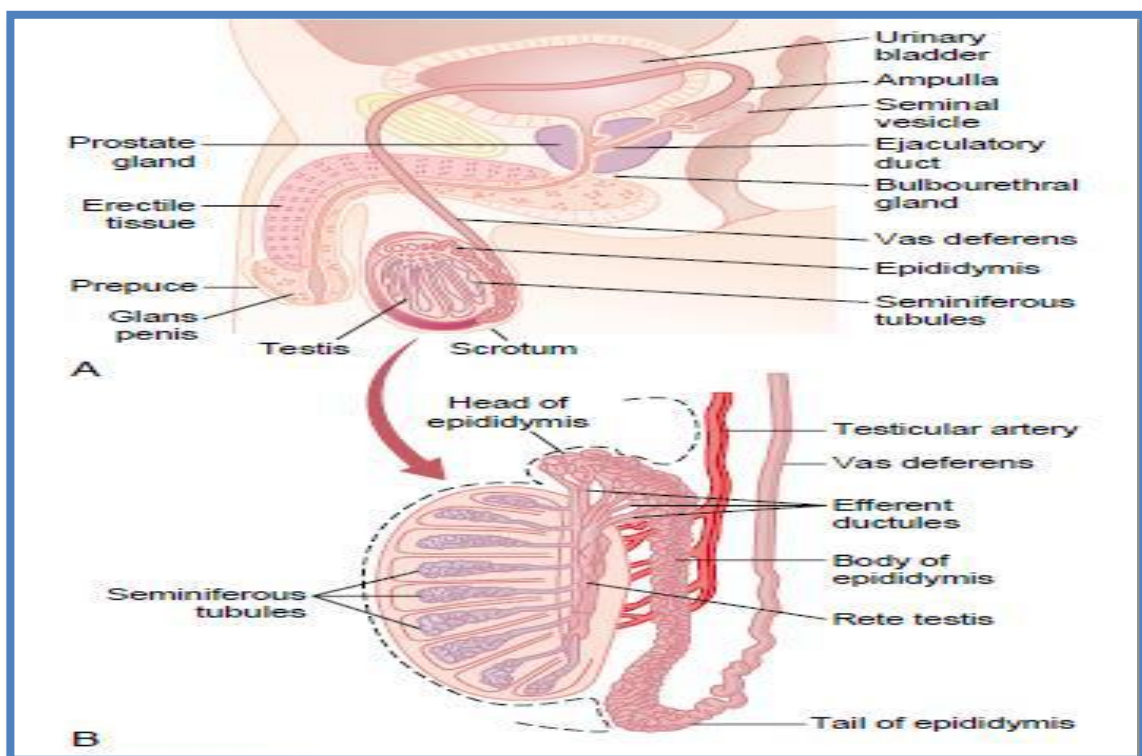


Testicle

The **testicle** or **testis** is the male reproductive gland in all animals, including humans. It is homologous to the female ovary. The functions of the **testes** are to produce both **sperm** and **androgens**, primarily **testosterone**. **Testosterone** release is controlled by the **anterior pituitary luteinizing hormone**; whereas sperm production is controlled both by the **anterior pituitary follicle-stimulating hormone** and **gonadal testosterone**.

Appearance

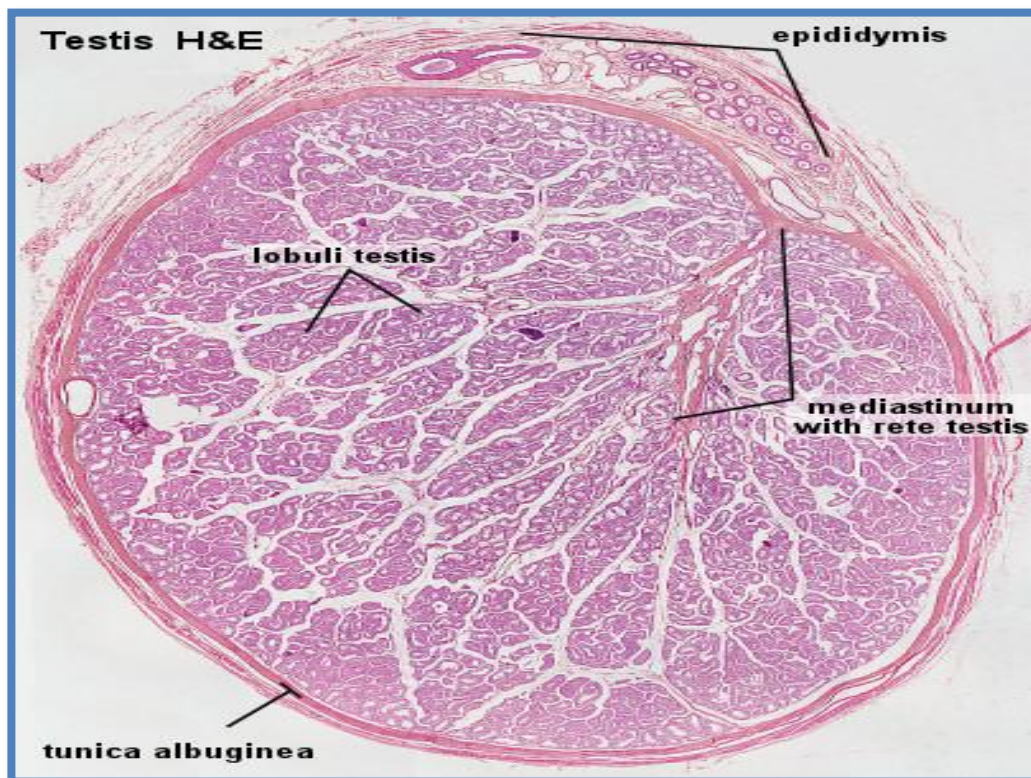
Males have two **testicles** of similar size contained within the **scrotum**, which is an extension of the abdominal wall. Scrotal asymmetry is not unusual: one **testicle** extends further down into the **scrotum** than the other due to differences in the anatomy of the vasculature.



Internal structure

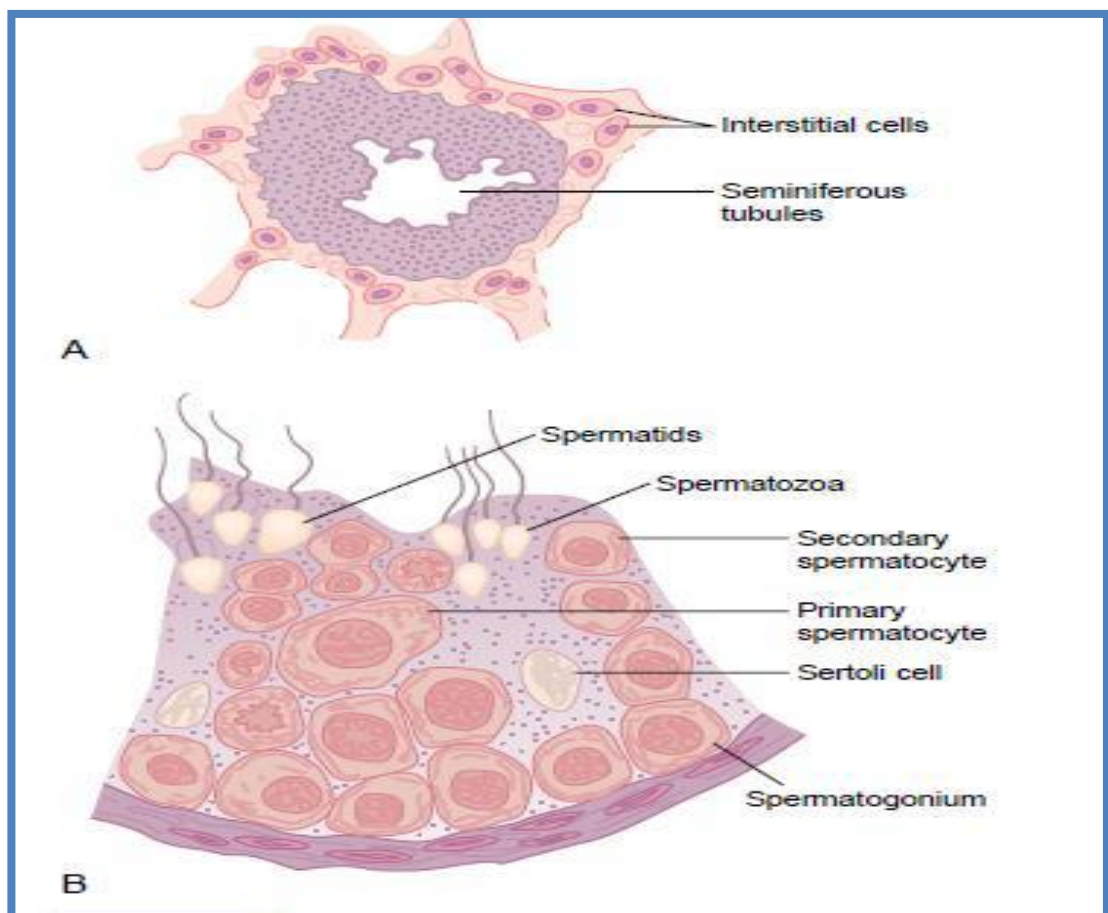
Duct system

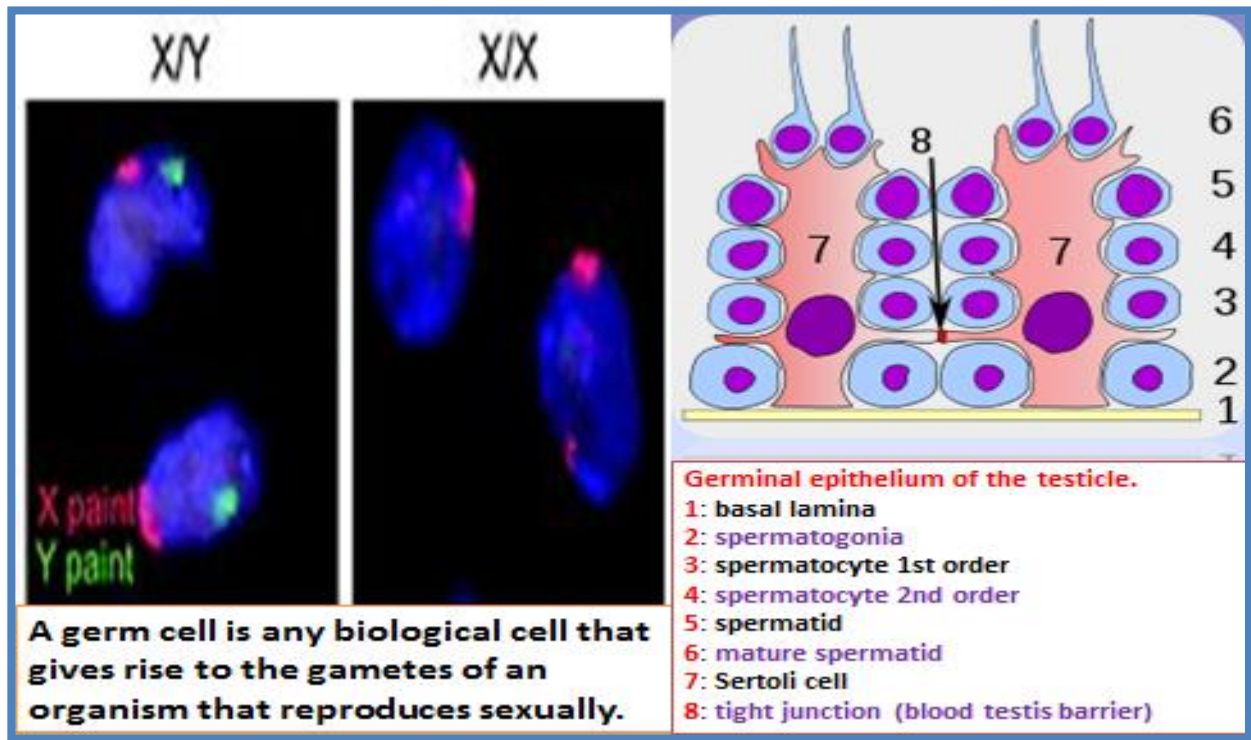
The **testes** are covered by a tough membranous shell called the **tunica albuginea**. Within the **testes** are very fine coiled tubes called **seminiferous tubules**. The tubules are lined with a layer of cells (**germ cells**) that develop from puberty through old age into **sperm cells** (also known as **spermatozoa** or **male gametes**). The developing sperm travel through the **seminiferous tubules** to the **rete testis** located in the **mediastinum testis**, to the **efferent ducts**, and then to the **epididymis** where newly created **sperm cells** mature (**spermatogenesis**). The sperm move into the **vas deferens**, and are eventually expelled through the **urethra** and out of the urethral orifice through muscular contractions.



Primary cell types within the seminiferous tubules

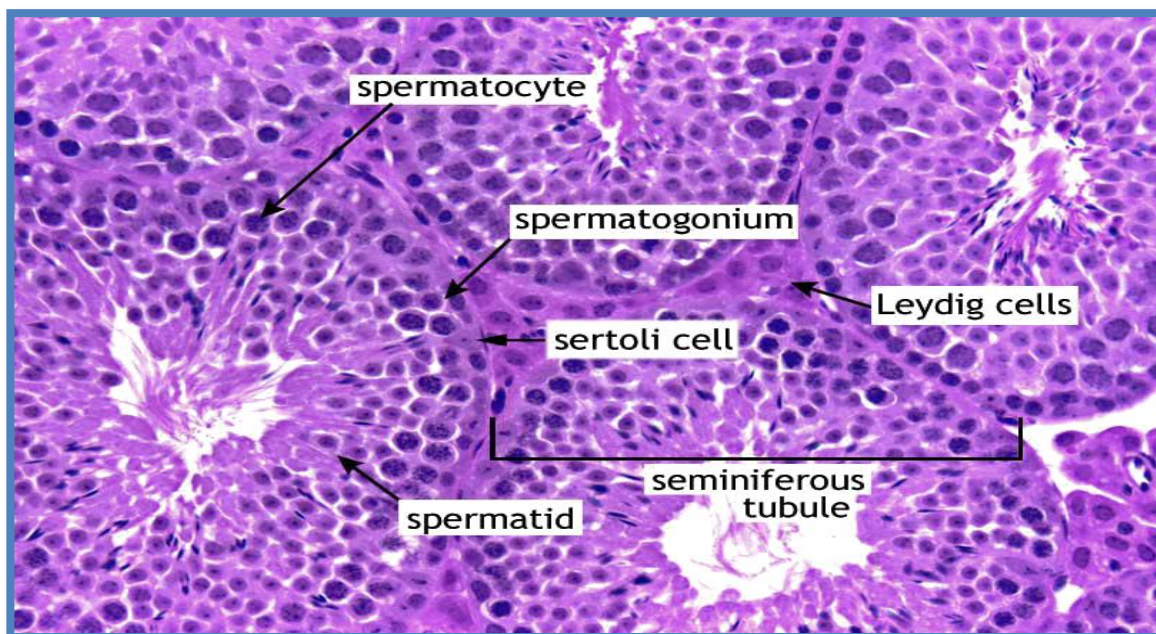
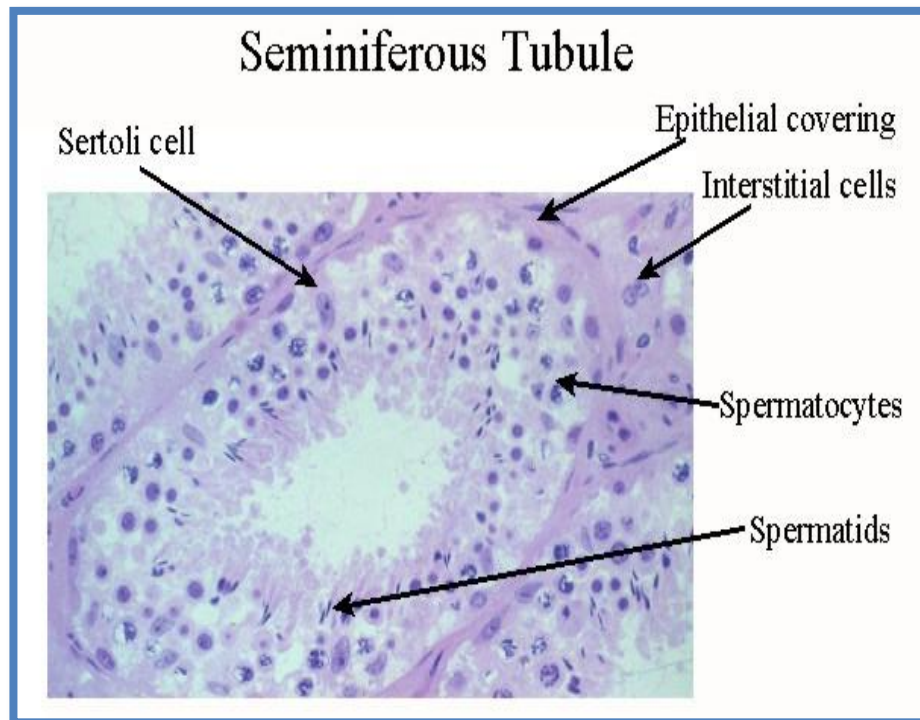
- **Germ cells** develop into **spermatogonia**, **spermatocytes**, **spermatids** and **spermatozoon** through the process of **spermatogenesis**. The gametes contain DNA for fertilization of an ovum
- **Sertoli cells** – the true epithelium of the seminiferous epithelium, critical for the support of **germ cell** development into **spermatozoa**. **Sertoli cells** secrete **inhibin**.
- **Peritubular myoid cells** surround the **seminiferous tubules**.





Primary cell types between tubules (interstitial cells)

- **Leydig cells** – cells localized between **seminiferous tubules** that produce and secrete **testosterone** and other **androgens** important for **sexual development** and **puberty**, **secondary sexual characteristics** like **facial hair**, **sexual behavior** and **libido**, supporting **spermatogenesis** and **erectile function**. Testosterone also controls testicular volume.
- **Also present are:**
 - ❖ **Immature Leydig cells**
 - ❖ **Interstitial macrophages** and **epithelial cells**

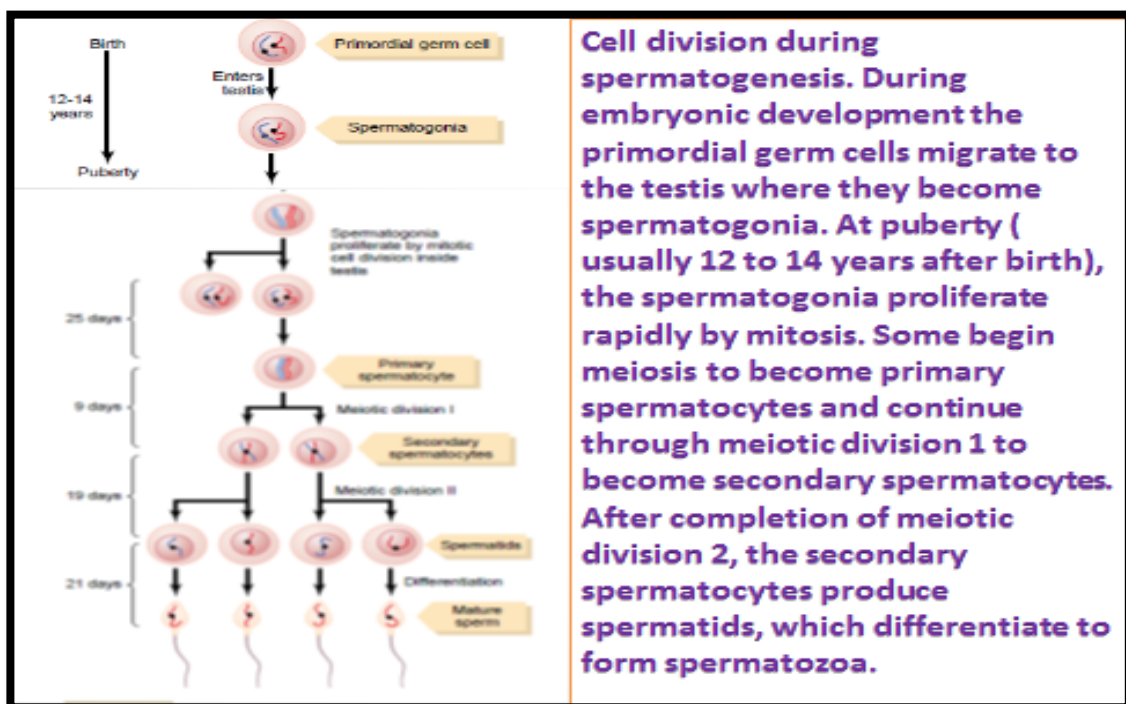


Layers

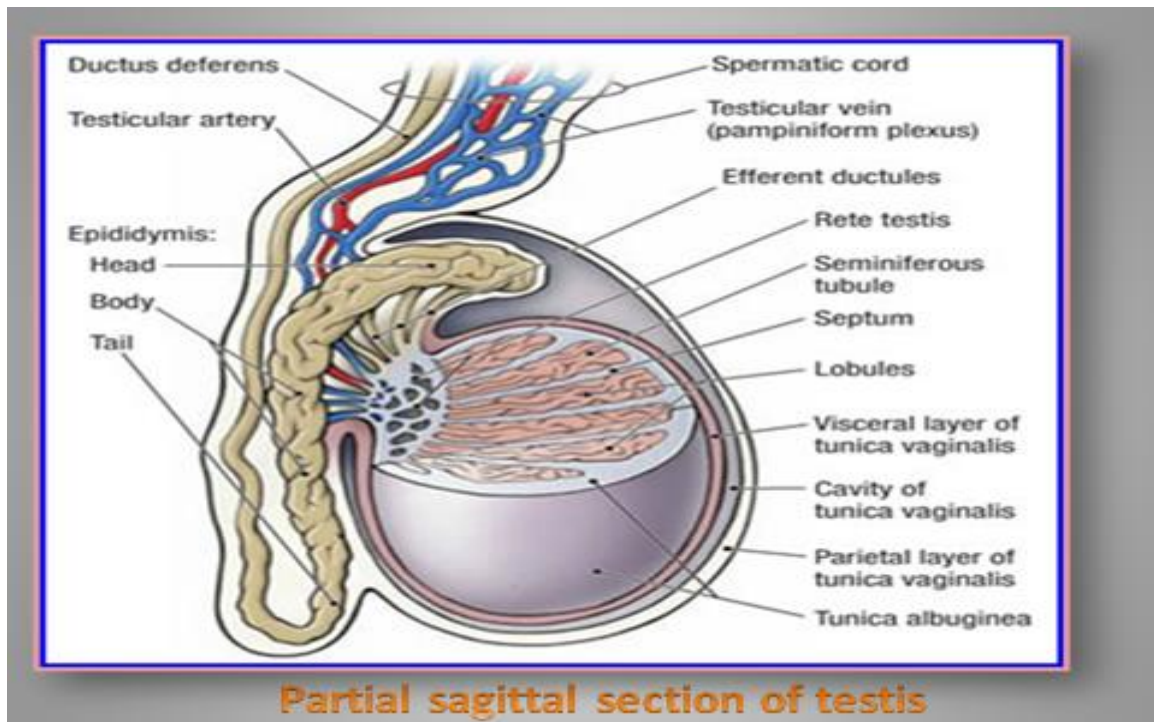
Many anatomical features of the adult **testis** reflect its developmental origin in the abdomen. The layers of tissue enclosing each **testicle** are derived from the layers of the anterior abdominal wall. Notably, the **cremasteric muscle** arises from the internal oblique muscle.

The blood-testis barrier

Large molecules cannot pass from the blood into the lumen of a **seminiferous tubule** due to the presence of tight junctions between adjacent **Sertoli cells**. The **spermatogonia** are in the basal compartment (deep to the level of the tight junctions) and the more mature forms such as primary and secondary **spermatocytes** and **spermatids** are in the **adluminal compartment**.



The function of the **blood-testis barrier** may be to prevent an auto-immune reaction. Mature **sperm** (and their antigens) arise long after immune tolerance is established in infancy. Therefore, since **sperm** are antigenically different from self tissue, a male animal can react immunologically to his own **sperm**. In fact, he is capable of making antibodies against them.



TEMPERATURE REGULATION

Spermatogenesis is enhanced at temperatures slightly less than core body temperature. The **spermatogenesis** is less efficient at lower and higher temperatures than **33 °C**. Because the **testes** are located outside the body, the smooth tissue of the scrotum can move them closer or further away from the body.

The temperature of the **testes** is maintained at **35** degrees Celsius (**95** degrees **Fahrenheit**) two degrees below the body temperature of **37** degrees Celsius (**98.6** degrees **Fahrenheit**). Higher temperatures affect **spermatogenesis**. There are a number of mechanisms to maintain the **testes** at the optimum temperature.

The **cremasteric muscle** is part of the spermatic cord. When this muscle contracts, the cord is shortened and the **testicle** is moved closer up toward the body, which provides slightly more warmth to maintain optimal testicular temperature. When cooling is required,

the **cremasteric muscle** relaxes and the **testicle** is lowered away from the warm body and is able to cool.

DEVELOPMENT

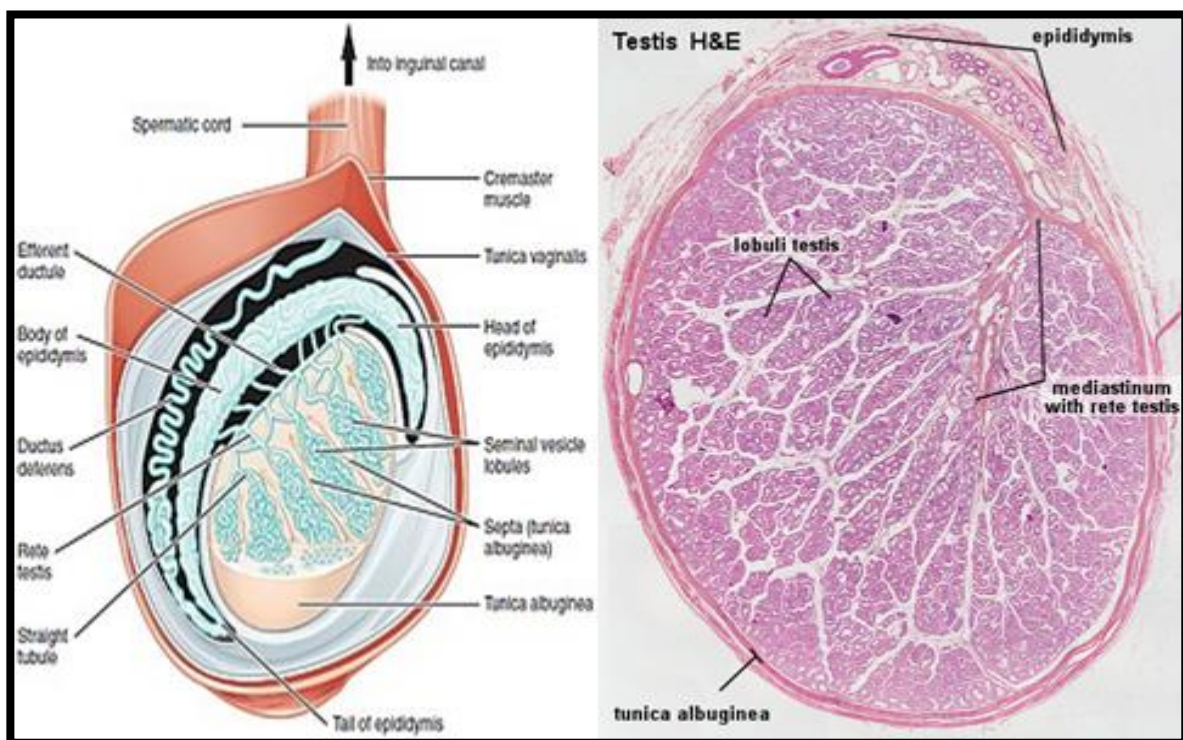
There are two phases in which the testes grow substantially; namely in embryonic and pubertal age.

1. Embryonic

During mammalian development, the gonads are at first capable of becoming either **ovaries** or **testes**. In humans, starting at about week **4** the gonadal rudiments are present within the intermediate **mesoderm** adjacent to the developing **kidneys**. At about week **6**, sex cords develop within the forming **testes**. These are made up of early **Sertoli cells** that surround and nurture the **germ cells** that migrate into the gonads shortly before sex determination begins. In males, the sex-specific gene **SRY** that is found on the **Y-chromosome** initiates sex determination by downstream regulation of sex-determining factors, which leads to development of the male phenotype, including directing development of the early **bipotential gonad** down the male path of development. **Testes** follow the "**path of descent**" from high in the posterior fetal abdomen to the **inguinal ring** and beyond to the **inguinal canal** and into the **scrotum**. In most cases both **testes** have descended by birth. In most other cases, only one testis fails to descend (**cryptorchidism**) and that will probably express itself within a year.

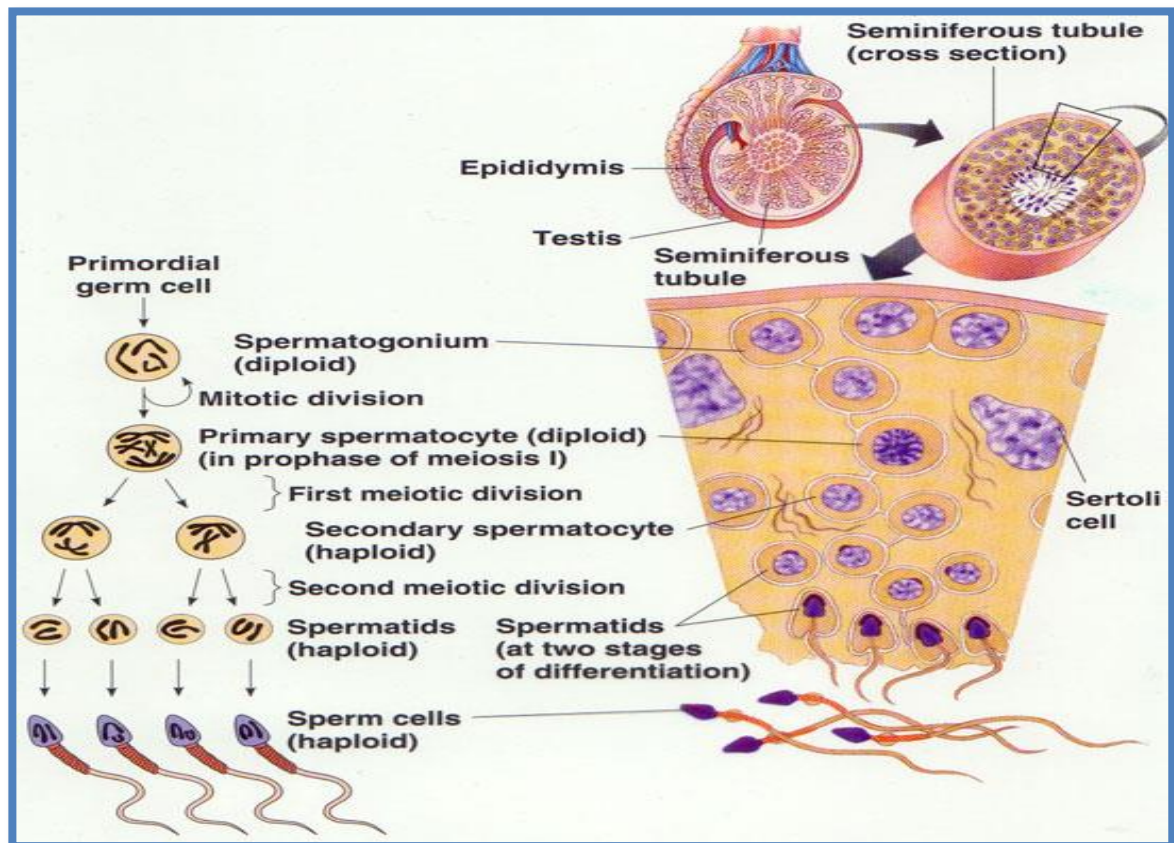
2. Pubertal

The **testes** grow in response to the start of **spermatogenesis**. Size depends on lytic function, **sperm** production, **interstitial fluid**, and **Sertoli cell fluid** production. After puberty, the volume of the **testes** can be increased by over **500%** as compared to the pre-pubertal size. **Testicles** are fully descended before one reaches puberty.



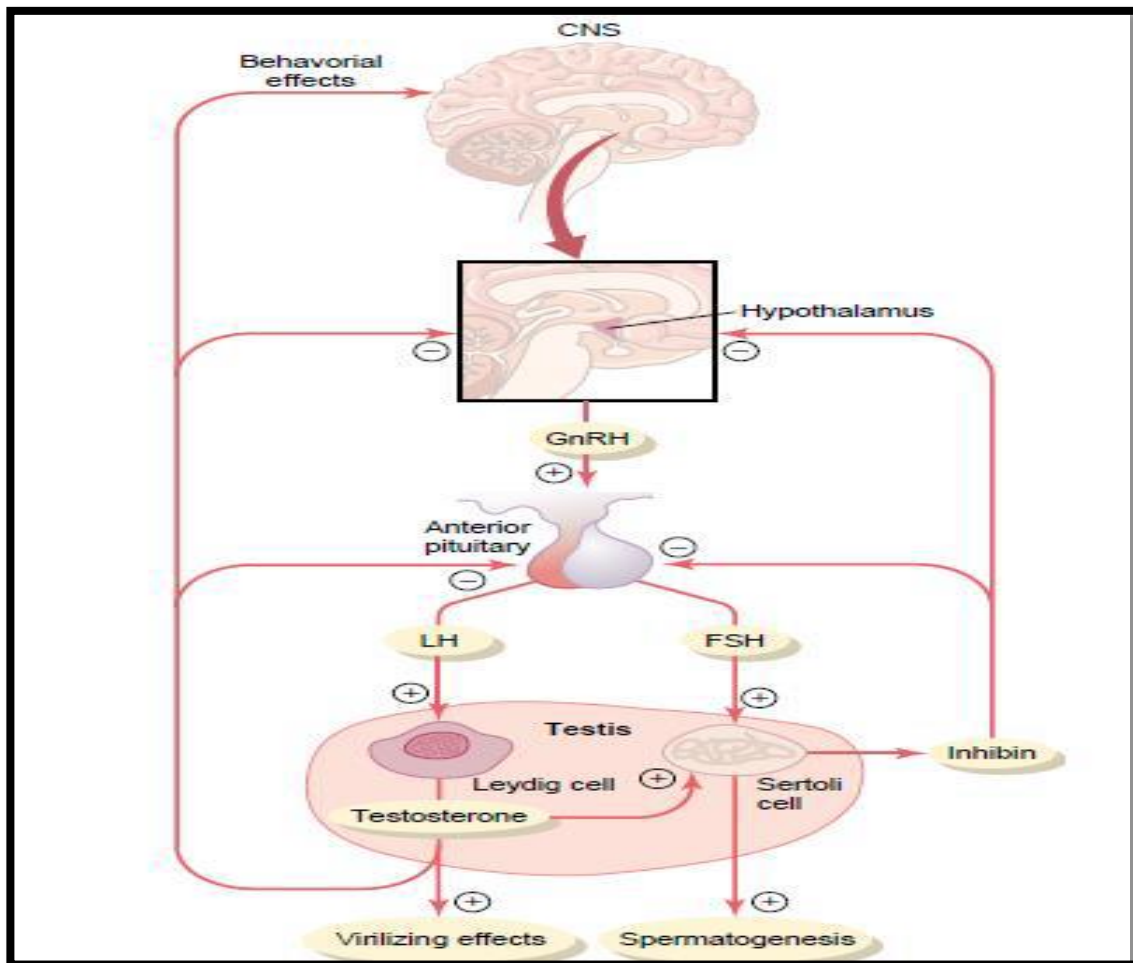
The **testes** of the male begin to produce **testosterone** at puberty in response to **LH**. **Testosterone** is the primary male sex hormone and an anabolic steroid, it plays a key role in the development of male reproductive tissues such as testes and prostate, it promotes maturation of the male reproductive organs, development of secondary sex characteristics (facial and body hair, also deepens the voice of a male at a certain age and maintaining healthy levels of muscle and bone mass) and production of sperm by the **testes**.

Testosterone is biosynthesized in several steps from cholesterol and is converted in the liver to inactive metabolites. It exerts its action through binding to and activation of the androgen receptor.

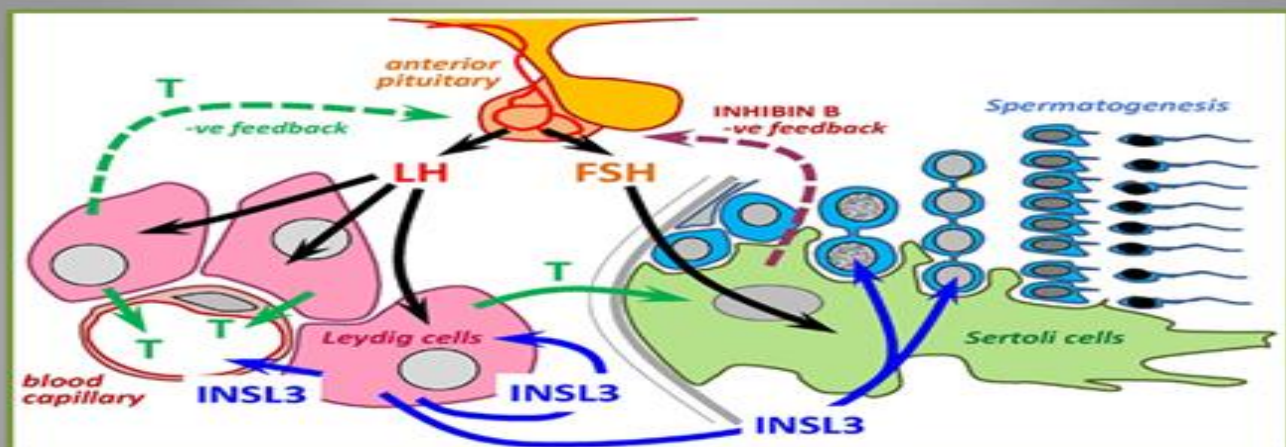


THE TESTES PRODUCE THREE MAIN HORMONES.

| Hormone | Produced by | Regulation | Action |
|--------------|---------------|--|---|
| Testosterone | Leydig Cells | GnRH from the Hypothalamus causes LH secretion from the Pituitary Gland which stimulates the Leydig Cells | This hormone controls and maintains the growth and functions of the reproductive organs. It enhances libido and is essential for spermatogenesis. |
| Inhibin | Sertoli Cells | GnRH from the Hypothalamus causes FSH secretion from the Pituitary Gland which stimulates the Sertoli Cells. | Prevents secretion of further FSH from the Pituitary Gland. |
| Oestradiol | Sertoli Cells | GnRH from the Hypothalamus causes FSH secretion from the Pituitary Gland which stimulates the Sertoli Cells | Converted from testosterone, this hormone's function is complex. It may prevent apoptosis of male germ cells. |



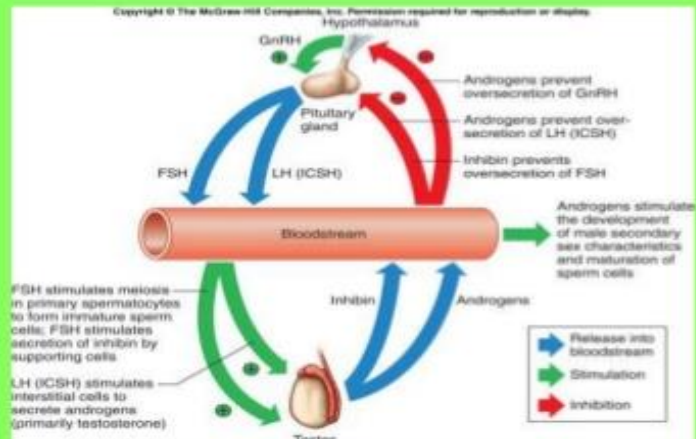
Feedback regulation of the hypothalamic-pituitary-testicular axis in males. Stimulatory effects are shown by \oplus and negative feedbacks inhibitory effects are shown by \ominus . GnRH, gonadotropin-releasing hormone; LH, luteinizing hormone; FSH, follicle-stimulating hormone.



The hypothalamic-pituitary-gonadal (HPG) axis

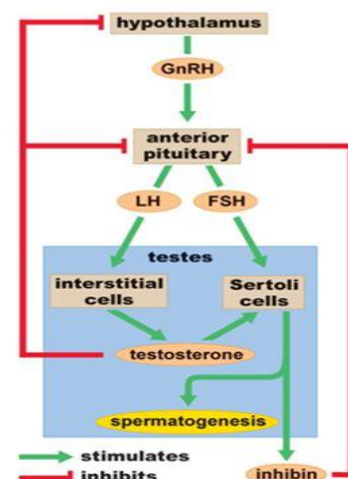
Regulation of Testosterone

- Hypothalamus regulates through negative feedback
- Increasing Concentration
 - Inhibits hypothalamus
 - stimulation of anterior pituitary gland decreases
 - As secretion of LH falls, testosterone release from cells decreases
- Decreasing Concentration
 - Hypothalamus stimulates anterior pituitary gland to release LH
 - Secretion causes interstitial cells to release testosterone



Testicular function is regulated by negative feedback

- Testosterone inhibits the release of GnRH, LH, and FSH
- The Sertoli cells are stimulated by FSH and testosterone to secrete inhibin, which inhibits FSH production by the anterior pituitary
- This complex feedback process maintains constant levels of testosterone and sperm production



Human Chorionic Gonadotropin Secreted by the Placenta During Pregnancy Stimulates Testosterone Secretion by the Fetal Testes

During pregnancy the hormone *human chorionic gonadotropin* (hCG) is secreted by the placenta and circulates both in the mother and in the fetus. This hormone has almost the same effects on the sexual organs as LH. During pregnancy, if the fetus is a male, hCG from the placenta causes the testes of the fetus to secrete testosterone. This testosterone is critical for promoting formation of the male sexual organs, as pointed out earlier.

Puberty and Regulation of Its Onset

Initiation of the onset of puberty has long been a mystery, but it has now been determined that *during childhood the hypothalamus does not secrete significant amounts of GnRH*. One of the reasons for this is that, during childhood, the slightest secretion of any sex steroid hormones exerts a strong inhibitory effect on hypothalamic secretion of GnRH. Yet, for reasons still not understood, at the time of puberty, the secretion of hypothalamic GnRH breaks through the childhood inhibition and adult sexual life begins.