<http://www.ncbi.nlm.nih.gov/pubmed/10687973>
113. El-Masri WS, et al. Long-term follow-up study of outcomes of bladder management in spinal cord injury patients under the care of the Midlands Centre for Spinal Injuries in Oswestry. *Spinal Cord*, 2012. 50(1): p. 14-21.
<http://www.ncbi.nlm.nih.gov/pubmed/21808256>
114. Menon EB, et al. Bladder training in patients with spinal cord injury. *Urology*, 1992. 40(5): p. 425-9.
<http://www.ncbi.nlm.nih.gov/pubmed/1441039>
115. Nijman RJ. Classification and treatment of functional incontinence in children. *BJU Int*, 2000. 85 Suppl 3: p. 37-42; discussion 45-6.
<http://www.ncbi.nlm.nih.gov/pubmed/11954196>
116. Singh R, et al. Bladder management methods and urological complications in spinal cord injury patients. *Indian J Orthop*, 2011. 45(2): p. 141-7.
<http://www.ncbi.nlm.nih.gov/pubmed/21430869>
117. Madersbacher H, et al. Conservative management in neuropathic urinary incontinence, in *Incontinence*, P. Abrams, et al., Editors. 2002, Health Publication: Plymouth: p. 697-754.
118. Fall M, et al. Electrical stimulation. A physiologic approach to the treatment of urinary incontinence. *Urol Clin North Am*, 1991. 18(2): p. 393-407.
<http://www.ncbi.nlm.nih.gov/pubmed/2017820>

119. Vodusek DB, et al. Detrusor inhibition induced by stimulation of pudendal nerve afferents. *Neurourol Urodyn*, 1986. 5: p. 381-9.
120. Bemelmans BL, et al. Neuromodulation by implant for treating lower urinary tract symptoms and dysfunction. *Eur Urol*, 1999. 36(2): p. 81-91.
<http://www.ncbi.nlm.nih.gov/pubmed/10420026>
121. Primus G, et al. Maximal external electrical stimulation for treatment of neurogenic or nonneurogenic urgency and/or urge incontinence. *Neurourol Urodyn*, 1996. 15(3): p. 187-94.
<http://www.ncbi.nlm.nih.gov/pubmed/8732985>
122. Opisso E, et al. Patient controlled versus automatic stimulation of pudendal nerve afferents to treat neurogenic detrusor overactivity. *J Urol*, 2008. 180(4): p. 1403-8.
<http://www.ncbi.nlm.nih.gov/pubmed/18710774>
123. Kabay SC, et al. Acute urodynamic effects of percutaneous posterior tibial nerve stimulation on neurogenic detrusor overactivity in patients with Parkinson's disease. *Neurourol Urodyn*, 2009. 28(1): p. 62-7.
<http://www.ncbi.nlm.nih.gov/pubmed/18837432>
124. Kabay S, et al. The clinical and urodynamic results of a 3-month percutaneous posterior tibial nerve stimulation treatment in patients with multiple sclerosis-related neurogenic bladder dysfunction. *Neurourol Urodyn*, 2009. 28(8): p. 964-8.
<http://www.ncbi.nlm.nih.gov/pubmed/19373898>
125. Pannek J, et al. [Neurogenic or idiopathic detrusor overactivity after failed antimuscarinic treatment: clinical value of external temporary electrostimulation]. *Urologe A*, 2010. 49(4): p. 530-5. [Article in German]
<http://www.ncbi.nlm.nih.gov/pubmed/20057991>
126. de Seze M, et al. Transcutaneous posterior tibial nerve stimulation for treatment of the overactive bladder syndrome in multiple sclerosis: results of a multicenter prospective study. *Neurourol Urodyn*, 2011. 30(3): p. 306-11.
<http://www.ncbi.nlm.nih.gov/pubmed/21305588>
127. Radziszewski K. Outcomes of electrical stimulation of the neurogenic bladder: results of a two-year follow-up study. *NeuroRehabilitation*, 2013. 32(4): p. 867-73.
<http://www.ncbi.nlm.nih.gov/pubmed/23867413>
128. Kajbafzadeh AM, et al. Efficacy of transcutaneous functional electrical stimulation on urinary incontinence in myelomeningocele: results of a pilot study. *Int Braz J Urol*, 2010. 36(5): p. 614-20.
<http://www.ncbi.nlm.nih.gov/pubmed/21044379>
129. McClurg D, et al. Neuromuscular electrical stimulation and the treatment of lower urinary tract dysfunction in multiple sclerosis--a double blind, placebo controlled, randomised clinical trial. *Neurourol Urodyn*, 2008. 27(3): p. 231-7.
<http://www.ncbi.nlm.nih.gov/pubmed/17705160>
130. McClurg D, et al. Comparison of pelvic floor muscle training, electromyography biofeedback, and neuromuscular electrical stimulation for bladder dysfunction in people with multiple sclerosis: a randomized pilot study. *Neurourol Urodyn*, 2006. 25(4): p. 337-48.
<http://www.ncbi.nlm.nih.gov/pubmed/16637070>
131. Hagerty JA, et al. Intravesical electrotherapy for neurogenic bladder dysfunction: a 22-year experience. *J Urol*, 2007. 178(4 Pt 2): p. 1680-3; discussion 1683.
<http://www.ncbi.nlm.nih.gov/pubmed/17707024>
132. Primus G, et al. Restoration of micturition in patients with acontractile and hypocontractile detrusor by transurethral electrical bladder stimulation. *Neurourol Urodyn*, 1996. 15(5): p. 489-97.
<http://www.ncbi.nlm.nih.gov/pubmed/8857617>
133. Lombardi G, et al. Clinical efficacy of intravesical electrostimulation on incomplete spinal cord patients suffering from chronic neurogenic non-obstructive retention: a 15-year single centre retrospective study. *Spinal Cord*, 2013. 51(3): p. 232-7.
<http://www.ncbi.nlm.nih.gov/pubmed/23147136>
134. Zempleni MZ, et al. Cortical substrate of bladder control in SCI and the effect of peripheral pudendal stimulation. *Neuroimage*, 2010. 49(4): p. 2983-94.
<http://www.ncbi.nlm.nih.gov/pubmed/19878725>
135. Lee YH, et al. The effect of semiconditional dorsal penile nerve electrical stimulation on capacity and compliance of the bladder with deformity in spinal cord injury patients: a pilot study. *Spinal Cord*, 2012. 50(4): p. 289-93.
<http://www.ncbi.nlm.nih.gov/pubmed/22231544>

136. Brusa L, et al. Effects of inhibitory rTMS on bladder function in Parkinson's disease patients. *Mov Disord*, 2009. 24(3): p. 445-8.
<http://www.ncbi.nlm.nih.gov/pubmed/19133657>
137. Centonze D, et al. Effects of motor cortex rTMS on lower urinary tract dysfunction in multiple sclerosis. *Mult Scler*, 2007. 13(2): p. 269-71.
<http://www.ncbi.nlm.nih.gov/pubmed/17439897>
138. Amend B, et al. Effective treatment of neurogenic detrusor dysfunction by combined high-dosed antimuscarinics without increased side-effects. *Eur Urol*, 2008. 53(5): p. 1021-8.
<http://www.ncbi.nlm.nih.gov/pubmed/18243516>
139. Cameron AP. Pharmacologic therapy for the neurogenic bladder. *Urol Clin North Am*, 2010. 37(4): p. 495-506.
<http://www.ncbi.nlm.nih.gov/pubmed/20955901>
140. Cameron AP, et al. Combination drug therapy improves compliance of the neurogenic bladder. *J Urol*, 2009. 182(3): p. 1062-7.
<http://www.ncbi.nlm.nih.gov/pubmed/19616807>
141. Thomas LH, et al. Treatment of urinary incontinence after stroke in adults. *Cochrane Database Syst Rev*, 2008(1): p. Cd004462.
<http://www.ncbi.nlm.nih.gov/pubmed/18254050>
142. Yeo L, et al. Urinary tract dysfunction in Parkinson's disease: a review. *Int Urol Nephrol*, 2012. 44(2): p. 415-24.
<http://www.ncbi.nlm.nih.gov/pubmed/21553114>
143. Andersson KE. Antimuscarinic mechanisms and the overactive detrusor: an update. *Eur Urol*, 2011. 59(3): p. 377-86.
<http://www.ncbi.nlm.nih.gov/pubmed/21168951>
144. Kennelly MJ, et al. Overactive bladder: pharmacologic treatments in the neurogenic population. *Rev Urol*, 2008. 10(3): p. 182-91.
<http://www.ncbi.nlm.nih.gov/pubmed/19628264>
145. Madersbacher H, et al. Neurogenic detrusor overactivity in adults: a review on efficacy, tolerability and safety of oral antimuscarinics. *Spinal Cord*, 2013. 51(6): p. 432-41.
<http://www.ncbi.nlm.nih.gov/pubmed/23743498>
146. Madhuvrata P, et al. Anticholinergic drugs for adult neurogenic detrusor overactivity: a systematic review and meta-analysis. *Eur Urol*, 2012. 62(5): p. 816-30.
<http://www.ncbi.nlm.nih.gov/pubmed/22397851>
147. Sakakibara R, et al. Dementia and lower urinary dysfunction: with a reference to anticholinergic use in elderly population. *Int J Urol*, 2008. 15(9): p. 778-88.
<http://www.ncbi.nlm.nih.gov/pubmed/18643858>
148. Yamaguchi O. Antimuscarinics and overactive bladder: other mechanism of action. *Neurourol Urodyn*, 2010. 29(1): p. 112-5.
<http://www.ncbi.nlm.nih.gov/pubmed/19693952>
149. Drake MJ, et al. Neurologic Urinary and Faecal Incontinence, In *Incontinence*, P. Abrams, et al., Editors. 2013, ICUD-EAU. p. 827-980.
150. Nicholas RS, et al. Anticholinergics for urinary symptoms in multiple sclerosis. *Cochrane Database Syst Rev*, 2009(1): p. Cd004193.
<http://www.ncbi.nlm.nih.gov/pubmed/19160231>
151. Reddy PP, et al. Long-term efficacy and safety of tolterodine in children with neurogenic detrusor overactivity. *J Pediatr Urol*, 2008. 4(6): p. 428-33.
<http://www.ncbi.nlm.nih.gov/pubmed/19013412>
152. Schulte-Baukloh H, et al. Urodynamic effects of propiverine in children and adolescents with neurogenic bladder: Results of a prospective long-term study. *Journal of Pediatric Urology*, 2012. 8(4): p. 386-392.
<http://www.ncbi.nlm.nih.gov/pubmed/21907623>
153. Bolduc S, et al. Prospective open label study of solifenacin for overactive bladder in children. *J Urol*, 2010. 184(4 Suppl): p. 1668-73.
<http://www.ncbi.nlm.nih.gov/pubmed/20728124>
154. A Study To Find Out How Fesoterodine Works In Children Aged 6 To 17 Years With Bladder Overactivity Caused By A Neurological Condition (NCT01557244), Pfizer. Last verified: January 2015.
<https://clinicaltrials.gov/ct2/show/NCT01557244>

155. Bennett N, et al. Can higher doses of oxybutynin improve efficacy in neurogenic bladder? *J Urol*, 2004. 171(2 Pt 1): p. 749-51.
<http://www.ncbi.nlm.nih.gov/pubmed/14713802>
156. Horstmann M, et al. Neurogenic bladder treatment by doubling the recommended antimuscarinic dosage. *Neurourol Urodyn*, 2006. 25(5): p. 441-5.
<http://www.ncbi.nlm.nih.gov/pubmed/16847942>
157. Menarini M, et al. Trospium chloride in patients with neurogenic detrusor overactivity: is dose titration of benefit to the patients? *Int J Clin Pharmacol Ther*, 2006. 44(12): p. 623-32.
<http://www.ncbi.nlm.nih.gov/pubmed/17190372>
158. Nardulli R, et al. Combined antimuscarinics for treatment of neurogenic overactive bladder. *Int J Immunopathol Pharmacol*, 2012. 25(1 Suppl): p. 35s-41s.
<http://www.ncbi.nlm.nih.gov/pubmed/22652160>
159. O'Leary M, et al. Effect of controlled-release oxybutynin on neurogenic bladder function in spinal cord injury. *J Spinal Cord Med*, 2003. 26(2): p. 159-62.
<http://www.ncbi.nlm.nih.gov/pubmed/12828295>
160. Stohrer M, et al. Propiverine compared to oxybutynin in neurogenic detrusor overactivity--results of a randomized, double-blind, multicenter clinical study. *Eur Urol*, 2007. 51(1): p. 235-42.
<http://www.ncbi.nlm.nih.gov/pubmed/16698176>
161. Verpoorten C, et al. The neurogenic bladder: medical treatment. *Pediatr Nephrol*, 2008. 23(5): p. 717-25.
<http://www.ncbi.nlm.nih.gov/pubmed/18095004>
162. Isik AT, et al. Trospium and cognition in patients with late onset Alzheimer disease. *J Nutr Health Aging*, 2009. 13(8): p. 672-6.
<http://www.ncbi.nlm.nih.gov/pubmed/19657549>
163. Ethans KD, et al. Efficacy and safety of tolterodine in people with neurogenic detrusor overactivity. *J Spinal Cord Med*, 2004. 27(3): p. 214-8.
<http://www.ncbi.nlm.nih.gov/pubmed/15478523>
164. Mahanta K, et al. Comparative efficacy and safety of extended-release and instant-release tolterodine in children with neural tube defects having cystometric abnormalities. *J Pediatr Urol*, 2008. 4(2): p. 118-23.
<http://www.ncbi.nlm.nih.gov/pubmed/18631906>
165. Grigoleit U, et al. Efficacy, tolerability and safety of propiverine hydrochloride in children and adolescents with congenital or traumatic neurogenic detrusor overactivity--a retrospective study. *Eur Urol*, 2006. 49(6): p. 1114-20; discussion 1120-1.
<http://www.ncbi.nlm.nih.gov/pubmed/16542772>
166. Madersbacher H, et al. Efficacy, tolerability and safety profile of propiverine in the treatment of the overactive bladder (non-neurogenic and neurogenic). *World J Urol*, 2001. 19(5): p. 324-35.
<http://www.ncbi.nlm.nih.gov/pubmed/11760781>
167. Madersbacher H, et al. Propiverine vs oxybutynin for treating neurogenic detrusor overactivity in children and adolescents: results of a multicentre observational cohort study. *BJU Int*, 2009. 103(6): p. 776-81.
<http://www.ncbi.nlm.nih.gov/pubmed/19007380>
168. Stohrer M, et al. Efficacy and tolerability of propiverine hydrochloride extended-release compared with immediate-release in patients with neurogenic detrusor overactivity. *Spinal Cord*, 2013. 51(5): p. 419-23.
<http://www.ncbi.nlm.nih.gov/pubmed/23338657>
169. Carl S, et al. Darifenacin is also effective in neurogenic bladder dysfunction (multiple sclerosis). *Urology*, 2006. 68 (suppl)(250).
170. Bycroft J, et al. The effect of darifenacin on neurogenic detrusor overactivity in patients with spinal cord injury. *Neurourol Urodyn* 2003. 22: p. A190.
<http://www.ics.org/Abstracts/Publish/41/000190.pdf>
171. Hassouna M. Comparative Study of the Efficacy and Safety of Muscarinic M3 Receptors Antagonists in the Treatment of Neurogenic Detrusor Overactivity (NCT00800462).
<http://clinicaltrials.gov/show/NCT00800462>
172. Astellas Pharma Inc. Clinical Study of Solifenacin Succinate in Patients With Bladder Symptoms Due to Spinal Cord Injury or Multiple Sclerosis (SONIC).
<http://clinicaltrials.gov/show/NCT00629642>
173. Zesiewicz T, et al. Solifenacin Succinate (VESIcare) for the Treatment of Overactive Bladder in Parkinson's Disease (URGE-PD).
<https://www.clinicaltrials.gov/show/NCT01018264>

174. Cartwright PC, et al. Efficacy and safety of transdermal and oral oxybutynin in children with neurogenic detrusor overactivity. *J Urol*, 2009. 182(4): p. 1548-54.
<http://www.ncbi.nlm.nih.gov/pubmed/19683731>
175. Kennelly MJ, et al. Efficacy and safety of oxybutynin transdermal system in spinal cord injury patients with neurogenic detrusor overactivity and incontinence: an open-label, dose-titration study. *Urology*, 2009. 74(4): p. 741-5.
<http://www.ncbi.nlm.nih.gov/pubmed/19628264>
176. Hill LA, Watson Pharmaceuticals. Safety and Efficacy Evaluation of Oxybutynin Topical Gel In Children With Neurogenic Bladder. (NCT01192568).
<https://clinicaltrials.gov/show/NCT01192568>
177. Fader M, et al. Intravesical atropine compared to oral oxybutynin for neurogenic detrusor overactivity: a double-blind, randomized crossover trial. *J Urol*, 2007. 177(1): p. 208-13; discussion 213.
<http://www.ncbi.nlm.nih.gov/pubmed/17162046>
178. Krause P, et al. Pharmacokinetics of intravesical versus oral oxybutynin in healthy adults: results of an open label, randomized, prospective clinical study. *J Urol*, 2013. 190(5): p. 1791-7.
<http://www.ncbi.nlm.nih.gov/pubmed/23669567>
179. Van Meel TD, et al. The effect of intravesical oxybutynin on the ice water test and on electrical perception thresholds in patients with neurogenic detrusor overactivity. *Neurourol Urodyn*, 2010. 29(3): p. 391-4.
<http://www.ncbi.nlm.nih.gov/pubmed/19787712>
180. Angulo J, et al. Tadalafil enhances the inhibitory effects of tamsulosin on neurogenic contractions of human prostate and bladder neck. *J Sex Med*, 2012. 9(9): p. 2293-306.
<http://www.ncbi.nlm.nih.gov/pubmed/22759598>
181. Behr-Roussel D, et al. Vardenafil decreases bladder afferent nerve activity in unanesthetized, decerebrate, spinal cord-injured rats. *Eur Urol*, 2011. 59(2): p. 272-9.
<http://www.ncbi.nlm.nih.gov/pubmed/21036463>
182. Kanai A, et al. Mechanisms of action of botulinum neurotoxins, beta3-adrenergic receptor agonists, and PDE5 inhibitors in modulating detrusor function in overactive bladders: ICI-RS 2011. *Neurourol Urodyn*, 2012. 31(3): p. 300-8.
<http://www.ncbi.nlm.nih.gov/pubmed/22275187>
183. Herschorn S, et al. A phase III, randomized, double-blind, parallel-group, placebo-controlled, multicentre study to assess the efficacy and safety of the beta(3) adrenoceptor agonist, mirabegron, in patients with symptoms of overactive bladder. *Urology*, 2013. 82(2): p. 313-20.
<http://www.ncbi.nlm.nih.gov/pubmed/23769122>
184. Nitti VW, et al. Urodynamics and safety of the beta(3)-adrenoceptor agonist mirabegron in males with lower urinary tract symptoms and bladder outlet obstruction. *J Urol*, 2013. 190(4): p. 1320-7.
<http://www.ncbi.nlm.nih.gov/pubmed/23727415>
185. Astellas Pharma Inc. A clinical study to assess the effect on pharmacokinetics of dosing mirabegron (YM178) and solifenacin simultaneously.
<https://www.clinicaltrials.gov/show/NCT01297192>
186. Barendrecht MM, et al. Is the use of parasympathomimetics for treating an underactive urinary bladder evidence-based? *BJU Int*, 2007. 99(4): p. 749-52.
<http://www.ncbi.nlm.nih.gov/pubmed/17233798>
187. Apostolidis A. Taming the cannabinoids: new potential in the pharmacologic control of lower urinary tract dysfunction. *Eur Urol*, 2012. 61(1): p. 107-9; discussion 109-11.
<http://www.ncbi.nlm.nih.gov/pubmed/21996529>
188. Gratzke C, et al. Effects of cannabior, a novel selective cannabinoid 2 receptor agonist, on bladder function in normal rats. *Eur Urol*, 2010. 57(6): p. 1093-100.
<http://www.ncbi.nlm.nih.gov/pubmed/20207474>
189. Kavia RB, et al. Randomized controlled trial of Sativex to treat detrusor overactivity in multiple sclerosis. *Mult Scler*, 2010. 16(11): p. 1349-59.
<http://www.ncbi.nlm.nih.gov/pubmed/20829244>
190. Takeda M, et al. Predictive factors for the effect of the alpha1-D/A adrenoceptor antagonist naftopidil on subjective and objective criteria in patients with neurogenic lower urinary tract dysfunction. *BJU Int*, 2011. 108(1): p. 100-7.
<http://www.ncbi.nlm.nih.gov/pubmed/21062392>

191. Yamanishi T, et al. Combination of a cholinergic drug and an alpha-blocker is more effective than monotherapy for the treatment of voiding difficulty in patients with underactive detrusor. *Int J Urol*, 2004. 11(2): p. 88-96.
<http://www.ncbi.nlm.nih.gov/pubmed/14706012>
192. Guttman L, et al. The value of intermittent catheterisation in the early management of traumatic paraplegia and tetraplegia. *Paraplegia*, 1966. 4(2): p. 63-84.
<http://www.ncbi.nlm.nih.gov/pubmed/5969402>
193. Lapidus J, et al. Clean, intermittent self-catheterization in the treatment of urinary tract disease. *J Urol*, 1972. 107(3): p. 458-61. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/5010715>
194. Wyndaele JJ. Intermittent catheterization: which is the optimal technique? *Spinal Cord*, 2002. 40(9): p. 432-7.
<http://www.ncbi.nlm.nih.gov/pubmed/12185603>
195. Prieto-Fingerhut T, et al. A study comparing sterile and nonsterile urethral catheterization in patients with spinal cord injury. *Rehabil Nurs*, 1997. 22(6): p. 299-302.
<http://www.ncbi.nlm.nih.gov/pubmed/9416190>
196. Matsumoto T, et al. Urinary tract infection in neurogenic bladder. *Int J Antimicrob Agents*, 2001. 17(4): p. 293-7.
<http://www.ncbi.nlm.nih.gov/pubmed/11295411>
197. Hudson E, et al. The 'no-touch' method of intermittent urinary catheter insertion: can it reduce the risk of bacteria entering the bladder? *Spinal Cord*, 2005. 43(10): p. 611-4.
<http://www.ncbi.nlm.nih.gov/pubmed/15852058>
198. Günther M, et al. [Effects of aseptic intermittent catheterisation on the male urethra]. *Der Urologe B*, 2001. 41(4): p. 359-361. [Article in German]
<http://link.springer.com/article/10.1007%2Fs001310170044>
199. Goepel M, et al. Der intermittierende Selbstkatheterismus - Ergebnisse einer vergleichenden Untersuchung. *Urologe [B]*, 1996. 36: p. 190-4. [Article in German]
200. Bakke A, et al. Physical predictors of infection in patients treated with clean intermittent catheterization: a prospective 7-year study. *Br J Urol*, 1997. 79(1): p. 85-90.
<http://www.ncbi.nlm.nih.gov/pubmed/9043503>
201. Waller L, et al. Clean intermittent catheterization in spinal cord injury patients: long-term followup of a hydrophilic low friction technique. *J Urol*, 1995. 153(2): p. 345-8.
<http://www.ncbi.nlm.nih.gov/pubmed/7815579>
202. Wyndaele JJ. Complications of intermittent catheterization: their prevention and treatment. *Spinal Cord*, 2002. 40(10): p. 536-41.
<http://www.ncbi.nlm.nih.gov/pubmed/12235537>
203. Woodbury MG, et al. Intermittent catheterization practices following spinal cord injury: a national survey. *Can J Urol*, 2008. 15(3): p. 4065-71.
<http://www.ncbi.nlm.nih.gov/pubmed/18570710>
204. Bennett CJ, et al. Comparison of bladder management complication outcomes in female spinal cord injury patients. *J Urol*, 1995. 153(5): p. 1458-60.
<http://www.ncbi.nlm.nih.gov/pubmed/7714965>
205. Chancellor MB, et al. Functional urethral closure with pubovaginal sling for destroyed female urethra after long-term urethral catheterization. *Urology*, 1994. 43(4): p. 499-505.
<http://www.ncbi.nlm.nih.gov/pubmed/8154071>
206. Chao R, et al. Fate of upper urinary tracts in patients with indwelling catheters after spinal cord injury. *Urology*, 1993. 42(3): p. 259-62.
<http://www.ncbi.nlm.nih.gov/pubmed/8379025>
207. Larsen LD, et al. Retrospective analysis of urologic complications in male patients with spinal cord injury managed with and without indwelling urinary catheters. *Urology*, 1997. 50(3): p. 418-22.
<http://www.ncbi.nlm.nih.gov/pubmed/9301708>
208. Mitsui T, et al. Is suprapubic cystostomy an optimal urinary management in high quadriplegics? A comparative study of suprapubic cystostomy and clean intermittent catheterization. *Eur Urol*, 2000. 38(4): p. 434-8.
<http://www.ncbi.nlm.nih.gov/pubmed/11025382>
209. Park YI, et al. A method to minimize indwelling catheter calcification and bladder stones in individuals with spinal cord injury. *J Spinal Cord Med*, 2001. 24(2): p. 105-8.
<http://www.ncbi.nlm.nih.gov/pubmed/11587416>

210. Weld KJ, et al. Influences on renal function in chronic spinal cord injured patients. *J Urol*, 2000. 164(5): p. 1490-3.
<http://www.ncbi.nlm.nih.gov/pubmed/11025689>
211. West DA, et al. Role of chronic catheterization in the development of bladder cancer in patients with spinal cord injury. *Urology*, 1999. 53(2): p. 292-7.
<http://www.ncbi.nlm.nih.gov/pubmed/9933042>
212. Hollingsworth JM, et al. Determining the noninfectious complications of indwelling urethral catheters: a systematic review and meta-analysis. *Ann Intern Med*, 2013. 159(6): p. 401-10.
<http://www.ncbi.nlm.nih.gov/pubmed/24042368>
213. Di Stasi SM, et al. Intravesical oxybutynin: mode of action assessed by passive diffusion and electromotive administration with pharmacokinetics of oxybutynin and N-desethyl oxybutynin. *J Urol*, 2001. 166(6): p. 2232-6.
<http://www.ncbi.nlm.nih.gov/pubmed/11696741>
214. Buysse G, et al. Intravesical oxybutynin for neurogenic bladder dysfunction: less systemic side effects due to reduced first pass metabolism. *J Urol*, 1998. 160(3 Pt 1): p. 892-6.
<http://www.ncbi.nlm.nih.gov/pubmed/9720583>
215. Haferkamp A, et al. Dosage escalation of intravesical oxybutynin in the treatment of neurogenic bladder patients. *Spinal Cord*, 2000. 38(4): p. 250-4.
<http://www.ncbi.nlm.nih.gov/pubmed/10822396>
216. Pannek J, et al. Combined intravesical and oral oxybutynin chloride in adult patients with spinal cord injury. *Urology*, 2000. 55(3): p. 358-62.
<http://www.ncbi.nlm.nih.gov/pubmed/10699610>
217. Geirsson G, et al. Clinical and urodynamic effects of intravesical capsaicin treatment in patients with chronic traumatic spinal detrusor hyperreflexia. *J Urol*, 1995. 154(5): p. 1825-9.
<http://www.ncbi.nlm.nih.gov/pubmed/7563356>
218. Giannantoni A, et al. Intravesical resiniferatoxin versus botulinum-A toxin injections for neurogenic detrusor overactivity: a prospective randomized study. *J Urol*, 2004. 172(1): p. 240-3.
<http://www.ncbi.nlm.nih.gov/pubmed/15201783>
219. Kim JH, et al. Intravesical resiniferatoxin for refractory detrusor hyperreflexia: a multicenter, blinded, randomized, placebo-controlled trial. *J Spinal Cord Med*, 2003. 26(4): p. 358-63.
<http://www.ncbi.nlm.nih.gov/pubmed/14992337>
220. Katona F, et al. [Intraluminal electrotherapy of various paralytic conditions of the gastrointestinal tract with the quadrangular current]. *Zentralbl Chir*, 1959. 84(24): p. 929-33. [Article in German] [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/13676705>
221. Kaplan WE. Intravesical electrical stimulation of the bladder: pro. *Urology*, 2000. 56(1): p. 2-4.
<http://www.ncbi.nlm.nih.gov/pubmed/10869607>
222. Ebner A, et al. Intravesical electrical stimulation--an experimental analysis of the mechanism of action. *J Urol*, 1992. 148(3): p. 920-4.
<http://www.ncbi.nlm.nih.gov/pubmed/1512860>
223. De Wachter S, et al. Quest for standardisation of electrical sensory testing in the lower urinary tract: the influence of technique related factors on bladder electrical thresholds. *Neurourol Urodyn*, 2003. 22(2): p. 118-22.
<http://www.ncbi.nlm.nih.gov/pubmed/12579628>
224. Katona F, et al. Intravesical transurethral electrotherapy in meningomyelocele patients. *Acta Paediatr Acad Sci Hung*, 1975. 16(3-4): p. 363-74.
<http://www.ncbi.nlm.nih.gov/pubmed/773096>
225. Nicholas JL, et al. Endovesical electrotherapy in treatment of urinary incontinence in spina-bifida patients. *Lancet*, 1975. 2(7948): p. 1276-7.
<http://www.ncbi.nlm.nih.gov/pubmed/54798>
226. Pugach JL, et al. Intravesical electrostimulation in pediatric patients with spinal cord defects. *J Urol*, 2000. 164(3 Pt 2): p. 965-8.
<http://www.ncbi.nlm.nih.gov/pubmed/10958718>
227. Reitz A, et al. European experience of 200 cases treated with botulinum-A toxin injections into the detrusor muscle for urinary incontinence due to neurogenic detrusor overactivity. *Eur Urol*, 2004. 45(4): p. 510-5.
<http://www.ncbi.nlm.nih.gov/pubmed/15041117>

228. Schurch B, et al. Botulinum toxin type a is a safe and effective treatment for neurogenic urinary incontinence: results of a single treatment, randomized, placebo controlled 6-month study. *J Urol*, 2005. 174(1): p. 196-200.
<http://www.ncbi.nlm.nih.gov/pubmed/15947626>
229. Cruz F, et al. Efficacy and safety of onabotulinumtoxinA in patients with urinary incontinence due to neurogenic detrusor overactivity: a randomised, double-blind, placebo-controlled trial. *Eur Urol*, 2011. 60(4): p. 742-50.
<http://www.ncbi.nlm.nih.gov/pubmed/21798658>
230. Ginsberg D, et al. Phase 3 efficacy and tolerability study of onabotulinumtoxinA for urinary incontinence from neurogenic detrusor overactivity. *J Urol*, 2012. 187(6): p. 2131-9.
<http://www.ncbi.nlm.nih.gov/pubmed/22503020>
231. Grosse J, et al. Success of repeat detrusor injections of botulinum a toxin in patients with severe neurogenic detrusor overactivity and incontinence. *Eur Urol*, 2005. 47(5): p. 653-9.
<http://www.ncbi.nlm.nih.gov/pubmed/15826758>
232. Haferkamp A, et al. Lack of ultrastructural detrusor changes following endoscopic injection of botulinum toxin type a in overactive neurogenic bladder. *Eur Urol*, 2004. 46(6): p. 784-91.
<http://www.ncbi.nlm.nih.gov/pubmed/15548448>
233. Dykstra DD, et al. Treatment of detrusor-sphincter dyssynergia with botulinum A toxin: a doubleblind study. *Arch Phys Med Rehabil*, 1990. 71(1): p. 24-6.
<http://www.ncbi.nlm.nih.gov/pubmed/2297305>
234. Petit H, et al. Botulinum A toxin treatment for detrusor-sphincter dyssynergia in spinal cord disease. *Spinal Cord*, 1998. 36(2): p. 91-4.
<http://www.ncbi.nlm.nih.gov/pubmed/9494997>
235. Schurch B, et al. Botulinum-A toxin as a treatment of detrusor-sphincter dyssynergia: a prospective study in 24 spinal cord injury patients. *J Urol*, 1996. 155(3): p. 1023-9.
<http://www.ncbi.nlm.nih.gov/pubmed/8583552>
236. Utomo E, et al. Surgical management of functional bladder outlet obstruction in adults with neurogenic bladder dysfunction. *Cochrane Database Syst Rev*, 2014. 5: p. Cd004927.
<http://www.ncbi.nlm.nih.gov/pubmed/24859260>
237. Chancellor MB, et al. Prospective comparison of external sphincter balloon dilatation and prosthesis placement with external sphincterotomy in spinal cord injured men. *Arch Phys Med Rehabil*, 1994. 75(3): p. 297-305.
<http://www.ncbi.nlm.nih.gov/pubmed/8129583>
238. Perkasch I. Use of contact laser crystal tip firing Nd:YAG to relieve urinary outflow obstruction in male neurogenic bladder patients. *J Clin Laser Med Surg*, 1998. 16(1): p. 33-8.
<http://www.ncbi.nlm.nih.gov/pubmed/9728128>
239. Reynard JM, et al. Sphincterotomy and the treatment of detrusor-sphincter dyssynergia: current status, future prospects. *Spinal Cord*, 2003. 41(1): p. 1-11.
<http://www.ncbi.nlm.nih.gov/pubmed/12494314>
240. Noll F, et al. Transurethral sphincterotomy in quadriplegic patients: long-term-follow-up. *Neurourol Urodyn*, 1995. 14(4): p. 351-8.
<http://www.ncbi.nlm.nih.gov/pubmed/7581471>
241. Derry F, et al. Audit of bladder neck resection in spinal cord injured patients. *Spinal Cord*, 1998. 36(5): p. 345-8.
<http://www.ncbi.nlm.nih.gov/pubmed/9601115>
242. Chancellor MB, et al. Long-term followup of the North American multicenter UroLume trial for the treatment of external detrusor-sphincter dyssynergia. *J Urol*, 1999. 161(5): p. 1545-50.
<http://www.ncbi.nlm.nih.gov/pubmed/10210393>
243. Seoane-Rodriguez S, et al. Long-term follow-up study of intraurethral stents in spinal cord injured patients with detrusor-sphincter dyssynergia. *Spinal Cord*, 2007. 45(9): p. 621-6.
<http://www.ncbi.nlm.nih.gov/pubmed/17211463>
244. Gajewski JB, et al. Removal of UroLume endoprosthesis: experience of the North American Study Group for detrusor-sphincter dyssynergia application. *J Urol*, 2000. 163(3): p. 773-6.
<http://www.ncbi.nlm.nih.gov/pubmed/10687974>
245. Wilson TS, et al. UroLume stents: lessons learned. *J Urol*, 2002. 167(6): p. 2477-80.
<http://www.ncbi.nlm.nih.gov/pubmed/11992061>
246. Polguer T, et al. [Treatment of detrusor-striated sphincter dyssynergia with permanent nitinol urethral stent: results after a minimum follow-up of 2 years]. *Prog Urol*, 2012. 22(17): p. 1058-63. [Article in French]
<http://www.ncbi.nlm.nih.gov/pubmed/23182120>

247. van der Merwe A, et al. Outcome of dual flange metallic urethral stents in the treatment of neuropathic bladder dysfunction after spinal cord injury. *J Endourol*, 2012. 26(9): p. 1210-5. <http://www.ncbi.nlm.nih.gov/pubmed/22519741>
248. Pannek J, et al. Clinical usefulness of the memokath stent as a second-line procedure after sphincterotomy failure. *J Endourol*, 2011. 25(2): p. 335-9. <http://www.ncbi.nlm.nih.gov/pubmed/20977372>
249. Abdul-Rahman A, et al. A 20-year follow-up of the mesh wallstent in the treatment of detrusor external sphincter dyssynergia in patients with spinal cord injury. *BJU Int*, 2010. 106(10): p. 1510-3. <http://www.ncbi.nlm.nih.gov/pubmed/20500511>
250. Bennett JK, et al. Collagen injections for intrinsic sphincter deficiency in the neuropathic urethra. *Paraplegia*, 1995. 33(12): p. 697-700. <http://www.ncbi.nlm.nih.gov/pubmed/8927407>
251. Block CA, et al. Long-term efficacy of periurethral collagen injection for the treatment of urinary incontinence secondary to myelomeningocele. *J Urol*, 2003. 169(1): p. 327-9. <http://www.ncbi.nlm.nih.gov/pubmed/12478183>
252. Schurch B, et al. Intraurethral sphincter prosthesis to treat hyporeflexic bladders in women: does it work? *BJU Int*, 1999. 84(7): p. 789-94. <http://www.ncbi.nlm.nih.gov/pubmed/10532973>
253. Barthold JS, et al. Results of the rectus fascial sling and wrap procedures for the treatment of neurogenic sphincteric incontinence. *J Urol*, 1999. 161(1): p. 272-4. <http://www.ncbi.nlm.nih.gov/pubmed/10037423>
254. Daneshmand S, et al. Puboprostatic sling repair for treatment of urethral incompetence in adult neurogenic incontinence. *J Urol*, 2003. 169(1): p. 199-202. <http://www.ncbi.nlm.nih.gov/pubmed/12478135>
255. Gormley EA, et al. Pubovaginal slings for the management of urinary incontinence in female adolescents. *J Urol*, 1994. 152(2 Pt 2): p. 822-5; discussion 826-7. <http://www.ncbi.nlm.nih.gov/pubmed/8022024>
256. Herschorn S, et al. Fascial slings and bladder neck tapering in the treatment of male neurogenic incontinence. *J Urol*, 1992. 147(4): p. 1073-5. <http://www.ncbi.nlm.nih.gov/pubmed/1552586>
257. Kakizaki H, et al. Fascial sling for the management of urinary incontinence due to sphincter incompetence. *J Urol*, 1995. 153(3 Pt 1): p. 644-7. <http://www.ncbi.nlm.nih.gov/pubmed/7861504>
258. Mingin GC, et al. The rectus myofascial wrap in the management of urethral sphincter incompetence. *BJU Int*, 2002. 90(6): p. 550-3. <http://www.ncbi.nlm.nih.gov/pubmed/12230615>
259. Athanasopoulos A, et al. Treating stress urinary incontinence in female patients with neuropathic bladder: the value of the autologous fascia rectus sling. *Int Urol Nephrol*, 2012. 44(5): p. 1363-7. <http://www.ncbi.nlm.nih.gov/pubmed/22821050>
260. Groen LA, et al. The AdVance male sling as a minimally invasive treatment for intrinsic sphincter deficiency in patients with neurogenic bladder sphincter dysfunction: a pilot study. *Neurourology Urodyn*, 2012. 31(8): p. 1284-7. <http://www.ncbi.nlm.nih.gov/pubmed/22847896>
261. Mehnert U, et al. Treatment of neurogenic stress urinary incontinence using an adjustable continence device: 4-year followup. *J Urol*, 2012. 188(6): p. 2274-80. <http://www.ncbi.nlm.nih.gov/pubmed/23083648>
262. Light JK, et al. Use of the artificial urinary sphincter in spinal cord injury patients. *J Urol*, 1983. 130(6): p. 1127-9. <http://www.ncbi.nlm.nih.gov/pubmed/6644893>
263. Sidi AA, et al. Comparison of artificial sphincter implantation and bladder neck reconstruction in patients with neurogenic urinary incontinence. *J Urol*, 1987. 138(4 Pt 2): p. 1120-2. <http://www.ncbi.nlm.nih.gov/pubmed/3656572>
264. Bauer SB. Long-term efficacy of artificial urinary sphincters in children. *J Urol*, 2008. 180(2): p. 441. <http://www.ncbi.nlm.nih.gov/pubmed/18550090>
265. Castera R, et al. 10-Year experience with artificial urinary sphincter in children and adolescents. *J Urol*, 2001. 165(6 Pt 2): p. 2373-6. <http://www.ncbi.nlm.nih.gov/pubmed/11371980>

266. Chartier Kastler E, et al. Treatment of neurogenic male urinary incontinence related to intrinsic sphincter insufficiency with an artificial urinary sphincter: a French retrospective multicentre study. *BJU Int*, 2011. 107(3): p. 426-32.
<http://www.ncbi.nlm.nih.gov/pubmed/20633005>
267. Elliott DS, et al. Mayo Clinic long-term analysis of the functional durability of the AMS 800 artificial urinary sphincter: a review of 323 cases. *J Urol*, 1998. 159(4): p. 1206-8.
<http://www.ncbi.nlm.nih.gov/pubmed/9507835>
268. Fulford SC, et al. The fate of the 'modern' artificial urinary sphincter with a follow-up of more than 10 years. *Br J Urol*, 1997. 79(5): p. 713-6.
<http://www.ncbi.nlm.nih.gov/pubmed/9158507>
269. Kryger JV, et al. Long-term results of artificial urinary sphincters in children are independent of age at implantation. *J Urol*, 2001. 165(6 Pt 2): p. 2377-9.
<http://www.ncbi.nlm.nih.gov/pubmed/11371981>
270. Viers BR, et al. Simultaneous augmentation cystoplasty and cuff only artificial urinary sphincter in children and young adults with neurogenic urinary incontinence. *J Urol*, 2014. 191(4): p. 1104-8.
<http://www.ncbi.nlm.nih.gov/pubmed/24060640>
271. Janknegt RA, et al. Electrically stimulated gracilis sphincter for treatment of bladder sphincter incontinence. *Lancet*, 1992. 340(8828): p. 1129-30.
<http://www.ncbi.nlm.nih.gov/pubmed/1359213>
272. Chancellor MB, et al. Gracilis muscle transposition with electrical stimulation for sphincteric incontinence: a new approach. *World J Urol*, 1997. 15(5): p. 320-8.
<http://www.ncbi.nlm.nih.gov/pubmed/9372585>
273. Chancellor MB, et al. Gracilis urethromyoplasty--an autologous urinary sphincter for neurologically impaired patients with stress incontinence. *Spinal Cord*, 1997. 35(8): p. 546-9.
<http://www.nature.com/sc/journal/v35/n8/abs/3100444a.html>
274. Donnhoo KK, et al. The Young-Dees-Leadbetter bladder neck repair for neurogenic incontinence. *J Urol*, 1999. 161(6): p. 1946-9.
<http://www.ncbi.nlm.nih.gov/pubmed/10332478>
275. Kropp KA, et al. Urethral lengthening and reimplantation for neurogenic incontinence in children. *J Urol*, 1986. 135(3): p. 533-6.
<http://www.ncbi.nlm.nih.gov/pubmed/3944902>
276. Salle JL, et al. Urethral lengthening with anterior bladder wall flap (Pippi Salle procedure): modifications and extended indications of the technique. *J Urol*, 1997. 158(2): p. 585-90.
<http://www.ncbi.nlm.nih.gov/pubmed/9224369>
277. Rawashdeh YF, et al. International Children's Continence Society's recommendations for therapeutic intervention in congenital neuropathic bladder and bowel dysfunction in children. *Neurourol Urodyn*, 2012. 31(5): p. 615-20.
<http://www.ncbi.nlm.nih.gov/pubmed/22532368>
278. Nagib A, et al. Successful control of selective anterior sacral rhizotomy for treatment of spastic bladder and ureteric reflux in paraplegics. *Med Serv J Can*, 1966. 22(7): p. 576-81.
<http://www.ncbi.nlm.nih.gov/pubmed/5966992>
279. Schneidau T, et al. Selective sacral rhizotomy for the management of neurogenic bladders in spina bifida patients: long-term followup. *J Urol*, 1995. 154(2 Pt 2): p. 766-8.
<http://www.ncbi.nlm.nih.gov/pubmed/7609174>
280. Young B, et al. Percutaneous sacral rhizotomy for neurogenic detrusor hyperreflexia. *J Neurosurg*, 1980. 53(1): p. 85-7.
<http://www.ncbi.nlm.nih.gov/pubmed/7411212>
281. Koldewijn EL, et al. Bladder compliance after posterior sacral root rhizotomies and anterior sacral root stimulation. *J Urol*, 1994. 151(4): p. 955-60.
<http://www.ncbi.nlm.nih.gov/pubmed/8126835>
282. Krasmik D, et al. Urodynamic results, clinical efficacy, and complication rates of sacral intradural deafferentation and sacral anterior root stimulation in patients with neurogenic lower urinary tract dysfunction resulting from complete spinal cord injury. *Neurourol Urodyn*, 2014. 33(8): p. 1202-6.
<http://www.ncbi.nlm.nih.gov/pubmed/24038405>
283. Singh G, et al. Intravesical oxybutynin in patients with posterior rhizotomies and sacral anterior root stimulators. *Neurourol Urodyn*, 1995. 14(1): p. 65-71.
<http://www.ncbi.nlm.nih.gov/pubmed/7742851>

284. Van Kerrebroeck PE, et al. Results of the treatment of neurogenic bladder dysfunction in spinal cord injury by sacral posterior root rhizotomy and anterior sacral root stimulation. *J Urol*, 1996. 155(4): p. 1378-81.
<http://www.ncbi.nlm.nih.gov/pubmed/8632580>
285. Kutzenberger JS. Surgical therapy of neurogenic detrusor overactivity (hyperreflexia) in paraplegic patients by sacral deafferentation and implant driven micturition by sacral anterior root stimulation: methods, indications, results, complications, and future prospects. *Acta Neurochir*, 2007. 97(Pt 1): p. 333-9.
<http://www.ncbi.nlm.nih.gov/pubmed/17691394>
286. Bhadra N, et al. Selective suppression of sphincter activation during sacral anterior nerve root stimulation. *Neurourol Urodyn*, 2002. 21(1): p. 55-64.
<http://www.ncbi.nlm.nih.gov/pubmed/11835425>
287. Kirkham AP, et al. Neuromodulation through sacral nerve roots 2 to 4 with a Finetech-Brindley sacral posterior and anterior root stimulator. *Spinal Cord*, 2002. 40(6): p. 272-81.
<http://www.ncbi.nlm.nih.gov/pubmed/12037708>
288. Schumacher S, et al. Extradural cold block for selective neurostimulation of the bladder: development of a new technique. *J Urol*, 1999. 161(3): p. 950-4.
<http://www.ncbi.nlm.nih.gov/pubmed/10022732>
289. Brindley GS. An implant to empty the bladder or close the urethra. *J Neurol Neurosurg Psychiatry*, 1977. 40(4): p. 358-69.
<http://www.ncbi.nlm.nih.gov/pubmed/406364>
290. Benard A, et al. Comparative cost-effectiveness analysis of sacral anterior root stimulation for rehabilitation of bladder dysfunction in spinal cord injured patients. *Neurosurgery*, 2013. 73(4): p. 600-8; discussion 608.
<http://www.ncbi.nlm.nih.gov/pubmed/23787880>
291. Martens FM, et al. Quality of life in complete spinal cord injury patients with a Brindley bladder stimulator compared to a matched control group. *Neurourol Urodyn*, 2011. 30(4): p. 551-5.
<http://www.ncbi.nlm.nih.gov/pubmed/21328472>
292. Wollner J, et al. Surgery Illustrated - surgical atlas sacral neuromodulation. *BJU Int*, 2012. 110(1): p. 146-59.
<http://www.ncbi.nlm.nih.gov/pubmed/22691023>
293. Kessler TM, et al. Sacral neuromodulation for neurogenic lower urinary tract dysfunction: systematic review and meta-analysis. *Eur Urol*, 2010. 58(6): p. 865-74.
<http://www.ncbi.nlm.nih.gov/pubmed/20934242>
294. Zhang YH, et al. Enveloping the bladder with displacement of flap of the rectus abdominis muscle for the treatment of neurogenic bladder. *J Urol*, 1990. 144(5): p. 1194-5.
<http://www.ncbi.nlm.nih.gov/pubmed/2146404>
295. Stenzl A, et al. Restoration of voluntary emptying of the bladder by transplantation of innervated free skeletal muscle. *Lancet*, 1998. 351(9114): p. 1483-5.
<http://www.ncbi.nlm.nih.gov/pubmed/9605805>
296. Gakis G, et al. Functional detrusor myoplasty for bladder acontractility: long-term results. *J Urol*, 2011. 185(2): p. 593-9.
<http://www.ncbi.nlm.nih.gov/pubmed/21168866>
297. Ninkovic M, et al. The latissimus dorsi detrusor myoplasty for functional treatment of bladder acontractility. *Clin Plast Surg*, 2012. 39(4): p. 507-12.
<http://www.ncbi.nlm.nih.gov/pubmed/23036300>
298. Braren V, et al. Laparoscopic bladder autoaugmentation in children. *Urol Clin North Am*, 1998. 25(3): p. 533-40.
<http://www.ncbi.nlm.nih.gov/pubmed/9728222>
299. Cartwright PC, et al. Bladder autoaugmentation: early clinical experience. *J Urol*, 1989. 142(2 Pt 2): p. 505-8; discussion 520-1.
<http://www.ncbi.nlm.nih.gov/pubmed/2746767>
300. Duel BP, et al. Alternative techniques for augmentation cystoplasty. *J Urol*, 1998. 159(3): p. 998-1005.
<http://www.ncbi.nlm.nih.gov/pubmed/9474216>
301. Poppas DP, et al. Laparoscopic laser assisted auto-augmentation of the pediatric neurogenic bladder: early experience with urodynamic followup. *J Urol*, 1996. 155(3): p. 1057-60.
<http://www.ncbi.nlm.nih.gov/pubmed/8583564>
302. Snow BW, et al. Bladder autoaugmentation. *Urol Clin North Am*, 1996. 23(2): p. 323-31.
<http://www.ncbi.nlm.nih.gov/pubmed/8659030>

303. Stohrer M, et al. Bladder auto-augmentation--an alternative for enterocystoplasty: preliminary results. *Neurourol Urodyn*, 1995. 14(1): p. 11-23. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/7742844>
304. Stohrer M, et al. Bladder autoaugmentation in adult patients with neurogenic voiding dysfunction. *Spinal Cord*, 1997. 35(7): p. 456-62.
<http://www.ncbi.nlm.nih.gov/pubmed/9232751>
305. Vainrib M, et al. Differences in urodynamic study variables in adult patients with neurogenic bladder and myelomeningocele before and after augmentation enterocystoplasty. *Neurourol Urodyn*, 2013. 32(3): p. 250-3.
<http://www.ncbi.nlm.nih.gov/pubmed/22965686>
306. Gough DC. Enterocystoplasty. *BJU Int*, 2001. 88(7): p. 739-43.
<http://www.ncbi.nlm.nih.gov/pubmed/11890246>
307. Greenwell TJ, et al. Augmentation cystoplasty. *BJU Int*, 2001. 88(6): p. 511-25.
<http://www.ncbi.nlm.nih.gov/pubmed/11678743>
308. Vajda P, et al. Histological findings after colocytoplasty and gastrocystoplasty. *J Urol*, 2002. 168(2): p. 698-701; discussion 701.
<http://www.ncbi.nlm.nih.gov/pubmed/12131353>
309. Chapple CR, et al. Surgery for detrusor overactivity. *World J Urol*, 1998. 16(4): p. 268-73.
<http://www.ncbi.nlm.nih.gov/pubmed/9775426>
310. Comer MT, et al. Reconstruction of the urinary bladder by auto-augmentation, enterocystoplasty, and composite enterocystoplasty. *Adv Exp Med Biol*, 1999. 462: p. 43-7. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/10599412>
311. Cranidis A, et al. Bladder augmentation. *Int Urogynecol J Pelvic Floor Dysfunct*, 2000. 11(1): p. 33-40.
<http://www.ncbi.nlm.nih.gov/pubmed/10738932>
312. Leng WW, et al. Enterocystoplasty or detrusor myectomy? Comparison of indications and outcomes for bladder augmentation. *J Urol*, 1999. 161(3): p. 758-63.
<http://www.ncbi.nlm.nih.gov/pubmed/10022679>
313. Niknejad KG, et al. Bladder augmentation techniques in women. *Int Urogynecol J Pelvic Floor Dysfunct*, 2000. 11(3): p. 156-69.
<http://www.ncbi.nlm.nih.gov/pubmed/11484743>
314. Oge O, et al. Urothelium-preserving augmentation cystoplasty covered with a peritoneal flap. *BJU Int*, 2000. 85(7): p. 802-5.
<http://www.ncbi.nlm.nih.gov/pubmed/10792156>
315. Siracusano S, et al. Laparoscopic bladder auto-augmentation in an incomplete traumatic spinal cord injury. *Spinal Cord*, 2000. 38(1): p. 59-61.
<http://www.ncbi.nlm.nih.gov/pubmed/10762200>
316. Westney OL, et al. Surgical procedures for the treatment of urge incontinence. *Tech Urol*, 2001. 7(2): p. 126-32.
<http://www.ncbi.nlm.nih.gov/pubmed/11383990>
317. Zhang F, et al. Sigmoidocolocystoplasty with ureteral reimplantation for treatment of neurogenic bladder. *Urology*, 2012. 80(2): p. 440-5.
<http://www.ncbi.nlm.nih.gov/pubmed/22857763>
318. Rubenwolf PC, et al. 15 years of continent urinary diversion and enterocystoplasty in children and adolescents: the Wurzburg experience. *BJU Int*, 2010. 105(5): p. 698-705.
<http://www.ncbi.nlm.nih.gov/pubmed/19832724>
319. Stein R, et al. Bladder augmentation and urinary diversion in patients with neurogenic bladder: non-surgical considerations. *J Pediatr Urol*, 2012. 8(2): p. 145-52.
<http://www.ncbi.nlm.nih.gov/pubmed/21493159>
320. O'Donnell WF. Urological management in the patient with acute spinal cord injury. *Crit Care Clin*, 1987. 3(3): p. 599-617.
<http://www.ncbi.nlm.nih.gov/pubmed/3332216>
321. Bennett JK, et al. Continent diversion and bladder augmentation in spinal cord-injured patients. *Semin Urol*, 1992. 10(2): p. 121-32. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/1636071>
322. Duckett JW, et al. Appendicovesicostomy (and variations) in bladder reconstruction. *J Urol*, 1993. 149(3): p. 567-9.
<http://www.ncbi.nlm.nih.gov/pubmed/8437267>

323. Kajbafzadeh AM, et al. Simultaneous Malone antegrade continent enema and Mitrofanoff principle using the divided appendix: report of a new technique for prevention of stoma complications. *J Urol*, 2001. 165(6 Pt 2): p. 2404-9.
<http://www.ncbi.nlm.nih.gov/pubmed/11371987>
324. Kawai K, et al. Tissue-engineered artificial urothelium. *World J Surg*, 2000. 24(10): p. 1160-2.
<http://www.ncbi.nlm.nih.gov/pubmed/11071451>
325. Liard A, et al. The Mitrofanoff procedure: 20 years later. *J Urol*, 2001. 165(6 Pt 2): p. 2394-8.
<http://www.ncbi.nlm.nih.gov/pubmed/11371985>
326. Moreno JG, et al. Improved quality of life and sexuality with continent urinary diversion in quadriplegic women with umbilical stoma. *Arch Phys Med Rehabil*, 1995. 76(8): p. 758-62.
<http://www.ncbi.nlm.nih.gov/pubmed/7632132>
327. Sekar P, et al. Comparison of long-term renal function after spinal cord injury using different urinary management methods. *Arch Phys Med Rehabil*, 1997. 78(9): p. 992-7.
<http://www.ncbi.nlm.nih.gov/pubmed/9305274>
328. Stein R, et al. Urinary diversion and orthotopic bladder substitution in children and young adults with neurogenic bladder: a safe option for treatment? *J Urol*, 2000. 163(2): p. 568-73.
<http://www.ncbi.nlm.nih.gov/pubmed/10647686>
329. Sylora JA, et al. Intermittent self-catheterization by quadriplegic patients via a catheterizable Mitrofanoff channel. *J Urol*, 1997. 157(1): p. 48-50.
<http://www.ncbi.nlm.nih.gov/pubmed/8976213>
330. Van Savage JG, et al. Transverse retubularized sigmoidovesicostomy continent urinary diversion to the umbilicus. *J Urol*, 2001. 166(2): p. 644-7.
<http://www.ncbi.nlm.nih.gov/pubmed/11458110>
331. Guillotreau J, et al. Prospective study of the impact on quality of life of cystectomy with ileal conduit urinary diversion for neurogenic bladder dysfunction. *Neurourol Urodyn*, 2011. 30(8): p. 1503-6.
<http://www.ncbi.nlm.nih.gov/pubmed/21674595>
332. Legrand G, et al. Functional outcomes after management of end-stage neurological bladder dysfunction with ileal conduit in a multiple sclerosis population: a monocentric experience. *Urology*, 2011. 78(4): p. 937-41.
<http://www.ncbi.nlm.nih.gov/pubmed/21820707>
333. Massaro PA, et al. Retubularization of the ileocystoplasty patch for conversion into an ileal conduit. *Can Urol Assoc J*, 2013. 7(7-8): p. E462-6.
<http://www.ncbi.nlm.nih.gov/pubmed/23914260>
334. Peterson AC, et al. Urinary diversion in patients with spinal cord injury in the United States. *Urology*, 2012. 80(6): p. 1247-51.
<http://www.ncbi.nlm.nih.gov/pubmed/23206770>
335. Vanni AJ, et al. Ileovesicostomy for the neurogenic bladder patient: outcome and cost comparison of open and robotic assisted techniques. *Urology*, 2011. 77(6): p. 1375-80.
<http://www.ncbi.nlm.nih.gov/pubmed/21146864>
336. Wiener JS, et al. Bladder augmentation versus urinary diversion in patients with spina bifida in the United States. *J Urol*, 2011. 186(1): p. 161-5.
<http://www.ncbi.nlm.nih.gov/pubmed/21575969>
337. Shapiro SR, et al. Fate of 90 children with ileal conduit urinary diversion a decade later: analysis of complications, pyelography, renal function and bacteriology. *J Urol*, 1975. 114(2): p. 289-95.
<http://www.ncbi.nlm.nih.gov/pubmed/1159925>
338. Atan A, et al. Advantages and risks of ileovesicostomy for the management of neuropathic bladder. *Urology*, 1999. 54(4): p. 636-40.
<http://www.ncbi.nlm.nih.gov/pubmed/10510920>
339. Cass AS, et al. A 22-year followup of ileal conduits in children with a neurogenic bladder. *J Urol*, 1984. 132(3): p. 529-31.
<http://www.ncbi.nlm.nih.gov/pubmed/6471190>
340. Hald T, et al. Vesicostomy--an alternative urine diversion operation. Long term results. *Scand J Urol Nephrol*, 1978. 12(3): p. 227-31.
<http://www.ncbi.nlm.nih.gov/pubmed/725543>
341. Schwartz SL, et al. Incontinent ileo-vesicostomy urinary diversion in the treatment of lower urinary tract dysfunction. *J Urol*, 1994. 152(1): p. 99-102.
<http://www.ncbi.nlm.nih.gov/pubmed/8201699>
342. Herschorn S, et al. Urinary undiversion in adults with myelodysplasia: long-term follow-up. *J Urol*, 1994. 152(2 Pt 1): p. 329-33.
<http://www.ncbi.nlm.nih.gov/pubmed/8015064>

343. [No authors listed]. The prevention and management of urinary tract infections among people with spinal cord injuries. National Institute on Disability and Rehabilitation Research Consensus Statement. January 27-29, 1992. *J Am Paraplegia Soc*, 1992. 15(3): p. 194-204.
<http://www.ncbi.nlm.nih.gov/pubmed/1500945>
344. Goetz LL, et al. International Spinal Cord Injury Urinary Tract Infection Basic Data Set. *Spinal Cord*, 2013. 51(9): p. 700-4.
<http://www.ncbi.nlm.nih.gov/pubmed/23896666>
345. Pannek J. Treatment of urinary tract infection in persons with spinal cord injury: guidelines, evidence, and clinical practice. A questionnaire-based survey and review of the literature. *J Spinal Cord Med*, 2011. 34(1): p. 11-5.
<http://www.ncbi.nlm.nih.gov/pubmed/21528621>
346. Deville WL, et al. The urine dipstick test useful to rule out infections. A meta-analysis of the accuracy. *BMC Urol*, 2004. 4: p. 4.
<http://www.ncbi.nlm.nih.gov/pubmed/15175113>
347. Hoffman JM, et al. Nitrite and leukocyte dipstick testing for urinary tract infection in individuals with spinal cord injury. *J Spinal Cord Med*, 2004. 27(2): p. 128-32.
<http://www.ncbi.nlm.nih.gov/pubmed/15162883>
348. Biering-Sorensen F, et al. Urinary tract infections in patients with spinal cord lesions: treatment and prevention. *Drugs*, 2001. 61(9): p. 1275-87.
<http://www.ncbi.nlm.nih.gov/pubmed/11511022>
349. Everaert K, et al. Urinary tract infections in spinal cord injury: prevention and treatment guidelines. *Acta Clin Belg*, 2009. 64(4): p. 335-40.
<http://www.ncbi.nlm.nih.gov/pubmed/19810421>
350. D'Hondt F, et al. Urinary tract infections in patients with spinal cord injuries. *Curr Infect Dis Rep*, 2011. 13(6): p. 544-51.
<http://www.ncbi.nlm.nih.gov/pubmed/21853416>
351. Jia C, et al. Detrusor botulinum toxin A injection significantly decreased urinary tract infection in patients with traumatic spinal cord injury. *Spinal Cord*, 2013. 51(6): p. 487-90.
<http://www.ncbi.nlm.nih.gov/pubmed/23357928>
352. Cardenas DD, et al. Hydrophilic catheters versus noncoated catheters for reducing the incidence of urinary tract infections: a randomized controlled trial. *Arch Phys Med Rehabil*, 2009. 90(10): p. 1668-71.
<http://www.ncbi.nlm.nih.gov/pubmed/19801054>
353. Waites KB, et al. Evaluation of 3 methods of bladder irrigation to treat bacteriuria in persons with neurogenic bladder. *J Spinal Cord Med*, 2006. 29(3): p. 217-26.
<http://www.ncbi.nlm.nih.gov/pubmed/16859225>
354. Lee BB, et al. Spinal-injured neuropathic bladder antisepsis (SINBA) trial. *Spinal Cord*, 2007. 45(8): p. 542-50.
<http://www.ncbi.nlm.nih.gov/pubmed/17043681>
355. Lee BS, et al. Methenamine hippurate for preventing urinary tract infections. *Cochrane Database Syst Rev*, 2012. 10: p. Cd003265.
<http://www.ncbi.nlm.nih.gov/pubmed/23076896>
356. Günther M, et al. Harnwegsinfektprophylaxe. Urinansäuerung mittels L-Methionin bei neurogener Blasenfunktionsstörung. *Urologe B*, 2002. 42: p. 218-220. [No abstract available] [Article in German]
357. Hachen HJ. Oral immunotherapy in paraplegic patients with chronic urinary tract infections: a double-blind, placebo-controlled trial. *J Urol*, 1990. 143(4): p. 759-62; discussion 762-3.
<http://www.ncbi.nlm.nih.gov/pubmed/2179584>
358. Salomon J, et al. Prevention of urinary tract infection in spinal cord-injured patients: safety and efficacy of a weekly oral cyclic antibiotic (WOCA) programme with a 2 year follow-up—an observational prospective study. *J Antimicrob Chemother*, 2006. 57(4): p. 784-8.
<http://www.ncbi.nlm.nih.gov/pubmed/16473921>
359. Darouiche RO, et al. Multicenter randomized controlled trial of bacterial interference for prevention of urinary tract infection in patients with neurogenic bladder. *Urology*, 2011. 78(2): p. 341-6.
<http://www.ncbi.nlm.nih.gov/pubmed/21683991>
360. Pannek J, et al. Usefulness of classical homoeopathy for the prevention of urinary tract infections in patients with neurogenic bladder dysfunction: a case series. *Indian J Res Homoeopathy*, 2014. 8: p. 31-36.
361. Rees PM, et al. Sexual function in men and women with neurological disorders. *Lancet*, 2007. 369(9560): p. 512-25.
<http://www.ncbi.nlm.nih.gov/pubmed/17292771>

362. Jungwirth A, et al. EAU Guidelines on Male Infertility. In: EAU Guidelines, edition presented at the 29th EAU Annual Congress, Stockholm, 2014. ISBN 978-90-79754-65-6.
<http://uroweb.org/guideline/male-infertility/>
363. Wespes E, et al. EAU guidelines on erectile dysfunction and premature ejaculation. In: EAU Guidelines, edition presented at the 29th EAU Annual Congress, Stockholm, 2014. ISBN 978-90-79754-65-6.
<http://uroweb.org/guideline/male-sexual-dysfunction/>
364. Annon JS. PLISSIT Therapy in Handbook of Innovative Psychotherapies. R. Corsini, Editor. 1981, Wiley & Sons: New York. p. 626-39.
365. Yuan J, et al. Comparative effectiveness and safety of oral phosphodiesterase type 5 inhibitors for erectile dysfunction: a systematic review and network meta-analysis. *Eur Urol*, 2013. 63(5): p. 902-12.
<http://www.ncbi.nlm.nih.gov/pubmed/23395275>
366. Lombardi G, et al. Ten years of phosphodiesterase type 5 inhibitors in spinal cord injured patients. *J Sex Med*, 2009. 6(5): p. 1248-58.
<http://www.ncbi.nlm.nih.gov/pubmed/19210710>
367. Lombardi G, et al. Treating erectile dysfunction and central neurological diseases with oral phosphodiesterase type 5 inhibitors. Review of the literature. *J Sex Med*, 2012. 9(4): p. 970-85.
<http://www.ncbi.nlm.nih.gov/pubmed/22304626>
368. Giuliano F, et al. Efficacy and safety of vardenafil in men with erectile dysfunction caused by spinal cord injury. *Neurology*, 2006. 66(2): p. 210-6.
<http://www.ncbi.nlm.nih.gov/pubmed/16434656>
369. Lombardi G, et al. Efficacy and safety of medium and long-term tadalafil use in spinal cord patients with erectile dysfunction. *J Sex Med*, 2009. 6(2): p. 535-43.
<http://www.ncbi.nlm.nih.gov/pubmed/19138363>
370. Rizio N, et al. Efficacy and satisfaction rates of oral PDE5is in the treatment of erectile dysfunction secondary to spinal cord injury: a review of literature. *J Spinal Cord Med*, 2012. 35(4): p. 219-28.
<http://www.ncbi.nlm.nih.gov/pubmed/22925748>
371. Soler JM, et al. Phosphodiesterase inhibitors in the treatment of erectile dysfunction in spinal cord injured men. *Spinal Cord*, 2007. 45(2): p. 169-73.
<http://www.ncbi.nlm.nih.gov/pubmed/16801935>
372. Fowler CJ, et al. A double blind, randomised study of sildenafil citrate for erectile dysfunction in men with multiple sclerosis. *J Neurol Neurosurg Psychiatry*, 2005. 76(5): p. 700-5.
<http://www.ncbi.nlm.nih.gov/pubmed/15834030>
373. Lombardi G, et al. Efficacy and safety of tadalafil for erectile dysfunction in patients with multiple sclerosis. *J Sex Med*, 2010. 7(6): p. 2192-200.
<http://www.ncbi.nlm.nih.gov/pubmed/20384939>
374. Xiao Y, et al. Sildenafil citrate for erectile dysfunction in patients with multiple sclerosis. *Cochrane Database Syst Rev*, 2012. 4: p. Cd009427.
<http://www.ncbi.nlm.nih.gov/pubmed/22513975>
375. Hussain IF, et al. Treatment of erectile dysfunction with sildenafil citrate (Viagra) in parkinsonism due to Parkinson's disease or multiple system atrophy with observations on orthostatic hypotension. *J Neurol Neurosurg Psychiatry*, 2001. 71(3): p. 371-4.
<http://www.ncbi.nlm.nih.gov/pubmed/11511713>
376. Raffaele R, et al. Efficacy and safety of fixed-dose oral sildenafil in the treatment of sexual dysfunction in depressed patients with idiopathic Parkinson's disease. *Eur Urol*, 2002. 41(4): p. 382- 6.
<http://www.ncbi.nlm.nih.gov/pubmed/12074807>
377. Safarinejad MR. Oral sildenafil in the treatment of erectile dysfunction in diabetic men: a randomized double-blind and placebo-controlled study. *J Diabetes Complications*, 2004. 18(4): p. 205-10.
<http://www.ncbi.nlm.nih.gov/pubmed/15207837>
378. Boulton AJ, et al. Sildenafil citrate for the treatment of erectile dysfunction in men with Type II diabetes mellitus. *Diabetologia*, 2001. 44(10): p. 1296-301.
<http://www.ncbi.nlm.nih.gov/pubmed/11692178>
379. Palmer JS, et al. Erectile dysfunction in patients with spina bifida is a treatable condition. *J Urol*, 2000. 164(3 Pt 2): p. 958-61.
<http://www.ncbi.nlm.nih.gov/pubmed/10958716>

380. Rendell MS, et al. Sildenafil for treatment of erectile dysfunction in men with diabetes: a randomized controlled trial. Sildenafil Diabetes Study Group. *Jama*, 1999. 281(5): p. 421-6.
<http://www.ncbi.nlm.nih.gov/pubmed/9952201>
381. Kaiho Y, et al. Optimization of sexual function outcome after radical prostatectomy using phosphodiesterase type 5 inhibitors. *Int J Urol*, 2013. 20(3): p. 285-9.
<http://www.ncbi.nlm.nih.gov/pubmed/23311962>
382. Chancellor MB, et al. Prospective comparison of topical minoxidil to vacuum constriction device and intracorporeal papaverine injection in treatment of erectile dysfunction due to spinal cord injury. *Urology*, 1994. 43(3): p. 365-9.
<http://www.ncbi.nlm.nih.gov/pubmed/8134992>
383. Cookson MS, et al. Long-term results with vacuum constriction device. *J Urol*, 1993. 149(2): p. 290-4.
<http://www.ncbi.nlm.nih.gov/pubmed/8426404>
384. Denil J, et al. Vacuum erection device in spinal cord injured men: patient and partner satisfaction. *Arch Phys Med Rehabil*, 1996. 77(8): p. 750-3.
<http://www.ncbi.nlm.nih.gov/pubmed/8702367>
385. Levine LA. External devices for treatment of erectile dysfunction. *Endocrine*, 2004. 23(2-3): p. 157-60.
<http://www.ncbi.nlm.nih.gov/pubmed/15146095>
386. Levine LA, et al. Vacuum constriction and external erection devices in erectile dysfunction. *Urol Clin North Am*, 2001. 28(2): p. 335-41, ix-x.
<http://www.ncbi.nlm.nih.gov/pubmed/11402585>
387. Bella AJ, et al. Intracavernous pharmacotherapy for erectile dysfunction. *Endocrine*, 2004. 23(2-3): p. 149-55.
<http://www.ncbi.nlm.nih.gov/pubmed/15146094>
388. Bodner DR, et al. The application of intracavernous injection of vasoactive medications for erection in men with spinal cord injury. *J Urol*, 1987. 138(2): p. 310-1.
<http://www.ncbi.nlm.nih.gov/pubmed/3599245>
389. Dinsmore WW, et al. Treating men with predominantly nonpsychogenic erectile dysfunction with intracavernosal vasoactive intestinal polypeptide and phentolamine mesylate in a novel auto-injector system: a multicentre double-blind placebo-controlled study. *BJU Int*, 1999. 83(3): p. 274-9.
<http://www.ncbi.nlm.nih.gov/pubmed/10233493>
390. Hirsch IH, et al. Use of intracavernous injection of prostaglandin E1 for neuropathic erectile dysfunction. *Paraplegia*, 1994. 32(10): p. 661-4.
<http://www.ncbi.nlm.nih.gov/pubmed/7831071>
391. Kapoor VK, et al. Intracavernous papaverine for impotence in spinal cord injured patients. *Paraplegia*, 1993. 31(10): p. 675-7.
<http://www.ncbi.nlm.nih.gov/pubmed/8259331>
392. Vidal J, et al. Intracavernous pharmacotherapy for management of erectile dysfunction in multiple sclerosis patients. *Rev Neurol*, 1995. 23(120): p. 269-71.
<http://www.ncbi.nlm.nih.gov/pubmed/7497173>
393. Bodner DR, et al. Intraurethral alprostadil for treatment of erectile dysfunction in patients with spinal cord injury. *Urology*, 1999. 53(1): p. 199-202.
<http://www.ncbi.nlm.nih.gov/pubmed/9886612>
394. Gross AJ, et al. Penile prostheses in paraplegic men. *Br J Urol*, 1996. 78(2): p. 262-4.
<http://www.ncbi.nlm.nih.gov/pubmed/8813925>
395. Kimoto Y, et al. Penile prostheses for the management of the neuropathic bladder and sexual dysfunction in spinal cord injury patients: long term follow up. *Paraplegia*, 1994. 32(5): p. 336-9.
<http://www.ncbi.nlm.nih.gov/pubmed/8058351>
396. Zermann DH, et al. Penile prosthetic surgery in neurologically impaired patients: long-term followup. *J Urol*, 2006. 175(3 Pt 1): p. 1041-4; discussion 1044.
<http://www.ncbi.nlm.nih.gov/pubmed/16469612>
397. Fode M, et al. Male sexual dysfunction and infertility associated with neurological disorders. *Asian J Androl*, 2012. 14(1): p. 61-8.
<http://www.ncbi.nlm.nih.gov/pubmed/22138899>
398. Lim TC, et al. A simple technique to prevent retrograde ejaculation during assisted ejaculation. *Paraplegia*, 1994. 32(3): p. 142-9.
<http://www.ncbi.nlm.nih.gov/pubmed/8008416>

399. Arafa MM, et al. Prostatic massage: a simple method of semen retrieval in men with spinal cord injury. *Int J Androl*, 2007. 30(3): p. 170-3.
<http://www.ncbi.nlm.nih.gov/pubmed/17298549>
400. DeForge D, et al. Fertility following spinal cord injury: a systematic review. *Spinal Cord*, 2005. 43(12): p. 693-703.
<http://www.ncbi.nlm.nih.gov/pubmed/15951744>
401. Kolettis PN, et al. Fertility outcomes after electroejaculation in men with spinal cord injury. *Fertil Steril*, 2002. 78(2): p. 429-31.
<http://www.ncbi.nlm.nih.gov/pubmed/12137889>
402. Ohl DA, et al. Electroejaculation versus vibratory stimulation in spinal cord injured men: sperm quality and patient preference. *J Urol*, 1997. 157(6): p. 2147-9.
<http://www.ncbi.nlm.nih.gov/pubmed/9146603>
403. Rutkowski SB, et al. A comprehensive approach to the management of male infertility following spinal cord injury. *Spinal Cord*, 1999. 37(7): p. 508-14.
<http://www.ncbi.nlm.nih.gov/pubmed/10438118>
404. Beretta G, et al. Reproductive aspects in spinal cord injured males. *Paraplegia*, 1989. 27(2): p. 113-8.
<http://www.ncbi.nlm.nih.gov/pubmed/2717193>
405. Brackett NL, et al. Application of 2 vibrators salvages ejaculatory failures to 1 vibrator during penile vibratory stimulation in men with spinal cord injuries. *J Urol*, 2007. 177(2): p. 660-3.
<http://www.ncbi.nlm.nih.gov/pubmed/17222653>
406. Sonksen J, et al. Ejaculation induced by penile vibratory stimulation in men with spinal cord injuries. The importance of the vibratory amplitude. *Paraplegia*, 1994. 32(10): p. 651-60.
<http://www.ncbi.nlm.nih.gov/pubmed/7831070>
407. Claydon VE, et al. Cardiovascular responses to vibrostimulation for sperm retrieval in men with spinal cord injury. *J Spinal Cord Med*, 2006. 29(3): p. 207-16.
<http://www.ncbi.nlm.nih.gov/pubmed/16859224>
408. Eklund MB, et al. Incidence of autonomic dysreflexia and silent autonomic dysreflexia in men with spinal cord injury undergoing sperm retrieval: implications for clinical practice. *J Spinal Cord Med*, 2008. 31(1): p. 33-9.
<http://www.ncbi.nlm.nih.gov/pubmed/18533409>
409. Brackett NL, et al. Treatment of infertility in men with spinal cord injury. *Nat Rev Urol*, 2010. 7(3): p. 162-72.
<http://www.ncbi.nlm.nih.gov/pubmed/20157304>
410. Dimitriadis F, et al. Erectile function and male reproduction in men with spinal cord injury: a review. *Andrologia*, 2010. 42(3): p. 139-65.
<http://www.ncbi.nlm.nih.gov/pubmed/20500744>
411. Schatte EC, et al. Treatment of infertility due to anejaculation in the male with electroejaculation and intracytoplasmic sperm injection. *J Urol*, 2000. 163(6): p. 1717-20.
<http://www.ncbi.nlm.nih.gov/pubmed/10799167>
412. Shieh JY, et al. A protocol of electroejaculation and systematic assisted reproductive technology achieved high efficiency and efficacy for pregnancy for anejaculatory men with spinal cord injury. *Arch Phys Med Rehabil*, 2003. 84(4): p. 535-40.
<http://www.ncbi.nlm.nih.gov/pubmed/12690592>
413. Taylor Z, et al. Contribution of the assisted reproductive technologies to fertility in males suffering spinal cord injury. *Aust N Z J Obstet Gynaecol*, 1999. 39(1): p. 84-7.
<http://www.ncbi.nlm.nih.gov/pubmed/10099757>
414. Brackett NL, et al. Semen quality of spinal cord injured men is better when obtained by vibratory stimulation versus electroejaculation. *J Urol*, 1997. 157(1): p. 151-7.
<http://www.ncbi.nlm.nih.gov/pubmed/8976239>
415. Brackett NL, et al. Semen retrieval in men with spinal cord injury is improved by interrupting current delivery during electroejaculation. *J Urol*, 2002. 167(1): p. 201-3.
<http://www.ncbi.nlm.nih.gov/pubmed/11743305>
416. Rutkowski SB, et al. The influence of bladder management on fertility in spinal cord injured males. *Paraplegia*, 1995. 33(5): p. 263-6.
<http://www.ncbi.nlm.nih.gov/pubmed/7630651>
417. Brackett NL, et al. Seminal plasma of spinal cord injured men inhibits sperm motility of normal men. *J Urol*, 1996. 155(5): p. 1632-5.
<http://www.ncbi.nlm.nih.gov/pubmed/8627840>

418. Ferreiro-Velasco ME, et al. Sexual issues in a sample of women with spinal cord injury. *Spinal Cord*, 2005. 43(1): p. 51-5.
<http://www.ncbi.nlm.nih.gov/pubmed/15303115>
419. Kreuter M, et al. Sexuality and sexual life in women with spinal cord injury: a controlled study. *J Rehabil Med*, 2008. 40(1): p. 61-9.
<http://www.ncbi.nlm.nih.gov/pubmed/18176739>
420. Kreuter M, et al. Sexual adjustment and quality of relationship in spinal paraplegia: a controlled study. *Arch Phys Med Rehabil*, 1996. 77(6): p. 541-8.
<http://www.ncbi.nlm.nih.gov/pubmed/8831469>
421. Kessler TM, et al. Sexual dysfunction in multiple sclerosis. *Expert Rev Neurother*, 2009. 9(3): p. 341-50.
<http://www.ncbi.nlm.nih.gov/pubmed/19271943>
422. Lew-Starowicz M, et al. Prevalence of sexual dysfunctions among women with multiple sclerosis. *Sex Disabil*, 2013. 31(2): p. 141-153.
<http://www.ncbi.nlm.nih.gov/pubmed/23704801>
423. Harrison J, et al. Factors associated with sexual functioning in women following spinal cord injury. *Paraplegia*, 1995. 33(12): p. 687-92.
<http://www.ncbi.nlm.nih.gov/pubmed/8927405>
424. Westgren N, et al. Sexuality in women with traumatic spinal cord injury. *Acta Obstet Gynecol Scand*, 1997. 76(10): p. 977-83.
<http://www.ncbi.nlm.nih.gov/pubmed/9435740>
425. Reitz A, et al. Impact of spinal cord injury on sexual health and quality of life. *Int J Impot Res*, 2004. 16(2): p. 167-74.
<http://www.ncbi.nlm.nih.gov/pubmed/14973522>
426. Sipski ML, et al. Sildenafil effects on sexual and cardiovascular responses in women with spinal cord injury. *Urology*, 2000. 55(6): p. 812-5.
<http://www.ncbi.nlm.nih.gov/pubmed/10840082>
427. Forsythe E, et al. Sexual rehabilitation of women with a spinal cord injury. *Spinal Cord*, 2006. 44(4): p. 234-41.
<http://www.ncbi.nlm.nih.gov/pubmed/16172622>
428. Fruhauf S, et al. Efficacy of psychological interventions for sexual dysfunction: a systematic review and meta-analysis. *Arch Sex Behav*, 2013. 42(6): p. 915-33.
<http://www.ncbi.nlm.nih.gov/pubmed/23559141>
429. Alexander M, et al. Spinal cord injuries and orgasm: a review. *J Sex Marital Ther*, 2008. 34(4): p. 308-24.
<http://www.ncbi.nlm.nih.gov/pubmed/18576233>
430. Sipski ML, et al. Sexual arousal and orgasm in women: effects of spinal cord injury. *Ann Neurol*, 2001. 49(1): p. 35-44.
<http://www.ncbi.nlm.nih.gov/pubmed/11198294>
431. Sipski ML, et al. Physiologic parameters associated with sexual arousal in women with incomplete spinal cord injuries. *Arch Phys Med Rehabil*, 1997. 78(3): p. 305-13.
<http://www.ncbi.nlm.nih.gov/pubmed/9084355>
432. McAlonan S. Improving sexual rehabilitation services: the patient's perspective. *Am J Occup Ther*, 1996. 50(10): p. 826-34.
<http://www.ncbi.nlm.nih.gov/pubmed/8947375>
433. Schopp LH, et al. Impact of comprehensive gynecologic services on health maintenance behaviours among women with spinal cord injury. *Disabil Rehabil*, 2002. 24(17): p. 899-903.
<http://www.ncbi.nlm.nih.gov/pubmed/12519485>
434. Sukumaran SC, et al. Polytherapy increases the risk of infertility in women with epilepsy. *Neurology*, 2010. 75(15): p. 1351-5.
<http://www.ncbi.nlm.nih.gov/pubmed/20938026>
435. Axel SJ. Spinal cord injured women's concerns: menstruation and pregnancy. *Rehabil Nurs*, 1982. 7(5): p. 10-5. [No abstract available]
<http://www.ncbi.nlm.nih.gov/pubmed/6921826>
436. Jackson AB, et al. A multicenter study of women's self-reported reproductive health after spinal cord injury. *Arch Phys Med Rehabil*, 1999. 80(11): p. 1420-8.
<http://www.ncbi.nlm.nih.gov/pubmed/10569436>
437. Baker ER, et al. Pregnancy in spinal cord injured women. *Arch Phys Med Rehabil*, 1996. 77(5): p. 501-7.
<http://www.ncbi.nlm.nih.gov/pubmed/8629929>

438. Baker ER, et al. Risks associated with pregnancy in spinal cord-injured women. *Obstet Gynecol*, 1992. 80(3 Pt 1): p. 425-8.
<http://www.ncbi.nlm.nih.gov/pubmed/1495699>
439. Cross LL, et al. Pregnancy, labor and delivery post spinal cord injury. *Paraplegia*, 1992. 30(12): p. 890-902.
<http://www.ncbi.nlm.nih.gov/pubmed/1287543>
440. Hughes SJ, et al. Management of the pregnant woman with spinal cord injuries. *Br J Obstet Gynaecol*, 1991. 98(6): p. 513-8.
<http://www.ncbi.nlm.nih.gov/pubmed/1873238>
441. Dannels A, et al. The perimenopause experience for women with spinal cord injuries. *SCI Nurs*, 2004. 21(1): p. 9-13.
<http://www.ncbi.nlm.nih.gov/pubmed/15176344>
442. Burns AS, et al. The management of neurogenic bladder and sexual dysfunction after spinal cord injury. *Spine (Phila Pa 1976)*, 2001. 26(24 Suppl): p. S129-36.
<http://www.ncbi.nlm.nih.gov/pubmed/11805620>
443. Cardenas DD, et al. Lower urinary changes over time in suprasacral spinal cord injury. *Paraplegia*, 1995. 33(6): p. 326-9.
<http://www.ncbi.nlm.nih.gov/pubmed/7644258>
444. Chen Y, et al. Current trend and risk factors for kidney stones in persons with spinal cord injury: a longitudinal study. *Spinal Cord*, 2000. 38(6): p. 346-53.
<http://www.ncbi.nlm.nih.gov/pubmed/10889563>
445. Ciancio SJ, et al. Urodynamic pattern changes in multiple sclerosis. *Urology*, 2001. 57(2): p. 239-45.
<http://www.ncbi.nlm.nih.gov/pubmed/11182328>
446. Elliott DS, et al. Recent advances in the management of the neurogenic bladder. *Urology*, 2000. 56(6 Suppl 1): p. 76-81.
<http://www.ncbi.nlm.nih.gov/pubmed/11114567>
447. Lawrenson R, et al. Renal failure in patients with neurogenic lower urinary tract dysfunction. *Neuroepidemiology*, 2001. 20(2): p. 138-43.
<http://www.ncbi.nlm.nih.gov/pubmed/11359083>
448. Perakash I. Long-term urologic management of the patient with spinal cord injury. *Urol Clin North Am*, 1993. 20(3): p. 423-34.
<http://www.ncbi.nlm.nih.gov/pubmed/8351768>
449. Polackwich AS, et al. Long-term followup after endoscopic treatment of vesicoureteral reflux with dextranomer/hyaluronic acid copolymer in patients with neurogenic bladder. *J Urol*, 2012. 188(4 Suppl): p. 1511-5.
<http://www.ncbi.nlm.nih.gov/pubmed/22910250>
450. Rashid TM, et al. Multiple sclerosis and the neurogenic bladder. *Phys Med Rehabil Clin N Am*, 1998. 9(3): p. 615-29.
<http://www.ncbi.nlm.nih.gov/pubmed/9894113>
451. Stöhrer M. Alterations in the urinary tract after spinal cord injury-diagnosis, prevention and therapy of late sequelae. *World J Urol* 1990. 7: p. 205-11.
452. Waites KB, et al. Compliance with annual urologic evaluations and preservation of renal function in persons with spinal cord injury. *J Spinal Cord Med*, 1995. 18(4): p. 251-4.
<http://www.ncbi.nlm.nih.gov/pubmed/8591072>
453. Wiedemann A, et al. Which clinical risk factors determine a pathological urodynamic evaluation in patients with multiple sclerosis? an analysis of 100 prospective cases. *World J Urol*, 2013. 31(1): p. 229-33.
<http://www.ncbi.nlm.nih.gov/pubmed/22227822>

5. CONFLICT OF INTEREST

All members of the Neuro-Urology Guidelines Panel have provided disclosure statements on all relationships that they have that might be perceived to be a potential source of a conflict of interest. This information is publically accessible through the EAU website. This guidelines document was developed with the financial support of the EAU. No external sources of funding and support have been involved. The EAU is a non-profit organisation, and funding is limited to administrative assistance and travel and meeting expenses. No honoraria or other reimbursements have been provided.