

HUMAN REPRODUCTION AND EMBRYONIC DEVELOPMENT

- Human beings reproduce sexually and are viviparous i.e. they give birth to young ones. The reproductive units in sexual reproduction are specialized cells called gametes.

PRIMARY SEX ORGANS

- Primary sex organs are those sex organs which produce gametes. They are also called gonads.
- They also produce steroid hormones essential for reproduction as well as the growth and development of the entire body.

SECONDARY SEX ORGANS

- Sex organs, glands and ducts which do not produce gametes and hormones but are otherwise essential for sexual reproduction are known as secondary sex organs.
- Secondary sex organs of human male reproductive system are vasa efferentia, epididymes, vas deferentia, ejaculatory ducts seminal vesicles, cowper's gland, urethra, prostate gland and penis.
- Secondary sex organs of human female reproductive system are fallopian tubes, uterus, vagina, external genitalia and mammary glands.

ACCESSORY OR EXTERNAL OR SECONDARY SEX CHARACTERS

- They are those external features which provide distinctiveness to the two sexes.
- Major accessory or external or secondary sex characters of human male are facial hair, body hair, more height, more muscles, broadening of shoulder, low pitched voice, narrow but strong pelvis
- Major secondary sex characters of human female are development of breasts, broader pelvis, rounded body contours, fat deposition in thighs, buttocks and face, high pitched voice, pubic hair and sterna breathing

PUBERTY

- Beginning of sexual maturity or ability to reproduce is known as puberty. It is the period when primary sex organs become functional and start secreting sex hormones which bring about developments of secondary sex organs and appearance of secondary sex characters.

- Puberty is characterized by rapid growth. The average age of onset for girls is 10 -14 years and for boys, 12 -15 years old
- In boys pubertal changes occur in response to testosterone, whereas in female child, estrogen is produced by ovarian follicles.

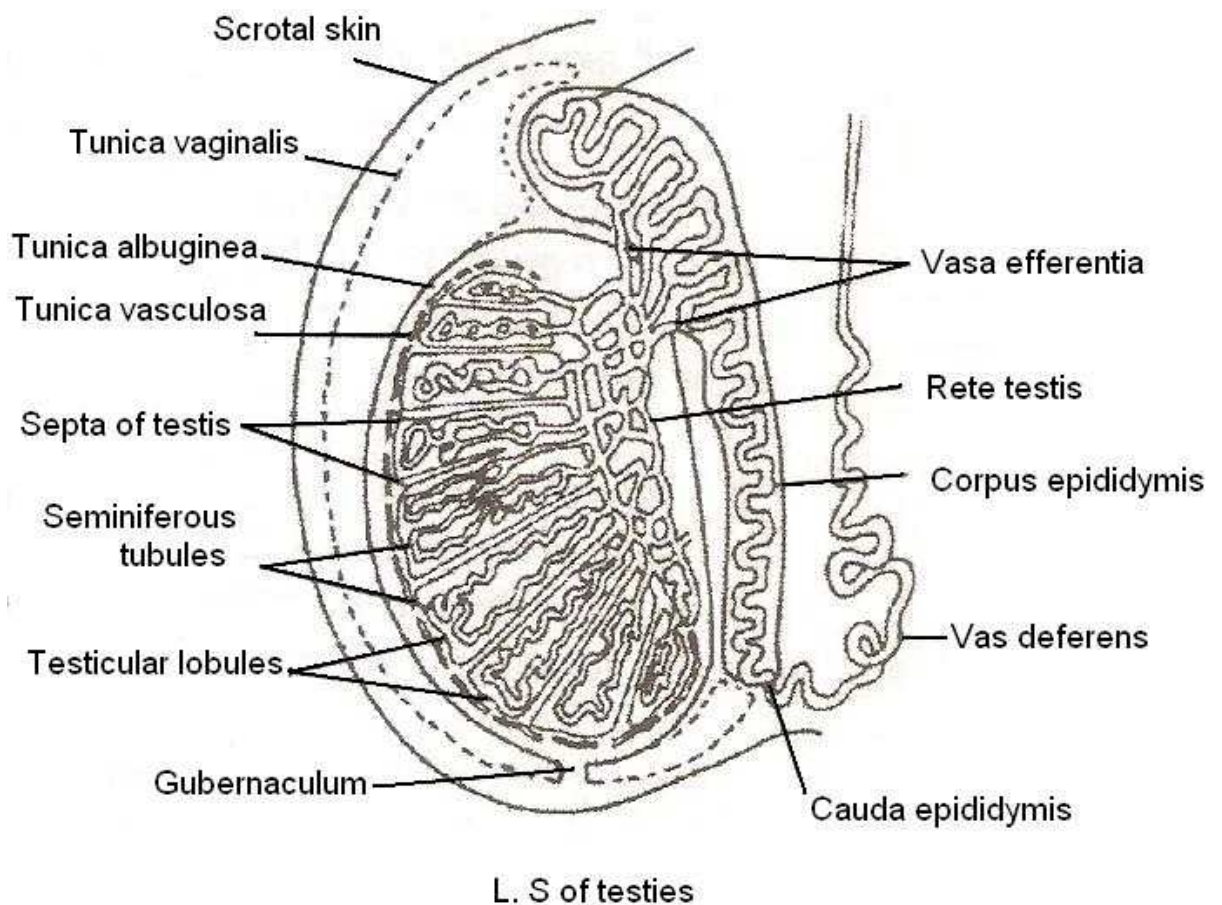
MALE REPRODUCTIVE SYSTEM

It is made of a pair of testes, scrotum, vasa efferentia, a pair of epididymes, a pair of vasa deferentia, a pair of seminal vesicles, a pair of ejaculatory ducts, urethra, prostate gland, a pair of Cowper's glands and penis

TESTES

- They are a pair of oval, pinkish primary sex organs of male reproductive system, each of with a size of 5cm (length), 3cm (thickness) and 2.5cm (width), weight about 12gm and lying obliquely in scrotum.
- During early foetal life, the testes develop in the abdominal cavity just below the kidney then they descend into the scrotum.
- The testis is surrounded by three layers
 - (i) Tunica vaginalis, a bilayer of peritoneum with narrow coelomic cavity filled with coelomic fluid. It helps the testis in frictionless sliding
 - (ii) Tunic albuginea, is the actual covering of the testes. The covering is made up of dense bluish with fibrous or collagenous connective tissue
 - (iii) Tunica vasculosa, is delicate loose connective tissue which lines the testicular lobules inner to tunica albuginea. It has rich supply of blood vessels.
- Tunica albuginea also projects inside testes to form a thick incomplete vertical column called mediastinum and a number of transverse septa. The septa produces 200 – 300 conical testicular lobules and each lobules is filled with connective tissue and 1-3 yellow convoluted seminiferous tubules. A total of 900 – 1000 yellowish seminiferous tubule occur in each testis. Each tubule is about 70-80 cm long. Seminiferous tubule join to form 20-30 short straight ducts called tubuli recti. Tubuli recti enter a network of channels called rete testes.
- The connective tissue lying in between seminiferous tubules are called Leydig's cell and it contain yellow pigment granules. Leydig cells are large polyhedral cells which have eccentric nucleus and small lipid containing vacuoles. They secrete testosterone and other androgens.

- The lining of seminiferous tubules is formed by a single layered germinal epithelium. The epithelium has two types of cells, spermatogenic (primary germ cells) and sertoli cells (supporting cells), Spermatogenic cells form 4-8 layers. The cells are destined to undergo spermatogenesis and form spermatozoa.
- Sertoli cells are large, elongated and pyramidal cells with bases resting on basal lamina (basement membrane) and apics projecting into the lumen of the seminiferous tubules. They secrete spermatogenic substances for nourishing and differentiation of cells undergoing spermatogenesis. They also secrete hormone inhibin for controlling FSH secretion.



SCROTUM

- It is a pouch of deeply pigmented skin arising from lower abdominal wall below the pubic symphysis and hanging between and in front of groin part of the thighs. Scrotal skin bears sebaceous glands that produce a characteristic odour, sweat glands and nerve endings.

- Wall of scrotum has three layers – outer wrinkled skin, connective tissue and smooth muscles. An internal septum scroti divides the scrotum into two sacs, each for one testis. Left testis lies a bit lower than the right one. Scrotum possesses a smooth involuntary muscle called dartos muscle. A testis rests in its chamber over pad called gubernaculum.
- The scrotum remains connected with the abdomen or pelvic cavity by the inguinal canals. The spermatic cord, formed from the spermatic artery, vein and nerve, bound together with connective tissue passes into the testis through inguinal canal.
- The testes develop in the abdominal cavity during the 7th month of gestation descend permanently into the respective scrotal sacs
- The scrotum acts as a thermoregulator, maintaining the testes at a temperature 2°C lower than that of the body. Movement of dartos muscle help in changing position of testes to keep them at proper temperature. When the body is chilled the smooth muscle contracts and brings the testes closer to the pelvic cavity to get warmth
- In some person testes fail to descend in scrotum. The condition is called cryptorchidism. It results in sterility

VAS EFFERENTIA

- Rete testis is connected to caput epididymis by 12-20 fine tubules called vasa efferentia
- Their lining epithelium is pseudostratified. It has large columnar ciliated cells and small nonciliated cells with endocytic activity, ciliated cells help in conducting sperms. Tubuli recti, rete testis and vas efferentia constitute an intratesticular genital duct system

EPIDIDYMES

- The epididymis is a mass of long narrow closely coiled tubule which lies along the inner side of each testis. coiling forms three parts – upper caput epididymis or head middle corpus epididymis or body and lower cauda epididymis or tail
- In the head of the epididymis, the sperms undergo physiological maturation, acquiring increased motility and fertilising capacity. In the tail of the epididymis sperms are stored before entering the vas deferens

- Epididymis is lined by pseudostratified epithelium that secretes nutrients required for maturation of spermatozoa. Non-ejaculated sperms are broken down after an interval.

VAS DEFERENS

- The vas deferens is a continuation of cauda epididymis which leaves the scrotal sac and enters the abdominal cavity. After entering abdomen it loops over the urinary bladder and dilates to spindle like ampulla for temporary storage of spermatozoa. It also conducts sperms.

EJACULATORY DUCTS

- They are shorter ducts of about 2cm length where male ejaculate is produced. Each duct is formed by joining of vas deferens and duct of seminal vesicle
- Ejaculatory ducts enter the prostate gland and join the prostatic urethra to produce a single urinogenital duct. In ejaculatory duct, the sperm mix up with secretion of seminal vesicles. The walls of ejaculatory ducts are muscular to quickly conduct the ejaculate through urinogenital duct.

URETHRA

- It arises from urinary bladder and is about 20cm long, differentiated into three regions
 - (i) A short proximal prostatic urethra which is surrounded by prostate glands.
 - (ii) A very short middle membranous urethra without any covering
 - (iii) A long distal penile urethra that passes through a penis
- The urethra has internal sphincter of smooth muscle fibres at its beginning and external sphincter of striated muscle fibres around its membranous part

PENIS

- It is male erectile copulatory organ which transfers semen into reproductive tract of female during sexual intercourse
- The penis contains three cylindrical masses of erectile tissue – two dorsal corpora cavernosa and one ventral corpus spongiosum
- The corpus spongiosum, which contains the penile urethra, is enlarged at the distal end of the penis to form glans penis. It is covered with smooth skin, foreskin.

SEMINAL VESICLES

- They are a pair of lobulated contorted musculoglandular sacs of 5cm length between urinary bladder and rectum. Their ducts join the vasa deferens to form ejaculatory ducts.
- They produce an alkaline secretion which forms 60% of the volume of semen. The secretion of seminal vesicles contains fructose, citrate, prostaglandins, inositol and clotting protein.
- Alkaline nature of the acidic environment of the male urethra which otherwise would inactivate and kill sperms
- The prostaglandins stimulate uterine contractions and thus may help the sperm to be moved towards female's oviducts, where fertilization takes place. The clotting protein help semen coagulate after ejaculate.

PROSTATE GLAND

- It is a large grayish to red pyramidal gland of 4cm width and 3cm height that encloses a part of urethra including its junction with ejaculatory ducts
- The produces a milky slightly alkaline secretion which forms 25% of the volume of semen. It possesses calcium, phosphate, bicarbonates, enzymes, clotting enzymes, profibrolysin and prostaglandins. Secretion of the prostate gland nourish and activates the spermatozoa to swim. Prostaglandins helps in liquefying cervical mucus and helps in liquefying cervical mucus and stimulating movement in the female tract.

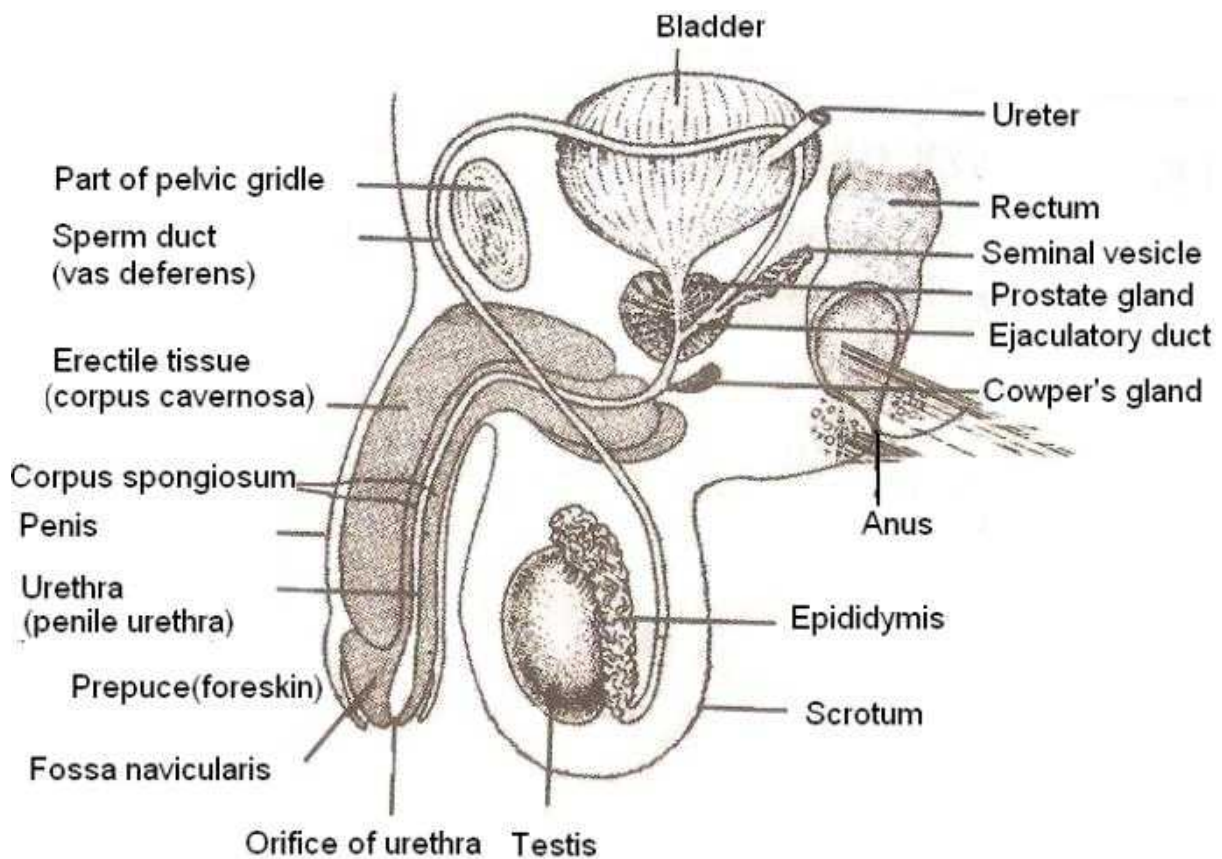
BULBOURETHRAL GLANDS

- A pair of small yellow pea seed sized lobulated tubalveolar glands, 4-5cm below prostate and opening into membranous urethra by separate ducts
- The secretion has abundant mucus for lubrication of reproductive tract. It neutralizes the urethra from remains of urine. Secretion of Cowper's gland is produced before the ejaculation of semen

SEMEN

- The secretion of accessory sex glands and mucus are added to sperms to form seminal fluid, or semen or seminal plasma
- It rich in fructose, calcium and certain enzymes. It has a pH of 7.35 to 7.5.

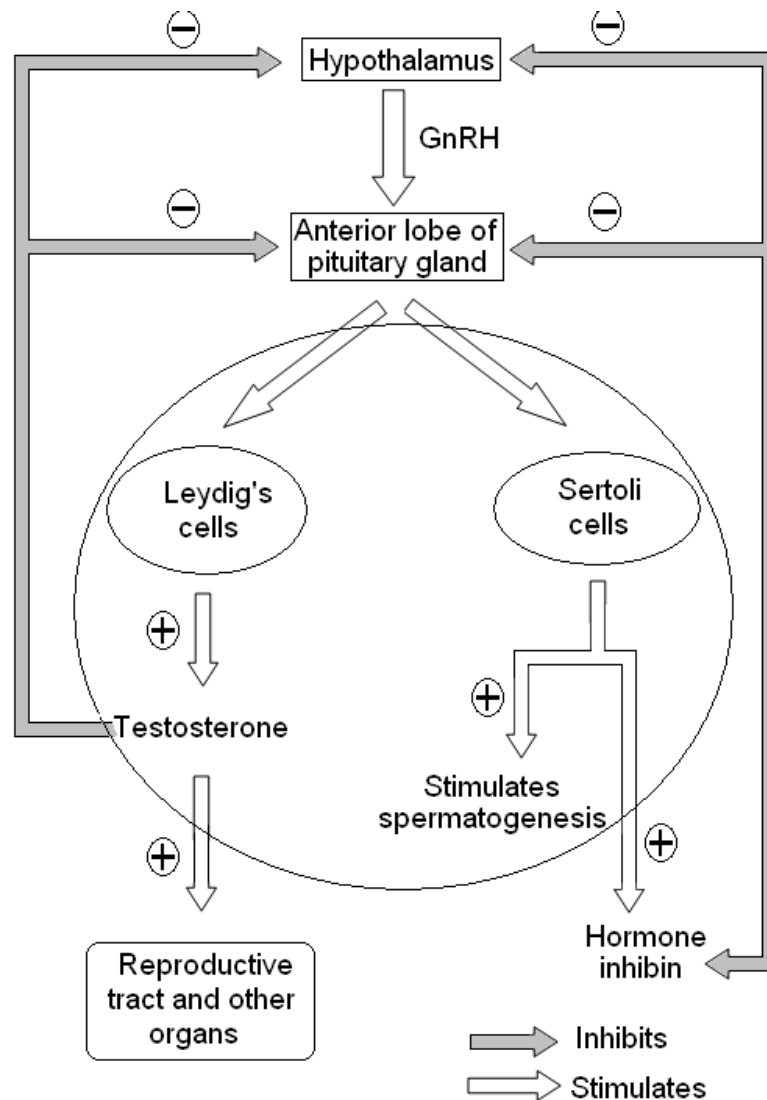
- It is ejected from the penis during ejaculation. A single ejaculation may contain 300 million sperms. It has many functions
 - (i) It provides a fluid medium for transmission of sperms into vagina of the female.
 - (ii) It nourishes and activate the sperms to keep them viable and motile
 - (iii) It neutralizes the acidity of the urine in the urethra of male and vagina of female to protect the sperms
 - (iv) It facilitates the sexual act by lubricating the reproductive tract of the female.



HORMONAL CONTROL OF MALE REPRODUCTIVE SYSTEM

- The growth, maintenance and functions of the male reproductive organs are under the hormonal control. GnRH (Gonadotropin releasing hormone) is secreted by hypothalamus. It stimulates the anterior lobe of pituitary gland to secrete and release LH and FSH. In male, LH is called interstitial cell stimulating hormone (ICSH) because it stimulates Leydig's cell of the testes to secrete androgens

- Testosterone is the principle androgen. The growth, maintenance and functions of secondary sex organs and functions of secondary sex organs and accessory glands are under the control of testosterone.
- FSH stimulates sertoli cells of testis to secrete an androgen binding protein (ABP) that concentrates testosterone in seminiferous tubule and also secrete a peptide inhibin which suppresses FSH synthesis. FSH acts directly on spermatogonia to stimulate sperm production.



COMMON DISORDERS OF MALE REPRODUCTIVE SYSTEM

- Prostatitis – It is inflammation of prostate generally caused by infection
- Benign prostatic hypertrophy (BPH) – This is the enlargement of prostate gland. It often causes in old age causing frequent night urination. Untreated BHP may lead to kidney damage.

- (iii) Prostate carcinoma – It is cancer of prostate which may grow unnoticed up to stage of metastasis
- (iv) Impotence – It is the inability of the adult male to achieve penile erection.
- (v) Sterility – Sperms are unable to fertilize the ovum due to low count or less motility.
- (vi) Cryptorchidism – It is a failure of one or both of the testicles to descend into scrotum. It is often results in pushing of an intestinal loop into scrotum, resulting in its abnormal size and discomfort.

FEMALE REPRODUCTIVE SYSTEM

The female reproductive system consists of a pair of ovaries, a pair of fallopian tubes, uterus, vagina, external genitalia and breasts

OVARIES

- They are a pair of almond shaped, solid, greyish pink gonads of the female. Each ovary is about 2-4 cm in length, 1.5cm in width and 1cm in the thickness. It is suspended from the dorsal body wall by a fold of peritoneum called mesovarium. It is held in position by ligaments which attach it to pelvic wall and uterus.
- Ovaries are differentiated into four parts
 - (i) Germinal epithelium is the outermost layer of the ovary which is formed of simple squamous and cuboidal cells
 - (ii) Tunica albuginea is poorly differentiated sheath of dense connective tissue that lies below the germinal epithelium and outside the cortex. Tunica albuginea provides greyish colour to ovary.
 - (iii) Cortex is a wide layer of connective tissue having a large number of spindle shaped fibroblasts, reticular fibres and ovarian follicles
 - (iv) Medulla is the central part of the ovary made of less dense connective tissue. Medulla is richly supplied with blood vessels. It has elastic fibres, smooth muscles.
- The ovarian medulla contains many rounded or oval bodies called ovarian follicles at various stage of development. The number of follicles in the two ovaries of young adult female is 1,20,000 – 1,60,000

FALLOPIAN TUBES (OVIDUCTS)

- They are pair of muscular and internally ciliated tubes of 10-12 cm length which lie horizontally over peritoneal cavity arising near and ending at uterus

- A fallopian tube is differentiated into four parts
 - (i) Infundibulum is a funnel shaped fimbriated free end of the oviduct. It opens into the body cavity by an aperture called ostium. Margin of the funnel bears motile, finger like projection called fimbriae. Fimbriae have folds lined with cilia that produce a slow current towards the ostium for passage of liberated ovum towards the inside of oviduct.
 - (ii) Ampulla is a curved dilated part of oviduct which is also the site of fertilization of ovum
 - (iii) Isthmus is the very short, narrow, thick walled, straight part of the oviduct which connects ampulla with lateral wall of uterus.
 - (iv) Uterine part is about 1cm long part that passes into uterine wall. Oviducal wall is composed of three layers – outer serosa of visceral peritoneum, middle muscularis and inner mucosa. Mucosa has an epithelium of ciliated and secretory columnar cells. Secretory cells produce a viscous secretion for nourishing and protection of ovum. Passage of ovum is facilitated by movement of cilia and muscular contractions of the wall.
- In most vertebrates both the ovaries and oviduct are functional. In birds the right ovary and right oviduct are atrophied

UTERUS

- It is the pyriform, hollow muscular thick walled but distensible median structure located above and behind urinary bladder that is meant for nourishing and development of foetus. For this, uterus is capable of tremendous enlargement. The empty uterus is 7.5cm, 5cm broad and 2.5cm thick
- Uterus is attached to pelvic wall by means of ligament
- Wall of uterus is differentiated into three layers
 - (i) Internal glandular endometrium
Endometrium has two parts, epithelium and lamina propia. Epithelium lines the luminal surface of uterus. It contains two types of columnar cell, ciliated and secretory. Lamina propia contains connective tissue with fibroblasts, tubular glands and blood vessels. Endometrium shows cyclic changes during menstrual cycle.
 - (ii) Myometrium is the middle layer of smooth muscles. It undergoes strong contractions during delivery of the baby
 - (iii) Perimetrium is an outer layer of uterus formed of either adventitia (connective tissue only) and serosa (connective tissue and mesothelium)

- Uterus is differentiated into three regions
 - (i) Fundus: Upper dome – shaped part above the opening of fallopian tube
 - (ii) Body : It is broad towards fundus and narrow down towards the cervix
 - (iii) Cervix: 2.5cm long narrow inferior extremity of uterus which protrudes into vagina. It is connected to the body by internal os. At birth it becomes irregularly bumpy. Upper part of uterus leans forwards. It is almost at right angle to vagina

VAGINA

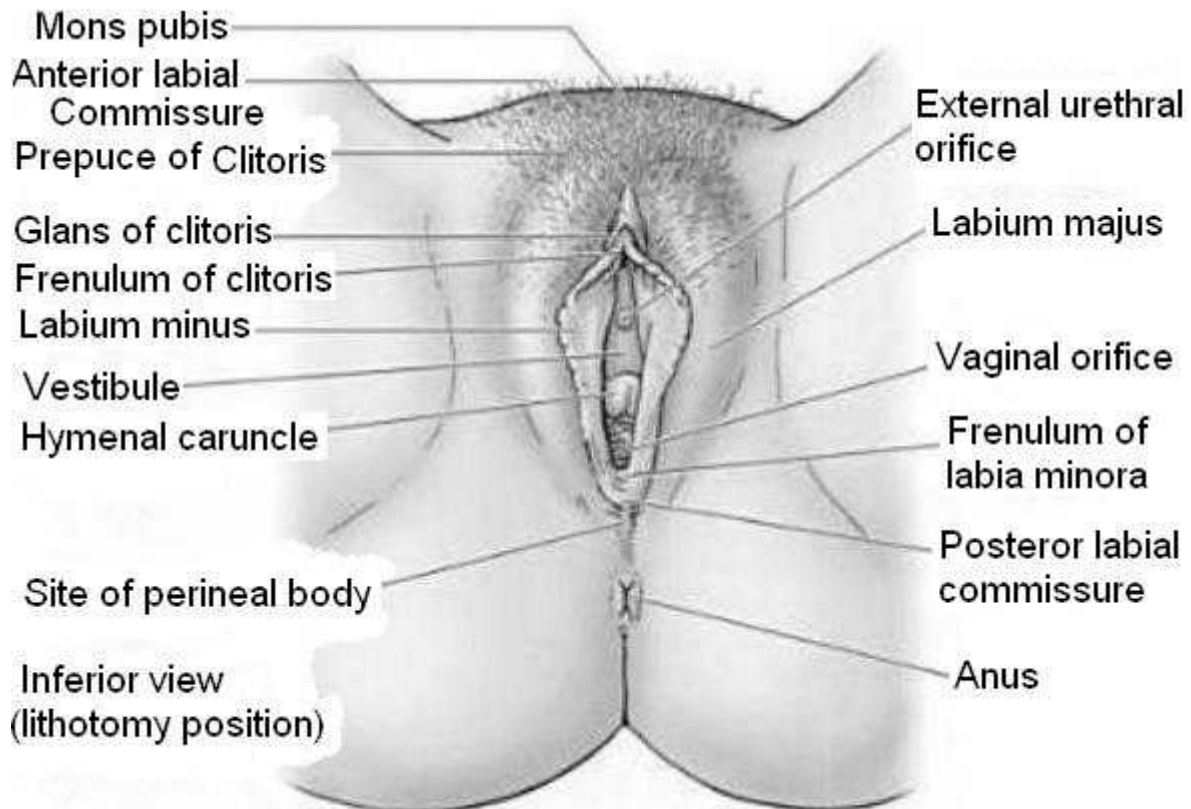
- It is an elastic muscular tube of 8-10cm length which functions as female copulatory organ, pathway for menstrual flow and birth canal. It is easily stretched
- The opening of the vagina is called vaginal orifice. Vaginal orifice is partially covered by a perforate membrane called hymen. It often gets ruptured during vigorous physical exercise or during sexual intercourse. In some it persists even after coitus

EXTERNAL GENITALIA

- The area having external genitalia is characterized by mons pubis on the upper side, perineum on the lower side and vestibule or depression in the centre
- Mons pubis is an eminence formed by fat over the pubic symphysis bones. Vestibule has urinary meatus with urethral opening on the upper side and vaginal orifice on the lower
- A small erectile organ, the clitoris, lies at the anterior junction of labia minora. It is homologous to the penis. It consists of a short shaft with erectile tissue. Its tip is round and of erectile tissue
- Vestibule is flanked by two pairs of fleshy folds of skin; the inner small, thin, moist labia minora which form clitoris in front and are connected behind by fourchette. They possess sebaceous gland. The outer larger, hair-covered labia majora. They also possess sebaceous gland.

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VESTIBULAR OR BARTHOLIN'S GLAND

- They are a pair of small tubulociner glands which open in the vestibule lateral to vaginal orifice. The secretion is thick viscid and alkaline for lubrication and counteracting urinary acidity. A number of small vestibular gland (paraurethral gland or gland of skene) are present on the either side of urethral opening.

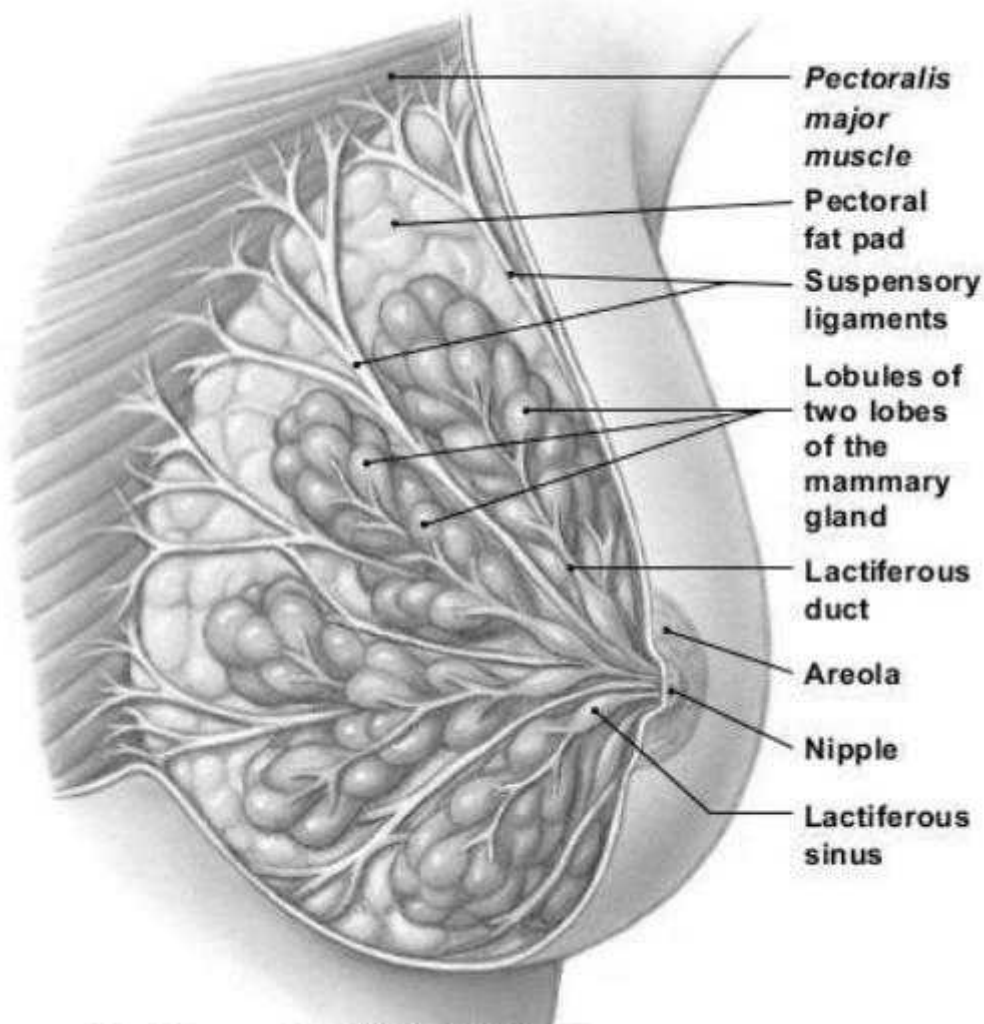
MAMMARY GLAND OR BREASTS

- Mammary glands or breasts are modified sweat glands that lie over the pectoral muscle which develops during puberty but becomes active only after child birth
- Each breast has a broad tip called nipple for the release of milk. A circular pigmented area called areola lies below it. Each breast contains 15-20 glandular lobes having secretory alveoli and each lobe is separated from each other by connective tissue and adipose tissue.
- The cells of alveoli produce milk which is stored in the cavities of alveoli. Alveoli opens into mammary tubules and then into mammary ducts.
- Mammary ducts forms mammary ampulla from which a lactiferous duct develops. Each lobe has a separate lactiferous duct and each opens at nipple by separate pores.

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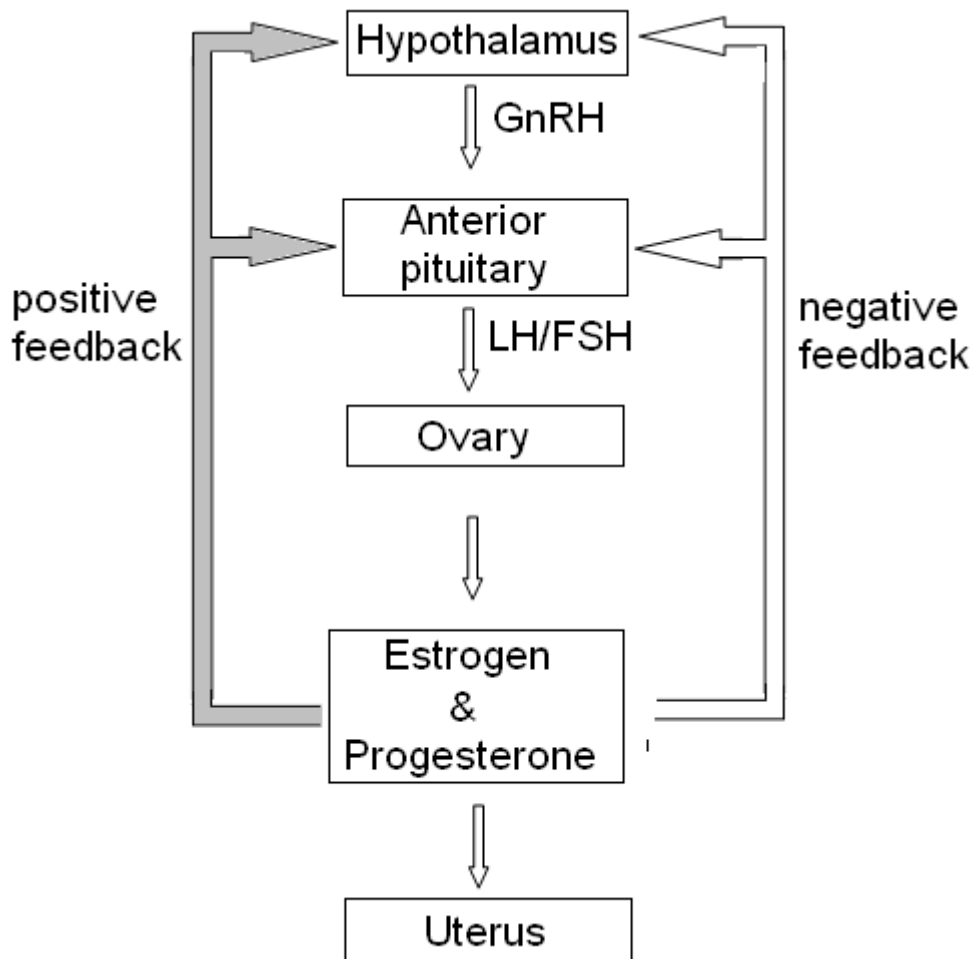
- During pregnancy, the glands grow under the influence of oestrogen and progesterone. On the infant's birth, the hormone prolactin stimulates the production of milk and hormone oxytocin causes release of milk as infant sucks the breast.



The Mammary Gland of the Left Breast.

HORMONAL CONTROL OF FEMALE REPRODUCTIVE SYSTEM

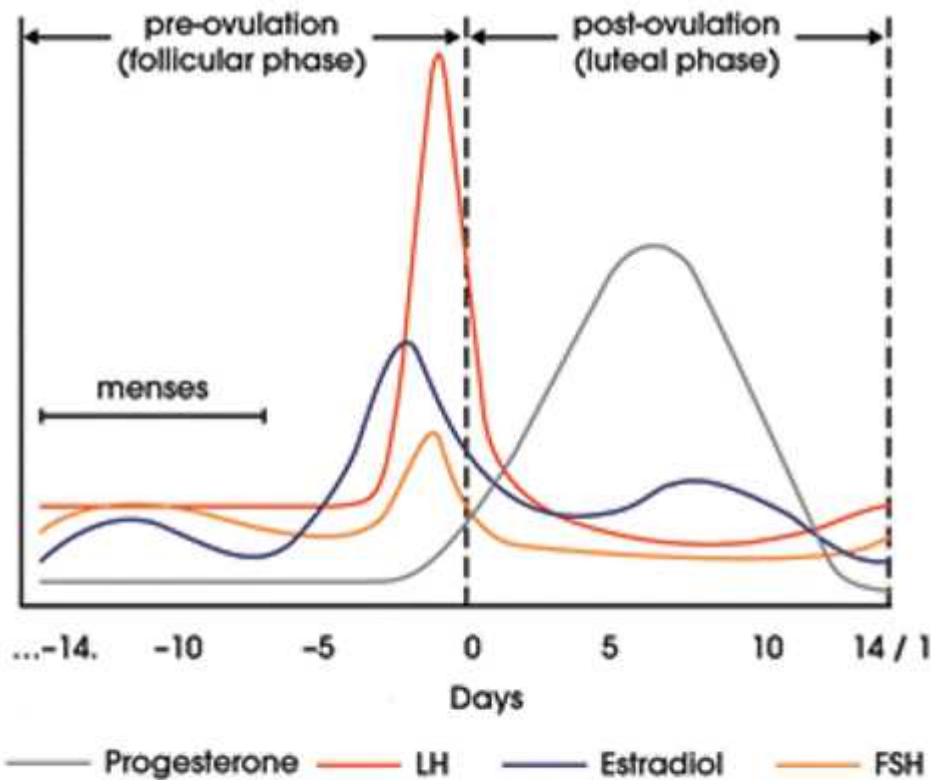
- GnRH secreted by hypothalamus stimulates the anterior lobe of pituitary gland to secrete LH and FSH. FSH stimulates the growth of ovarian follicles and also increase the development of egg(oocyte with the follicle to complete the meiosis I to form secondary oocyte. FSH also stimulates formation of oestrogen.
- Lutenising hormone (LH) stimulates the corpus luteum to secrete progesterone.
- Rising level of progesterone inhibits release of GnRH which in turn inhibits the production of FSH, LH and progesterone.



MENSTRUAL CYCLE

- It is a series of cyclic changes that occurs in the reproductive tract of human females with periodicity of 28 days, right from menarche to menopause except during period of pregnancy.
- The first menstrual period is called menarche. This occurs between the age of 12 to 15 years. Menopause is stoppage of menstrual period. It occurs between 45-55 years.
- Menstrual cycle is controlled by gonadotropin and ovarian hormones.
- Menstrual cycle has following phases:-
 - I. Menstrual phase
 - This phase lasts for about 4 days.
 - If fertilization do not occur progesterone secreted by persistent corpus luteum inhibits the release of LH from pituitary.
 - Reduction of LH causes subsequent fall in progesterone level.

- The uterine lining due to deficiency of progesterone causing rupture of blood vessels and leads to discharge of blood, a serous fluid, cell debris and mucosal fragments.
 - Lowered levels of progesterone causes release of FSH from the anterior pituitary. This initiates a new cycle.
- II. Follicular phase (proliferative phase)
- In the presence of FSH released by anterior pituitary, 6-12 ovarian follicles begin enlargement through proliferation of their granulosa cells to secrete estrogen. Estrogen stimulates the proliferation of the endometrium of uterine wall.
 - After about a week of development only one ovarian follicle continues growth while the rest degenerates.
 - Towards the end of proliferative phase the endometrium becomes about 3mm thick.
- III. Ovulatory phase
- High level of both LH and estrogen cause rapid growth of graafian follicle.
 - Graafian follicle rises to the surface of the ovary, produce a protuberance or stigma. The stigma ruptures and the ovum surrounded by corona radiata comes out in a viscous fluid. The process is called ovulation. It occurs after about 14 days.
- IV. Luteal or secretory phase
- Under the influence of LH the empty graafian follicle continues growth.
 - The follicular cells are converted into lutein cells by deposition of yellowish lipid inclusions. The phenomenon is called luteinisation. The ruptured graafian follicle is now called corpus luteum. It secretes progesterone.
 - Both LH and progesterone help in further growth and thickening of endometrium. Thickness of endometrium becomes 5-6mm.
 - Progesterone inhibits uterine movements as well as proliferation of new ovarian follicles. The phase lasts for about 10 days.



The "Normal" Pattern of Hormonal Changes
that Regulate the Ovarian Cycles

- If puberty starts early then it is known as precocious puberty.
- Dysmenorrhoea is painful menses.
- Menorrhagia is excessive menstruation.
- Oligomenorrhoea is infrequent menstruation.
- Amenorrhoea is non-occurrences of menses.

DISORDER OF THE HUMAN FEMALE REPRODUCTIVE SYSTEM

- Breast cancer: Breast cancer increases after menopause. The standard treatment for breast cancer is mastectomy (removal of breasts).
- Cervical cancer: It is relatively slow growing cancer. Cervical cancer may be treated by radiation or surgery.
- Oophorocystosis: Ovarian cysts are fluid filled tumors of the ovary. Such cysts sometimes rupture and regresses during pregnancy.
- Ectopic pregnancy: It is implantation of embryo at a place other than uterus, generally oviduct.
- Oophoritis: It is inflammation of ovary, usually caused by an infection.

(vi) Endometriosis: It is growth of endometrial tissue outside the uterus.

Treatment is usually hormone therapy or surgery.

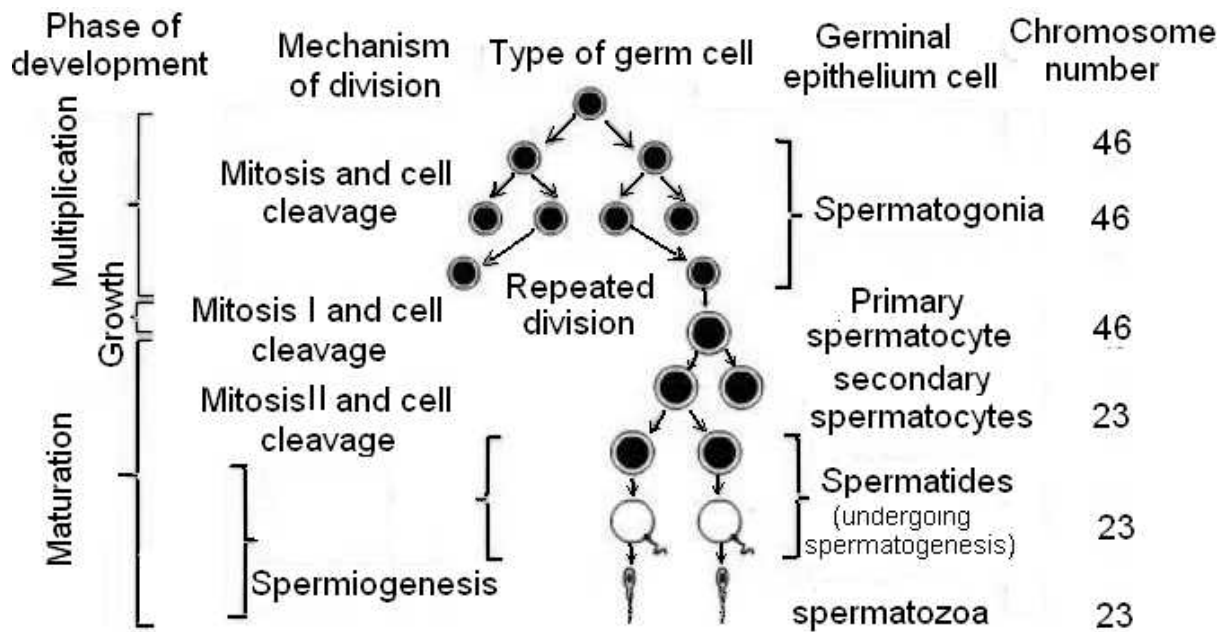
(vii) Infertility: Infertility in women is inability to become pregnant.

SPERMATOGENESIS

- Spermatogenesis is the process of formation of haploid spermatozoa (sperms) from diploid spermatogonia inside the testis of the male.

Formation of spermatids

- (i) Multiplication phase: At sexual maturity, the undifferentiated germ cells divide several times by mitosis to produce a large number of spermatogonia (2N) are of two types:-
Type A spermatogonia function as stem cells and produce more spermatogonia.
Type B spermatogonia divide mitotically to form primary spermatocytes. This process is called spermatocytogenesis.
- (ii) Growth phase: Primary spermatocytes grow to become almost double in size.
- (iii) Maturation phase: Each diploid primary spermatocyte undergoes meiosis I to form two haploid secondary spermatocytes. Each secondary spermatocyte divides by meiosis II, giving rise to two haploid spermatids. The spermatids become partially embedded in Sertoli cells for nourishment and support.
- (iv) Spermiogenesis: The transformation of spermatids into spermatozoa is called spermiogenesis. The different changes which occur during spermiogenesis are
 - a. Formation of acrosome by Golgi apparatus
 - b. Elongation and condensation of nucleus.
 - c. Separation of centrioles.
 - Spermatogenesis requires about 64 days.
 - The number of sperms produced by adult male per day is 10^{12} - 10^{13} .
Sertoli cells act as phagocytes. They consume the residual cytoplasm discarded during spermiogenesis.
 - After maturation of spermatozoa, they get detached from Sertoli cells. The process is called spermiation. The released sperms are stored in epididymis up to one month.

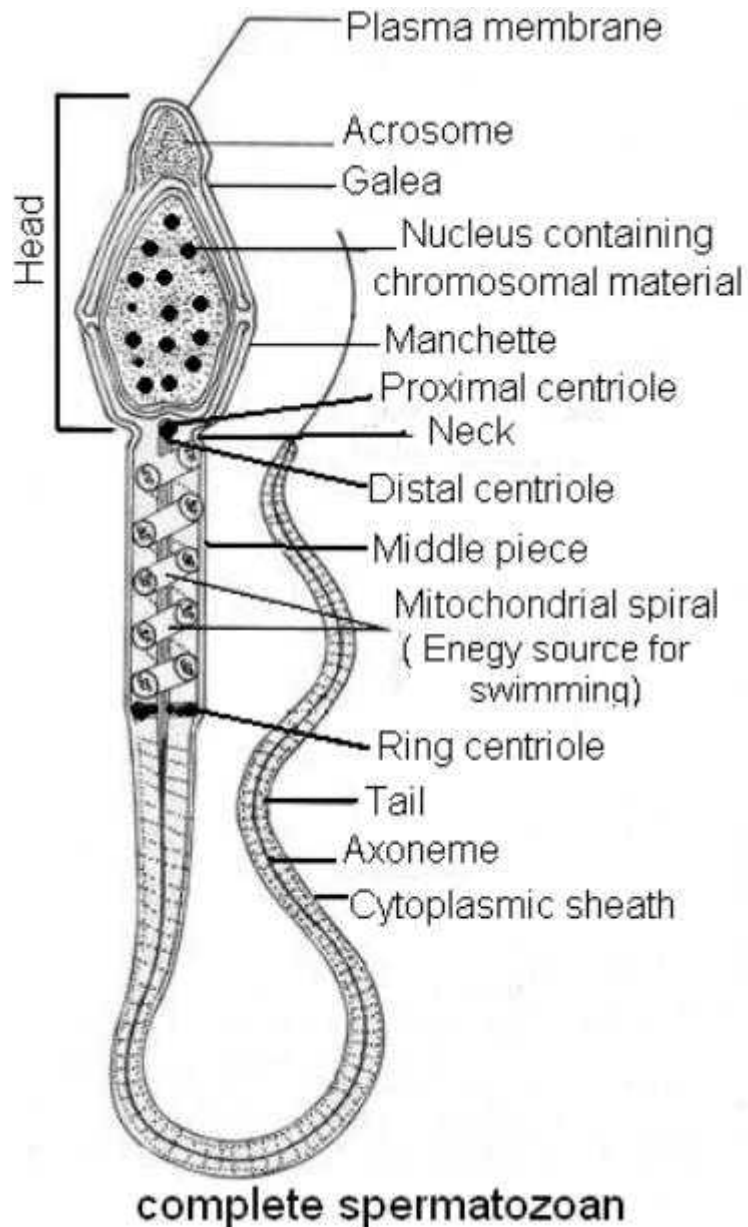


SPERM

- Human sperm is dart-like flagellate structure of 60 μm length and maximum breadth of 3.5μm.
- It has four parts:
 - (i) Head: Knob-like structure but flat terminal part, 4-5 μm long. It has two components acrosome and nucleus. Cytoplasm is nearly absent. Acrosome forms a covering over the anterior two third of head. It is derived from golgi apparatus from spermatid. It contains lysins like hyaluronidase and proteolytic enzymes. Surface contains compatibility protein bindin for attaching to receptors of egg. Nucleus is dense mass of chromatin having protamines. These may be one or more less dense areas called nuclear vacuoles. On the outside is present a double membrane head cap.
 - (ii) Neck: It is a short narrow part between head and middle piece which contains two centrioles, unconnected proximal centriole and distal centriole attached to filament that passes into middle piece.
 - (iii) Middle piece: It is a cylindrical part 5-7μm long and 1μm in breadth. It has axial filament surrounded by 10-14 spiral turns of mitochondria and bearing towards the end of a ring centriole. Mitochondria provides energy for swimming. All are embedded in a thin sheath of cytoplasm.
 - (iv) Tail: It is narrow vibratile long part about 50μm in length. The tail is broad at beginning but narrows down gradually.

SIGNIFICANCE OF SPERMATOGENESIS

- (i) During spermatogenesis, one spermatogonium produce four sperms
- (ii) Sperms have half the number of chromosomes
- (iii) During meiosis "I" crossing over takes place which brings variation



OOGENESIS

Oogenesis is the process of formation of functional haploid ova from the diploid germinal cells in the ovary

It consists of three phases

(i) Multiplication phase

- During foetal development, certain cells of the germinal epithelium of ovary which are larger than others function as germ cells. They undergo repeated mitotic divisions to produce undifferentiated germ cells oogonia or egg mother cells (2N)
- The oogonia multiply by mitotic divisions and become primary oocyte
- Other oogonia forms a regular layer, the follicular epithelium around the primary oocyte to protect and nourish it. The structure thus formed, is called primary ovarian follicle.

(ii) Growth phase

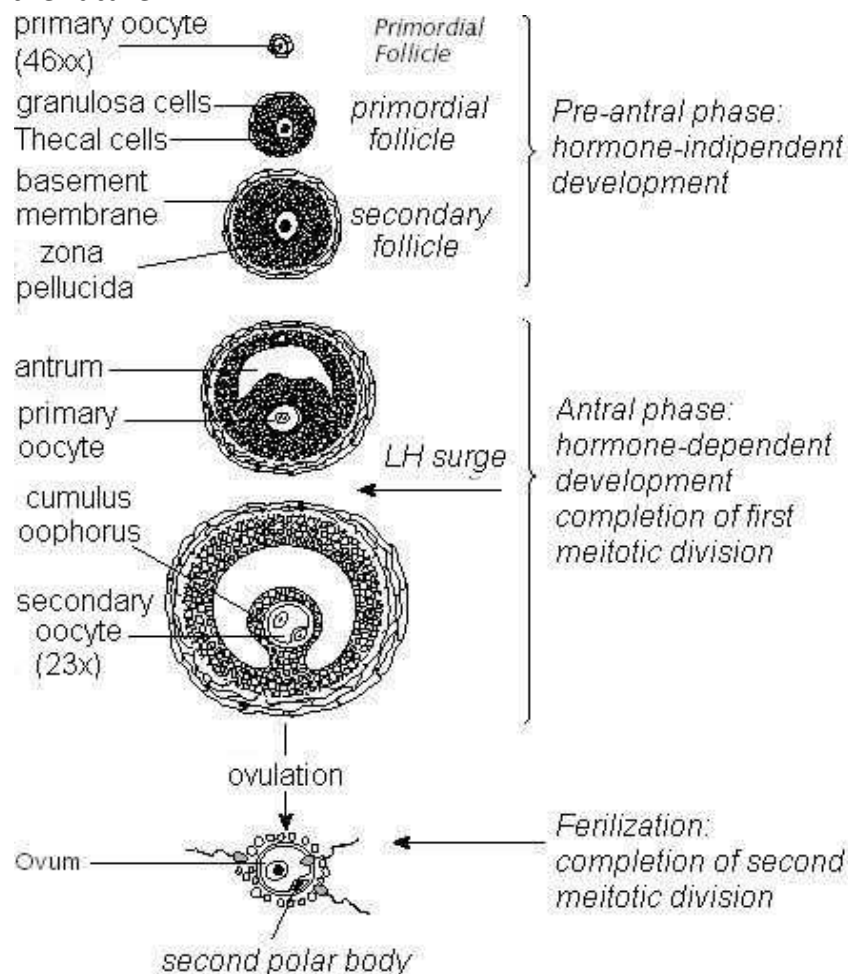
- This phase of the primary oocyte is very long. The oogonium grows into a large primary oocyte by taking food from the surrounding follicle cells, it happens after puberty
- Meiosis begins in the primary oocytes soon after their formation. However, the oocytes are arrested in the early part of meiotic prophase "I". This is the first resting stage. They undergo a round of DNA synthesis and chromosome pairing takes place, but meiosis does not proceed further until years later

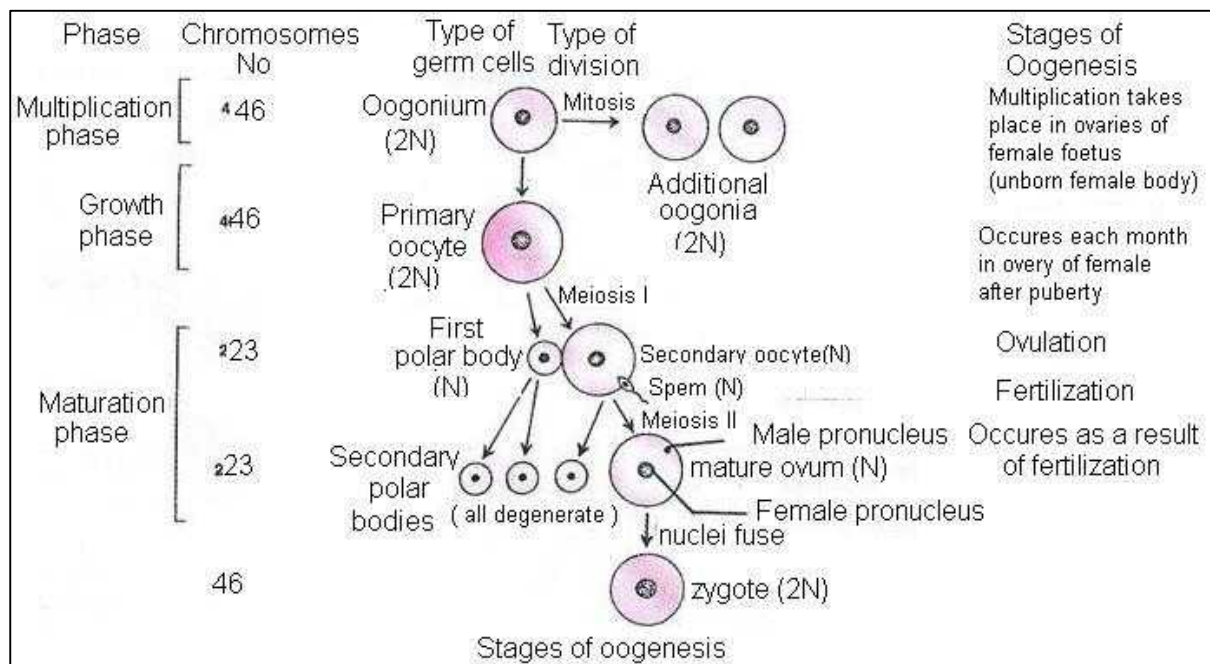
(iii) Maturation phase

- After attainment of puberty, follicles and their contained oocytes resume development. Follicular sheath differentiates into layers of granulosa cells. Another layer called theca, primary follicle is transformed into two layers. Granulosa cells secrete fluid that causes the development of a cavity or antrum around the primary oocyte. The stage is called tertiary follicle.
- Primary oocyte grows further and completes meiosis "I". It produces a large secondary oocyte and a small polar body. Both are haploid. The polar body has a very small amount of cytoplasm but whole chromosome set. The bulk of nutrient rich cytoplasm is retained in the secondary oocyte.
- The follicle grows to maximum size and is called graafian follicle. Zona pellucida develops around secondary oocyte

- The secondary oocyte proceeds with meiosis II but division gets arrested in metaphase II stage due to accumulation of metaphase promoting factor (MPF). This is the second resting stage
- It is in this stage of oocyte that the ovum is shed during ovulation. It passes into oviduct, where in ampulla part, cell cycle will resume only after the entry of sperm. It triggers the breakdown of MPF and promotes synthesis of anaphase promoting complex(APC) meiosisII is completed.
- As secondary polar body is extruded. The first polar body is extruded. The first polar body divide to form two second polar bodies. Thus form one oogonium, one ovum and three polar bodies are formed. In human females the polar body does not divide further. The oocyte is now changed into ovum or ootid. The ovum is actual female gametes. The polar bodies take no part in reproduction and soon degenerates due to lack of cytoplasm and food. The formation of non-functional polar bodies enables the egg to get rid of excess chromosomes. The unequal cytoplasmic division enables the ovum to retain the whole of the cytoplasm of the primary oocyte in it for the development of

the future





SIGNIFICANCE OF OOGENESIS

- (i) One oogonium produces one ovum and three polar bodies
- (ii) It helps to retain sufficient amount of cytoplasm in the ovum which is essential for the development of early embryo
- (iii) During meiosis first crossing over takes place which brings about variation

OVUM

- Human egg or ovum is noncleidoic (without shell) and alecithal (absence of yolk) rounded female gamete having diameter of about 100µm
- The ovum possesses three coverings – inner plasma membrane, middle glycoprotein zona pellucida and outer cellular corona radiata with radially elongated scattered cells held in mucopolysaccharide.
- Zona pellucida carries compatibility receptor proteins collectively called as fertilizing. In between plasma membrane and zona pellucida is perivitelline space in which one polar body is present towards animal pole. The opposite end is vegetal pole. Cytoplasm of ovum is called ooplasm. It has a large nucleus or germinal vesicle.
- Typical nucleus or pronucleus is formed at the time of fertilization
- Ectoplasm possesses mucopolysaccharides granules and microtubules. Mucopolysaccharides granules or cortical granules are extruded membrane and zona pellucida for preventing entry of another sperm. Endoplasm has

mitochondria, golgi apparatus, ribosomes, RNA, fat droplets, glycogen particles and proteins.

TYPES OF OVUM

On the basis of amount of yolk, eggs are classified as:

- (i) Alecithal – No yolk, example : human egg
 - (ii) Microlecithal – Small amount of yolk example : Sea urchin
 - (iii) Mesolecithal – Moderate amount of yolk, example frog and other amphibian egg
 - (iv) Macrolecithal or polylecithal – Large amount of yolk example : reptilian and avian eggs
- On the basis of distribution of yolk, eggs are classified as:-
 - (i) Isolecithal or Homolecithal _ Having homogenously distributed yolk, example : Protochordates and echinoderms.
 - (ii) Hetrolecithal – egg with unevenly distributed yolk
 - (iii) Telolecithal – Having yolk concerned in one half

Example : amphibian eggs

- (iv) Centrolecithal – yolk is concentrated in centre and cytoplasm is peripheral example – insect eggs
 - (v) Discoidal or Meiolecithal – Almost the whole of the egg is occupied by the yolk except a small disc, example : eggs of birds and reptiles
- On the basis of presence and absence of shell eggs are differentiated into cleidoic (surrounded with water proof shell e.g. birds and reptiles) and noncleidoic (shell absent)

FERTILIZATION IN HUMANS

- It is fusion of male and female gametes to form zygote. In human being fertilisation is internal. Human beings are viviparous
- Here the embryo is retained and nourished inside the uterus of the female by means of an attachment called placenta. At one time only a single ovum is released in human females from one of the two ovaries towards the middle of ovarian/ menstrual cycle. It passes into fallopian tubes and rests inside ampulla for some time. The journey time is 12-24 hours
- Human male produces 300-400 millions sperms per ejaculation. They are deposited in vagina during coitus. The process of deposition of sperms in the

female genital tract is called insemination. A number of them are demobilized or eat but a number of them remain functional and undergo capacitation (sperm activation) that provides them the ability to fertilise an ovum. Capacitation requires 6-10 hrs.

- Capacitation consists of three process
 - (i) Neutralisation of inhibitory factors present in semen
 - (ii) Weakening of covering membrane of acrosome head by dissolution of cholesterol
 - (iii) Entry of Ca^{2+} into sperms which changes sperm movement from undulation to whiplash motion
- The activated sperm being to pass into uterus and from there to oviducts. Viscous fluid secreted by female genital tract further enhances sperm motility. A number of sperms reach the ampulla part of oviduct where the egg rests temporarily
- Fertilization involves following steps:
 - (i) Approximation of sperm and ovum
Sperm can remain motile for 24-48 hrs. They swim at the rate of 1.5-3 mm/min. They are able to reach the ampulla part of female genital tract partly by contraction of uterus and fallopian tubes stimulated by prostaglandins (in male semen) and oxytocin (often formed in females). The movements are powerful within 5 minutes. After reaching an ovum, one sperm comes to lie against it. It releases lysine from its acrosomal region. Hyalouronidase and corona penetrating enzymes as well as dissolves cells of corona radiata. The sperm head now reaches zona pellucida where receptors protein fertilizing helps in attachment to specific protein of sperm. It is compatibility reaction.
 - (ii) Acrosome reaction
In contact with zona pellucida, acrosome covering degenerates. The contained enzymes are released. Acrosin or zona lysine dissolves zona pellucida in area of contact.
 - (iii) Egg reaction
A small protuberance or fertilization cone develops from the surface of ovum in the region of animal pole.
 - (iv) Penetration of sperm
Sperm head established contact with the lateral surface of fertilization cone. It produces a weak depolarization and Ca^{2+} wave. Plasma membranes of the two, dissolve. Contents of head, neck and middle sperm enter

ooplasm. Tail is left outside. Fertilization cone subsides. Cortical granules are extruded. They convert plasma membrane and deactivate sperm receptor of zona pellucida. A perivitelline space is created between it and zona pellucida. This prevents entry of a second sperm.

(v) Activation of ovum

Ovum undergoes meiosis II and extrudes a secondary polar body. It is now the actual ovum or female gametes.

(vi) Fusion of sperm and egg

The envelope of the sperm and egg pronuclei degenerates to form 'synkaryon'. The act is called karyogamy or syngamy. The proximal centriole brought by sperm helps form the spindle for the division of synkaryon (cleavage nucleus). Fertilized egg is also called zygote. It immediately begins cleavage.

SIGNIFICANCE OF FERTILIZATION

- (i) It restores the diploid number of chromosomes, characteristics of species i.e. 46 in human being
- (ii) Fertilization initiates cleavage

CLEAVAGE

- Cleavage is a series of rapid mitotic divisions of the zygote, characterized by absence of growth of daughter cells, which convert the single celled zygote into a multicellular structure called blastula (blastocyst)
- Cleavage differ from mitosis in the respect that
 - (i) There is no growth phase between successive division.
 - (ii) The size of cells gradually decreases
 - (iii) The metabolism becomes fast
 - (iv) There is rapid DNA replication
 - (v) High consumption of oxygen
- Types of cleavage
 - (i) Holoblastic: When whole of the egg is divided, it is found in microlecithal and mesolecithal egg. It may further be..
 - (a) Equal – When both the blastomers are equal Example – Amphioxus
 - (b) Unequal – When the blastomers are unequal in size. Example- frog
 - (ii) Meroblastic: When a part of the egg is divided. It is found in polylecithal eggs. It may be discoidal (e.g. birds) or superficial (e.g. insects)

- Planes of cleavage include meridional, vertical, equatorial and transverse. Patterns can be radial (sponges, coelenterates, some echinoderms like star fish) biradial, spiral (flatworms, annelids, non-cephalopod mollusks) bilateral (nematods, cephalopods, fishes, amphibians, reptiles, birds) and rotational (placental mammals)

MORULA

Early cleavage produces a solid ball of cells called morula

BLASTULA

- Multicellular ball like embryo produced at the end of cleavage and usually having a fluid filled blastocoels, is called blastula
- It is of the following types
 - (i) Stereoblastula (solid blastula): It is blastula without blastocoels. E.g. Nereis
 - (ii) Coeloblastula : A blastula with a prominent blastocoels. e.g. frog
 - (iii) Discoblastula : A blastula having a many layered disc of blastomeres above the yolk. It develops as a result of meroblastic divisions in polylecithal eggs. E.g. Hen
 - (iv) Superficial blastula (Periblastula) : A blastula having a single layer of blastomeres around the central yolk. E.g. insects

GASTRULATION

- Sum total of all the processes which convert a solid or hollow ball of cells or many layered disc of blastula into two or three germinal layers of gastrula is called gastrulation
- Gastrulation takes place by the migratory or formative or morphogenetic movements of blastomeres from the surface of blastula to the proper position in the gastrula.
- These movements are classified into
 - (a) Epiboly: Growth of one part over another like prospective ectoderm over the rest except blastocoels.
 - (b) Emboly: Morphogenetic movements like migration of ectoderm, mesoderm and notochord cells from surface to interior. The emboly may occur by way of
 - (i) Involution : Rolling of cells into interior
 - (ii) Invagination : Infolding
 - (iii) Ingression: New cells migrating into blastocoels

- (iv) Delamination : Formation blastocoels is obliterated and a new cavity archenteron is formed which in future alimentary canal of the animal. Blastopore is opening of archenteron.

ORGANOGENESIS

The development of tissues and organs from the three germ layer is called organogenesis

MORPHOGENESIS

The assumption of shape, size and other morphological features by embryo is called morphogenesis

DIFFERENTIATION

It is the formation of different types of cells, which become different in size, form, chemical composition and perform different functions

FATE OF THREE GERMINAL LAYERS

Ectoderm

Central nervous system, nerves, retina, lens, cornea of eyes. Conjunctiva, ciliary and iridial muscles, lining of nasal chambers. Labyrinth, epidermis, cutaneous, glands, hair, nails, claws, hypophysis, adrenal medulla, salivary glands and enamel of teeth

Mesoderm

Dermis of skin, connective tissue, muscles, notochord, skeleton, blood, heart, blood vessel, adrenal cortex, urino – genitals system except part of urinary bladder, lining of coelom, spleen and eyes.

Endoderm

Digestive glands, liver, pancreas, middle ear, Eustachian tubes, lining of urinary bladder, respiratory system, adenophypophysis thymus, parathyroid and thyroid glands, lining of vagina and urethra, prostate.

FOETAL MEMBRANES

They are extra embryonic membranes that provide protection and nourishment to foetus. Foetal membranes are of four types

- (i) Chorion – Outer foetal membrane that also takes part in formation of placenta.
- (ii) Amnion – Inner foetal membrane that invests the embryo and forms a space called amniotic cavity. It is filled with fluid called amniotic fluid. Amnion protects the foetus from shock
- (iii) Allantois – sac – like, develops from gut of embryo, supplies blood vessels to placenta. In reptiles and birds, it helps in respiration nutrition and excretion
- (iv) Yolk sac – Membraneous sac attached to embryo near allantois, having yolk in egg laying animals and forms corpuscles in mammals till liver takes over

EMBRYO FORMATION IN HUMAN BEINGS

MORULA FORMATION

- Soon after fertilization, the zygote begins cleavage or segmentation. Cleavage consists of early mitotic divisions of fertilized egg without involving growth of daughter cells. There is rapid synthesis of new DNA and increased oxygen consumption. Surface –volume and nucleo-cytoplasmic ratio increases. The cells formed after cleavage are called blastomers. Cleavage is simple and holoblastic in humans as there is no yolk.
- The first cleavage is animal – vegetal axis or primary axis. It is slow and is completed within 30 hours of fertilization
- One of the two blastomers, is however slightly larger. Hence, the first cleavage is holoblastic and unequal. Second cleavage is at right angle to the first one. It takes about 30 hours and is completed slightly earlier in the larger blastomere so that a transitional 3-celled stage appears. Subsequent divisions are rapid and occur in different planes. They produce a solid ball of blastomers called morula. Phase of compaction ensues in 8-celled stage. Morula has almost the same size as that of fertilized egg due to presence of zona pellucida
- Morula has 16-32 cells. The cells are compacted and of two types, outer slightly smaller peripheral cells with tight junctions than the inner mass of cells with gap junctions. During the cleavage stage the embryo descends in the fallopian

tube slowly due to feeble fluid produced by epithelial secretion and cilia. Ultimately it reach the uterus. It takes 4-6 days. Corona radiate dissolved away during this period.

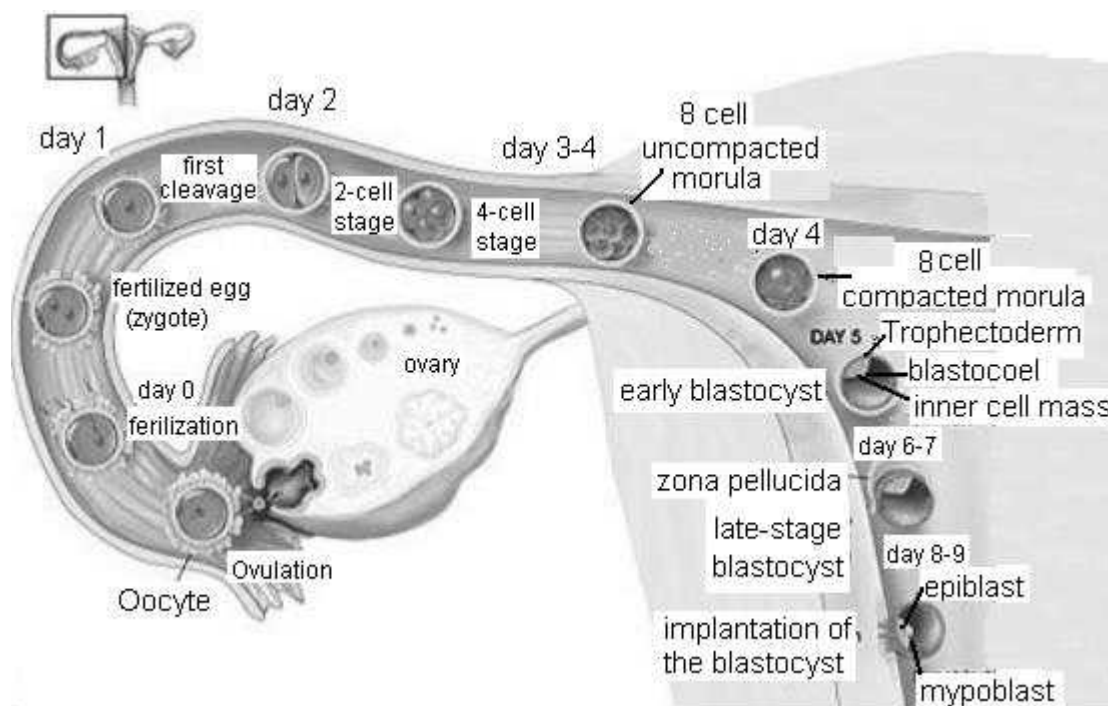
BLASTULATION

- In uterus, the endometrial cells become full of nutrients which are also secreted into uterine cavity. As the young embryo reaches uterus, its outer cells begins to absorb nourishment and grow while covered by zona pellucida
- The outer cells enlarge, flatten and form trophoblast. Trophoblast pours as fluid towards interior producing a cavity called blastocoels or blastocyst. It is equivalent to blastula of othe animals. The size of blastocyst is roughly three times the size of morula.
- Trophoblast then separates from inner cells except at one point called embryonic pole. The inner cells now occur at one side and called inner cell now occur at one side and called inner cell mass or embryonal knob as the latter is to form the body of embryo. Trophoblast cells in contact with inner mass are called cell of Rauber. Embryonic pole is also called animal pole. The opposite end of blastocyst is called abembryoic pole
- Blastocyst stage is completed after about 5 days of fertilization. Trophoblast later becomes two – layered, outer syncytiotrophoblast and inner cytophoblast. It secretes hCG (human chorionic gonadotrophin). Forms villi for implantation and later on produces chorion, amnion and foetal part of placenta

IMPLANTATION

- It is a choring or embedding of blastocyst into endomerium of uterus. Implantation begins about 7th day after fertilization of ovum. It takes about process to be completed
- Blastocyst comes in contact with the endometrium in the region of embryonal knob or embryonic disc
- The surfacea cells of trophoblast secrete lytic enzymes which causes corrosion of endometrial lining. They also give rise to figure – like outgrowths called chorionic villi and uterine tissue becomes interdigitated. Villi not only help in fixation but also absorption of nourishment

- The blastocyst sink in the pit formed in endometrium then get completed buried in the endometrium that grows around it. The embedded blastocyst forms villi all around it to obtain nourishment.
- Implantation causes nutrient enrichment, enlargement of cells and vascular endometrium. Vascular endometrium (deciduas of pregnancy) are stomal cells which have accumulated glycogen and lipid in their distended cytoplasm. The interglandular tissue increases in quantity.
- They may offer nutrition which is engulfed by the syntial trophoblast but they have been regulated as defensive mechanism
- Trophoblast covering secretes hormone called human chorionic gonadotropin (hCG). The hormone can be detected in the urine of woman within day after implantation
- hCG maintain the corpus luteum beyond its normal life. It continues to secrete pregnancy which prevents menstruation and maintains uterine lining in nutrient rich state.
- Progesterone induces cervical glands to secrete viscous mucus for filling the cervical canal to form a protective plug. By the 16th week of pregnancy, placenta produces enough progesterone and the corpus luteum regresse.



PLACENTA

- Placenta is a temporary organ found only in mammals during gestation period and it is composed of cells derived from two different organisms, the foetus and mother
- Placenta is connection between the foetal membranes and uterine wall.

Formation of placenta

- The outer surface of the chorion in human develops a number of finger like projections known as chorionic villi, which grow into the tissue of the uterus. These villi, penetrate the tissue of the uterine wall in which they are embedded to make up the organ known as placenta by means of which the developing embryo obtains nutrients, oxygen and gets rid of carbon dioxide and metabolic wastes
- A fully formed human placenta is reddish – brown disc. Its foetal surface is smooth and has the umbilical cord. The allantois gives rise to umbilical cord. Umbilical cord has two umbilical arteries (small diameter) and two umbilical veins (large diameter)
- Umbilical arteries convey oxygen poor blood from the foetus to placenta and umbilical veins carry oxygen rich blood from the placenta to the foetus
- The blood of foetus in the capillaries of the chorionic villi comes in close contact with the mother's blood in the tissue between the villi, but are always separated by a membrane, through which substances must diffuse
- The maternal and foetal blood are not in direct contact in the placenta because
 - (i) Two may be incompatible
 - (ii) The pressure of maternal blood is far too high for the foetal blood vessels
 - (iii) There must be a check on passage of harmful material into foetal blood

Functions of placenta

- (i) Nutritive organ – Food materials from the mother's blood into the foetal blood through the placenta
- (ii) Digestive organ – Trophoblast of placenta digests proteins before passing them into the foetal blood.
- (iii) Respiratory organ – Oxygen diffuses from the maternal blood into the foetal blood through the placenta. Carbon dioxide diffuses from the foetal

blood into the maternal blood also through the placenta for elimination by the mother's lungs. Foetal haemoglobin has a greater affinity for oxygen than adult haemoglobin

- (iv) Excretory organ – Nitrogenous waste such as urea, pass form the foetal blood into the maternal blood via placenta for elimination by mother's kidney
- (v) Endocrine organ – Placenta secretes some hormone such as estrogens, progesterone, human chorionic gonadotropin (hCG), human placental lactogen (hPL) chorionic thyrotropin, chorionic corticotrophin and relaxin. The hCG stimulates and maintains the corpus luteum to secrete progesterone until the end of pregnancy. The hPL stimulates the growth of the mammary glands during pregnancy. Relaxin facilitates parturition by softening the connective tissue of pubic symphysis
- (vi) Storage organ – The placenta stores glycogen for the foetus before liver is formed

PARTURITION

- It is the process of giving birth to a baby . The physical activities in parturition like uterine and abdominal contractions dilation of cervix and passage of baby are collectively called labour
 - Labour is accompanied by localized sensation of discomfort or agony called labour pains.
 - Parturation is controlled by complex neuroendocrine mechanism. Signals originate from fully formed foetus and placenta. They cause mild uterine contraction called foetal ejection reflex. It is accompanied by rise in estrogen to progesterone ratio, increase in oxytocin receptors in uterine muscles, increase in level of oxytocin secretion by both mother and foetus and stretching of uterian musculature
- (i) Dilation stage
- Uterine contractions begin from top. They occur once every 30 minutes. Contractions forces the baby towards cervix. The intervals between successive contraction decreases about every 1-3 minutes. Contractions are accompanied by pain caused by compression of blood vessels and uterine muscles
- Oxytocin induce contraction and more oxytocin secretion. The strength of uterine contraction continues to increase due to stimulatory reflex. As the baby is pushed down in uterus, its head come to lie against cervix which

therefore gets dilated and stretched, A similar dilation also occurs in vagina. The first stage of labour continues for 6-12 hours. It culminates in rupturing of amniotic membrane. The amniotic fluid flows out

(ii) Expulsion stage

The intensity of uterine and abdominal contractions increases. The foetus passes out through cervix and vagina with head in forward direction in normal deliveries. It causes intense labour pain. Expulsion stage takes about 20-60 minutes. Umbilical cord is tied and cut off close to navel

(iii) After birth

Within 10-45 minutes of the delivery of baby the placenta separates from uterus and is expelled out due to series of strong uterine contractions In neonate there is a change in respiratory and circulatory system. The switch over is initiated by gaseous hormone nitric oxide (NO). Lungs expand and infant starts breathing. Blood flow through umbilical cord, foramen ovale ceases. It starts passing through heart, aorta and pulmonary arteries

GESTATION PERIOD IN SOME MAMMALS

SrNO	Mammal	Days
1	Mare	335
2	Ass	365 -370
3	Cow	282-270
4	Cat	63
5	Dog	60-63
6	Elephant	624
7	Goat	148
8	Horse	335-340
9	Whale	330-365
10	Rats	22
11	Guinea pig	68
12	Lion	105-115
13	Monkey (Rhesus)	164
14	Mouse	19-20
15	Human beings	266 -280 days
16	Rabbit	32
17	Sheep	148
18	Tiger	155
19	Swine (pig)	114

IMPORTANT DEVELOPMENTAL CHANGES IN HUMAN EMBRYO

1. Week 1
 - Fertilization
 - Cleavage starts about 24 hours after fertilization
 - Cleavage to form a blastocyst after fertilization
 - More than 100 cells
 - Implantation 6-9 days after fertilization
2. Week 2
 - The three primary germ layers develop
3. Week 3
 - Woman will not have period. These may be first sign of pregnancy
 - Beginnings of backbone
 - Neural tube develops, the beginning of brain and spinal cord
4. Week 4
 - Heart, blood vessels, blood and gut start forming
 - Umbilical cord developing
5. Week 5
 - Brain developing
 - 'Limb buds' a small swelling which are the beginnings of arms and legs
 - Heart is a large tube and starts to beat, pumping blood. This can be seen on an ultrasound scan
6. Week 6
 - Eyes and ears start to form
7. Week 7
 - All major internal organs developing
 - Face, forming
 - Eyes have some colour
 - Mouth and tongue develop
 - Beginnings of hands and feet
 - 2.5 cm long
8. By week 12
 - Foetus fully formed with all organs, muscles, bone, toes and fingers
 - Sex organs well developed
 - Foetus reaches 7.5cm in height and about 14g weight
9. By week 20

- Hair beginning to grow, including eyebrows and eyelashes
- Fingerprints developed
- Fingernails and toenails growing
- Firm hand grip
- Between 16 and 20 weeks baby usually felt moving for first time

10. Week 24

- Eyelids open
- Foetus measures about 32 cm and weighs about 650gm

11. Week 26

Has a good chance of survival if born prematurely

12. By week 28

- Baby moving vigorously
- Responds to touch and noise
- Swallowing amniotic fluid

13. By week 30

Usually lying head down ready by birth

Foetus is about 43cm long and its weight is about 1800gm

14. By week 40

Birth

Generally child is about 50cm long and weighs about 3300gm

LACTATION

- Production of milk in the female's breasts following the birth of a young one in mammals is called lactation

Preparation of breast (mammogenesis)

During pregnancy, the breast enlarges due to growth of mammary glands

Synthesis and secretion from the breast alveoli (lactogenesis)

Secretion and storage of milk begins after birth of the young one, usually within 24 hours under the influence of prolactin

When the estrogen and progesterone are withdrawn following delivery, prolactin begins its milk secretory activity in previously fully developed mammary glands

Ejection of milk

The actual release of milk called milk letdown, requires the presence of oxytocin, which brings about contraction of smooth muscles of the ducts

within the mammary gland. Secretion of prolactin and oxytocin depends on suckling stimuli produced by the nursing infant on the nipples of breasts

Maintenance of lactation (galactopoiesis)

For maintenance of lactation, suckling is important

Milk pressure reduces the rate of production and hence periodic breast feeding is necessary to relieve the pressure which is in turn maintains the secretion.

- After birth, the breast first release is not milk, but colostrums for 2 or 3 days. It is a thin, yellowish, fluid called foremilk which is rich in protein, antibodies but low in fat.
- Human milk consists of water, fat, casein, lactose, mineral salts and vitamins. A nursing woman secretes 1 to 2 litres of milk per day

DEVELOPMENTAL DISPRDERS

AMNIONITIS

It is inflammation of amnion, usually resulting from premature rupture of amnion

ABORTION

It is giving birth to an embryo or foetus at the stage of about 20 weeks of gestation

TERATOGENY

Teratogens are certain agents or drugs that cause abnormal development in developing embryo/foetus. It may cause malformation in developing embryo.

ECTOPIC PREGNANCY

- The developmental site of foetus is other than the uterus like fallopian tube or cervix
- The growth of foetus may cause tube to rupture and bleed and may lead to miscarriage
- The condition is diagnosed by ultrasound and foetus may be removed by laparoscopy before damage is done to fallopian tube.