

SO1: The menstrual cycle (AMBOS)

Menarche: start of menstruation

Menopause: cessation thereof

Menses: Bleeding (menstrual) → controlled by HPA.

Small Δ in Hormonal Balance = menstrual irregularities

Δ's ≠ necessarily pathological
 Abnormal menses = Δ's in $\left. \begin{matrix} \text{frequency} \\ \text{intensity} \\ \text{onset} \end{matrix} \right\}$ AUB.
 } Amenorrhoea.
 } Dysmenorrhoea.

PMS → NB (premenstrual syndrome)

Physiology of menstrual cycle

(+) & (-) feedback loops

Follicular phase

Hypothalamus = release GnRH in pulses

GnRH → Ant pit. → LH & FSH

LH & FSH → Ovaries → Graafian follicle

Day before LH ↑ = one follicle becomes dominant!

Remaining follicles regress as (FSH ↓) & Estrodiol peak → (+)-feedback

↳ LH surge

Ovulation = LH surge induces ovulation → Mature oocyte released from dominant follicle

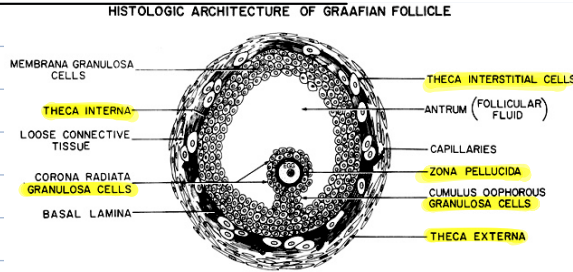
↳ Corpus luteum prod. progesterone = LH inhibition. [progesterone → LH]

Luteal phase:

Falling LH lvs cause degeneration of corpus luteum = ↓

progesterone
 Estrodiol

∴ Endometrium & maintained.

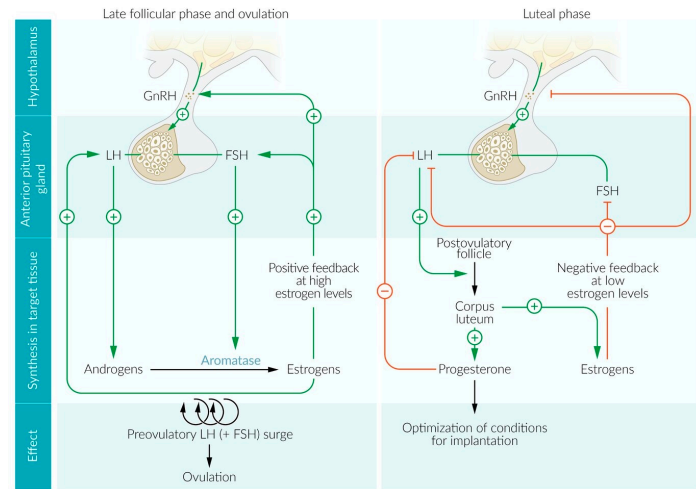
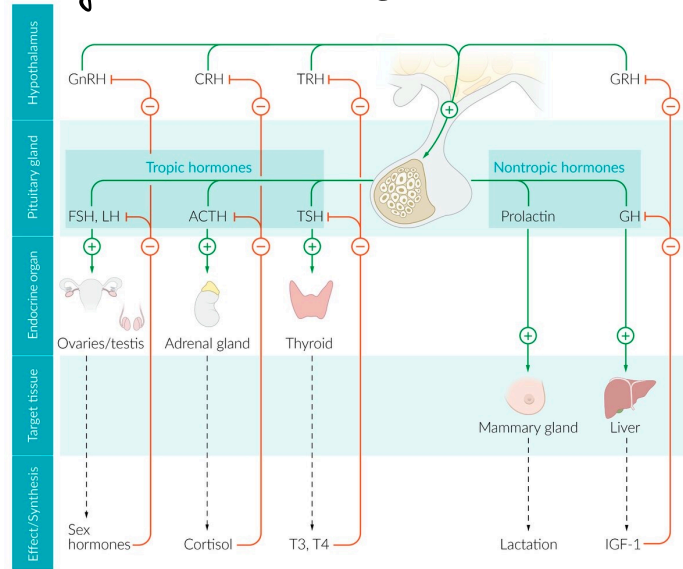


Theca cells → progesterone & Androstenedione

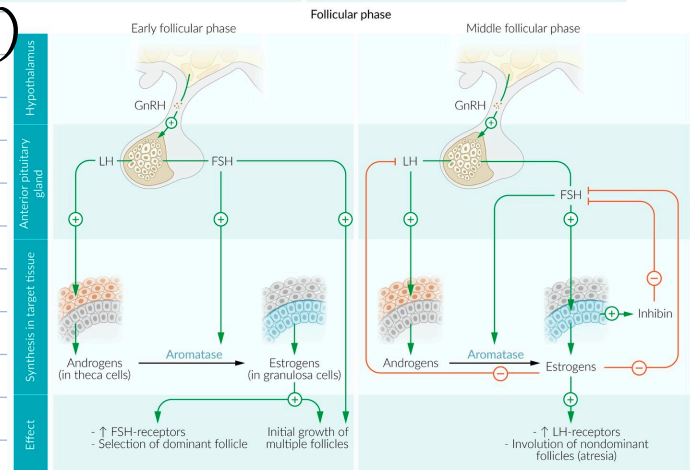
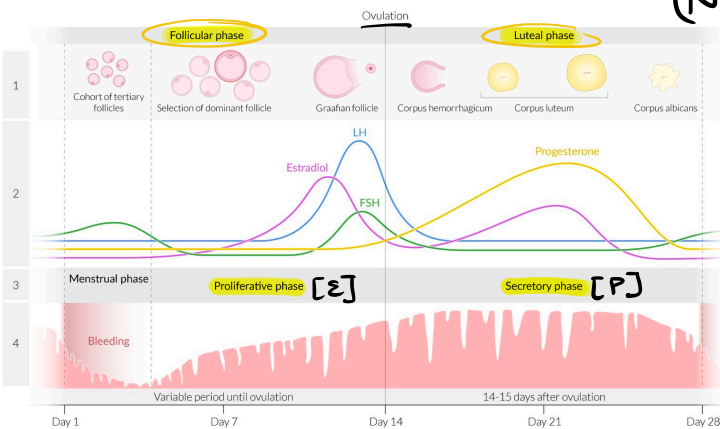
Granulosa cells → Estrodiol & inhibin B

(-) feedback

↳ ↓ FSH Release



(NB)



Normal menstrual cycle = 24-38 days.

Day 1 = first day of menstrual cycle.

Menses = 3-7 days (\pm 35-50 ml)

Menses = o's in ovaries (ovarian cycle)
= o's in uterus (uterine cycle)

} cycles relate to one another

Ovarian o's regulated by HPA.

Uterine o's regulated by ovarian o's - low estradiol = FSH \uparrow LH \downarrow = menstruation
- High estradiol = \uparrow FSH/LH = thickening of endometrium.

- E \uparrow P released by corpus luteum = development of endometrium
- E \uparrow P i secretion of FSH \uparrow LH = prevent further follicular development.

Menstrual cycle changes					
Cycle	Duration	Phases	Description	Mechanism	Histological changes [2]
Ovarian cycle	1-14 days	Follicular phase	<ul style="list-style-type: none"> From the first day of menses to the day before the LH surge Accounts for most of the variability in the length of the menstrual cycle Follicle growth speeds up during the 2nd week of this phase. 	<ul style="list-style-type: none"> FSH stimulates the development of several follicles in the ovaries \rightarrow granulosa cells of follicles produce estrogen \rightarrow estrogen suppresses the release of FSH via negative feedback loop Selection of a dominant follicle (Graafian follicle) Positive feedback loop: high estrogen levels \rightarrow FSH release \rightarrow LH surge \rightarrow ovulation <i>since granulosa cells releasing so much oestrogen.</i> 	<ul style="list-style-type: none"> During the secondary follicle stage, cuboidal granulosa cells continue to proliferate. Multiple layers of thick granulosa cells surround an eccentrically located oocyte. Most secondary follicles become atretic, but usually only one becomes a dominant follicle.
	14-15 days	Luteal phase	<ul style="list-style-type: none"> From the day of the LH surge to the beginning of the next menses 	<ul style="list-style-type: none"> In ovulation, the Graafian follicle ruptures, releasing the oocyte. Following ovulation, the granulosa cells produce LH receptors \rightarrow LH-induced transformation of the Graafian follicle into the corpus luteum \rightarrow progesterone production \rightarrow inhibition of LH release (progesterone increase indicates that ovulation has occurred) If no pregnancy occurs, the corpus luteum regresses. 	<ul style="list-style-type: none"> During the tertiary follicle stage, the oocyte is surrounded by the corona radiata and floats in follicular fluid. The Graafian follicle moves to the surface of the ovary, where it ruptures and the secondary oocyte is released. The empty tertiary follicle collapses. The corpus luteum atrophies.
	3-7 days	Menses	<ul style="list-style-type: none"> Menstrual bleeding occurs in this phase (usually 14 days after ovulation). 	<ul style="list-style-type: none"> Absence of pregnancy \rightarrow resolution of corpus luteum \rightarrow \downarrow progesterone concentration \rightarrow vasospasms in the uterine spiral arteries \rightarrow bleeding To actively delay menses, administer progesterone in the second half of the menstrual cycle. 	<ul style="list-style-type: none"> Contraction of spiral arteries in the stratum functionalis layer \rightarrow ischemia \rightarrow degeneration of the functionalis layer \rightarrow sloughing off of the functional layer of the endometrium
Uterine cycle	~10 days	Proliferative phase	<ul style="list-style-type: none"> Characterized by the growth of the endometrium under the influence of estrogens 	<ul style="list-style-type: none"> Growing follicles produce estrogen (granulosa cells express aromatase, which converts androgens to estrogens) \rightarrow proliferation of the endometrium 	<ul style="list-style-type: none"> Proliferation of endometrial epithelial cells (cells show high mitotic activity) Endometrial glands become straight, tubular, and lined by simple columnar epithelium. Stromal cells start to divide, enlarge, and accumulate glycogen. Uterine spiral arteries start to regenerate and extend two-thirds of the way into the endometrium.
	10-14 days	Secretory phase	<ul style="list-style-type: none"> The functional layer of the endometrium is prepared for implantation under the influence of progesterone. 	<ul style="list-style-type: none"> Progesterone promotes endometrial differentiation \rightarrow preparation of the functional layer of the endometrium for oocyte implantation \uparrow Cervical mucus secretion (prevents spermatozooids from entering uterus) \uparrow Basal body temperature In the absence of pregnancy: \downarrow progesterone levels \rightarrow apoptosis of the functional layer of the endometrium \rightarrow menstruation 	<ul style="list-style-type: none"> Intracellular subnuclear vacuoles Increased endometrial gland tortuosity Glycogen-rich secretions Edematous stromal cells Uterine spiral arteries extend the full length of the endometrium.

