

# Egyptian Clinical Practice Guidelines for Male Sexual Dysfunction

## Preliminary pages

### Copyright page

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## Acknowledgement

### **i** Guidelines Development Group (GDG) of Male Sexual Dysfunction committee.

1. Prof Abdel Rahman M. Zahran, MD Faculty of Medicine, Alexandria University
2. Prof AbdelNasser El Gamasy, Tanta University
3. Prof Ahmed Aly Morsy, Cairo University
4. Prof Ahmed El Taher, MD Faculty of Medicine, Assuit University
5. Prof Ahmed I. El-Sakka, MD (Chief) Faculty of Medicine, Suez Canal University
6. Prof Ahmed M. Al Adl, MD Faculty of Medicine, Benha university
7. Prof Emad A. Salem, MD Faculty of Medicine, Zagazig University
8. Prof Hisham Hammouda, Assiut University
9. Prof Khaled Mohyelden, MD Faculty of Medicine, Fayoum University
10. Prof Magdy S. El-Bahnasawy, MD Faculty of Medicine, Mansoura University
11. Prof Mamdouh M. El-Hawy MD Faculty of Medicine, Minia University
12. Prof Mohamed Ahmed Shalaby, Assiut University
13. Prof Mohamed Rafik El Halaby, Ain Shams University
14. Prof Mohamed Sherif Mourad, Ain Shams University (Chair of Panel)
15. Prof Mohammed Abdel-Rassoul, MD Faculty of Medicine, Cairo University
16. Prof Raouf M. Seyam King Faisal specialist hospital, Saudi

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## List of Abbreviations

<b>i</b>	<b>Abbreviation</b>	<b>Description</b>
	<b>AIPE</b>	<i>Arabic Index of Premature Ejaculation</i>
	<b>ART</b>	<i>Assisted reproductive technology</i>
	<b>AUA</b>	<i>American Urological Association</i>
	<b>BSSM</b>	<i>British Society for Sexual Medicine</i>
	<b>CCH</b>	<i>Collagenase Clostridium histolyticum</i>
	<b>CDU</b>	<i>Color duplex ultrasonography</i>
	<b>DE</b>	<i>Delayed Ejaculation</i>
	<b>DM</b>	<i>Diabetes mellitus</i>
	<b>EAU</b>	<i>European Association of Urology</i>
	<b>ED</b>	<i>Erectile dysfunction</i>
	<b>EDV</b>	<i>End diastolic velocity</i>
	<b>EE</b>	<i>Electo-ejaculation</i>
	<b>EMA</b>	<i>European Medicines Agency</i>
	<b>eNOS</b>	<i>Endothelial nitric oxide synthase</i>
	<b>ESWT</b>	<i>Extracorporeal Shockwave Therapy</i>
	<b>FDA</b>	<i>Food and drug administration</i>
	<b>FSH</b>	<i>Follicular Stimulating Hormone</i>
	<b>ICI</b>	<i>Intracavernous injection</i>
	<b>ICSI</b>	<i>Intracytoplasmic sperm injection</i>
	<b>IELT</b>	<i>Intravaginal ejaculation latency time</i>
	<b>IHD</b>	<i>Ischaemic heart disease</i>
	<b>IIEF</b>	<i>International Index of Erectile Function</i>
	<b>iNOS</b>	<i>Inducible nitric oxide synthase</i>
	<b>ISSM</b>	<i>International Society of Sexual Medicine</i>
	<b>IVF</b>	<i>In-vitro fertilization</i>
	<b>LH</b>	<i>Luteinizing Hormone</i>
	<b>LUTS</b>	<i>Lower urinary tract symptoms</i>
	<b>nNOS</b>	<i>Neuronal nitric oxide synthase</i>
	<b>NPT</b>	<i>Nocturnal penile tumescence</i>
	<b>PD</b>	<i>Peyronie's disease</i>
	<b>PDE5Is</b>	<i>Phosphodiesterase type 5 inhibitors</i>
	<b>PE</b>	<i>Premature ejaculation</i>
	<b>PEDT</b>	<i>Premature Ejaculation Diagnostic Tool</i>
	<b>PGE1</b>	<i>Prostaglandin E 1</i>
	<b>PP</b>	<i>Penile prosthesis</i>
	<b>PSA</b>	<i>Prostate-specific antigen</i>
	<b>PSV</b>	<i>Peak systolic velocity</i>
	<b>PVS</b>	<i>Penile vibratory stimulation</i>
	<b>QoL</b>	<i>Quality of life</i>

<b>RI</b>	<i>Resistive index</i>
<b>RP</b>	<i>Radical prostatectomy</i>
<b>SCD</b>	<i>Sickle cell disease</i>
<b>SCI</b>	<i>Spinal cord injury</i>
<b>SCs</b>	<i>Stem cells</i>
<b>SSRIs</b>	<i>Selective serotonin reuptake inhibitors</i>
<b>TA</b>	<i>Tunica albuginea</i>
<b>TGF-β1</b>	<i>Transforming growth factor beta 1</i>
<b>VED</b>	<i>Vacuum erection device</i>
<b>VOD</b>	<i>Veno-occlusive dysfunction</i>

## Glossary

- i** **1- Erectile Dysfunction (ED):** The persistent or recurrent inability to attain and maintain an erection sufficient to permit satisfactory sexual performance.
- 2- Premature Ejaculation:** Ejaculation that always or nearly always occurs prior to or within about one minute of vaginal penetration (lifelong PE) or a clinically significant and bothersome reduction in latency time, often to about three minutes or less (acquired PE).
- 3- Delayed Ejaculation:** Marked delay in ejaculation or marked infrequency or absence of ejaculation on almost all or all occasions (75-100% of the times) of partnered sexual activity without the individual desiring delay persisting for at least 6 months and causing significant distress to the individual.
- 4- Peyronie’s Disease (PD)** is a symptomatic disorder characterized by a constellation of penile symptoms and signs, such as penile pain, curvature, shortening, narrowing, hinge deformity, and palpable plaque with subsequent ED.
- 5- Priapism** is a persistent penile erection for more than four hours and not related to sexual stimulation or relieved by ejaculation. Priapism carries high risk of structural damage to the cavernosal tissue which may lead to permanent ED.
- 4- Anejaculation:** The complete absence of ejaculation either antegrade or retrograde. Caused by failed seminal emission from the seminal vesicles, prostate, and ejaculatory ducts into the urethra. In true anejaculation, there is normal orgasmic sensation and is always associated with central or peripheral nervous system dysfunction or with drugs.
- 5- Painful Ejaculation:** is a condition in which the patient may feel variable degrees of pain during or after ejaculation involving the penis, scrotum, and perineum.
- 6- Haemospermia:** is the presence of blood in the seminal fluid ejaculate. The condition causes anxiety and may indicate underlying pathology in many cases.

## Executive Summary

### **i** Scope of the guidelines

Sexual health-related issues are wide-ranging and of importance to the overall health and sense of well-being for couples and families, and to the social and economic development of communities and countries. Erectile dysfunction (ED) and disorders of ejaculation are frequent encounters in male sexual medicine in the Middle East with the association of different risk factors and medical comorbidities in Arab region countries. Pharmacological therapies have completely changed the diagnostic and therapeutic approach to ED. This article integrates recent international guidelines with local experience and highlights the apparent lack of congruency between available treatment and communication, cultural, and gender norms of Middle East populations that may inhibit treatment seeking.

The Egyptian Urological Association (EUA) Male Sexual Dysfunction Guidelines aims to present the contemporary evidence for medical practice in Egypt for the diagnosis and treatment of patients suffering from sexual dysfunction.

### **Recommendations of the Male Sexual Dysfunction**

<b>Recommendations of Erectile Dysfunction:</b>	
1.	Obtain a thorough medical and sexual history for all patients. <b>(Strong)</b>
2.	Use a validated questionnaire especially Arabic version (if available) related to ED to assess all sexual function domains and the effect of a specific treatment modality. <b>(Strong)</b>
3.	Perform physical examination in the initial assessment of men with ED to identify underlying medical conditions and comorbid genital disorders that may be associated with ED <b>(Strong)</b> .
4.	Assess routine laboratory tests, including glucose-lipid profile and total testosterone, to identify and treat any reversible risk factors and lifestyle factors that can be modified. <b>(Strong)</b> .
5.	Consider specific diagnostic tests in the initial evaluation only in the presence of "Indications for specific diagnostic tests" <b>(Strong)</b> .
6.	Ensure Including changes in diet, increased physical activity, stop smoking, improve overall health at or before treatment of erectile dysfunction. <b>(Strong)</b>
7.	Inform patients regarding approved PDE5Is, including discussion of benefits and risks/burdens. <b>(Strong)</b> .
8.	Use PDE5Is as first-line therapy. The dose should be titrated to provide optimal efficacy. <b>(Strong)</b>
9.	Consider early rehabilitation programs (use of PDE5I and VED) post-RP may improve erectile function <b>(Strong)</b> .
10.	Inform patients that PDE5Is may be more effective if combined with testosterone therapy when indicated. <b>(Strong)</b> .
11.	Assess patients for, inadequate/incorrect prescriptions, poor sexual stimulation, and fat meals when not advised <b>(Conditional)</b> .

12. Discuss benefits and risks/burdens regarding the use of VED, especially in well-informed older patients with infrequent sexual intercourse and comorbidity requiring non-invasive, drug-free management of ED <b>(Conditional)</b>
13. Perform an in-office injection test. Home therapy after positive office ICI test <b>(Conditional)</b> .
14. Alprostadil (PGE1) is the best agent however its cost is a limitation. <b>(Conditional)</b>
15. Use low intensity shockwave treatment (LI-SWT) in patients not candidate for oral vasoactive treatment or non-responders to PDE5Is <b>(Conditional)</b>
16. Intracavernosal stem cell therapy should be considered investigational for treatment of ED <b>(Conditional)</b>
17. <i>Intracavernosal Platelet Rich Plasma</i> should be considered investigational for ED treatment <b>(Conditional)</b>
18. <b>Botulinum Neurotoxin A (BoNT-A)</b> : Should be considered investigational for treatment of ED <b>(Conditional)</b> .
19. Surgery should be reserved for men in whom less invasive reversible treatment has not succeeded or is contraindicated or undesirable. <b>Strong</b>
20. Arterial revascularization surgery is offered only to select patients with ED who meet strict clinical and radiographic criteria for surgical success. <b>(Strong)</b>
21. Vascular surgery for veno-occlusive dysfunction is no longer recommended. <b>Strong</b>
22. Use implantation of a penile prosthesis as third-line therapy if other treatments fail or based upon patient preference <b>Strong</b> .
<b>Recommendations of Premature Ejaculation (PE)</b>
23. Obtain medical and sexual history to diagnose and classify PE, which should include assessment of intravaginal ejaculatory latency time (IELT) (self-estimated), perceived control, distress, and interpersonal difficulty due to the ejaculatory dysfunction. <b>Strong</b>
24. Perform physical examination in the initial assessment of PE to identify anatomical abnormalities that may be associated with PE or other sexual dysfunctions, particularly erectile dysfunction (ED). <b>Strong</b> .
25. Use the patient-reported outcomes tools: Premature Ejaculation Diagnostic Tool (PEDT) and Arabic Index of Premature Ejaculation (AIPE) in daily clinical practice. <b>(Conditional)</b>
26. Laboratory or neuro-physiological tests are not routine. They should only be directed by specific findings from history or physical examination. <b>Strong</b> .
27. Define the subtype of PE and discuss patient's expectations thoroughly before starting any treatment. <b>Strong</b> .
28. Treat the underlying cause (e.g., ED, prostatitis, LUTS, anxiety, hyperthyroidism) as the initial goal for patients with acquired PE. <b>Strong</b> .
29. Consider pharmacotherapy as the first-line treatment for patients with lifelong PE i.e. dapoxetine <b>Strong</b> .
30. The use of off-label topical anaesthetic agents i.e. the lidocaine/prilocaine spray is suggested as a viable alternative to oral treatment with SSRIs. <b>(Conditional)</b>
31. Use psychological/behavioural therapies in combination with pharmacological treatment in the management of acquired PE. <b>(Conditional)</b> .

32. Use various behavioural techniques in treating variable and subjective PE <b>(Strong)</b> .
33. The on-demand Tramadol is a weak alternative to SSRIs. <b>(Conditional)</b> .
34. PDE5Is alone or in combination with other therapies in patients with PE (without ED) may be used. <b>(Conditional)</b> .
<b>Recommendations for Delayed Ejaculation (DE)</b>
35. Perform a thorough analysis of the complaint to exclude misdiagnosed other sexual dysfunctions stressing on anorgasmia <b>Strong</b> .
36. Obtain a detailed medical and sexual history to exclude risk factors (medications especially SSRIs, antipsychotics, drug abuse, DM, depression, LUTS, etc) <b>Strong</b> .
37. Define if DE is lifelong or acquired, global or situational. <b>Strong</b> .
38. Assess intravaginal ejaculatory latency time (IELT) (self-estimated) <b>(Conditional)</b> .
39. Include physical examination in the initial assessment of DE to identify hypogonadism or anatomical abnormalities that may be associated with DE or other sexual dysfunctions, particularly erectile dysfunction <b>Strong</b> .
40. Request post-coital first voided urine sample to exclude retrograde ejaculation <b>Strong</b> .
41. Use specific questionnaires, specialized laboratory tests and radiologic investigation when indicated only. <b>(Conditional)</b> .
42. If acquired DE, consider stopping or modifying underlying incriminated drug regimen. <b>Strong</b> .
43. Improving erectile function and maximizing stimulation may trigger ejaculation. <b>(Conditional)</b> .
44. Psychosexual therapy can be particularly helpful in primary DE. <b>(Conditional)</b>
45. Testosterone replacement in hypogonadal patients may improve DE. <b>(Conditional)</b>
46. Cabergoline and bupropion could be beneficial for some cases of delayed ejaculation. <b>(Conditional)</b> .
47. Use PDE5I treatment significantly improved ejaculation and orgasm <b>Strong</b> .
48. Sympathetic $\alpha 1$ receptor agonists may help ejaculation with variable success rates in non-SCI patients. <b>(Conditional)</b>
49. Use penile vibratory stimulation or electro-ejaculation for sperm retrieval in patients with fertility issues and SCI. <b>Strong</b> .
<b>Recommendations for Peyronie's Disease (PD)</b>
50. Obtain a detailed history with specific emphasis on various characteristics of PD, such as onset, duration, course, pain, deformity, and ED. <b>(Strong)</b>
51. Perform physical examination, include assessment of palpable plaques, penile length, extent of curvature (self-photograph, or pharmacological-induced erection). <b>Strong</b> .
52. Do not use specific PD questionnaire, ultrasound measurement of plaque size in everyday clinical practice. <b>(Conditional)</b> .
53. Perform proper pre-operative counselling including the available treatment options and the known benefits and risks of each treatment, and the patient expectation will reduce post treatment patient dissatisfaction. <b>(Strong)</b>

54. Use conservative treatment in patients not fit for surgery or when surgery is not acceptable to the patient. <b>(Conditional)</b> .
55. Consider that intralesional collagenase injection has shown some outcome benefits in PD management. <b>(Strong)</b> .
56. Offer extracorporeal shockwave treatment in the active stage of the disease may alleviate penile pain. Do not use extracorporeal shockwave treatment to improve penile curvature and reduce plaque size. <b>(Conditional)</b> .
57. Offer penile traction devices and vacuum devices may reduce penile deformity and increase penile length. <b>(Conditional)</b> .
58. Do not use oral treatment with vitamin E and tamoxifen for significant reduction in penile curvature or plaque size. <b>(Strong)</b> .
59. Do not offer other oral treatments in chronic phase of PD (acetyl esters of carnitine, pentoxifylline, colchicine). <b>(Conditional)</b> .
60. Perform surgery only when PD has been stable for at least three months (without pain or deformity deterioration), which is usually the case after twelve months from the onset of symptoms. <b>Strong</b> .
61. Assess penile length, curvature severity, erectile function (including response to pharmacotherapy in case of ED) and patients' expectations prior to surgery. <b>Strong</b> .
62. Use tunical shortening procedures, especially plication techniques as the first treatment option for PD with adequate penile length, curvature < 60°, absence of special deformities (hourglass, hinge) and adequate erection. <b>Strong</b> .
63. Use grafting techniques for patients with PD with less than adequate penile length, curvature > 60°, presence of special deformities (hourglass, hinge) and adequate erection. <b>(Strong)</b> .
64. Use penile prosthesis implantation, with or without any additional procedure (modelling, plication, relaxing parallel incisions, grafting), in PD patients with ED not responding to pharmacotherapy. <b>Strong</b> .
<b>Recommendations for Priapism</b>
65. Obtain thorough history, is important in making diagnosis, etiology and type of priapism. <b>Strong</b>
66. Perform physical examination of the genitalia, the perineum and the abdomen. <b>Strong</b> .
67. Include laboratory investigations, complete blood count, coagulation profile and arterial blood gases. <b>Strong</b> .
68. Perform color duplex ultrasound of the penis and perineum for the differentiation between ischemic and non-ischemic priapism. <b>Strong</b> .
69. Use magnetic resonance imaging of the penis to predict smooth muscle viability in prolonged ischemic priapism. <b>(Strong)</b> .
70. Perform selected pudendal arteriogram when embolization is planned for the management of non-ischemic priapism. <b>Strong</b> .
71. Start management of ischaemic priapism as early as possible (within four to six hours) and follow a stepwise approach. <b>Strong</b> .
72. First, decompress the corpora cavernosa by penile aspiration until fresh red blood is obtained. <b>(Conditional)</b> .
73. Proceed to the next step, which is ICI of a sympathomimetic drug, in priapism that persists despite aspiration. <b>Strong</b> .

74. Repeat injections and aspiration for at least up to 1 hour prior to proceeding with surgical intervention in patients presenting with a priapism of less than 24 hours. <b>Strong.</b>
75. Consider more immediate surgical intervention in ischemic priapism of extended durations (typically greater than 72h), is unlikely to resolve with ICI therapy alone. <b>Strong.</b>
76. Perform distal shunt surgical procedures. Result of proximal procedures in case of failure is questionable. <b>Strong.</b>
77. Consider insertion of a penile prosthesis only if priapism episode is > 36 hours, or in cases for which all other interventions have failed. <b>Strong.</b>
<b>Recommendations for the treatment of non-ischemic priapism</b>
78. Non-ischaemic priapism is not an emergency, perform definitive management at the discretion of the treating physician. <b>(Conditional)</b>
79. Perform superselective arterial embolization, using temporary material for recurrent nonischaemic priapism <b>Strong</b>
80. Repeat the procedure with temporary or permanent material for recurrent nonischaemic priapism following selective arterial embolization. <b>(Conditional)</b>
81. Reserve selective surgical ligation of a fistula as a final treatment option when embolization has failed. <b>(Conditional).</b>
<b>Recommendations for the treatment of Stuttering priapism</b>
82. Treatment of Stuttering priapism, manage each acute episode similar to that for ischaemic priapism. <b>(Conditional).</b>
83. Use hormonal therapies (mainly gonadotropin-receptor hormone agonists or antagonists) and/or anti-androgens for the prevention of future episodes in patients with frequent relapses of stuttering priapism. Do not use them before sexual maturation is reached. <b>(Conditional)</b>
84. Initiate treatment with phosphodiesterase type 5 inhibitors in stuttering priapism only when the penis is in its flaccid state. <b>(Conditional).</b>
85. Use digoxin, $\alpha$ -adrenergic agonists, baclofen, gabapentin, or terbutaline only in patients with very frequent and uncontrolled relapses stuttering priapism. <b>(Conditional).</b>
86. Use intracavernous self-injections at home of sympathomimetic drugs until ischaemic priapism has been alleviated. <b>(Conditional).</b>

# Main body

## Introduction, purpose, scope, and audience

### **i** Introduction

Strategies for diagnosis and treatment of male sexual problems should consider the sociocultural factors that influence diagnosis and treatment seeking and engagement behaviours necessary for successful outcomes. Specifically, the detrimental effects of sexual problems on quality of life and the potential benefits of proper diagnosis and treatment should be more widely communicated to diminish the social disgrace associated with sexual problems and their management.

Erectile dysfunction (ED) and premature ejaculation (PE) are the two main complaints in male sexual medicine in the Middle East (1-2). Pharmacological therapies have completely changed the diagnostic and therapeutic approach to ED (3,4). The prevalence of ED is 20–90% among patients with different risk factors and medical comorbidities in Arab region countries and severe ED in patients in this region could be attributed to: (1) the high prevalence of risk factors; (2) the poor control of those risk factors; (3) the delay in seeking medical advice; and (4) the non-compliance with treatment (1-2). Unfortunately, in Arab countries there are no firm data on the true prevalence of sexual dysfunction. This prompted several investigators in the region to conduct research to identify the magnitude of the current problem (1-2).

This article integrates recent international guidelines with local experience and also highlights the apparent lack of congruency between available treatment and communication, cultural, and gender norms of Middle East populations that may inhibit treatment seeking. We clarified in our recent publication that strategies for diagnosis and treatment should consider the sociocultural factors that influence diagnosis and treatment seeking and engagement behaviours necessary for successful outcomes. Specifically, the detrimental effects of sexual problems on quality of life and the potential benefits of proper diagnosis and treatment should be more widely communicated to diminish the social disgrace associated with sexual problems and their management (5).

### **Sexual dysfunction issues unique to our region:**

- **Infertility and sexual dysfunction:** Infertility is negatively linked to sexuality in couples seeking assisted reproductive technology (ART), suggesting the need for integrated management of psychosexual problems (6). Unique to infertile couples in Egypt, like the Arab and Muslim world, the option of donor insemination is not accepted. The challenge of unsuccessful fertility issues in Egyptians may even further have a detrimental effect on the couple's sexual function. Infertility and sexual dysfunction are associated (7). Lack of sexual awareness and education contribute to this problem. Psychosexual management is warranted in these couples.
- **Unconsummated marriage:** A specific situation urologist face in our region is unconsummated marriage. It is a social challenge for the man to deal with his wife's virginity on the wedding night. Such stress may lead to performance anxiety and failure, accumulating into a full-blown ED situation in an otherwise healthy young man. Unconsummated marriage might occur in men with normal erection due to other causes as premature ejaculation, performance anxiety, lack of desire, hypogonadism, lack of knowledge, social pressure, and

female factors (8,9). The most common female factor was vaginismus (10). Particular to our regions, male lack of sexual desire may be related to consanguinity (11).

- **Polygamy, motives, and sexual dysfunction:** Egypt is among the countries where polygamy is legal (12). Polygamy has a psychosexual impact on the first wife, impacting intimacy with her husband and negatively affecting the dynamics of the family that is peculiar to these parts of the world (13). Non-monogamous female drive to sex includes coping mechanisms to keep the partner, maintain self-esteem, and seek higher levels of sexual pleasure (14). Men seek polygamy for a variety of reasons. For example, in a Turkish study, men reported that they had a second wife because of decreased satisfaction of sexual desires by a wife, falling in love with the second wife, and incompatibility with the first wife (15). In the Asian community, a prevalent polygamous practice has many underlying factors (16). These include prestige, economic advantage, social customs, and exposure to commercial sex. While polygamy may negatively affect wives and children, a couple of studies showed that polygamous men have less ED, less premature ejaculation, lower depression scores, and higher sexual satisfaction (15,17).

Other male sexual problems include **Premature Ejaculation:** Ejaculation that always or nearly always occurs prior to or within about one minute of vaginal penetration (lifelong PE) or a clinically significant and bothersome reduction in latency time, often to about three minutes or less (acquired PE). **Delayed Ejaculation:** Marked delay in ejaculation or marked infrequency or absence of ejaculation on almost all or all occasions (75-100% of the times) of partnered sexual activity without the individual desiring delay persisting for at least 6 months and causing significant distress to the individual. **Peyronie's Disease** (PD) is a symptomatic disorder characterized by a constellation of penile symptoms and signs, such as penile pain, curvature, shortening, narrowing, hinge deformity, and palpable plaque with subsequent ED. **Priapism** is a persistent penile erection for more than four hours and not related to sexual stimulation or relieved by ejaculation. Priapism carries high risk of structural damage to the cavernosal tissue which may lead to permanent ED.

## **Purpose**

*The Urologic Egyptian Guidelines on Male Sexual Dysfunction aim to present the contemporary evidence for medical practice in Egypt for the diagnosis and treatment of patients suffering from sexual dysfunction.*

## **Scope**

The Urologic Egyptian Guidelines on Male Sexual Dysfunction help and guide clinical practitioners to have knowledge of the incidence, pathophysiology, and strategies for diagnosis and treatment of male sexual problems. This document integrates recent international guidelines with local experts' opinions based on Egyptian healthcare and socioeconomic circumstances. It also reflects the opinions of experts in Sexual Dysfunction and represents state-of-the art references for all clinicians, as of the publication date.

### ***Target audience***

The target audience refers to those that deliver or implement the recommendations as well as health policymakers and other stakeholders involved in the adoption, adaptation, and transfer of health policies. The target audience of the guideline should not be misunderstood with the beneficiaries of the interventions or target population described in the guideline.

- **Urologists**
- **Dermatologists and Andrologists**
- **Family medicine and general practitioners**
- **Gynaecologists, psychiatrists and endocrinologists**

## Methods

**i** A comprehensive search for guidelines was undertaken to identify the most relevant guidelines to consider for adaptation.

Inclusion/exclusion criteria followed in the search and retrieval of guidelines to be adapted:

- Selecting only evidence-based guidelines (guideline must include a report on systematic literature searches and explicit links between individual recommendations and their supporting evidence)
- Selecting only national and/or international guidelines
- Specific range of dates for publication (using Guidelines published or updated 2015 and later)
- Selecting peer reviewed publications only
- Selecting guidelines written in English language
- Excluding guidelines written by a single author not on behalf of an organization in order to be valid and comprehensive, a guideline ideally requires multidisciplinary input
- Excluding guidelines published without references as the panel needs to know whether a thorough literature review was conducted and whether current evidence was used in the preparation of the recommendations

The following characteristics of the retrieved guidelines were summarized in a table:

- Developing organisation/authors
- Date of publication, posting, and release
- Country/language of publication
- Date of posting and/or release
- Dates of the search used by the source guideline developers

All retrieved Guidelines were screened and appraised using AGREE II instrument ([www.agreetrust.org](http://www.agreetrust.org)) by at least two members. the panel decided a cut-off point or rank the guidelines (any guideline scoring above 50% on the rigour dimension was retained).

### **Evidence assessment.**

According to WHO handbook for Guidelines we used the GRADE (Grading of Recommendations, Assessment, Development and Evaluation) approach to assess the quality of a body of evidence, develop and report recommendations (18, 19). GRADE methods are used by WHO because these represent internationally agreed standards for making transparent recommendations. Detailed information on GRADE is available on the following sites:

- GRADE working group: <http://www.gradeworkinggroup.org>
- GRADE online training modules: <http://cebgrade.mcmaster.ca/>
- GRADE profile software: <http://ims.cochrane.org/revman/gradepr>

### **Table 1 Quality of evidence in GRADE**

Quality level	Definition
<b>High</b>	We are very confident that the true effect lies close to that of the estimate of the effect.
<b>Moderate</b>	We are moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.
<b>Low</b>	Our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.
<b>Very low</b>	We have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

GRADE: Grading of Recommendations Assessment, Development and Evaluation.

**Table 2 Significance of the four levels of evidence**

Quality	Definition	Implications
High	The guideline development group is very confident that the true effect lies close to that of the estimate of the effect	Further research is very unlikely to change confidence in the estimate of effect
Moderate	The guideline development group is moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different	Further research is likely to have an important impact on confidence in the estimate of effect and may change the estimate
Low	Confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the true effect	Further research is very likely to have an important impact on confidence in the estimate of effect and is unlikely to change the estimate
Very low	The group has very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of the effect	Any estimate of effect is very uncertain

**Table 3: Factors that determine How to upgrade or downgrade the quality of evidence**

<b>Downgrade in presence of</b>	<b>Upgrade in presence of</b>
<b>Study limitations</b> -1 Serious limitations -2 Very serious limitations	<b>Dose-response gradient</b> +1 Evidence of a dose-response gradient
<b>Consistency</b> -1 Important inconsistency	<b>Direction of plausible bias</b> +1 All plausible confounders would have reduced the effect
<b>Directness</b> -1 Some uncertainty -2 Major uncertainty	<b>Magnitude of the effect</b> +1 Strong, no plausible confounders, consistent and direct evidence
<b>Precision</b> -1 Imprecise data	+2 Very strong, no major threats to validity and direct evidence
<b>Reporting bias</b> -1 High probability of reporting bias	

**The strength of the recommendation.**

The strength of a recommendation communicates the importance of adherence to the recommendation.

**Strong recommendations.**

With strong recommendations, the guideline communicates the message that the desirable effects of adherence to the recommendation outweigh the undesirable effects. This means that in most situations the recommendation can be adopted as policy.

**Conditional recommendations.**

These are made when there is greater uncertainty about the four factors above or if local adaptation has to account for a greater variety in values and preferences, or when resource use makes the intervention suitable for some, but not for other locations. This means that there is a need for substantial debate and involvement of stakeholders before this recommendation can be adopted as policy.

**When not to make recommendations.**

When there is lack of evidence on the effectiveness of an intervention, it may be appropriate not to make a recommendation.

Databases searched included four resource categories:

1. Four international guidelines and recommendations, namely European Association of Urology [EAU], American Urological Association Guidelines [AUA], British Society for Sexual Medicine [BSSM], International Society of Sexual Medicine [ISSM] (20 – 24).
2. Review of several guides, reviews, statements, recommendations, and standards (23 – 25).
3. Relevant Egyptian publications.

4. A panel of 10 high-calibre urologists and andrologists representing different universities, institutions and private practice in Egypt.

Adaptation of the Egyptian cultural aspects, the level of urologists' capabilities and the availability of well-equipped hospitals were considered in the methodology of diagnosis and different treatment modalities.

## Recommendations

### **i** Recommendations for the diagnosis of Erectile Dysfunction:

Recommendations (20-24, 28-41)	GRADE Level of certainty	Strength Rating
1- Obtain a thorough medical and sexual history for all patients.	High (28-32)	(Strong)
2- Use a validated questionnaire especially Arabic version (if available) related to ED to assess all sexual function domains and the effect of a specific treatment modality.	High (28- 30)	(Strong)
3- Perform physical examination in the initial assessment of men with ED to identify underlying medical conditions and comorbid genital disorders that may be associated with ED.	High (33 – 34)	(Strong)
4- Assess routine laboratory tests, including glucose-lipid profile and total testosterone, to identify and treat any reversible risk factors and lifestyle factors that can be modified.	High (35-39)	(Strong)
5- Consider specific diagnostic tests in the initial evaluation only in the presence of "Indications for specific diagnostic tests".	High (20-24, 35-41)	(Strong)

Recommendations for treatment of ED	GRADE Level of certainty	Strength Rating
6- Ensure Including changes in diet, increased physical activity, stop smoking, improve overall health at or before treatment of erectile dysfunction.	High (5, 42,43)	Strong
7- Inform patients regarding approved PDE5s, including discussion of benefits and risks/burdens.	High (20-24, 42, 44-48)	Strong
8- Use PDE5s as first-line therapy. The dose should be titrated to provide optimal efficacy.	High (20-24, 42, 44-48)	Strong
9- Consider early rehabilitation programs (use of PDE5I and VED) post-RP may improve erectile function.	Moderate (20-24, 49-56)	Strong
10- <b>Erectile Dysfunction and hypogonadism:</b> Inform patients that PDE5s may be more effective if combined with testosterone therapy when indicated.	Moderate (20-24, 57,58)	Strong
11- <b>PDE5s failure in patients with ED:</b> Assess patients for, inadequate/incorrect prescriptions, poor sexual stimulation, and fat meals when not advised.	Low (20-24, 44-48, 59-61)	Conditional
12- Discuss benefits and risks/burdens regarding the use of VED, especially in well-informed older patients with infrequent sexual intercourse and comorbidity requiring non-invasive, drug-free management of ED.	Low (20-24, 62, 63)	Conditional
13- Perform an in-office injection test. Home therapy after positive office ICI test	Low (20-24, 57, 64)	Conditional
14- Alprostadil (PGE1) is the best agent however its cost is a limitation.	Low (57)	Conditional
15- Use low intensity shockwave treatment (LI-SWT) in patients not candidate for oral vasoactive treatment or non-responders to PDE5s.	Low (20-24, 65-75)	Conditional
16- Intracavernosal stem cell therapy should be considered investigational for treatment of ED	Low (20-24)	Conditional
17- Intracavernosal Platelet Rich Plasma should be considered investigational for ED treatment	Low (20, 76-83)	Conditional

<b>18- Botulinum Neurotoxin A (BoNT-A):</b> Should be considered investigational for treatment of ED	Low (20, 84-87)	Conditional
<b>Recommendations for Surgical treatment</b>		
<b>19-</b> Surgery should be reserved for men in whom less invasive reversible treatment has not succeeded or is contraindicated or undesirable.	High (20-24)	Strong
<b>20-</b> Arterial revascularization surgery is offered only to select patients with ED who meet strict clinical and radiographic criteria for surgical success.	Moderate (88-90)	Strong
<b>21-</b> Vascular surgery for veno-occlusive dysfunction is no longer recommended.	High (20-24, 91)	Strong
<b>22-</b> Use implantation of a penile prosthesis as third-line therapy if other treatments fail or based upon patient preference.	High (92-100)	Strong

**Clinical Indicators for monitoring:**

- 1. Thorough medical and sexual history using a validated questionnaire especially Arabic version.**
- 2. Focused physical examination.**
- 3. Testosterone and lipid profile.**
- 4. Consider specific diagnostic tests when indicated.**

## Recommendation for assessment and management of Premature Ejaculation

Recommendation	GRADE Level of certainty	Strength Rating
<b>23-</b> Obtain medical and sexual history to diagnose and classify PE, which should include assessment of intravaginal ejaculatory latency time (IELT) (self-estimated), perceived control, distress, and interpersonal difficulty due to the ejaculatory dysfunction.	High (101-103)	Strong
<b>24-</b> Perform physical examination in the initial assessment of PE to identify anatomical abnormalities that may be associated with PE or other sexual dysfunctions, particularly erectile dysfunction (ED).	High (101-103)	Strong
<b>25-</b> Use the patient-reported outcomes tools: Premature Ejaculation Diagnostic Tool (PEDT) and Arabic Index of Premature Ejaculation (AIPE) in daily clinical practice.	Low (104)	Conditional
<b>26-</b> Laboratory or neuro-physiological tests are not routine. They should only be directed by specific findings from history or physical examination.	High (101-103)	Strong
<b>27-</b> Define the subtype of PE and discuss patient's expectations thoroughly before starting any treatment.	High (101-103)	Strong
<b>28-</b> Treat the underlying cause (e.g., ED, prostatitis, LUTS, anxiety, hyperthyroidism) as the initial goal for patients with acquired PE	High (104)	Strong
<b>29-</b> Consider pharmacotherapy as the first-line treatment for patients with lifelong PE i.e. dapoxetine	High (104-106)	Strong
<b>30-</b> The use of off-label topical anesthetic agents i.e. the lidocaine/prilocaine spray is suggested as a viable alternative to oral treatment with SSRIs.	Moderate (104-106)	Conditional
<b>31-</b> Use psychological/behavioural therapies in combination with pharmacological treatment in the management of acquired PE	Low (104-106)	Conditional
<b>32-</b> Use various behavioural techniques in treating variable and subjective PE	Moderate (104-106)	Strong
<b>33-</b> The on-demand Tramadol is a weak alternative to SSRIs.	Low (104-106)	Conditional
<b>34-</b> PDE5Is alone or in combination with other therapies in patients with PE (without ED) may be used.	Low (104-106)	Conditional

### Clinical Indicators for monitoring:

- 1. Medical and sexual history to diagnose and classify PE, use the patient-reported outcomes tools.**
- 2. Focused physical examination.**
- 3. Routine laboratory with seminal fluid culture and sensitivity to exclude underlying cause in patients with acquired PE**

### Recommendations for assessment of Delayed Ejaculation (107-109)

Recommendation	GRADE Level of certainty	Strength Rating
<b>35-</b> Perform a thorough analysis of the complaint to exclude misdiagnosed other sexual dysfunctions stressing on anorgasmia	High 107	Strong
<b>36-</b> Obtain a detailed medical and sexual history to exclude risk factors (medications especially SSRIs, antipsychotics, drug abuse, DM, depression, LUTS, etc)	High 107	Strong
<b>37-</b> Define if DE is lifelong or acquired, global or situational.	High 107	Strong
<b>38-</b> Assess intravaginal ejaculatory latency time (IELT) (self-estimated)	Low (108-109)	Conditional
<b>39-</b> Include physical examination in the initial assessment of DE to identify hypogonadism or anatomical abnormalities that may be associated with DE or other sexual dysfunctions, particularly erectile dysfunction	High (108-109)	Strong
<b>40-</b> Request post-coital first voided urine sample to exclude retrograde ejaculation.	High (108-109)	Strong
<b>41-</b> Use specific questionnaires, specialized laboratory tests and radiologic investigation when indicated only.	Low (108-109)	Conditional
<b>42-</b> If acquired DE, consider stopping or modifying underlying incriminated drug regimen.	High (108-109)	Strong
<b>43-</b> Improving erectile function and maximizing stimulation may trigger ejaculation.	Low (108-109)	Conditional
<b>44-</b> Psychosexual therapy can be particularly helpful in primary DE.	Low (108-109)	Conditional
<b>45-</b> Testosterone replacement in hypogonadal patients may improve DE.	Low (108-109)	Conditional
<b>46-</b> Cabergoline and bupropion could be beneficial for some cases of delayed ejaculation.	Low (108-109)	Conditional
<b>47-</b> Use PDE5I treatment significantly improved ejaculation and orgasm.	High (107-109)	Strong
<b>48-</b> Sympathetic $\alpha 1$ receptor agonists may help ejaculation with variable success rates in non-SCI patients.	Low (108-109)	Conditional
<b>49-</b> Use penile vibratory stimulation or electro-ejaculation for sperm retrieval in patients with fertility issues and SCI.	High (108-109)	Strong

#### Clinical Indicators for monitoring:

1. **Medical and sexual history with intravaginal ejaculatory latency time (IELT).**
2. **Focused physical examination.**
3. **Define if DE is lifelong or acquired, global or situational.**
4. **Specialized laboratory tests and radiologic investigation when indicated only.**

### Recommendations for evaluation and management of Peyronie's Disease (PD):

Recommendations	GRADE Level of certainty	Strength Rating
50. There is currently no international standard evaluation and treatment for PD and a detailed history should be obtained with specific emphasis on various characteristics of PD, such as onset, duration, course, pain, deformity, ED.	Moderate (20-24, 110-113)	Strong
51. Physical examination, include assessment of palpable plaques, penile length, extent of curvature (self-photograph, or pharmacological-induced erection).	High (20-24, 110-113)	Strong
52. Do not use specific PD questionnaire, ultrasound measurement of plaque size in everyday clinical practice.	Low 114	Conditional
53. Proper pre-operative counselling including the available treatment options and the known benefits and risks of each treatment, and the patient expectation will reduce post treatment patient dissatisfaction.	Moderate (86,114,115,116)	Strong
54. Use conservative treatment in patients not fit for surgery or when surgery is not acceptable to the patient.	Low (114,115,116)	Conditional
55. Intralesional collagenase injection has shown some outcome benefits in PD management.	Moderate (116)	Strong
56. Extracorporeal shockwave treatment may only be offered in the active stage of the disease to alleviate penile pain. Do not use extracorporeal shockwave treatment to improve penile curvature and reduce plaque size.	Low (115)	Conditional
57. Use penile traction devices and vacuum devices to reduce penile deformity and increase penile length.	Low (114)	Conditional
58. Do not use oral treatment with vitamin E and tamoxifen for significant reduction in penile curvature or plaque size.	High (20-24, 114)	Strong
59. Do not offer other oral treatments in chronic phase of PD (acetyl esters of carnitine, pentoxifylline, colchicine).	Low (114)	Conditional
60. Perform surgery only when PD has been stable for at least three months (without pain or deformity deterioration), which is usually the case after twelve months from the onset of symptoms.	High (20-24, 117,118)	Strong
61. Prior to surgery, assess penile length, curvature severity, erectile function (including response to pharmacotherapy in case of ED) and patient expectations.	High (20-24, 117,118)	Strong
62. Use tunical shortening procedures, especially plication techniques as the first treatment option for PD with adequate penile length, curvature < 60°, absence of special deformities (hourglass, hinge) and adequate erection.	High (20-24, 117,118)	Strong
63. Use grafting techniques for patients with PD with less than adequate penile length, curvature > 60°, presence of special deformities (hourglass, hinge) and adequate erection.	Moderate (20-24, 117,118)	Strong
64. Use penile prosthesis implantation, with or without any additional procedure (modelling, plication, relaxing parallel incisions, grafting), in PD patients with ED not responding to pharmacotherapy.	High (20-24, 117,118)	Strong

#### **Clinical Indicators for monitoring:**

1. **Medical and sexual history.**
2. **Focused physical examination (self-photograph, or pharmacological-induced erection).**
3. **Penile length, curvature severity, and erectile function.**

## Recommendations for diagnosis of ischemic priapism

Recommendation	GRADE Level of certainty	Strength Rating
65. Obtain thorough history, is important in making diagnosis, etiology and type of priapism.	High (119-122)	Strong
66. Perform physical examination of the genitalia, the perineum and the abdomen.	High (119-122)	Strong
67. Include laboratory investigations, complete blood count, coagulation profile and arterial blood gases.	High (119-122)	Strong
68. Perform color duplex ultrasound of the penis and perineum for the differentiation between ischemic and non-ischemic priapism.	High (119-122)	Strong
69. Use magnetic resonance imaging of the penis to predict smooth muscle viability in prolonged ischemic priapism.	Moderate (119-122)	Strong
70. Perform selected pudendal arteriogram when embolization is planned for the management of non-ischemic priapism.	High (119-122)	Strong

## Recommendations for the treatment of ischemic priapism:

Recommendations	GRADE Level of certainty	Strength Rating
71. Start management of ischaemic priapism as early as possible (within four to six hours) and follow a stepwise approach.	High (119-122)	Strong
72. First, decompress the corpora cavernosa by penile aspiration until fresh red blood is obtained.	Low (119)	Conditional
73. Proceed to the next step, which is ICI of a sympathomimetic drug, in priapism that persists despite aspiration.	High (119-122)	Strong
74. Repeat injections and aspiration for at least up to 1 hour prior to proceeding with surgical intervention in patients presenting with a priapism of less than 24 hours.	High (119-122)	Strong
75. Consider more immediate surgical intervention in ischemic priapism of extended durations (typically greater than 72h), is unlikely to resolve with ICI therapy alone.	High (119-122)	Strong
76. Perform distal shunt surgical procedures. Result of proximal procedures in case of failure is questionable.	High (119-122)	Strong
77. Consider insertion of a penile prosthesis only if priapism episode is > 36 hours, or in cases for which all other interventions have failed.	High (119-122)	Strong

## Recommendations for the treatment of non-ischemic priapism

Recommendations	GRADE Level of certainty	Strength Rating
78. Perform definitive management at the discretion of the treating physician, because non-ischaemic priapism is not an emergency.	Low (123-126)	Conditional
79. Perform superselective arterial embolization, using temporary material.	High (20-24)	Strong
80. Repeat the procedure with temporary or permanent material for recurrent nonischaemic priapism following selective arterial embolization.	Low (123-126)	Conditional
81. Reserve selective surgical ligation of a fistula as a final treatment option when embolization has failed.	Low (123-126)	Conditional

### Recommendations for the treatment of Stuttering priapism

Recommendations	GRADE Level of certainty	Strength Rating
<b>82.</b> Treatment of Stuttering priapism, manage each acute episode similar to that for ischaemic priapism.	Low (119-121)	Conditional
<b>83.</b> Use hormonal therapies (mainly gonadotropin-receptor hormone agonists or antagonists) and/or anti-androgens for the prevention of future episodes in patients with frequent relapses of stuttering priapism. Do not use them before sexual maturation is reached.	Low (119-121)	Conditional
<b>84.</b> Initiate treatment with phosphodiesterase type 5 inhibitors in stuttering priapism only when the penis is in its flaccid state.	High (119)	Conditional
<b>85.</b> Use digoxin, $\alpha$ -adrenergic agonists, baclofen, gabapentin, or terbutaline only in patients with very frequent and uncontrolled relapses stuttering priapism.	Low (119-121)	Conditional
<b>86.</b> Use intracavernous self-injections at home of sympathomimetic drugs until ischaemic priapism has been alleviated.	Low (119-121)	Conditional

### **Clinical Indicators for monitoring:**

- 1. Medical and sexual history.**
- 2. Focused physical examination.**
- 3. laboratory investigations, complete blood count, coagulation profile and arterial blood gases.**
- 4. Color duplex ultrasound of the penis and perineum**

## Update of Guidelines

**i** These guidelines will be updated whenever there is new evidence.

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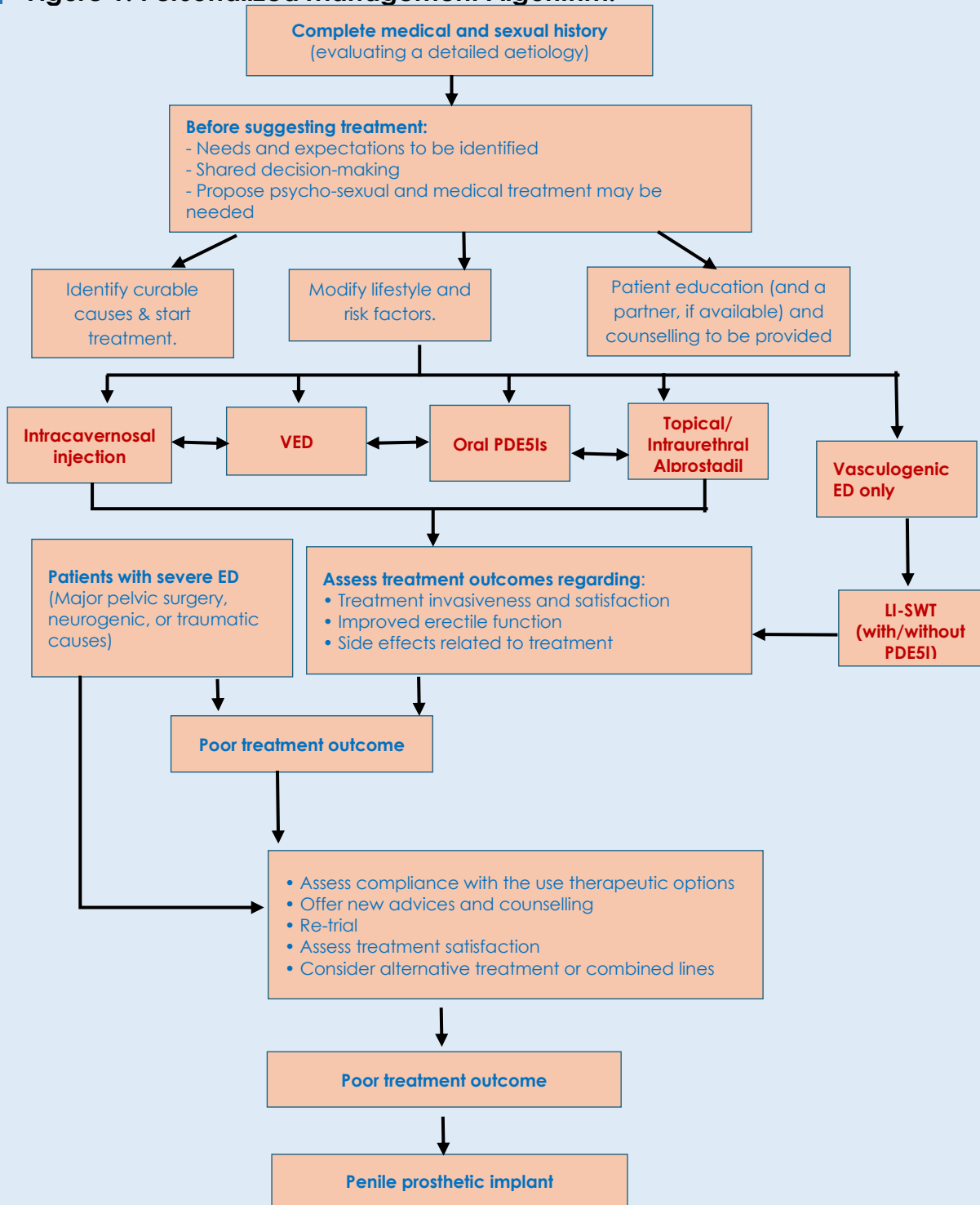
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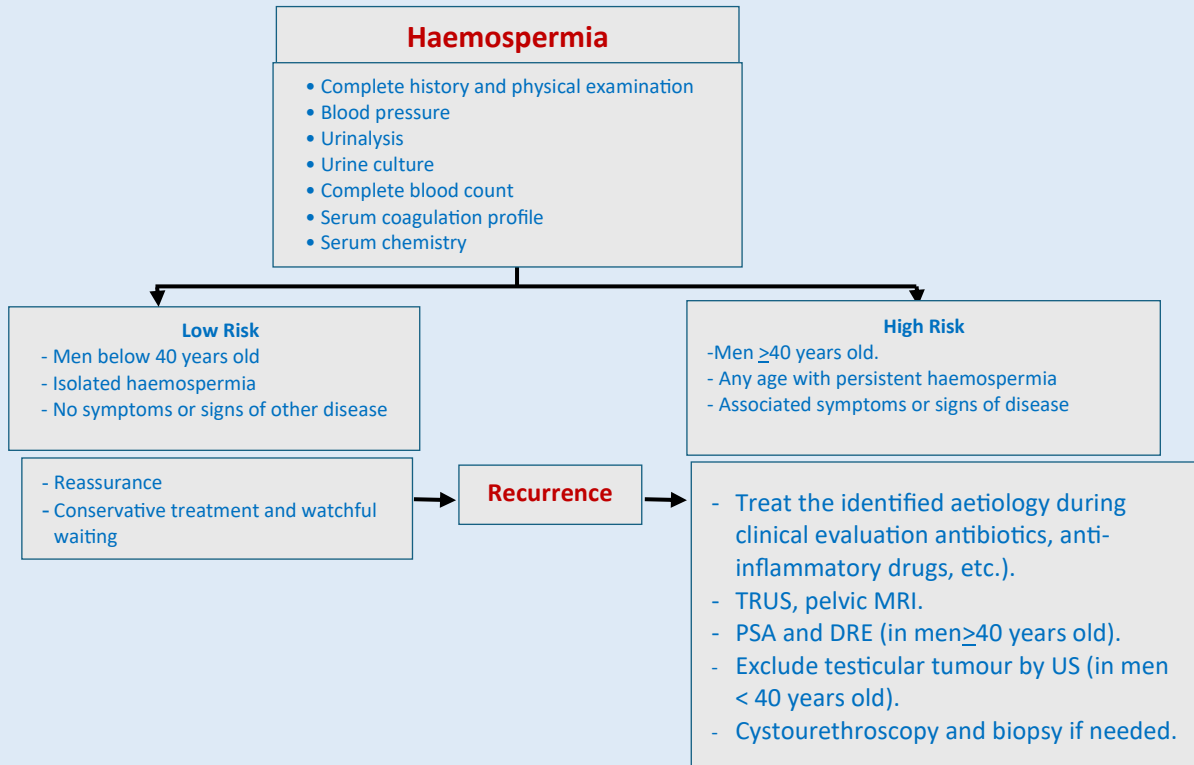
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**i** Figure 1: Personalized Management Algorithm:



**Figure 2: Management algorithm for Haemospermia**



**Table 4: Comparison of the properties of phosphodiesterase type 5 inhibitors (PDE5i) (42)**

Property	Sildenafil	Tadalafil	Vardenafil	Avanafil
TMAX	30–120min	30–360min	30–120min	30–45min
Terminal half life	4h	17.5h	4h	6–17h
Available doses	25mg,50mg,100mg	5mg,10mg,20mg	5mg,10mg,20mg	50mg,100mg,200mg
Max dose	100mg	20mg	20mg	200mg
Efficacy	Each offer similar efficacy			
Side effects (5 most common)	Headache, flushing, dyspepsia, nasal congestion, alteration in color vision	Headache, dyspepsia, back pain, myalgia, nasal congestion	Headache, flushing, rhinitis, dyspepsia, sinusitis	Headache, flushing, rhinitis, dyspepsia, sinusitis
Use with $\alpha$ -blockers.	- Concomitant use of selective $\alpha$ -blockers does not present a risk for significant hypotension - There is a risk of significant hypotension when using non-selective $\alpha$ -blockers			
Contraindications	- Regularly or intermittent use of organic nitrates. - Known hypersensitivity to any component of the tablet			
Dose adjustments that may be needed	<ul style="list-style-type: none"> <li>• Patients aged &gt; 65 years.</li> <li>• Hepatic impairment</li> <li>• Renal impairment</li> <li>• Concomitant use of potent cytochrome P450 3A4 inhibitors (e.g. ritonavir, cobicistat and erythromycin)</li> <li>• Concomitant use of cimetidine with sildenafil</li> </ul>			
TMAX = time to maximum plasma concentration.				

**Table 5: Clinical History, Physical Examination, Laboratory Investigations and Radiologic Assessment in Different Types of Priapism (119,120).**

Variant	History and clinical examination	Penile blood appearance	Penile blood gas findings	Color Duplex ultrasonography findings
<b>Ischemic priapism</b>	Tender and rigid corpora cavernosa	Corpus cavernosum testing: blood is hypoxic and dark in color	pO <sub>2</sub> <30 mmHg pCO <sub>2</sub> > 60mmHg pH <7.25	Minimal or absent blood flow
<b>Nonischemic priapism</b>	Perineal or penile trauma; non tender, partially tumescent corpora cavernosa	Corpus cavernosum testing: blood is oxygenated and red	pO <sub>2</sub> >90 mmHg pCO <sub>2</sub> < 40mmHg pH =7.4 similar to normal arterial blood)	Blood flow is normal to high in velocity
<b>Stuttering (recurrent) priapism</b>	Similar attacks	Corpus cavernosum testing: blood is hypoxic and dark in color	Blood gases: pO <sub>2</sub> <30 mmHg; pCO <sub>2</sub> >60 mmHg pH <7.25	Minimal or absent blood flow during acute priapism; normal blood flow otherwise
pCO <sub>2</sub> , partial pressure of carbon dioxide; pO <sub>2</sub> , partial pressure of oxygen.				

**Table 6: Percutaneous distal shunts, open distal shunts, open proximal shunts, and vein anastomoses/shunts**

		Example	Technique
Distal shunts	Percutaneous distal shunts	Winter (corporoglanular)	shunt large biopsy needle is inserted through glans
		Ebbehoj (corporoglanular)	shunt #11 blade scalpel is percutaneously passed
		T shunt (corporoglanular shunt)	Modified Ebbehoj using #10 blade scalpel and introducing the scalpel rotating it inside 90°
	Open distal shunt	Al-Ghorab	A 1 cm incision is made distal to coronal sulcus with excision of 5 × 5 mm cone segment of distal tunica albuginea from each corporal body
		Burnett 'snake' maneuver	Modification of Al-Ghorab shunt. A Hegar dilator is used to evacuate ischemic blood through a distal tunical window
Proximal shunts	Open proximal shunt	Quackels or Sacher (corporospongiosal) shunt	In lithotomy position, bulbocavernosus muscle is dissected from corpus spongiosum and 1 cm staggered ellipses of tissue are incised/excised from spongiosal/corporal bodies, and the defects anastomosed together
	Corporo saphenous vein or superficial/deep dorsal vein shunts	Grayhack shunt	The saphenous vein is ligated and anastomosed with corpora cavernosa
		Barry shunt	The superficial or deep dorsal vein is ligated and anastomosed to the corpora cavernosa