



Physiological changes in pregnancy

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References: 437 Lectures And Notes, 436 teamwork

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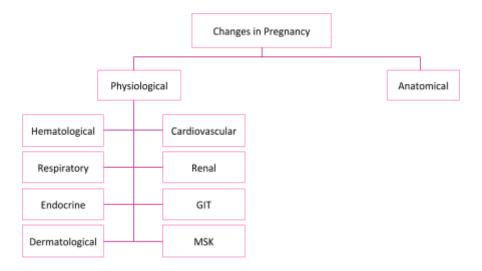
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Objectives:

- 1. Symptoms and physical findings of each organ system
- 2. Physiologic versus pathologic changes
- 3. Diagnostic tests and interpretations during physiological changes







This lecture talks about **physiological** findings, so whenever we say increase or decrease it means within the normal limits.

Hematological changes

Hypercoagulable pregnancy is a hypercoagulable state! More risk for DVTs and PEs. (pregnancy and 6 weeks of postpartum period)

- High Estrogen "which is the main factor" & Vascular stasis. there is a decrease in vascular resistance and vasodilation that contribute to stasis.
- Increased risk for thromboembolic disease
 - Increase in fibrinogen, all coag factors VII, VIII, IX, and von Willebrand Factor except II, V, XII,
 - Fall in protein S and sensitivity to APC that's why you never test for thrombophilia during pregnancy which can be misleading. you have to wait postpartum 2-3 months until everything goes back to normal.
- Fall in platelets. Normally (non-pregnant) 150000-300000, but in pregnancy it is about 160000 (lower normal) but if it's less than 150000 this is pathological and factor XI and XIII.
- Increase in WBC (same concept as the platelets).
- Hemoglobin:
 - Stays the same (12-16 g/dL) initially. mainly in the first 12 weeks.
 - May drop down to 10 g/dL and still be normal physiological anemia. But Hb of 8 is not normal!
 - Normal pregnancy Hgb is 10-14 g/dL. 9 is not normal
- Decrease Hematocrit (Hct): 38-47%.
 - Normal pregnancy Hct (Hematocrit) is 32-42 later in pregnancy.

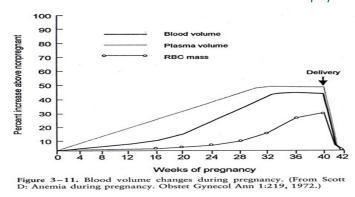




So **important** to differentiate between pathological and physiological anemia. if the women present for the 1st time with hemoglobin= 13 then after 2 months= 11 that consider physiological anemia. so hemoglobin 10 or more consider physiological anemia. and always keep in mind that pregnant lady may develop iron deficiency anemia.

Physiologic anemia of pregnancy

- Plasma volume increases 50-70 % (Beginning by the 6th week), the main reason.
- RBC mass increases 20-35 % (Beginning by the 12th week).
- Note the difference in the increase between the two components, the plasma increases way more than the RBC and that results in hemodilution and not polycythemia for example.
- Hemodilution. Plasma volume increases and that is what causes physiological anemia.



The graph clearly shows when physiological anemia happens. Plasma volume will start to increase earlier than RBCs mass. The blood volume will hugely increase in the 2nd trimester due to both plasma and RBCs increase. After delivery everything goes back to normal.

Hematological change in summary

- Decreases in:
 - Haemoglobin concentration & Haematocrit.
 - Plasma folate concentration. That is why we supply women with folic acid. Its imp bc plasma volume increases cause hemodilution and to prevent certain congenital anomalies such as neural tube defect and congenital heart disease.
- Increases in:
 - White cell count. increases in pregnancy. usually in non-pregnant ladies less than 11. but in pregnancy 12, 12.5, 13 and nothing to worry about. but if women show up with WBC= 15 or 20 thats a pathological increase, or if come with fever and WBC= 13 its pathological.
 - Erythrocyte sedimentation rate & Fibrinogen. higher in pregnancy

A useless test in pregnancy is D-dimer when you are suspecting DVT because it also will be high physiologically.







Normal changes in heart sounds during pregnancy

- Increased loudness of both s1 and s2. because of increase blood and plasma volume
- Increased splitting of mitral and tricuspid components of s1.
- No constant changes in s2.
- Loud s3 by 20 weeks' gestation. especially in the 2nd trimester.
- <5% with s4.</p>
- >95% develop systolic murmur which disappears after delivery (Ejection systolic murmur especially in 3rd trimester at left sternal border due to increased CO passing through the aortic and pulmonary valves).
- 20% have a transient diastolic murmur. Diastolic murmurs are never normal in pregnancy and must be investigated.
- 10% develop continuous murmurs due to increased mammary blood flow, so it's not coming from the heart, it comes from mammary vessels b/c of increased blood flow to the breast. you have to know where it's coming from.

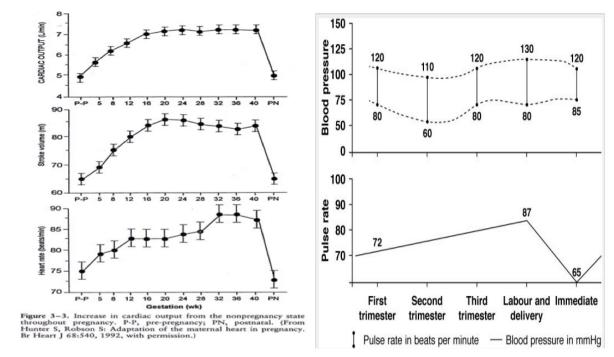
Cardiovascular changes

- Heart rate increases (10-20%). For example, the normal heart rate is between 60-100 Bt/min. And the average of young non-pregnant women is 70, so the pregnant women could have an average of 90-95 (higher than usual but within the normal range). But if the HR of a pregnant woman is 120 this is abnormal and **pathological**.
- Stroke volume increases (10%).
- Cardiac output increases (30-50%).
- Mean arterial pressure decreases (10%). especially in the 2nd trimester.
 - Systolic and diastolic values both decline early in the first trimester, reaching a nadir by 24–28 weeks, then they gradually rise toward term but never return quite to prepregnancy baseline. Arterial blood pressure is never normally elevated in pregnancy Central venous pressure (CVP) is unchanged with pregnancy.
- Peripheral resistance decreases (35%) وتقدر و تثبتو المعلومة هذي بأن ضغط المرأة الحامل ما يزيد في (35%) الأوضاع الطبيعية مع إن حجم الدم زاد مرة! معناها مقاومة الأوعية الدموية قلّت لأن الضغط يتكون من كمية الدم الأوضاع الطبيعية مع إن حجم الام زاد مرة! معناها مقاومة الأوعية الدموية

Blood pressure = Vascular resistance + Blood volume.







Left graph: The graph shows how high HR **(but still under 100!)** is in the pregnancy and how immediately after delivery it drops back to normal as soon as the placenta is out and the uterus contracts.

Right upper graph: Notice the slight BP drop in the 2nd trimester which may lead to <u>postural</u> <u>hypotension</u>. very common.

Right lower graph: Notice the slight increase within the normal range. As discussed before.

Signs & Symptoms of Normal Pregnancy that may Mimic Heart Disease:

Signs	 Peripheral edema dig more in history because there is a fine line between physiological and pathological causes such as pre-eclampsia and heart failure. JVD increases a little especially in the 3rd trimester. Arrhythmia is a pathological not a physiological finding.
Symptoms	 Reduced exercise tolerance. Dyspnea dig in history! There is a huge difference between someone who is short of breath at rest and someone who's short of breath after walking.
Auscultation	S3 gallop.Systolic ejection murmur.
Chest X-ray	Change in heart position & sizeIncreased vascular markings.
EKG	 Nonspecific ST-T wave changes. All the changes are slight and not that significant Axis deviation. LVH.





Renal change

Change in filter

Renin – stimulated by progesterone

- Also made by placenta.
- Angiotensinogen → Angiotensin I → Angiotensin II → Aldosterone Distal tubule (and that's what causes edema and water retention).
 - Net absorption of Na+.
 - Excretion of K+.
 - Water retention: 6-8 liters. most of the time water retention in pregnancy is due to renal reasons not cardio. So the total reabsorption and excretion is changed.

Increased renal blood flow due to increased cardiac output

- 50-75% increase.
- GFR 50% increase.
- Decreased Albumin = lower colloid oncotic pressure.bc of increasing of plasma volume = lower colloid oncotic pressure, which this decreasing of albumin leads also to fluid retention and peripheral edema

Other urinary tract change

- Ureteral dilation / hydroureter. hydroureter is very common, and it happens in one side than the other bc uterus presses in one kidney more than the other
 - The smooth muscle of the renal pelvis and ureters relaxes due to the effect of prostaglandin, causing their dilatation. This increase the capacity of the renal pelvis and ureters and increase the chances of urinary infection.
 - Later exacerbation by uterine obstruction.
 - Urinary stasis causing mild hydronephrosis.
- Dilation of the pelvis and calyces.
- Increased kidney size, due to increase in renal blood flow. This doesn't reverse until 3
 months postpartum.

The urinary tract and renal function

Renal changes

- Blood flow increases (60-75%).
- Glomerular filtration increases (50%).





- Clearance of most substances is enhanced including iodine.
- Plasma creatinine, urea and urate are reduced.
- Glycosuria is normal, due to enhanced clearance of substances. The tubular reabsorption threshold falls from 195 to 155 mg/dL.
- Proteinuria—Urine protein remains unchanged.

Respiratory system



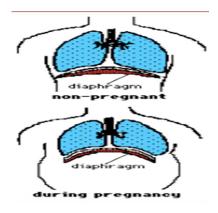
 Diaphragm pushed up and the lung capacity is changed. And different sternal angle and chest diameter.

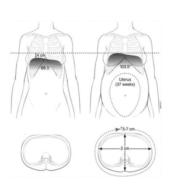
Consumption:

- Increase in needed oxygen.
- O₂ consumption Increases 15-20%.
- 50% of this increase is required by the uterus.
- Despite increase in oxygen requirements, with the increase in Cardiac Output
 and increase in alveolar ventilation oxygen consumption exceeds the
 requirements. Therefore, arteriovenous oxygen difference falls and arterial PCO₂
 falls. The rise in Vt produces a respiratory alkalosis with a decrease in Pco2 and
 an increase in pH. An increased renal loss of bicarbonate helps compensate,
 resulting in an alkalotic urine. especially in the third trimester.
- Stimulation: Progesterone stimulation.

Other change

- Increase in Subcostal angle and chest circumference and diameter
- Mucosal hyperemia
- Mild respiratory alkalosis
- Total lung capacity decrease by 15%
- Tidal volume (Vt): increases with pregnancy to 40% (30 40%). It is the only lung volume that does not decrease with pregnancy.
- Minute ventilation (Ve): increases up to 40% (30 40%) its the product of respiratory rate (RR) and Vt. RR remains unchanged with Vt increasing.
- Residual volume (RV): decreases up to 20% due to upward displacement of intraabdominal contents.









Endocrine glands

1- Pituitary gland

- Pituitary size increases by 100% making it susceptible to ischemic injury (Sheehan syndrome) from postpartum hypotension following postpartum bleeding.
- FSH and LH أحتاج أحفر بويضات جديدة تكبر ألريدي عندي حمل FSH and LH
- ACTH, Thyrotrophin, melanocyte hormone (causing skin pigmentation) and prolactin
 ↑.
- Prolactin level ↑ until the 30th week of pregnancy then more slowly to term.
 Sometimes the pregnant lady may tell you her breast is secreting milk and that is completely normal so just reassure the patient.

2- Adrenal gland

- Adrenal gland size is unchanged
- Total corticosteroids ↑ progressively to term. This will ↑ the tendency of pregnant women to develop abdominal striae, glycosuria and hypertension. hypertension is never normal in pregnancy but it slightly increase from baseline

3- Thyroid gland

- Enlarges during pregnancy, occasionally to twice its normal size. This is mainly due to colloid deposition caused by a lower plasma level of iodine, consequent on the increased ability of the kidneys to excrete during pregnancy.
- Oestrogen stimulates or increased secretion of thyroxine in binding globulin.
- Both T3 and T4 levels rise. This rise will not indicate hyperthyroidism.

Other endocrine

- Pancreas:
 - Carbohydrate metabolism.
 - Increased Insulin resistance.
 - Those changes are due to Human placental lactogen, cortisol.

Thyroid Function:

Increased thyroid binding globulin TIBG (via liver) resulting in Increased total T4 and T3. However, free levels is unchanged. This is because HCG suppresses TSH from the pituitary. why? because they share the alpha subunit. so especially in the 1st trimester when HCG increases dramatically, TSH will be low which is normal. We just repeat the test after the first trimester.

Adrenal function:

Free plasma cortisol is elevated bc CRH from placenta stimulates ACTH.





Gastrointestinal tract

- Slowed GI motility:
 - Constipation very common mainly due to decreased GI motility
 - early satiety due to the compression of the uterus on the abdomen.
- Relaxation of LES causing GERD (Heartburn) the most common GI symptom. due to high progesterone levels. Mostly in the 3rd trimester.
- Nausea / vomiting: Often proportional to HCG level usually in the 1st trimester because
 of increased HCG. women with twins will have more vomiting.
- Liver / gallbladder:
 - Biliary stasis, cholesterol saturation leading to more stones.
 - Bile salt buildup which may lead to itching (this is pathological and dangerous to the baby and the mum so any pregnant complain of itching check the bile).
 - Coagulation factors.
 - Increased binding proteins (thyroid, steroid, vitamin D)
- Digestive system slows due to progesterone.
- Nausea and vomiting.
- Ptyalism: increase salivation. especially in the 1st trimester
- Hemorrhoids due to the mechanical pressure (constipation). very common. we don't treat it, we just treat the symptoms. We treat when there is severe pain or complication.
- Displacement of the stomach and intestines.
- Appendix can be displaced to reach the right flank.
- Gastric emptying and intestinal transit times are
 - Delayed secondary to hormonal and mechanical factors.
- Pyrosis "heartburn" is common due to the reflux of secretions.
- Vascular swelling of the gums a lot of women will tell you they bleed when they brush their teeth cuz of the hypervolemia of the gums
- Gastric motility decreases and emptying time increases, this increase in stomach residual volumes, predisposes to aspiration pneumonia with general anesthesia at delivery





Dermatological changes a lot of it is not reversible

Skin change	Definition		
Striae gravidarum Starts red and then white	"Stretch marks" that develop in genetically predisposed women on the abdomen and buttocks. To minimize the effect of striae always tell women to moisturize.		
Linea nigra	Increased pigmentation of the lower abdominal midline from the pubis to the umbilicus.		
Chloasma or melasma gravidarum	Blotchy pigmentation of the nose and face.		
Spider angiomata and palmer erythema	From increased skin vascularity		
Chadwick sign	Bluish or purplish discoloration of the vagina and cervix as a result of increased vascularity.		

MSK changes

- Musculoskeletal consequence that ensues result of hormonal changes and weight gain (28lbs ave) include:
 - Force across a joint is increased up to two-fold
 - Joint laxity (Relaxin) in the anterior and posterior longitudinal ligaments of the lumbar spine put strain on the lumbar spine.
 - There is widening and increased mobility of the sacroiliac joints and pubic symphysis to facilitate the baby's passage through the birth
 - o canal.
 - A significant increase in the anterior tilt of the pelvis occurs, with increased use of hip extensor, abductor, and ankle plantar flexor muscles





Weight change

- there is an increase weight of approximately 12.5kg at term. During the 1st trimester: 1 kg, during 2nd trimester: 5 kg and during 3rd trimester: 5 kg
- the main increase occurs in the 2nd half of the pregnancy (0.5 kg/week)

Weight.. Where do the pregnancy Kilos go?

Where do the pregnancy Kilos go?				
Maternal stores of nutrients and muscle development	3 Kg			
Increased body fluid	2 Kg			
Increased blood	1.5 - 2 Kg			
Breast growth	600g			
Enlarged uterus	1 Kg			
Amniotic fluid	1 Kg			
Placenta	600g			
Baby	3.4 - 4 Kg			
Total	11 - 16 Kg			

Pre-pregnancy BMI	Weight gain in kilograms	Weight gain in pounds
Underweight (under 18.5 BMI)	12.5-18	28-40
Normal weight (18.5-25 BMI)	11.5-16	25-35
Overweight (25-30 BMI)	7-11.5	15-25
Obese	5-9	11-20

Anatomical change

Genital tract change

- A. Uterus: Uterine muscles grow to 15 times than pre-pregnancy length.
- Uterine weight increases from 50 g before pregnancy to 950 g at term.
- In the early weeks of pregnancy the growth is by hyperplasia and more partially by hypertrophy of the muscle fibers.
- By 20 weeks growth ceases and the uterus expands by distension.
- The uterine blood vessels also undergo hypertrophy and become increasingly coiled in the first half of pregnancy but no further growth after that.
- The lower uterine segment is that part of the lower uterus and upper cervix lying between the line of attachment of the peritoneum of the utero vesical pouch superiorly and the histological internal os interiorly.





B. The Cervix

- Becomes softer and swollen in pregnancy, with the result that columnar epithelium lining the cervical canal becomes exposed to the vaginal secretions.
- Prostaglandins act on the collagen fibres, especially in the last week of pregnancy. At the some time collagenase is released from leukocytes, which also helps in breaking down collagen. The cervix becomes softer and more easily dilatable the so called ripening of the cervix.

C. Vagina

- The vaginal mucosa becomes thicker, the vaginal muscle hypertrophy.
- There is an alteration in the composition of the connective tissue, with the result that the vagina dilates more easily to accommodate the fetus during delivery.
- Oestrogen → desquamation of the superficial vaginal mucosal cells with ↑ in vaginal discharge when pathogenesis enter the vagina (candida, trichomonas) they will flourish rapidly. you always want to detect candida because you have to treat it.

Breast changes

Summary

Hematology	 RBC mass increase by ~25% and plasma volume by ~50% (not proportional)that will lead to dilutional physiological anemia hg>10 with no symptoms. WBC,ESR increase. coagulation profile changes(increase in factors 6,7,10 and fibrinogen,and decrease level of protein C,S) pregnancy is hypercoagulable state.
Cardiovascular	 Normal changes in heart sounds:loud S1,S2 with systolic murmurs that disappear after delivery. increase in heart rate(20%),stroke volume(10%) and cardiac output(50%). postural hypotension due to decrease peripheral vascular resistance.
Others	 Increase O₂ consumption. Glucosuria. Increase in total T3,T4 the free is unchanged. Increase insulin resistant.





MCQs

1- which of the following is normal during pregnancy:

A- hemoglobin of 8.

B- glucosuria

C-grade 6 diastolic murmurs.

D-proteinuria.

2- In a normal cardiovascular adaptation during pregnancy. Which of the following tend to decrease:

A- Blood volume

B- Cardiac output.

C- Heart rate.

D- Peripheral resistance

3- How much does the Cardiac output increases during pregnancy?

A- 25%.

B- 50%.

C- 100%.

D- 78%.

4- in normal pregnancy are there any Ptt,pt or INR changes?

A- Yes

B- No

Answers: 1- B. 2- D. 3- B. 4 - B.