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IPS Textbook of Sexuality and Sexual Medicine



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Mrugesh Vaishnav
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JAYPEE

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CHAPTER

3 Anatomical and Physiological Aspects of Sexuality

Shobit Garg, Veena Tejan

"Sex is something you do. Sexuality is something you are."

–Anna Freud

INTRODUCTION

Human sexuality involves sexual attraction to another person, which is mostly seen toward the opposite sex, some to the same sex, or some having both and not being attracted to anyone in a sexual manner (asexuality).[1]

Human sexuality is mainly determined by the anatomy, physiology, the culture (in which a person lives), the relationships, and developmental experiences. Human sexuality encompasses the perception of being male or female, the owned private thoughts, the fantasies as well as the behavior.[2]

Sexuality has always been a subject of significance to the medical community. In the classical era, Hippocrates considered the clitoris as the site of sexual arousal in females. In the middle ages, Islamic physicians suggested that coitus interruptus is a form of birth control. During the Victorian era, intellectuals such as Havelock Ellis and Richard von Krafft-Ebing presented diverging views on sexual behavior.[3] During the same period, Sigmund Freud developed his innovative theories on childhood sexuality, libido, and the effects of the sexual impulses on the human behavior and psychopathology.[2] In the modern era, Alfred Kinsey (1938) attempted to replace the conventional ideas of normal sexual behavior with a new definition with biological under-pinning's, i.e., "nearly all the so-called sexual perversions fall within the range of biological normality".[4]

Dr Alfred Kinsey initiated a large-scale survey research on human sexual behavior. The results were published in the two books firstly titled "Sexual Behavior in the Human Male" and secondly "Sexual Behavior in the Human Female", which were published in 1948 and 1953, respectively.[5]

William Masters and Virginia Johnson (in 1966) published a book detailing the results of their observations of nearly 700 people and their physiological responses during sexual behavior.[6] Unlike Kinsey, who used personal interviews and surveys to collect data, Masters and Johnson observed people having intercourse in a variety of positions, and then masturbating, manually or with the aid of a device.[6,7] Based on these observations, Masters and Johnson divided the sexual response cycle (SRC) into four phases, i.e., excitement, plateau, orgasm, and resolution. These four phases are fairly similar in men and women.[8]

The work of William Masters and Virginia Johnson and the development of drugs that prevent contraception, aiding erection, and hormones (replacement) that decrease with menopause and aging contributed to the development of an era of sexual liberality.

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GROSS ANATOMY

Female Reproductive System

The most visible structure of female sexual anatomy is the vulva. The primary functions of the vulva are pleasure and protection. The vulva is composed of the female's *external sex organs* **(Fig. 1)**. It includes many parts:

Mons Pubis

The "mons pubis" is the round shape, hair-bearing area of skin, and adipose tissue over the pubic symphysis (and adjacent pubic bone). The mons pubis is relatively flat and hairless before puberty, becomes prominent with coarse hair through adolescence, and atrophies slightly after menopause.

Labia Majora

Lateral boundaries of vulva are formed by two longitudinal folds of skin, labia majora, which does not fuse posteriorly and merge into the neighboring skin almost parallel to each other. Externally, each labium has a pigmented surface covered with hairs. Internally, the labia has smooth pink surface with large sebaceous follicles.

Labia Minora

Between the labia majora, there lies two small cutaneous folds, devoid of fat named "labia minora". Each labium minora bifurcates anteriorly. The upper layer of each side passing above the clitoris forms the hood or prepuce while lower layer passing below the clitoris form the frenulum of the clitoris.

Minor and Major Vestibular Glands

The major vestibular glands (VGs) (also known as Bartholin's glands) are located just to the left and right of the vagina. These promote lubrication and aid in sexual intercourse. The minor VGs—also called Skene's glands, are on the wall of the vagina and are associated with female ejaculation, and mythologically associated with the G-spot.

Clitoris

The "clitoris" is an erectile structure. It has a glans, body, and a root. The glans is a small

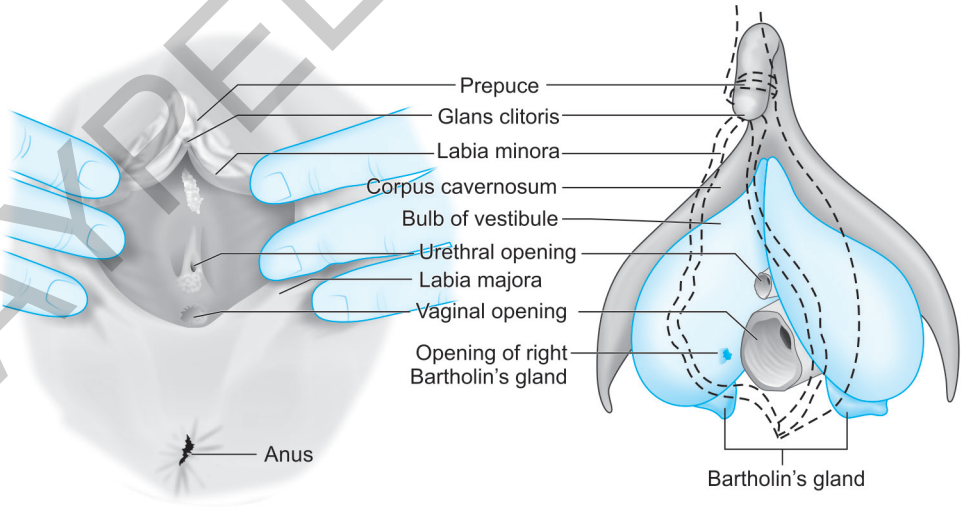

Fig. 1: The superficial features of the perineum.

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and round tubercle of spongy erectile tissue placed at the end of the body. Its epithelium has highly sensitivity and is important in sexual responses. The body can be palpated through the skin.

Vagina

The vagina is a fibromuscular tube which is lined by nonkeratinized stratified epithelium. The upper end of the vagina is surrounded by the lower end of uterine cervix. The annular recess between the vagina and cervix is known as the "fornix". The fornix has different parts: anterior, posterior, right, and left lateral, but they are continuous. The anterior and posterior walls of the vagina are approximately 7.5 cm and 9 cm long, respectively.

Table 1 details arterial supply, venous drainage, lymphatic drainage, and nerve supply to the female external genitals.

The internal sex organs of the female reproductive system (functions to store, transport ovum cells, and produce hormones) include:

- *The uterus*: Where fetal development occurs.
- *The ovaries*: They house the ova and produce estrogen, progesterone, and testosterone (small amounts).
- *The Fallopian tubes*: Where fertilization occurs.

The primary functions of these internal sex organs are to store, transport, and keep ovum cells healthy; and produce hormones.

Male Reproductive System

The male reproductive system consists of the testes, spermatic cord, accessory glands, and external genitalia.

Testis

Testis is responsible for sperm and testosterone production. In adults, they typically measure 4–5 cm in length, 2–3 cm in breadth, and 3–4 cm in anteroposterior diameter;[9] their weight varies between 12 and 20 g. Average testicular volume ranges from 15 to 25 mL.[10] They are enclosed in a tough capsule which is made up of three layers: (1) An innermost tunica vasculosa; (2) An intermediate tunica albuginea; (3) An outer tunica vaginalis. Epididymis has its attachment on the posterior surface of

TABLE 1: Arterial supply, venous drainage, lymphatic drainage, and nerve supply to the female external genitals.

Arterial supply	***Venous drainage***	***Innervation***	***Lymphatic drainage***
Female external genital area is supplied superiorly by superficial and deep external pudendal artery and inferiorly by internal pudendal artery	Venous drainage of the vulval skin is via external pudendal veins to the long saphenous vein. Clitoris is drained via internal pudendal vein and long saphenous veins	Nerve supply of the vulva is by the pudendal nerve (S2, S3, and S4). Ilioinguinal nerve (L1) supplies the anterior third of the labium major. The pudendal nerve supplies the posterior two-thirds. The lateral aspect is supplied by perineal branch of the posterior cutaneous nerve of the thigh (S2)	Lymphatic drainage of external genitalia is into deep inguinal nodes

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TABLE 2: Arterial supply, venous drainage, lymphatic drainage, and nerve supply to the testis.

Arterial supply	*Venous drainage*	*Lymphatic drainage*	*Innervation*
		Testis	
Approximately two-thirds of testicular blood supply is from testicular artery branch of abdominal aorta, vas deferens, and ductus deferens by vasal artery and approximately one-third of testicular supply is by the cremasteric arteries, branch of the inferior epigastric artery	Testicular veins emerge posteriorly from the testis and drain the epididymis, and unite to form richly anastomotic channels (surrounding the testis) known as *pampiniform plexus*. This vascular arrangement permits the exchange of heat and small molecules, facilitating in maintenance of lower testicular temperatures	Lymphatic drainage of testicles is in aortocaval nodes	Testis is innervated by the nerve fibers arising from the 10th and 11th thoracic spinal segments, relaying via the renal and aortic plexuses[11]
		Seminal vesicle	
Primary blood supply to seminal vesicle is from branch of umbilical artery	Venous drainage is provided by the vesiculodeferential veins and the inferior vesical plexus	The seminal vesicles receive preganglionic parasympathetic input from the pelvic nerve, and sympathetic (postganglionic) and parasympathetic (preganglionic) input from the hypogastric nerve	Lymphatic drainage occurs via internal iliac nodes

testis. **Table 2** details arterial supply, venous drainage, lymphatic drainage, and nerve supply to the testis.[11]

Spermatic Cord

It consists of epididymis, vas deferens, paradidymis, and ejaculatory ducts.

Epididymis: The epididymis lies posterior, and laterally (slightly), to the testis. Anatomically, it is divided into three sections: (1) Caput (head); (2) Corpus (body); (3) Cauda (tail).

Vas deferens: The vas deferens (ductus deferens) is a tubular structure and transports sperm from the epididymis to the urethra. It also serves absorptive and secretory functions.[12] It arises from the cauda epididymis. The vas is tortuous for 2–3 cm and culminates at the ejaculatory duct.

Paradidymis: The paradidymis is a small collection of convoluted tubules. These are located anteriorly in the spermatic cord just above the caput epididymis.

Ejaculatory ducts: The duct of the seminal vesicle unites with the ampulla of the vas deferens to form two paired ejaculatory ducts. Each duct is approximately 2 cm in length.

Accessory Glandular Structures

Seminal vesicles: The vas deferens gives two paired outpouchings, the seminal vesicles

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located at the base of prostate. **Table 2** details arterial supply, venous drainage, lymphatic drainage, and nerve supply to the seminal vesicle.

External Genitalia

Penis: The penis consists of root and a pendulous body. Root is attached in the perineum, and the pendulous body is entirely enveloped in skin. The penile shaft contains three erectile columns—the paired corpora cavernosa and the corpus spongiosum; the urethra; and the investing fasciae, blood vessels, and nerves associated with these structures **(Fig. 2)**. The foreskin covering the glans is called as "prepuce". Prepuce is a loose fold of retractable skin attached to the ventral surface of the glans at the frenulum.

Microstructure of penis: The corpora cavernosa is spongy erectile tissue which is composed of interconnected endothelium-lined sinusoids that are separated by smooth muscle trabeculae. These smooth muscle trabeculae are surrounded by elastic fibers, collagen, and loose areolar tissue. The tunica albuginea consists of an inner circular and outer longitudinal layer of elastic fibers arranged in a latticed network. The corpora cavernosa is surrounded by tunica albuginea.

Venous drainage from the corpora cavernosa originates in the trabeculae between the peripheral sinusoids and the tunica albuginea before exiting the corpora via the emissary veins. The emissary veins travel between the two layers of the tunica albuginea, and exit the outer layer in an oblique fashion. The outer layer compresses the emissary veins when the penis becomes engorged with blood, preventing venous outflow, and thereby maintains the erection. The structure of the corpus spongiosum is almost similar to that of the corpora cavernosa. **Table 3** details arterial supply, venous drainage, lymphatic drainage, and nerve supply to the penis.

Scrotum: The scrotum has multiple layers of tissues. These include skin, Dartos muscle, external spermatic, cremasteric, and internal spermatic fasciae. Scrotal skin is sensitive to temperature changes as it is richly innervated by sensory nerves that respond to stimulation of the skin and hairs. **Table 3** has mentioned arterial supply, venous drainage, lymphatic drainage, and nerve supply to the scrotum.

NEUROANATOMY OF HUMAN SEXUALITY

The neuroanatomy concerning human sexuality is complex phenomenon and depends upon interaction of several brain regions, confounding neurotransmitters and signals from the peripheral nervous system. Some of the important brain areas are discussed here which have been implicated in human sexuality.

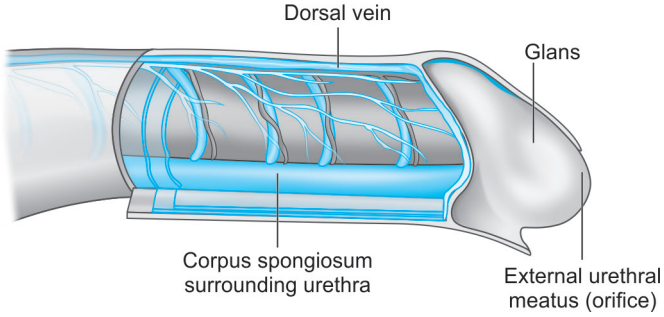

Fig. 2: Layers of penis and main structures.

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TABLE 3: Arterial supply, venous drainage, lymphatic drainage, and nerve supply to the penis.

Arterial supply	*Venous drainage*	*Lymphatic drainage*	*Innervation*
		Penis	
Blood supply to the corporal bodies is derived from the internal pudendal artery, a branch of the internal iliac artery. The common penile artery has three main branches: (1) The bulbourethral artery; (2) The dorsal penile artery; (3) The cavernous (deep and cavernosal) artery	• Penile blood is carried by any of three venous systems: (1) Superficial; (2) Intermediate; or (3) Deep. • The penile shaft skin and prepuce drain blood in superficial venous system. • The intermediate drainage system receives blood from the glans, corpus spongiosum, and the distal two-thirds of the penis • Deep venous drainage receives blood from the proximal third of the penis	The lymphatic drainage of the penile and perineal skin drains to the superficial inguinal nodes, glans penis to the deep inguinal and external iliac nodes, and erectile tissue and penile urethra to the internal iliac nodes	• Parasympathetic input to the penis (which is excitatory) originates in first four sacral segments of the spinal cord and travels via the pelvic splanchnic nerves (nervi erigentes) and to the pelvic plexus **(Fig. 3)**, where preganglionic fibers synapse on postganglionic neurons that give rise to the cavernous nerve. • Sympathetic input to the penis (inhibitory) originates from the 11th and 12th thoracic, and first lumbar segments of the spinal cord, and travels through the sympathetic trunk before descending to the pelvic plexus
		Scrotum	
Scrotal skin is supplied by the branches of the femoral artery, the internal pudendal artery, and the inferior epigastric artery	Venous drainage follows the arterial network	The lymphatic drainage from the scrotum is routed to the ipsilateral superficial inguinal nodes	Anterior third of the scrotum is innervated by the ilioinguinal and genitofemoral nerves and posterior two-thirds is by the perineal and posterior femoral cutaneous nerves

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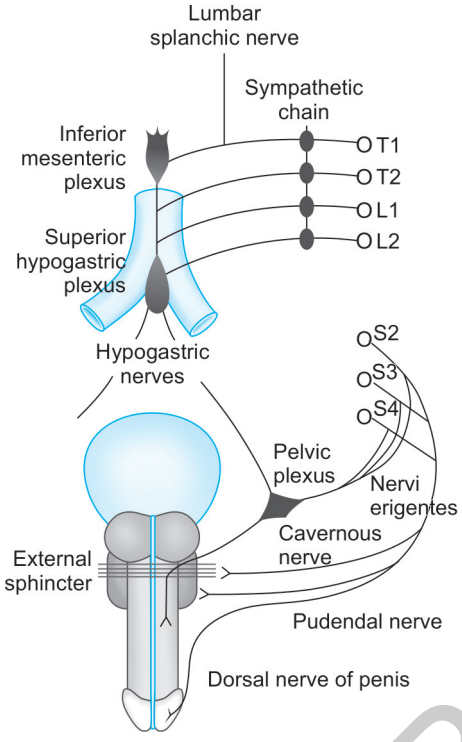

Fig. 3: Nerve supply to the penis.

Hypothalamus

In hypothalamus, hypothalamic releasing factors (HRF) inhibit or stimulate the secretion of pituitary hormones, influencing hunger, thirst, body temperature, circadian cycles, and last but not the least sexual drive.[13]

The hypothalamus has a role in penile erection. The erection function is derived by the two nuclei: dorsomedial hypothalamic nucleus (DMHN) and ventromedial hypothalamic nucleus (VMHN). The DMHN projects into the mesencephalic reticular formation. Both the DMHN and VMHN synapse and reach the lumbosacral autonomic centers (involved in penile erection) via the dorsolateral funiculus of the spinal cord.[14] In bottom-up control manner, hypothalamic nuclei also receive information from the genital regions.[15] Additionally, paraventricular hypothalamic nucleus (PVHN) is shown to be involved in crucial sexual activities such as copulation and orgasm.[16] Moreover, recently deep sexual identity has been correlated to bilateral hypothalamic activation in recent fMRI study.[14]

Amygdala

The amygdala has been implicated in processing mental (and emotional) aspects of sexual-arousal. Amygdala has widespread cortical and subcortical connections and thus may play an important role in sexual drive.[17] After understanding emotional relevance, stimuli are conveyed from amygdala to prefrontal cortex (PFC) and orbitofrontal cortices (OFC). Also projections to both to hypothalamus and nucleus accumbens (NAc) (key areas regulating sexual behavior) are activated. Bilateral lesions of the amygdala have been responsible for abnormal sexual behavior states such as those of Kluver–Bucy syndrome.[18] Lastly, the activation of the amygdala has shown a positive correlation with the erotic stimuli and with the consequent sexual-arousal.[19]

Cortical Areas

Several cortical areas are involved in the conscious processing of sexual stimuli. These areas also help in regulating the appropriate responses to sexual desire.[17] Here, we look at some of the cortical areas playing an important role in sexual behavior.

Prefrontal and Orbitofrontal Cortices

It is well known that these frontal regions play a critical role in moral affiliations, self-awareness, and self-other relations.[20,21] Sexual inhibition is an adaptive response serving

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as the critical social endpoint.[22] Sexual inhibition represents an approach–avoidance conflict, where the reward expectation drives the desire, but aversive consequences blunt the initiation of behavior.[23] These inhibitory systems are located in the PFC. PFC may inhibit the activation of excitatory mechanisms and subsequently shifting attention to nonsexual stimuli situations. Similarly, during high sexual arousal activity like genital stimulation, there is a steady decrease in ventromedial PFC and medial temporal cortex activity.[24,25] Moreover, OFC regions play a role in pleasant body representation and euphoric feelings during peak of sexual cycle. Finally, both PFC and OFC via its connections to subcortical structures (reward system) are responsible for the cognitive filtering of sexual behavior.[26]

Cingulate Cortex

The cingulate cortex is divided into anterior, a middle, and a genual subregion. The anterior cingulate cortex (ACC) is further divided into two areas (functionally segregated), i.e., ventral anterior cingulate cortex (vACC) and dorsal anterior cingulate cortex (dACC) regions. Numerous studies point has revealed the significant role of the ACC in sexual behavior. ACC activation in response to erotic stimuli suggests its involvement in processing sexual stimuli (in divergent contexts). Sequentially, postactivation ACC would augment decision-making through its efferent outputs to motor-related areas and periaqueductal gray.[27] Dorsal anterior cingulate activation, in turn, has been attributed to play an important role in modulation of skeleto-motor activities (which characterize sexual arousal) and the professed urge to act.[17] But, activity in subgenual region of cingulated cortex has been reported to negatively correlate with arousal.[19] Lastly, increased cingulate cortex activation has been observed during erection in males and during orgasm in females.[28]

Insula

The insula is a center of the salience network processing.[29] Insula processes the sensorial stimuli and relays them to other cortical areas.[29] Thus, it would facilitate attention and working memory.[29] Insula (akin to cingulate cortex) conveys the integrated information to brainstem structures (to regulate autonomic responses) and to the motor areas facilitating the suitable responses to sexual stimuli.[30]

The insula and its role in male sexual behavior has been supported by various functional neuroimaging studies. Role of insula in perceptual processing of penile inputs, onset and recognition of erection, and sustain penile response to erotic stimuli have been mentioned.[31] Anterior and posterior insula have been found to be active during the desire and arousal phase respectively.[25] As mesolimbic dopaminergic projections reach both the insula and ACC (as well as NAc), the reward processing related to sex-related stimuli is further reinforced.[32]

NEUROCHEMISTRY AND SEXUAL FUNCTION

Within the past decade, research has directed increasing attention to the neurobiology of sexual function. Various key neurochemicals involved in the sexuality are briefly mentioned in **Table 4**.

PHYSIOLOGY AND THE SEXUAL RESPONSE CYCLE

The brain and other sex organs respond to sexual stimuli in a universal fashion known as the SRC.[8] The SRC is composed of four phases:

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TABLE 4: Role of neurotransmitters and hormones in sexuality.

Serotonin	• 5-hydroxytryptamine (5-HT) is released in the brain by raphe nuclei.[32] *Caudal* raphe nuclei axonal efferents terminate in the cerebellar cortex, deep cerebellar nuclei, and the spinal cord. Whereas, *rostral* raphe nuclei axonal efferents terminate in the striatum, thalamus, hypothalamus, NAc (nucleus accumbens), hippocampus, amygdale, cingulate gyrus, and neocortex[33] • 5-HT has very importantly an inhibitory effect on sexual function.[34] That is why selective serotonin reuptake inhibitor class (SSRI) can impair all the stages in sexual response cycle, i.e., desire, erection, and ejaculatory/orgasmic function • 5-HT1A receptors which are present in the dorsal horn and dorsal gray matter commissure, may modulate ejaculation via spinal processing of sensory information to the brain[35]
Dopamine	• Ventro-striatal DA has a crucial role in the motoric aspect of copulation. But, it does not have any active role during precopulatory exposure to receptive partner. Dopamine in the striatum disembarks the brakes leading to disinhibition of motor cortex and elicitng movements[35,36] • The prosexual role of dopamine is certainly stamped by the fact that dopaminergic drugs have been known to enhance male sexual function clinically • What physiological sexual changes take place when increased DA levels from MPAO? – *Mild*: Disinhibition of genital reflexes (via D2 receptor family) – *Moderate*: Enhance copulatory behavior due to parasympathetically mediated erections (via D1 receptors) – *Large*: Promote sympathetically mediated ejaculation but would inhibit erections[35,36]
Norepinephrine (NE)	• Adrenergic activity makes penis flaccid and detumescence. Inhibition of the α1 adrenergic receptors present in the human penile tissue creates an erection. Studies reporting the effects of drugs that act on NE receptors indicate that the monoamine is important in male sexual function. NE levels balance the negative impact of 5-HT on human sexuality as per homeostatic principle. Therefore, the newer classes of antidepressant like desvenlafaxine that act on NE neurotransmission have lower incidence of sexual dysfunction than SSRI[37-40] • Acetylcholine (Ach) in conjunction with the vasoactive intestinal peptide (VIP) has positive physiological role in penile erection. The human corpus cavernosum is richly innervated by cholinergic nerves. Ach via its endogenous activity relaxes the smooth muscles (corpus cavernosum) and increase in the blood flow into the penile tissue[39]
Nitric oxide (NO)	• Nitric oxide has an important role in penile and clitoral vasocongestion and tumescence. Increased NO production leads to release of guanylate cyclase which leads to of conversion guanosine triphosphate to cGMP. Due to cGMP, smooth muscles of penis relax and there is increased blood flow to penis.[41] Partial or total disruption of this process may lead of erectile dysfunction • Sildenafil being a phosphordiesterase-5 inhibitor prolongs the action of cGMP and reduces erectile dysfunction[42]

Contd...

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Opioids	Opioids abuse generally results in loss of libido, erectile dysfunction, and delayed orgasms.[43,44] Long-term opioids usage may lead to range of sexual issues like mild loss of libido to complete loss of sexual functions. Opiate withdrawal results in increased frequency of morning erections, night falls, and a slow return to normal sexual drive. The exact mechanism is not clear, but persistent opioid intake leads to decrease in circulating luteinizing hormone (LH) and testosterone levels which results in sexual impairment[45]
Histamine	Cimetidine being a histamine antagonist produces decrease in libido and erectile dysfunction as per case reports. Also, psychogenic impotence has shown to reverse when histamine was injected in corpus cavernosum in males.[46] Histamine antagonist may reduce the uptake of testosterone[47]
Sex hormones	• Androgens have a crucial role in sexual function. They are required for penile tissue growth and development, and also erectile function maintenance • Sexual hormones by CNS integration of somatic and autonomic sexual systems play an important role in sexual arousal • Prolactin mediates gratification after sexual acts, although its higher levels could produce impotence and attenuate sexual[48]

(cGMP: cyclic guanosine monophosphate; CNS: central nervous system; DA: dopamine; MPOA: medial preoptic area; 5-HT1A: 5-hydroxytryptamine 1A)

1. Excitement

Activation of the sympathetic branch of the autonomic nervous system defines the excitement phase; heart rate and breathing accelerates, along with increased blood flow to the penis, vaginal walls, clitoris, and nipples.

2. Plateau

Blood flow, heart rate, and breathing intensify during the plateau phase. During this phase, often referred to as "foreplay," females experience an orgasmic platform—the outer third of the vaginal walls tightening—and males experience a release of preseminal fluid-containing healthy sperm cells.

3. Orgasm

The shortest but most pleasurable phase is the orgasm phase. After reaching its climax, neuromuscular tension is released and the hormone oxytocin floods the bloodstream resulting in emotional bonding. Although the rhythmic muscular contractions of an orgasm are temporally associated with ejaculation, this association is not necessary because orgasm and ejaculation are two separate physiological processes.

4. Resolution

The body returns to a prearoused state in the resolution phase. Males enter a refractory period of being unresponsive to sexual stimuli. The length of this period depends on age, level of intimacy with a partner, and novelty. Because females do not have a refractory period, they have a greater potential of having multiple orgasms.

CONCLUSION

A basic understanding of anatomy and physiology is essential for physicians dealing with various cases in sexual medicine. Anatomical and physiological changes occurring during an individual's lifetime also impact sexual health and functioning. Understanding the nervous system's role in sexual functioning is helpful as it plays

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the most crucial role in all aspects of sexual functioning. Many patients presenting for consultations in OPDs have several myths and misconceptions related to normal anatomical and physiological processes of sexual organs. A lack of sex education and the general availability of misinformation causes such situations.

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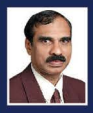

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