



THE UNIVERSITY OF
TENNESSEE
HEALTH SCIENCE CENTER.

Maternal Adaptations of Pregnancy

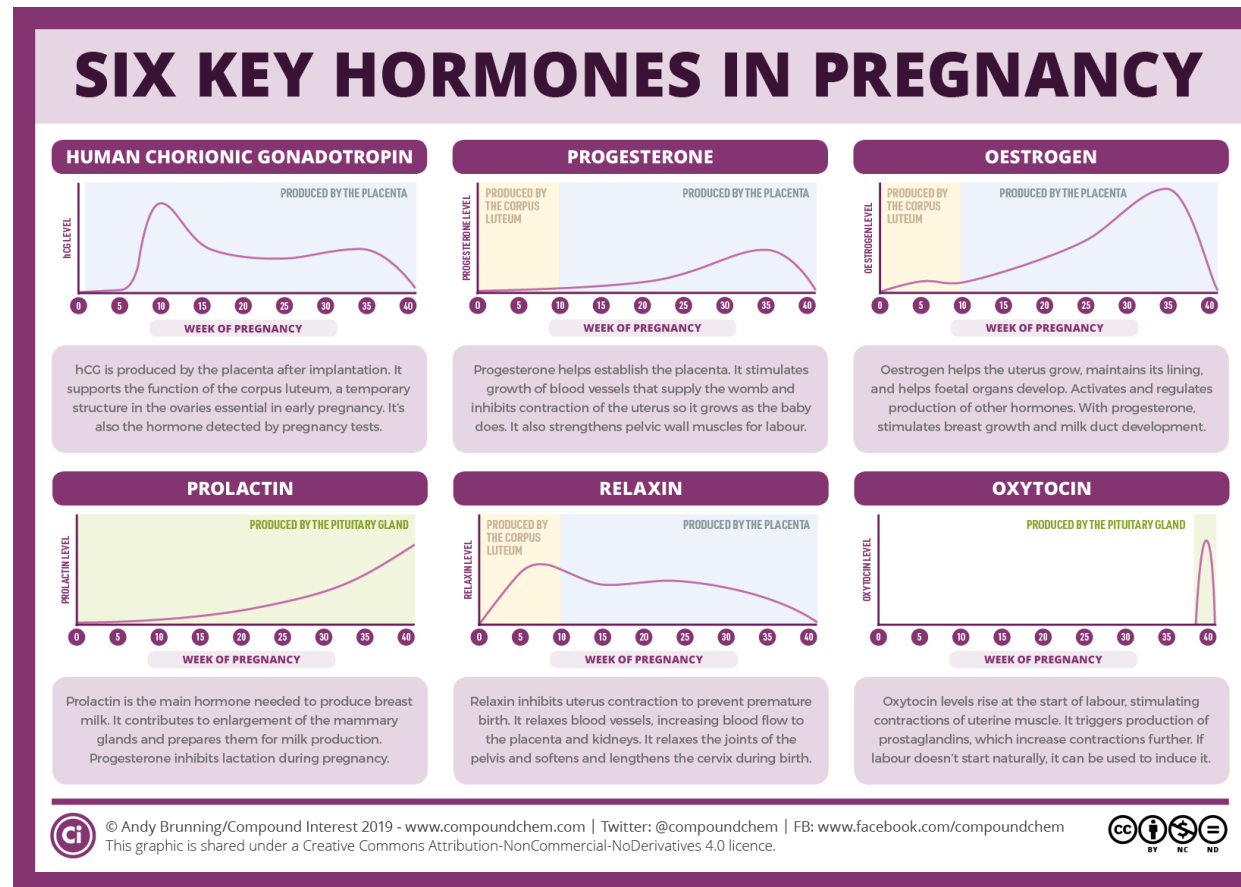
Anna Joy Rogers, MD, DrPH

Importance

- Transformations in nearly every organ system
- Fundamental to caring for pregnant women
- Misinterpreting normal adaptations can lead to unnecessary testing
- Help understand maternal symptoms and complications

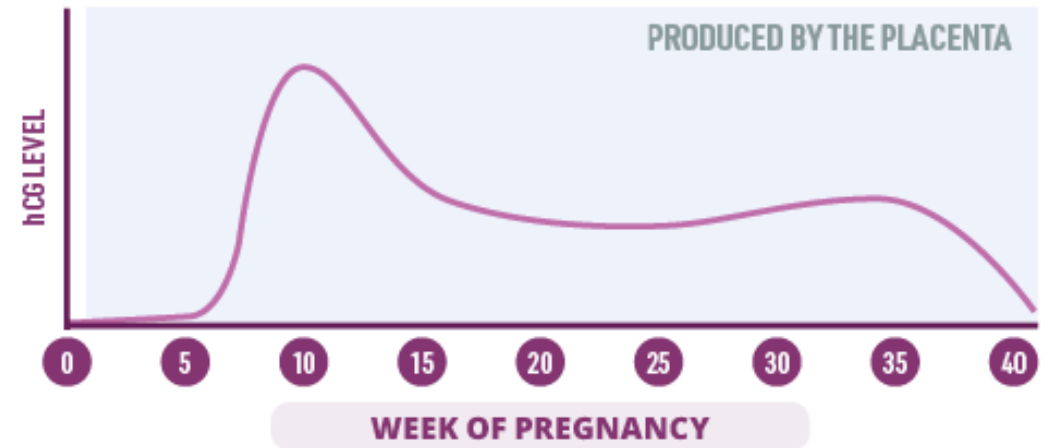


Hormonal Adaptations



Hormonal: Human chorionic gonadotrophin

- Source: placental syncytiotrophoblasts
 - Detectable ~8 days post fertilization
 - Peaks at ~8-11 weeks gestation
- Functions:
 - Maintains corpus luteum
 - Promotes uterine angiogenesis
 - Suppresses maternal immune response to embryo



- Clinical relevance:
 - Marker of abnormal pregnancies, multiple gestation

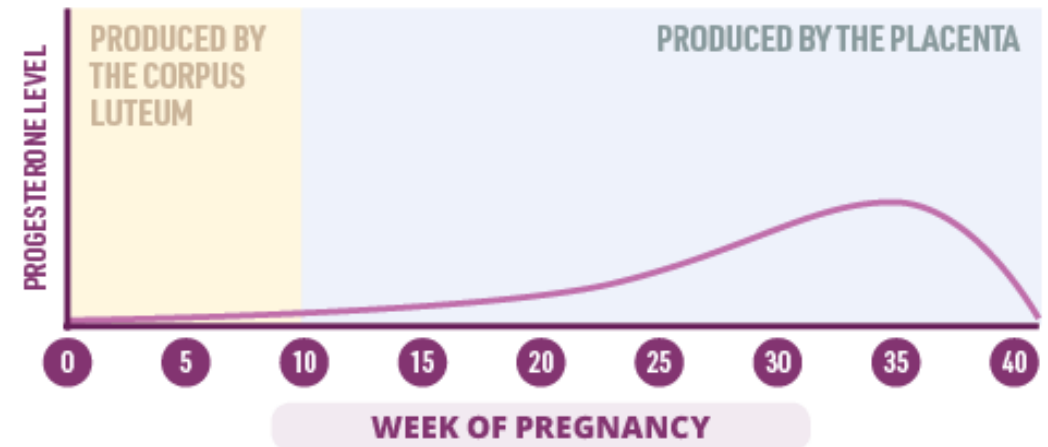
Hormonal: Progesterone

- Source:

- Corpus luteum (first 8-10 weeks)
- Placenta

- Functions:

- Decidualization of stromal cells
- Promotes lobules and alveoli development in breast
- Relaxes smooth muscle
 - Maintains uterine quiescence
 - GI (constipation, GERD)
 - Musculoskeletal changes

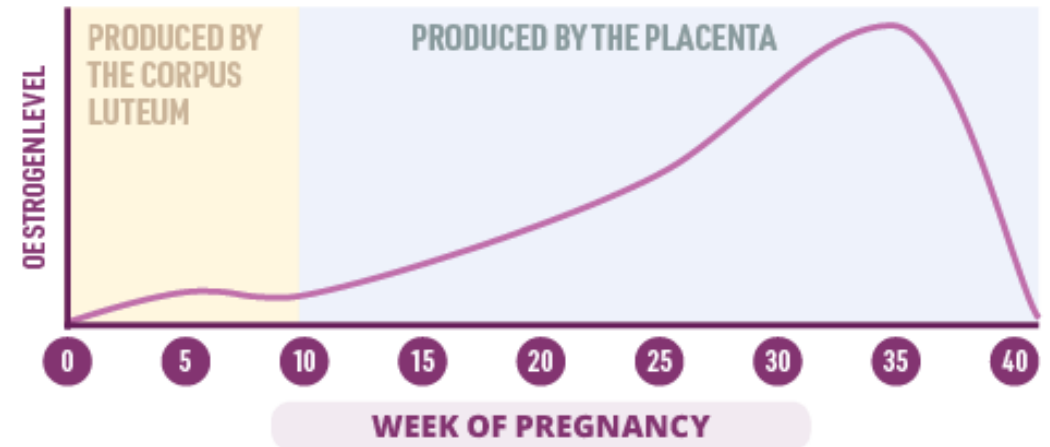


- Clinical relevance:

- Marker for early pregnancy failure
- Progesterone supplementation in shortened cervix
- Luteal phase deficiency support

Hormonal: Estrogen

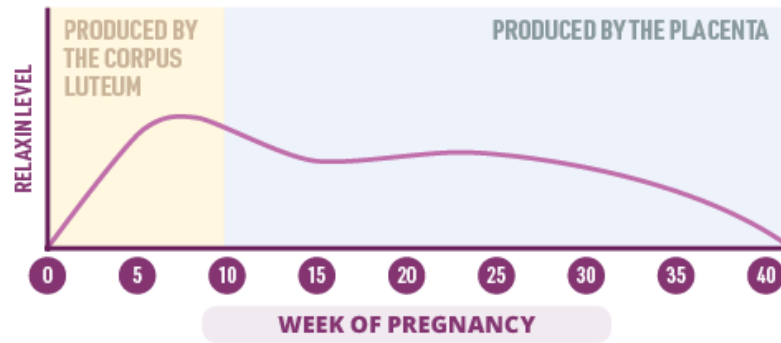
- Source:
 - Corpus luteum (first 8-10 weeks)
 - Placenta
- Functions:
 - Stimulates proliferation of uterine stromal cells, glands, and vasculature
 - Promotes proliferation and branching of breast ductal system
 - Supports maternal immune tolerance of the fetus



- Clinical relevance:
 - Low levels of estriol in quad screen associated with Trisomy 18 & 21
 - High levels are associated with hyperpigmentation

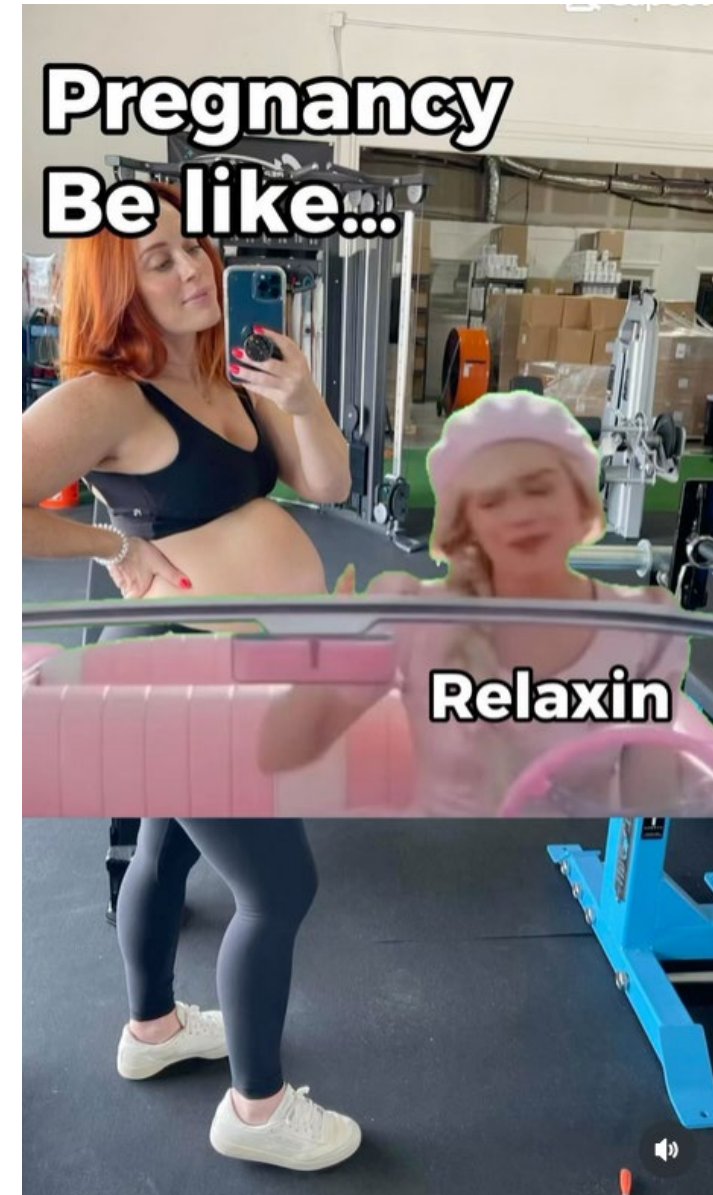
Hormonal: Relaxin

- Source:



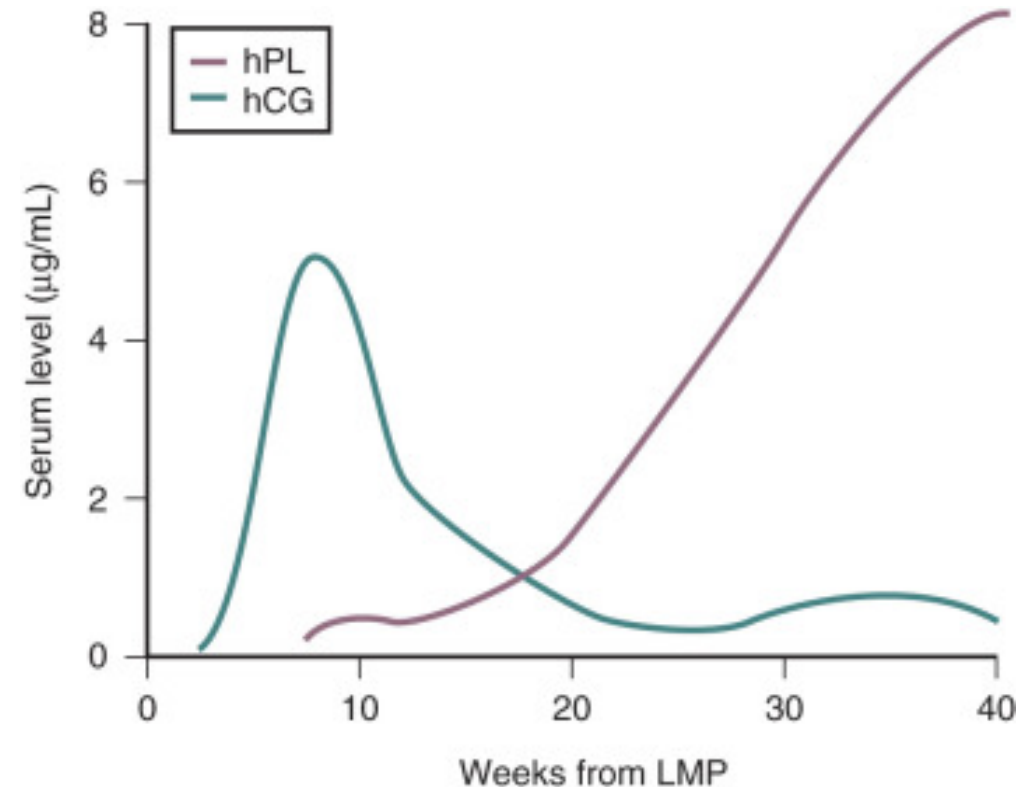
- Functions:

- Remodels reproductive tract connective tissue
- Increases renal blood flow
- Increases ligament laxity



Hormonal: Human placental lactogen

- Source:
 - Placental syncytiotrophoblasts
- Functions:
 - Induces insulin resistance in mother to ensure glucose availability for fetus
 - Promotes lipolysis as an alternate maternal energy source
- Clinical relevance:
 - Contributes to development of gestational diabetes



Jerome F. Strauss III, Sam A. Mesiano. Chapter 41 - Placental Production of Peptide, Steroid, and Lipid Hormones, Maternal-Fetal and Neonatal Endocrinology Physiology, Pathophysiology, and Clinical Management 2020, Pages 685-706

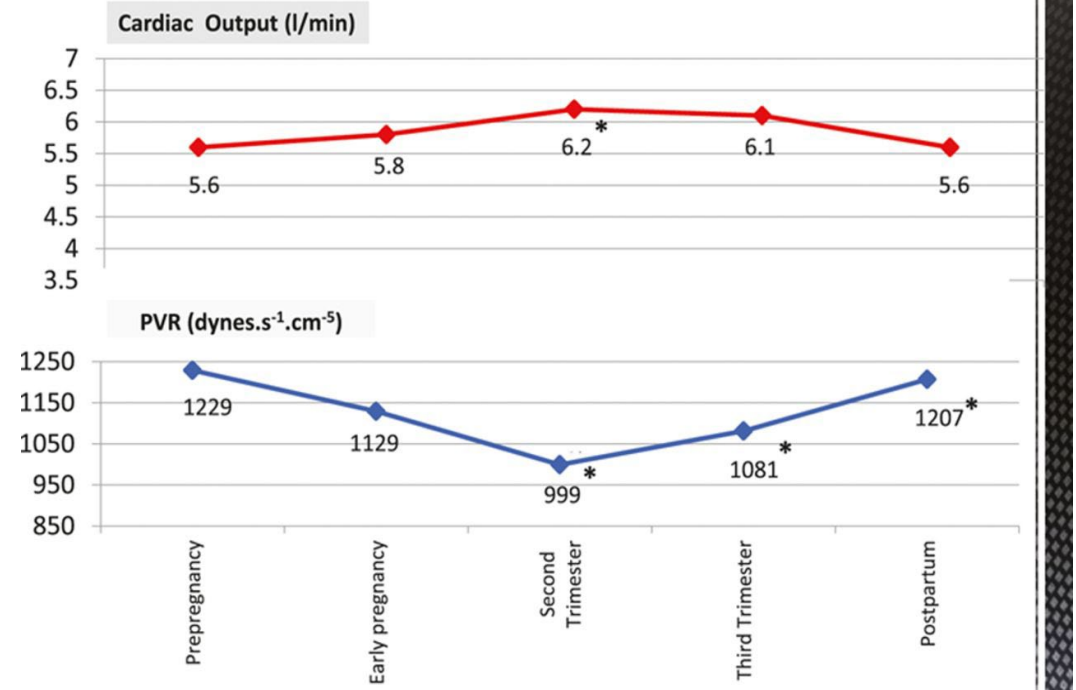
Cardiovascular

- Cardiac output

- From 4.5 L/min → 6.0 L/min
- Begins to increase as early as 5 weeks' gestation
- ↑ 30–50% by mid-pregnancy
- Peaks ~20–24 weeks

- Peripheral vascular resistance

- Nadir in second trimester
- BP rises toward baseline by term
- Due to progesterone and relaxin



Cardiovascular Physiology of Pregnancy. Monika Sanghavi, and John D. Rutherford. Circulation Volume 130, Number 12

Cardiovascular

- Vasodilation

- Promotes uterine blood flow
- Increased risk of peripheral edema, varicosities, supine hypotension

- Inferior vena cava compression

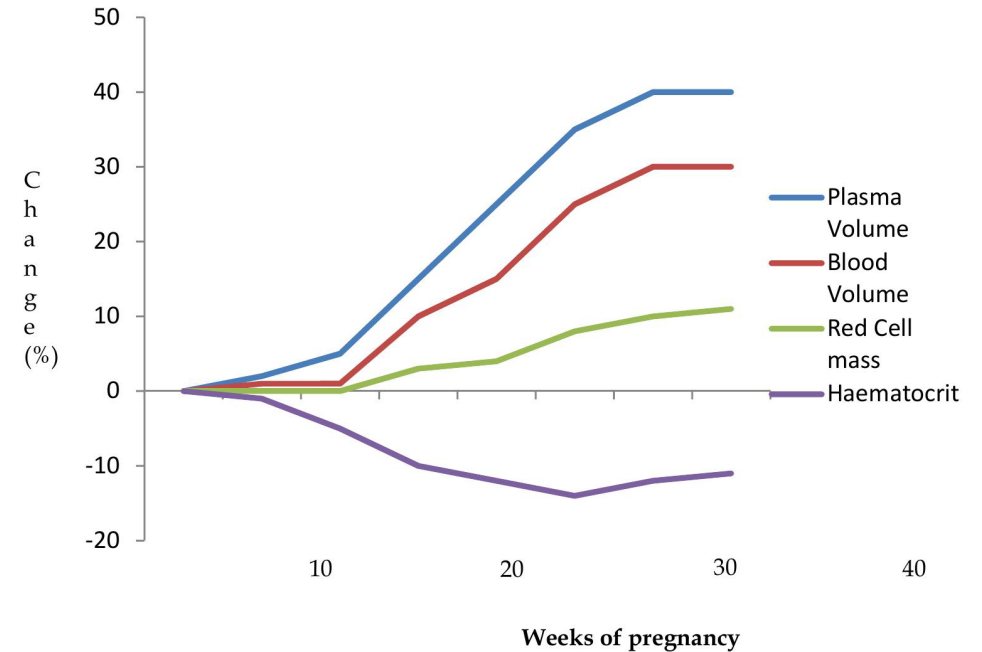
- Uterine enlargement compresses IVC when supine
- Relieved by left lateral position

- Heart sounds

- Increased blood volume, CO, SV, HR lead to hyperdynamic circulation
- Enhanced flow causes turbulent flow across aortic and pulmonic valves
- Physiologic:
 - Grade $\leq 2/6$ systolic ejection murmur
- Pathologic
 - Diastolic component
 - Pansystolic or late systolic quality
 - Grade $\geq 3/6$
 - Associated symptoms: dyspnea,

Hematologic

- Plasma volume
 - Increases by ~40–50% above baseline
 - Begins at 6 weeks; Peaks by 32 weeks
- Red blood cell mass
 - Increases ~20–30%
 - ↑ erythropoietin
 - Less than plasma volume increase
 - Physiologic anemia
 - Importance of iron intake



Ezechi Oliver and Kalejaiye Olufunto. Management of Anaemia in Pregnancy. Anemia. 2012
DOI: 10.5772/28646

Trimester	1 st	2 nd	3 rd
Hemoglobin lower limit of normal (g/dL)	11.0	10.9	11.0

Coagulation

- **Procoagulants**
 - ↑ Fibrinogen, Factor VII, VIII, X
- **Anticoagulants**
 - ↓ Protein S
- **Fibrinolytics**
 - ↓ Plasminogen activator inhibitors
- **Prothrombotic state**
 - Reduce hemorrhage risk at delivery
 - Increased risk of VTE

Coagulation factor	Change from non-pregnant state
Antithrombin III	No change
Plasma fibrinogen (factor I)	↑
Factor II	No change
Factor V	No change
Factor VII	↑
Factor VIII	↑
Factor IX	No change
Factor X	↑
Free protein S	↓
Plasminogen activator inhibitor 1	↑
Plasminogen activator inhibitor 2	↑
Protein C	No change
von Willebrand factor	↑

<https://miscarriagehopedesk.org/blood-clotting-disorders-miscarriage/>

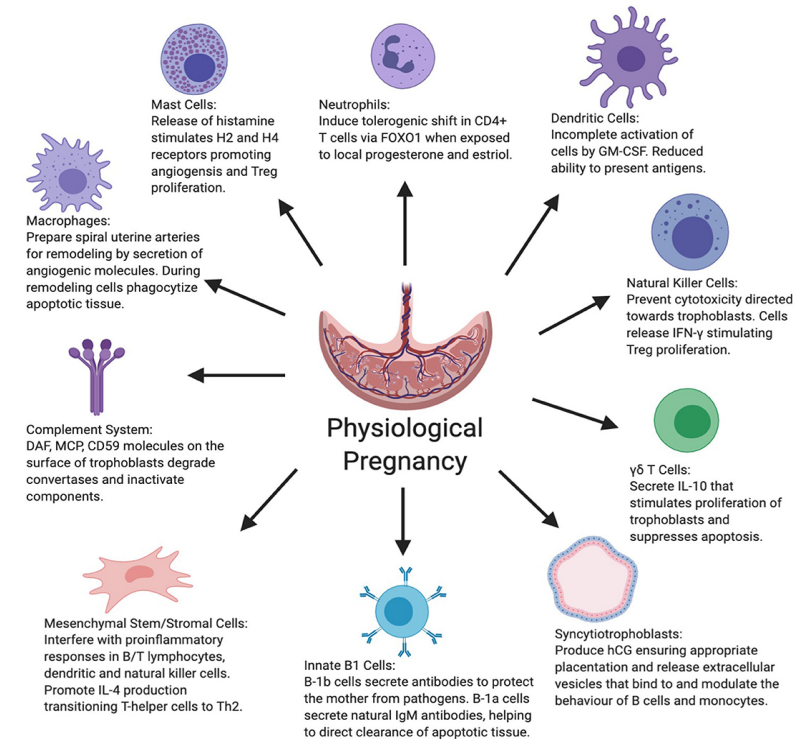
Immunologic

- **Innate immunity**

- ↑ Innate immune activity (NØ, MØ, NK cells)
- ↑ Inflammatory cytokines (e.g. IL-6, TNF-α)
- Enhanced inflammatory responses to infections
 - Higher vascular permeability, leukocyte adhesion, coagulation activation

- **Adaptive immunity**

- ↓ Cell-mediated immunity
- ↑ Humoral immunity
- Varied effect on autoimmune diseases
 - Improved rheumatoid arthritis & multiple sclerosis
 - Worsened lupus & asthma



Anuman I, Pienaar D, Suvakov S, Simic TP, Garovic VD, McClements L. Mechanisms of Key Innate Immune Cells in Early- and Late-Onset Preeclampsia. *Front Immunol.* 2020 Aug 18;11:1864. doi: 10.3389/fimmu.2020.01864. PMID: 33013837; PMCID: PMC7462000.

Respiratory

- Increased CO₂ Elimination

- ↑ Minute ventilation
 - Respiratory rate unchanged
 - ↑ Tidal volume (30–40%)
- Normal to have chronic respiratory alkalosis
 - ↓ PaCO₂, ↑ pH
- A "normal" CO₂ in pregnancy is abnormal and may indicate impending respiratory failure

- Increased O₂ Consumption

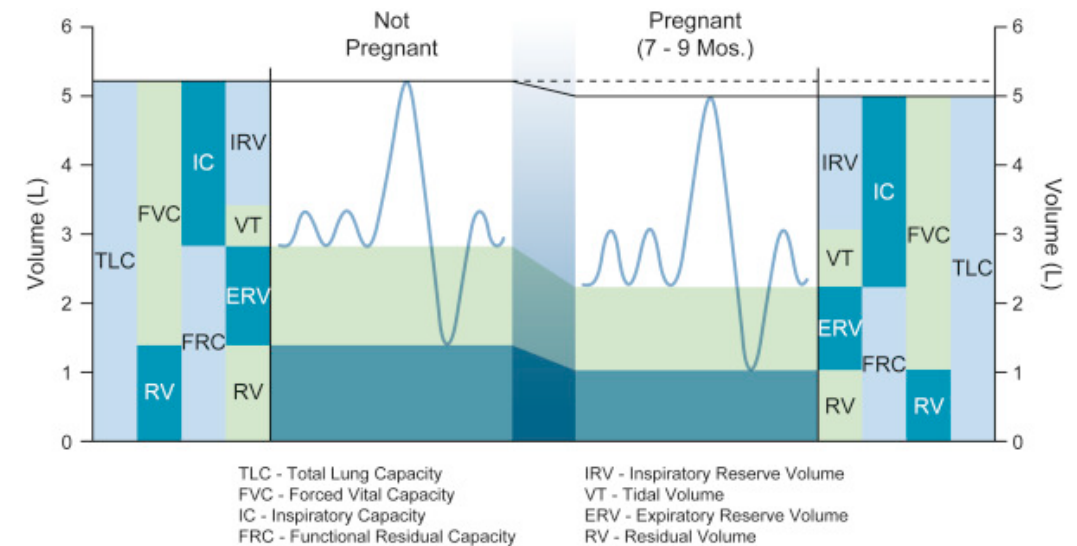
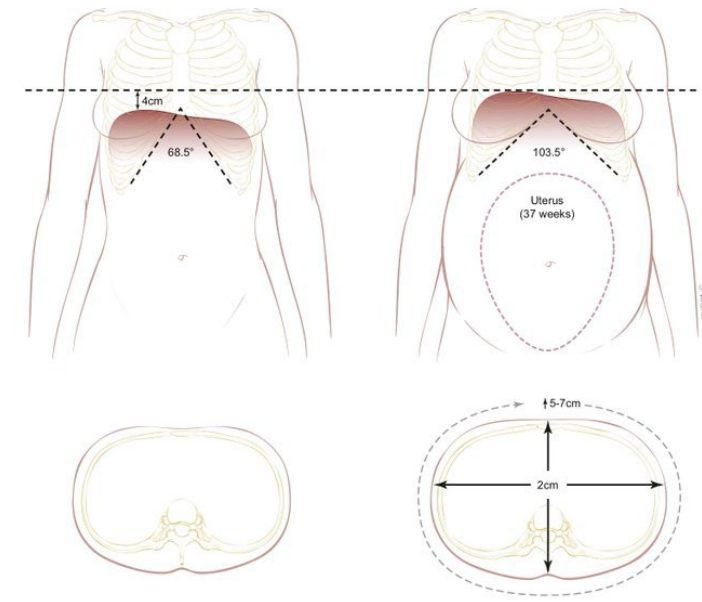
- Driven by metabolic demands of fetus, uterus, and maternal organs
- ↓ respiratory reserve
- More rapid desaturation during apnea or anesthesia

Table 3 Arterial blood gas measurement alterations in pregnancy			
Arterial blood gas measurement	1st trimester	3rd trimester	Nonpregnant
pH	7.42–7.46	7.43	7.4
PaO ₂ (mm Hg)	105–106	101–106	93
PaCO ₂ (mm Hg)	28–29	26–30	37
Serum HCO ₃ (mEq/L)	18	17	23

Sarah Dotters-Katz. Treatment Considerations for COVID-19 in Pregnancy. SMFM Webinar

Respiratory

- ↓ **Functional residual capacity**
 - Volume of air remaining in the lungs at the end of a passive exhalation
 - Due to elevated diaphragm by 4-5cm
 - FRC decreases by ~20% (400–500 mL) by the third trimester
 - Lower respiratory reserve
 - Increased risk of hypoxemia & rapid desaturation during airway obstruction, sedation, or anesthesia



Hegewald MJ, Crapo RO. Respiratory physiology in pregnancy. Clin Chest Med. 2011 Mar;32(1):1-13. 10.1016/j.ccm.2010.11.001.

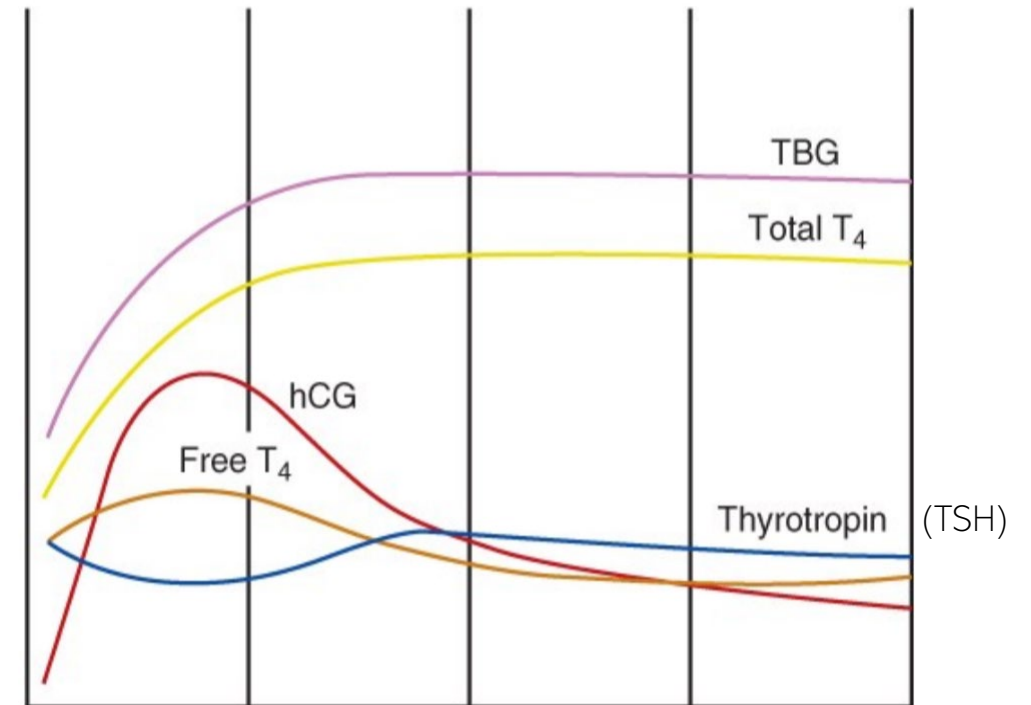
Endocrine

- **Thyroid**

- Estrogen ↑ hepatic production of thyroid binding globulin
 - Elevated bound T4 values
 - No change in free T4 and T3 levels
 - Fetus depends on maternal T4 until 20 weeks
- Increased basal metabolic rate

- **Structural similarity of TSH and hCG**

- High serum hCG levels can cause thyroid stimulation
- α -subunits are identical
- β -subunits differ amino acid sequence



Editors. In: Cunningham F, Leveno KJ, Dashe JS, Hoffman BL, Spong CY, Casey BM. eds. *Williams Obstetrics*, 26e. McGraw Hill; 2022. Accessed July 14, 2025.
<https://accessmedicine.mhmedical.com/content.aspx?bookid=2977§ionid=263812626>

Endocrine

- Hypothyroidism

- 2-10/1,000 pregnancies
- Risks of miscarriage, preeclampsia, preterm birth, stillbirth, neurodevelopment
- Anti-TPO Ab rarely cross placenta
- Treatment: Levothyroxine

- Hyperthyroidism

- 2-7/1,000 pregnancies
- Risks of preeclampsia, heart failure, thyroid storm, preterm birth, stillbirth, miscarriage
- TSH-binding Ab may cross placenta
- Treatment: Propylthiouracil (T1)
 - Methimazole or PTU (T2, T3)

“Universal testing for thyroid disease in pregnancy is not recommended. Indicated testing of thyroid function should be performed in women with a personal or family history of thyroid disease, type 1 diabetes mellitus, or clinical suspicion of thyroid disease.”

ACOG PB 223: Thyroid Disease in Pregnancy

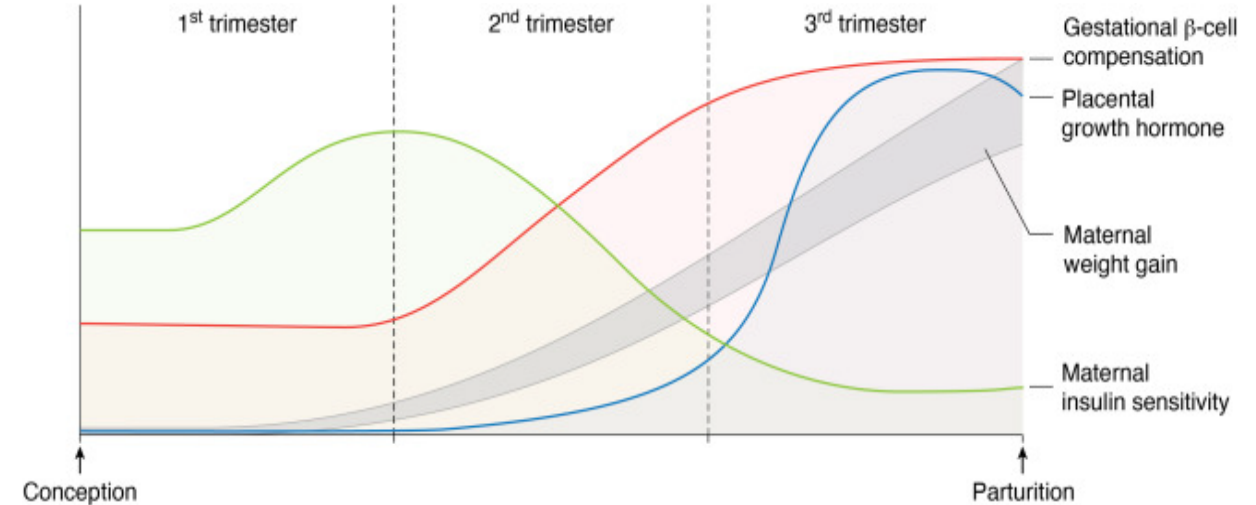
Endocrine

- Pituitary gland

- ↑ Prolactin prepares breasts for lactation
- Oxytocin surges in labor
- Placental growth hormone
 - ↑ metabolism to increase availability of nutrients for fetus

- Adrenal gland

- ↑ Cortisol: Glucose for fetus
- ↑ Aldosterone: sodium retention and plasma volume expansion



Taofeek O. Usman, Goma Chhetri, Hsuan Yeh, H. Henry Dong. Beta-cell compensation and gestational diabetes. Journal of Biological Chemistry, Volume 299, Issue 12, 2023, 105405, <https://doi.org/10.1016/j.jbc.2023.105405>.

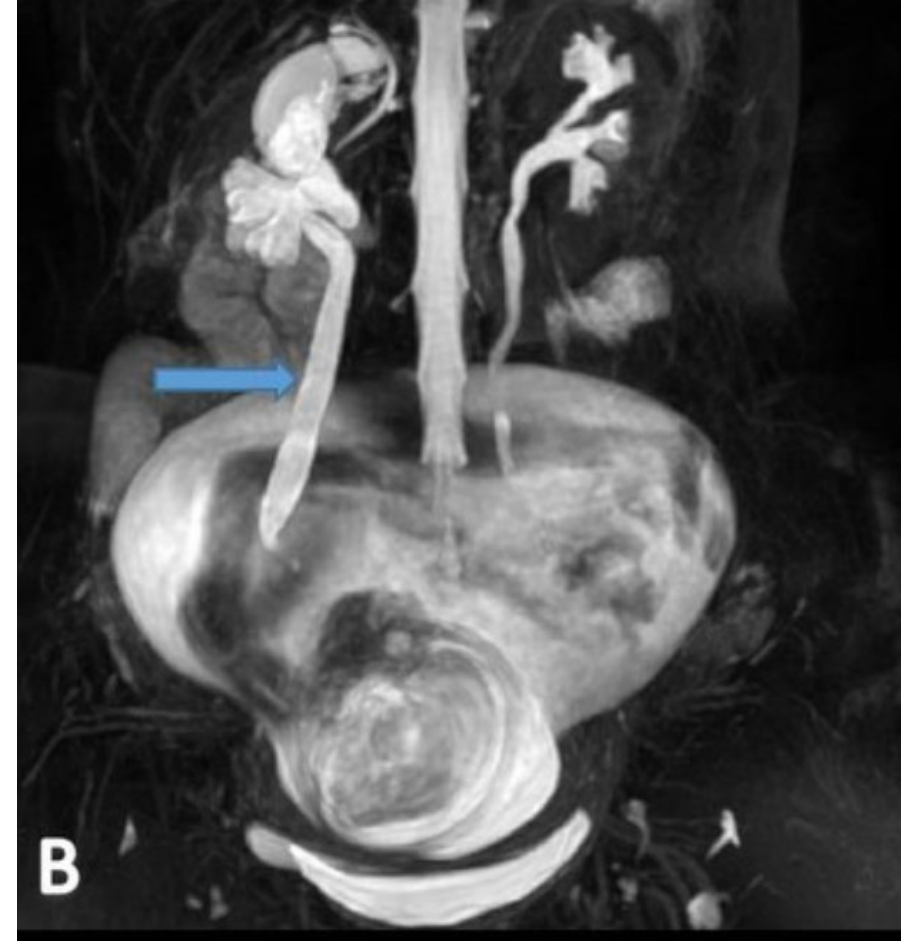
- Pancreas

- ↑ Insulin resistance
- Driven by human placental lactogen, cortisol, progesterone, and placental growth hormone

Renal

- Anatomic changes

- Kidneys: enlarge by 1-1.5cm
- Hydronephrosis: dilated renal calices, more prominent on the right
- Hydroureters: Ureters dilate and exhibit decreased peristalsis
- Bladder: compression by uterus



Kadam, Dipali, Saurabh Patil, Avinash Dhok, and Meenal Jain. "MR urography in evaluating obstructive uropathy: one stop shop." *International Surgery Journal* 6, no. 3 (2019): 944. <http://dx.doi.org/10.18203/2349-2902.isj20190829>

Renal

- ↑ Glomerular filtration rate
 - Due to ↑ renal plasma flow and vasodilation
 - Serum creatinine and BUN decrease
 - Normal Cr: ~0.4–0.6 mg/dL
 - 1.0 may indicate renal dysfunction
 - Mild proteinuria is normal
 - (<300 mg/day)
 - Mild glycosuria is normal
- Renin-Angiotensin-Aldosterone System (RAAS) Activation
 - Supports sodium and water retention
 - Maintains plasma volume expansion
 - Lower osmolality and thirst threshold leads to increased water intake and mild hyponatremia

Gastrointestinal

- Decreased gastric motility

- ↑ Nausea and emesis
- ↓ Gastric emptying
 - Bloating, acid reflux
 - NPO status prior to procedures
- ↑ Intestinal transit time
 - Due to progesterone, gravid uterus, iron supplementation
 - Constipation
 - Hemorrhoids from straining

- Gallbladder and biliary tract

- Gallbladder stasis
- ↑ Cholesterol saturation → increased risk of gallstones

- Liver

- ↑ Alkaline phosphatase (produced by the placenta)
- AST, ALT, bilirubin unchanged
- Serum albumin ↓ due to hemodilution

Genital Tract

- Uterus

- Growth from ~70 g at conception to ~1100 g at term
 - Hypertrophy of muscle fibers, not hyperplasia

- Vagina

- ↑ Glycogen in epithelial cells metabolized by *Lactobacillus* to lactic acid
 - Acidic pH (3.5–6.0)
 - Protects against pathogens

- Cervix

- By 4-6 weeks, increased edema, vascularity, hypertrophy and hyperplasia of glands
 - Chadwick sign (bluish coloration)
- Endocervical glands proliferate → form the mucus plug
 - Protect from ascending infection

Musculoskeletal & Skin

- Posture

- Lumbar lordosis to maintain center of gravity
- Gait change due to pelvic ligament laxity, driven by relaxin and progesterone
- Shift in weight-bearing

- Abdominal wall

- Striae gravidarum
- Diastasis recti

- Skin

- Hyperpigmentation due to ↑ melanocyte-stimulating hormone and estrogen
- Face, areolae, Linea nigra, perineum
- Spider angiomas, palmar erythema

- Hair

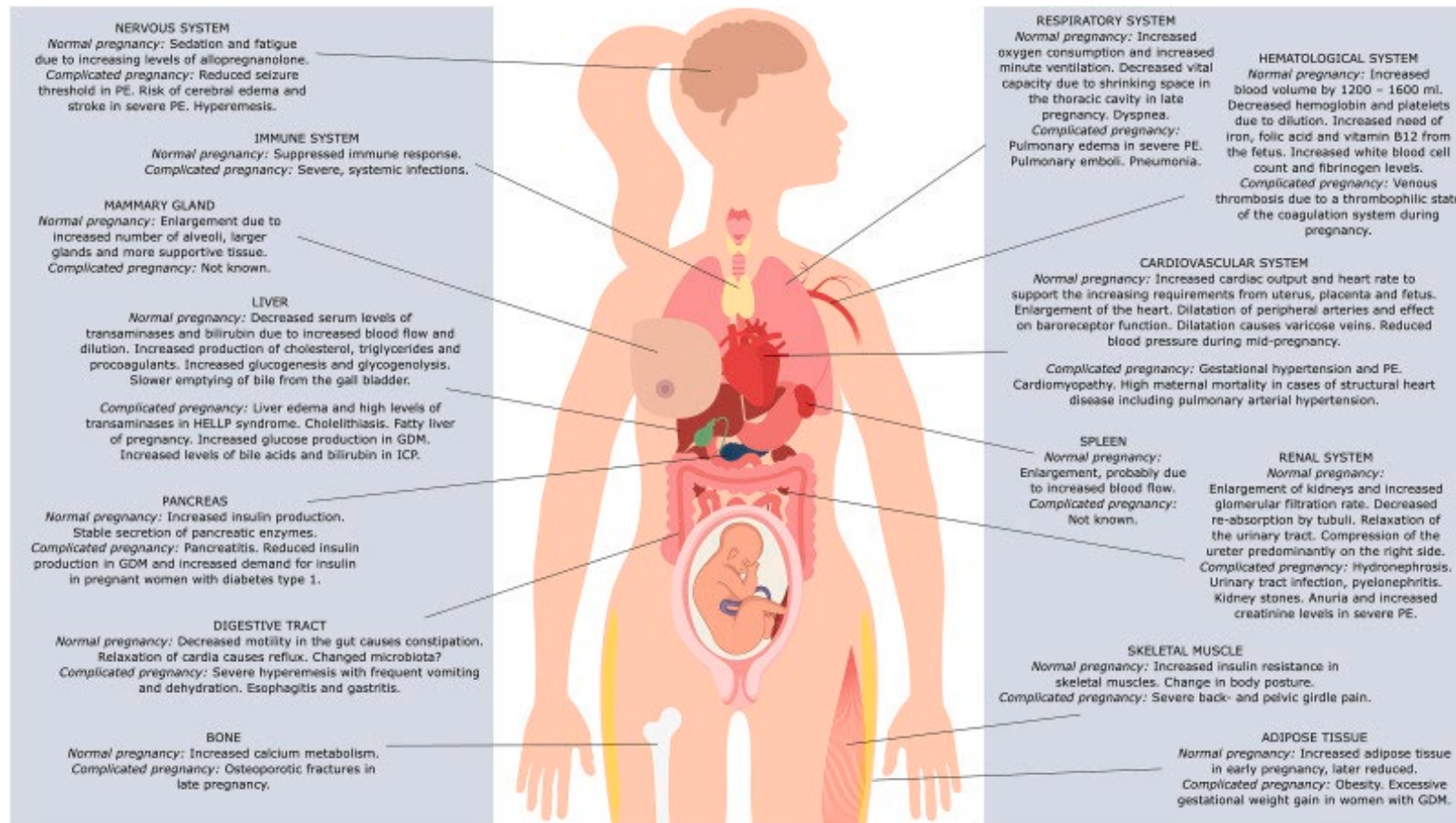
- Hair growth ↑ during pregnancy (anagen phase prolonged)
- Telogen effluvium: postpartum hair loss 2–4 months after delivery



<https://drmaryamzamani.com/blog/what-is-melasma/>.

<https://veincenter.doctor/child-dermatologist-for-facial-spider-veins/>

Questions?



Vinnars MT, Bixo M, Damdimopoulou P. Pregnancy-related maternal physiological adaptations and fetal chemical exposure. Mol Cell Endocrinol. 2023 Dec 1;578:112064. doi: 10.1016/j.mce.2023.112064. Epub 2023 Sep 6. PMID: 37683908.

Thank you!
(Time for RELAXIN on the beach...)

