

TSH ↓  
T<sub>3</sub> ↑  
T<sub>4</sub> ↑

⇒ Hyperthyroidism (1ry) → Graves  
Graves  
toxic nodules

↑  
↑  
↑

⇒ Hyperthyroidism (2ry) → TSH-secreting adenoma.  
pit

---

TSH ↑  
T<sub>3</sub> ↓  
T<sub>4</sub> ↓

⇒ Hypothyroidism (1ry) → Hashimoto. (anti TPO)

↓  
↓  
↓

⇒ Hypothyroidism (2ry) → pitu. tumor.

# Thyroid development

The fetal thyroid arises from an outpouching of the foregut at the base of the tongue (foramen cecum).

It migrates to its normal location over the thyroid cartilage by 8-10 wk of gestation.

The thyroid bilobed shape is recognized by 7 wk of gestation.

Thyroglobulin synthesis occurs from 4 wk, iodine trapping occurs by 8-10 wk, and thyroxine (T4) and, to a lesser extent, triiodothyronine (T3) synthesis and secretion occur from 12 wk of gestation

Maturation of the hypothalamic-pituitary-thyroid axis occurs over the second half of gestation, but normal feedback relationships are not mature until 1-3 mo of postnatal life

\* premature at risk of transient ↓



Bind to  $\left\{ \begin{array}{l} \text{TBG} \\ \text{albumin} \end{array} \right.$

## Thyroid changes at birth

Maternal T4 plays a role in fetal development, especially that of the brain, before the synthesis of fetal thyroid hormone begins.

At birth, there is a surge of TSH that peaks by 12 hrs of age up to the next 2-5 days

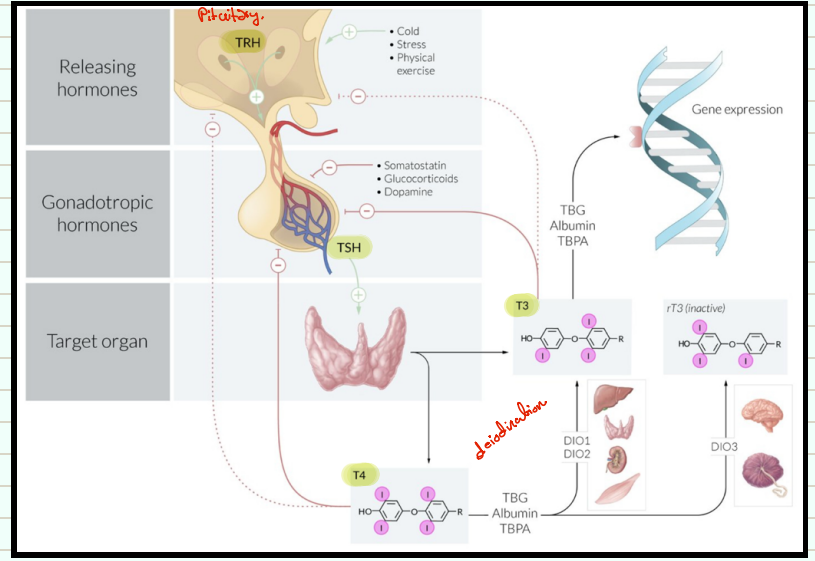
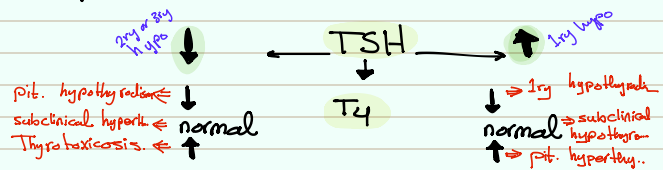
Followed by a rise in T4 and T3 levels, which peak during the first day of life and then slowly fall until 1 week - 2 weeks of life

Important for interpretation of newborn screening tests  
 Prematures and sick babies have the same changes with lesser quantity

Can cross the placenta  
 -> اسهل ما في المشيمة  
 -> اسهل ما في المشيمة  
 asymptomatic

## #Thyroid H. actions:-

- ↑ oxidative metabolism ⇒ O<sub>2</sub> consumption
- ↑ growth & development ⇒ BMR
- ↑ Essential for normal myelination ⇒ glucose metabolism
- ↑ Augmentation of cardiac function ⇒ fat metabolism
- ↑ Important for normal reproductive function



\* surge shortly after birth! 12hr!

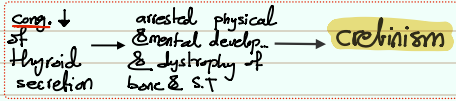
# Hypothyroidism

## # Congenital hypothyroidism

\* preventable cause of intellectual disability.

\* Mostly → primary. ♂:♀ 2:1.

# RF :- Multiple gestation, prematurity, babies of old age mother, Asian & hispanic, Down syndrome.



### Clinical manifestations:

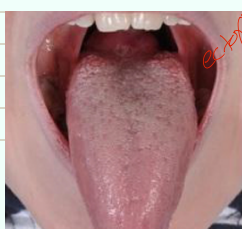
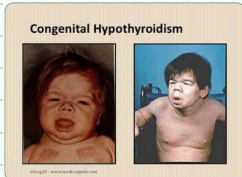
- Respiratory distress  $\Delta D$  1-3
- Large posterior fontanelle
- Abdominal distention
- Lethargy and poor feeding
- Prolonged jaundice (may be the earliest sign)
- edema
- Umbilical hernia
- Mottled skin
- Constipation
- Dry skin
- Hoarse cry/cry little, sleep much, poor appetite

# By age 3-6 months the clinical picture is fully developed

- Growth is stunted, extremities are short. Head size normal or increased.
- Open posterior fontanelle Dx: Down
- The mouth is kept open and large tongue protruding
- Dentition is delayed
- Neck is short and thick
- Myxedema of the eyelids, dorsum of hands and ext genitalia
- Carotenemia: yellow skin white sclera, skin is dry and scaly little perspiration
- Scalp is thickened hair is coarse brittle and scanty
- Low hair line
- Broad Hands, short fingers

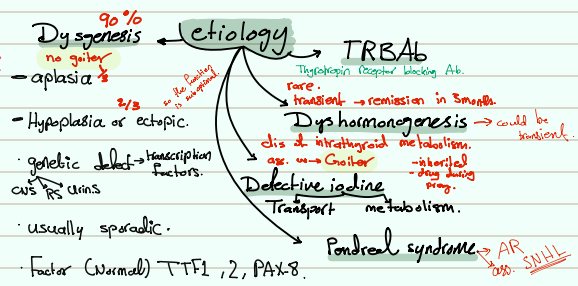
- Bone age delay
- Brain maturation
- thyroid hormone is essential for bone and brain maturation
- Development is retarded

- Muscles hypotonia with pseudohypertrophy



Mostly non-\*

acc & ectopic



The 7 P's of congenital hypothyroidism are Pot-bellied, Pale, Puffy-faced, Protruding umbilicus, Protuberant tongue, Poor brain development, and Prolonged neonatal jaundice

## # Most infants at birth ⇒ asymptomatic.

weight & length → normal  
HC → slightly ↑  
Screening is crucial!  
Scaly skin, no cry, weeks or month later

- Hints
- acrocyanosis
  - hypothermia
  - > 42 wks (gest)
  - Birth weight > 4kg
  - vis. anomalies  $\leq 2$

T<sub>4</sub>, T<sub>3</sub> ↓ low  
TSH ↑ > 100 mU/L

# Does it cause precocious puberty? No  
# Thyroxin affect <sup>only</sup> child head Proportion

Thyroid US  
Scintigraphy  
but treatment should not be delayed to obtain such imaging before 2 weeks.

- Thyroglobulin
- ↑: nonates Graves & subacute Endemic goiter.
- Skull xray
- ECG

## ؟ قبل الفحص #

- Initial blood Thyroxine assay + TSH assay follow up. IF T<sub>4</sub> < 10th percentile.
- TSH assay
- Simultaneous T<sub>4</sub> & TSH

## Treatment is

- Sodium Thyroxine T<sub>4</sub>
- New born → 10-15 ug/kg
  - Childhood → 3-5 ug/kg
  - Adults → 2 ug/kg

3:6

delay bone age

# # Acquired hypothyroidism

## Etiology

1. Lymphocytic thyroiditis most common
2. Autoimmune polyglandular syndrome
3. Subtotal thyroidectomy
4. Histiocytic infiltration
5. Secondary medication or irradiation

May develop during first year of treatment:

- Deterioration school work
- Poor sleeping habit
- Restlessness
- Short attention span
- Behavioral problems

# # Hashimoto #

♀ > ♂ x 4-7 times.  
 insidious onset after 6y  
 peak → adolescence.

MC presentation → Growth retardation  
 Goiter.

Most are euthyroid

Associated auto immune diseases:  
 DM1, Adrenal insufficiency, hypoparathyroidism,  
 pernicious anemia, vitiligo, alopecia, congenital rubella  
 Higher incidence in Turner and trisomy 21

## Clinical features

- Early-stage
  - Primarily asymptomatic
  - **Goiter**: nontender or painless, rubbery thyroid with moderate and symmetrical enlargement
  - **Hashitoxicosis** may occur: transient thyrotoxicosis due to follicular rupture of hormone-containing thyroid tissue that manifests with signs of hyperthyroidism (e.g., irritability, heat intolerance, diarrhea)
- Late-stage
  - Thyroid may be normal-sized or small if extensive fibrosis has occurred.
  - **Signs of hypothyroidism** (e.g., cold intolerance, constipation, fatigue)

T<sub>3</sub> normal ↑ TSH.  
 T<sub>4</sub>

- Antithyroid peroxidase
- Antithyroglobulin ab.
- Thyroid U/S. → scattered (hypoechoogenicity).
- Thyroid scan.
- Biopsy (definitive)

- Levothyroxin.  
 if eu → Follow up  
 6-12 months.

if goit. normal TSH → No Tx.

# # Congenital hyperthyroidism #

Fetal tachycardia.  
 Mother → Graves.

Findings	Hypothyroidism	Hyperthyroidism
Metabolic	Intolerance to cold Weight gain Decreased appetite	Intolerance to heat Weight loss Increased appetite
Neuropsychiatric	Excessive fatigue	Irritability, restlessness
Periorbital findings	Periorbital edema (due to generalized myxedema)	Periorbital edema, lid lag, and exophthalmos (in Graves ophthalmopathy)
Cardiovascular	Bradycardia, decreased cardiac output	Tachycardia, palpitations, hypertension
Skin	Cold, dry	Warm, moist
Gastrointestinal	Constipation	Hyperdefecation
Musculoskeletal	Cramps	Osteopathy
Edema	Generalized myxedema (initially pretibial)	Pretibial myxedema (in Graves disease)
Myopathy	* Delayed osseous maturation. * precocious puberty.	Proximal
Reproduction	♀ Menstrual disorders ♂ Decreased libido, infertility	
Hair		Hair loss

# Hyperthyroidism.

# # Graves #

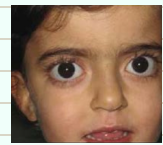
♀ > ♂  
 5:1  
 gradual onset

Enlargement → spleen  
 Thyroid LN  
 Peripheral lymphocytosis.  
 \* TRSAB  
 \* TRBAR  
 aet → HLA-B8 HLA-DR3.  
 Addison BMI MGs Celiac  
 SLE RA arthritis vitiligo  
 pernicious anemia ITP.

earliest sign → emotional disturbances.

# # Thyroid crisis (storm)

Acute onset  
 Hyperthermia  
 Severe tachycardia  
 Restlessness  
 Delirium coma death



Clinical manifestations cont
<b>GRAVES DISEASE</b>
• Tremor
• Voracious appetite with loss of weight
• Goiter found in most pt
• Exophthalmos noticeable but mild lid lag. Although 50-75% of children with Graves disease have some eye finding, the symptoms are much milder than in adults.
• Skin smooth and flushed, excessive sweating
• Muscular weakness
• Tachycardia palpitations dyspnea cardiomegaly atrial fibrillation rare complication
• Mitral regurgitation
• Systolic B/P and pulse increase 90
• Advanced skeletal maturation and craniosynostosis (neonatal Graves) <small>subvs cond. vs only.</small>
• Note: Eye manifestations are much less common in children than adults

Differential diagnosis (cont)  
 If precocious puberty, café au lait spot and polyostotic fibrous dysplasia → McCune-Albright  
 Generalized thyroid hormone unresponsiveness (Thyroxine resistance)  
 High level T3 but levels TSH elevated or normal  
 Exogenous thyroid hormone, T4 TSH same as Graves but thyroglobulin is very low.



## ✓ Medical therapies

- Radioactive iodine ablation  
 effective / safe  
 ↓ hypo / anorexia
- Subtotal thyroidectomy  
 ↓ hypo parathyroidism  
 ↓ vocal cord paralysis  
 ↓ recurrence

Propylthiouracil PTU  
 ✓ pregnancy / lactating  
 Methimazole  
 ✓ Peds  
 Transient leukopenia  
 = arthralgia rash

β adrenergic agents  
 propranolol.  
 0.5-2 mg/kg.

Congenital hyperthyroidism	Treatment of Congenital hyperthyroidism
• Presence of RBC • Small liver size • Bone density: hyperostosis • Hypocalcaemia • Rapidly progressive exophthalmos • Tachycardia tachypnoea high Temp • Failure to gain weight • Jaundice • Cardiovascularly • Congenital hyperostosis • May die if not treated properly.	• Carbimazole • Propranolol • Light iodine: 10mg 3x per day and iodine in water / reduce iodine in formula • Full thyroidectomy: 1-2 mo 1 day increasing iodine intake and inhibiting the enzyme thyroid peroxidase • Full dose PTU (200mg) 4 times a day • Support for heart failure

\* TRAb  
 To diagnose & tx of Graves not others  
 ↑ maximum  
 ↑ hyper...

# subclinical hyperthyroidism  
 ↑ TSH  
 normal T<sub>4</sub>