A practical guide on managing erectile dysfunction



Based on the 2017 British Society for Sexual Medicine (BSSM) guidelines on the management of erectile dysfunction in men¹

What is erectile dysfunction (ED)?

- ED is the persistent inability to attain and/or maintain an erection sufficient for satisfactory sexual performance
- ED is caused by various vascular, neuronal, hormonal and metabolic factors, mediated by endothelial and smooth-muscle dysfunction
- Although most causes of ED are physical, some are due to psychosexual issues; nevertheless, all patients with ED should have a history, examination and investigations performed, even if a psychological cause is suspected
- ED is a cardiovascular (CV) risk factor, posing a risk equivalent to that of current, moderate smoking
- ED is also an important marker for future CV events, with symptoms occurring some 3-5 years before an event^{2,3}
- The physical and psychosocial effects of ED can significantly affect the quality of life of patients and their partners⁴

Who is at risk?

- The risk factors for ED are similar to those for cardiovascular disease (CVD):2,3
 - Older age
 - Sedentary lifestyle
 - Obesity
 - Dyslipidaemia
 - Metabolic syndror

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- Diabetes
- Smoking

What are the other benefits of case-finding ED in practice?

- Increasing awareness regarding the availability of safe and effective oral drugs for ED,5-7 has led to more men seeking help for this condition, which facilitates the early detection of:
 - Diabetes (ED may be the first symptom in up to 20% of men)8,9
 - Dyslipidaemia (may not require treatment according to primary prevention guidelines, but may be a major reversible component in ED)9
 - Occult cardiac disease (in an otherwise asymptomatic man, ED may be a marker for underlying coronary artery disease)9
 - Testosterone deficiency (TD; a reversible cause of ED that may not require specific ED treatment, and which also has other long-term health implications)10
 - Associated lower urinary tract symptoms (LUTS)/benign prostatic hyperplasia (BPH) (ED and LUTS severity are closely related, and treatments for one condition may beneficially or adversely affect the other)8,11

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History taking

- **Obtain** a detailed description of the problem, including: - Symptom duration
 - Predisposing, precipitating and maintaining factors (if identified)
 - Any subsequent investigations
 - Previous/current treatment interventions and response
 - Reported tumescence, rigidity and quality of morning, spontaneous, masturbatory and/or partner-related erections
 - Sexual desire
 - Ejaculatory timing, control and orgasmic dysfunction
 - Previous erectile capacity
 - Any personal issues regarding sexual aversion or pain
 - Any partner issues, such as low sexual desire, menopause or gynaecological pain

- **Record** concurrent medical, psychiatric and surgical history, current relationship status, history of sexual partners and relationships, alcohol intake, smoking status and recreational drug use
- **Consider** the use of validated questionnaires, (such as the IIEF, shorter version of the SHIM, IPSS or AMS scale) to assess sexual function domains and response to therapy
- Note any issues regarding sexual orientation and gender identity

AMS – Aging Males' Symptoms, IIEF – International Index of Erectile Function, IPSS – International Prostate Symptom Score, SHIM – Sexual Health Inventory for Men

Physical examination

- **Measure** heart rate, blood pressure, abdominal circumference, weight,⁹ and height if body mass index required
- **Perform** a genital examination, particularly with pain of sudden onset, deviation of the penis during tumescence, symptoms of TD, or other urological symptoms (past or present)
- **Conduct** a DRE of the prostate if there are genitourinary or protracted secondary ejaculatory symptoms

DRE – digital rectal examination

Investigations

- **Check** serum lipids, fasting plasma glucose and/or glycated haemoglobin
- Measure serum TT in the morning (before 11 am), in the fasting state. If low (TT <8 nmol/L) or borderline (TT 8–12 nmol/L), repeat with serum LH and prolactin.¹² FT has a greater correlation with clinical symptoms of TD¹³ (FT and bioavailable testosterone can be calculated from TT, SHBG and albumin; an online FT calculator and downloadable app, sponsored by the Primary Care Testosterone Advisory Group (PCTAG), can be found at http://www.pctag.uk/testosterone-calculator/). If levels are abnormal, consider specialist referral
- **Consider** specialist investigations in men who:
- Wish to know the aetiology of their ED
- Have an arterial abnormality on Doppler ultrasound
- Have a history of trauma
- Are young and:
 - Have always had trouble obtaining/maintaining an erection
 - Have a primary CV abnormality
 - Have suspected primary venous leakage
 - Are being considered for surgical intervention
- Have an abnormality of the penis or testes
- Have not responded to medical therapy and may want surgical treatment
- **Consider** measuring PSA if clinically indicated, and certainly before commencing testosterone therapy and at 3, 6, and 12 months afterwards¹⁴
- **Consider** thyroid function tests

FT - free testosterone, LH - luteinising hormone, PSA - prostate-specific antigen, SHBG - sex hormone-binding globulin, TT - total testosterone

Diagnosing and managing ED in primary care

- The primary objective in the management of ED is to enable the man or couple to enjoy a satisfactory sexual experience
- When managing ED, consider not only the efficacy and safety of the different treatments, but also patient and partner preference, and all the factors that may influence this
- It is of paramount importance to use the opportunity to manage any previously undiagnosed comorbidities that present following the patient assessment, and to treat to target any existing conditions and make lifestyle modifications, where necessary



ED therapies

PDE5

- ~25–50% of men fail to respond within 12 months16
- Rates are higher in men with T2DM or post-RP16
- Inadequate prescribing/instruction is the main cause of treatment failure
- Daily/frequent dosing regimens may salvage men who've failed with on-demand therapy
- Correction of testosterone levels <10.4 nmol/L may salvage non-responders
- Nitrates may be safely discontinued (with a cardiologist's approval), to aid therapy¹⁷
- Co-administration with antihypertensives may increase the drop in blood pressure¹⁸

Contraindications include:*19-22

- Use of nitrates in any form, guanylate cyclase stimulators, potent CYP3A4 inhibitors
- Loss of vision in one eye due to NAION
- Severe renal/hepatic impairment
- Hypotension

Dose adjustments may be required in:*19-22

- Renal or hepatic impairment
- Concomitant use of CYP3A4 inhibitors
- Apply caution in:*19-22
- Patients receiving alpha-blockers and those with anatomical penile deformities or a predisposition to priapism
- Possible adverse effects include:*19-22
- Headache
- Dizziness
- Flushing
- Dyspepsia
- Nasal congestion

Intra-urethral alprostadil

- · Less invasive than injection therapy but works for only ~30-60% of patients^{35,3}
- Higher doses of MUSE (500/1000 µg) are usually required³⁵

Contraindications include:37

- Anatomical deformity of the penis
- Predisposition to priapism

Apply caution in:37

- Men whose partners could be pregnant (a condom must be used)
- Possible adverse effects include:37
- Penile pain
- Haematoma
- Headache
- Dizziness
- Hypotension
- Muscle spasm

Vacuum erection devices

- Highly effective, regardless of ED aetiology^{23–25}
- Reported satisfaction rates vary between 35%26 to 84%27
- Can be a useful adjunct to PDE5I/injection therapy post-RP, and to salvage treatment failures
- Work best if the man and his partner receive sufficient instruction, and have positive attitudes towards their use

Contraindications include:

- Bleeding disorders
- Concurrent anticoagulant therapy
- Possible adverse effects include:29
- Bruising
- Local pain

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June 2016.

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Eur Urol 2017;72:1000-11.

- Failure to ejaculate
- Coldness of the penis

Intracavernous injections

- Include alprostadil and aviptadil+phentolamine (A/P)
- Alprostadil is effective in >70% of men,³⁰ but compliance rates may be low³¹
- A/P has similar efficacy to alprostadil, with less painful injections³³
- Unlike alprostadil, A/P injections need to be accompanied by some form of sexual stimulation for an optimal erection
- Apply caution in:*33,3
- Men at increased risk of bleeding (anticoagulants are contraindicated with A/P)
- Contraindications include:*33,34
- Predisposition to priapism
- Penile implants
- Possible adverse effects include:*33,34
- · Bruising and haematoma at injection site
- Penile pain (alprostadil)
- Flushing of face/trunk (A/P)

Alprostadil cream

- Store alprostadil cream (Vitaros) in the refrigerator
- Apply 300 µg in 100 mg (3 mg/g) into the urethral meatus at room temperature
- · Improved tumescence of the glans may be useful in men post-penile prosthesis¹
- Has been shown to produce a 2.5-point increase in IIEF and a 15% relative increase in successful intercourse attempts³⁸

Contraindications include:39

- Anatomical deformity of the penis
- Predisposition to priapism

Apply caution in:39

 Men whose partners could be pregnant (a condom must be used)

Baltaci S et al. Treating erectile dysfunction with a vacuum

tumescence device: a retrospective analysis of acceptance and satisfaction. Br J Urol 1995;76:757-60.

tumescence device: a retrospective analysis of acceptance

Porst H. The rationale for prostaglandin E1 in erectile failure: a survey of worldwide experience. J Urol 1996;155:802-15.

Shah PJR et al. Injection therapy for the treatment of erectile

Mulhall JP et al. The causes of patient dropout from penile self-injection therapy for impotence. J Urol 1999;162:1291-4.

dysfunction: a comparison between alprostadil and a combination of vasoactive intestinal polypeptide and phentolamine mesylate. Curr Med Res Opin

Caverject 10 μg powder for solution for injection SmPC. Pfizer Ltd. March 2017.

Invicorp 25 μg/2 mg solution for injection SmPC. Evolan Pharma AB. September 2017.

Guay AT et al. Clinical experience with intraurethral alprostadil (MUSE) in the treatment of men with erectile dysfunction. A retrospective study. Medicated urethral system for erection. Eur Urol 2000;38:671-6.

NICE. Erectile dysfunction: alprostadil cream. Available at: https://www.nice.org.uk/advice/esnm50/chapter/Key-points-from-the-evidence (Accessed June 2018).

Copyright © British Society for Sexual Medicine. June 2018

Vitaros 3 mg/g cream SmPC. Ferring Pharmaceuticals Ltd. October 2017.

MUSE 1000 μg urethral stick SmPC. MEDA Pharmaceuticals. December 2013.

Fulgham PF et al. Disappointing initial results with transurethral alprostadil for erectile dysfunction in a urology practice setting. J Urol 1998;160:2041-6.

Brock G et al. Safety and efficacy of vardenafil for the treatment of men with erectile dysfunction after radical retropubic prostatectomy. J Urol 2003;170:1278-83.

29. Baltaci S et al. Treating erectile dysfunction with a vacuum

and satisfaction. Br J Urol 1995;76:757-60.

- Possible adverse effects include:39
- Penile/genital pain/erythema
- Urethral pain

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2007;23:2577-83.

*See the individual Summary of Product Characteristics for full details, because these vary between products. CYP3A4 – cytochrome P450 3A4, NAION – non-arteritic anterior ischaemic optic neuropathy, RP – radical prostatectomy, T2DM – type 2 diabetes mellitus

References

- Hackett G et al. British Society for Sexual Medicine Guidelines on the Management of Erectile Dysfunction in Men - 2017. J Sex Med 2018;15:430-57.
- 2
- Thompson IM et al. Erectile dysfunction and subsequent cardiovascular disease. JAMA 2005;294:2996-3002. Vlachopoulos CV et al. Prediction of cardiovascular events and all-cause mortality with erectile dysfunction: a systematic review and meta-analysis of cohort studies. 3.
- Circ Cardiovasc Qual Outcomes 2013;6:99-109. Feldman HA et al. Impotence and its medical and psychosocial correlates: results of the Massachusetts Male Aging Study. J Urol 1994;151:54-61. 4.
- Brock GB et al. Efficacy and safety of tadalafil for the treatment of erectile dysfunction: results of integrated analyses. J Urol 2002;168:1332-6. Goldstein I et al. Oral sildenafil in the treatment of erectile 5.
- 6 dysfunction. Sildenafil Study Group. N Engl J Med 1998;338:1397-404.
- Hellstrom WJ et al. Sustained efficacy and tolerability of vardenafil, a highly potent selective phosphodiesterase 7. type 5 inhibitor, in men with erectile dysfunction: results of a randomized, double-blind, 26-week placebo-controlled pivotal trial. Urology 2003;61:8-14.
- Rosen R et al. The multinational Men's Attitudes to Life Events and Sexuality (MALES) study: I. Prevalence of erectile dysfunction and related health concerns in the general population. Curr Med Res Opin 2004;20:607-17. 8.
- Jackson G et al. The assessment of vascular risk in men with erectile dysfunction: the role of the cardiologist and general physician. Int J Clin Pract 2013;67:1163-72. 9
- 10. Pye SR et al. Late-onset hypogonadism and mortality in aging men. J Clin Endocrinol Metab 2014;99:1357-66. Rosen R et al. Lower urinary tract symptoms and male 11.
- sexual dysfunction: the multinational survey of the aging male (MSAM-7). Eur Urol 2003;44:637-49.
- 12. Dean JD et al. The International Society for Sexual Medicine's process of care for the assessment and management of testosterone deficiency in adult men J Sex Med 2015;12:1660-86.

- 13. Antonio L et al. Low free testosterone is associated with hypogonadal signs and symptoms in men with normal total testosterone. J Clin Endocrinol Metab 2016;101:2647-57.
- Hackett G et al. The British Society for Sexual Medicine guidelines on adult testosterone deficiency with statements for UK practice. J Sex Med 2017;14:1504-23.
- 15. Dunn ME. Restoration of couple's intimacy and relationship vital to re-establishing erectile function. J Am Osteopath Assoc 2004;104:S6-S10.
- Kirana PS et al. (eds.) The EFS and ESSM Syllabus of Clinical Sexology. Amsterdam: Medix Publishers; 2013.
- Jackson G et al. Successful withdrawal of oral long-acting nitrates to facilitate phosphodiesterase type 5 inhibitor use J Sex Med 2005;2:513-6.
- McMahon C. Efficacy and safety of daily tadalafil in men with erectile dysfunction previously unresponsive to on-demand tadalafil. J Sex Med 2004;1:292-300. 19. Cialis 10 mg film coated tablet SmPC. Eli Lilly and Company.

Spedra 100 mg tablets SmPC. A. Menarini Farmaceutica Internazionale SRL. November 2017.

dysfunction: use and results. World J Urol 1997;15:78-82. 24. Levine LA et al. Vacuum constriction and external erection

dysfunction: a long-term, prospective study of patients with mild, moderate, and severe dysfunction. Urology 1999;

Levitra 10 mg orodispersible tablets SmPC. Bayer PLC. December 2017.

21. Viagra 100 mg film coated tablets SmPC. Pfizer Ltd.

23. Lewis RW et al. External vacuum therapy for erectile

devices in erectile dysfunction. Urol Clin North Am 2001;28:335-41.

25. Dutta TC et al. Vacuum constriction devices for erectile

26. Corona G et al. Meta-analysis of the results of testosterone

therapy on sexual function based on the IIEF scores.