ENDOCRINE BLOCK

PHYSIOLOGY TEAM 431



THE THYROID GLAND - 2

REGULATION OF HORMONES SECRETION

It is regulated by the hypothalamic-pituitary axis.



- Glycoprotein in nature.
- Secreted from Anterior pituitary.
- Regulate metabolism , secretion and growth of thyroid gland (trophic effect).

Action of TSH:

- Increase proteolysis of the thyroglobulin
- Increase pump activity
- Increase iodination of tyrosine
- Increase coupling reaction
- Trophic effect
- ✓ TSH secretion started at 11-12 of gestational weeks
- ✓ TSH + receptor → activation of adenylyl cyclase via Gs protein → \uparrow cAMP → \uparrow activation of protein kinase → multiple phosphorylation → secretion and thyroid growth.

Table 9-8 Factors Affecting Thyroid Hormone Secretion				
Inhibitory Factors				
I [–] deficiency				
Deiodinase deficiency				
Excessive I ⁻ intake (Wolff- Chaikoff effect)				
Perchlorate; thiocyanate (inhibit Na ⁺ -I ⁻ cotransport)				
Propylthiouracil (inhibits peroxidase enzyme)				
Decreased TBG levels (e.g., liver disease)				

Hyperthyroidism (Over activity of the thyroid gland)

- Activity of gland:
 - ✓ 5 10 times increase in secretion
 - ✓ 2 3 times increase in size

🗷 Women : men ratio (8:1)

Causes:

1. Graves' disease:

- An autoimmune disorder.
- Increased circulating level of thyroid- stimulating Immunoglobulins (TSI).
- 95%.
- 4 8 times more common in women than men.

2. Thyroid gland tumor:

- 95% is benign.
- 5% is malignant.
- History of head and neck irradiation and family history.

3. Exogenous T3 and T4

- Maybe by mistake by taking drugs that increase T4 and T3
- Rarely cause

4. Excess TSH secretion:

- Diseases of the hypothalamus (TRH).
- Diseases of the pituitary (TSH).

Diagnosis:

- ✓ Based on the Signs and Symptoms:
 - 1. Goiter in 95% (increase in thyroid gland size)
 - 2. Skin:
 - Smooth, warm and moist.
 - Heat intolerance. (Abnormal sensitivity to a hot environment)
 - Night sweating.

3. Musculoskeletal:

Muscle atrophy (thyroxin catabolic (breakdown) protein which leads to Muscle atrophy).

4. Neurological:

- Tremor.
- Enhanced reflexes.
- Irritability.
- 5. Cardiovascular:
 - Arrhythmias.
 - Hypertension.

- Increase heart rate.
- Increase stroke volume.

6. **G.I tract:**

- Weight loss.
- Diarrhea.
- 7. Renal function:
 - \uparrow Glomerular filtration rate.

8. Exophthalmos:

- (The cause is unknown but it is because of swelling of retroocular tissue)
- Anxious staring expression.
- Protrusion of eye balls.
- 9. Others:
 - Menstrual cycle disturbance



Based on the Investigations:

Serum measurement of T3, T4 and TSH

	Т3,Т4	TSH
primary hyperthyroidism	high	low
secondary hyperthyroidism	high	high

Treatment:

- Medical therapy:
 - E.g. Propylthiouracil (Antithyroid agent)
 - Usually for 12-18 months course
 - With 3-4 monthly monitoring

• Surgery:

- Subtotal thyroidectomy
- Indication for surgery:
 - 1. Relapse after medical treatment
 - 2. Drug intolerance
 - 3. Cosmetic.(Surgery that modifies or improves the appearance of a physical feature)
 - 4. Suspected malignancy

Hypothyroidism

- ✓ Under activity of the thyroid gland
- ✓ more in woman (30- 60 years).

Causes:

- 1. Inherited abnormalities of thyroid hormone synthesis :
 - Peroxidase defect
 - Iodide trapping defect
 - Thyroglobulin defect

2. Endemic Colloid Goiter:

- Table Salt (iodized salt) reduced the incidence
- \downarrow lodide $\rightarrow \downarrow$ Hormone formation $\rightarrow \uparrow$ TSH
 - \rightarrow \uparrow Thyroglobulin \rightarrow \uparrow Size (10-times)

3. Idiopathic Nontoxic Colloid Goiter:

- There is no iodine deficiency.
- It has the symptoms of mild Thyroditis
- Inflammation \rightarrow \uparrow Cell damage \rightarrow \downarrow Hormone secretion \rightarrow \uparrow TSH \rightarrow \uparrow Activity of normal cells \rightarrow \uparrow Size
- 4. Gland destruction (surgery)
- 5. Pituitary diseases or tumor
- 6. Hypothalamus diseases or tumor

Diagnosis

- Based on the Signs and Symptoms
 - 1. Skin:
 - Dry Skin
 - Cold Intolerance. (Abnormal sensitivity to a cold environment)

2. Musculoskeletal:

- 个Muscle Bulk
- \downarrow In Skeletal Growth
- Muscle Sluggishness
- Slow relaxation after contraction.



3. Neurological:

- Slow Movement
- Impaired Memory
- Decrease Mental Capacity

4. Cardiovascular:

- ↓Heart rate
- ↓Stroke volume
- 5. **G.I tract:**
 - Constipation
 - Increase Weight

6. Renal function:

- \downarrow Glomerular filtration rate

7. Myxoedema:

- An edematous appearance throughout the body
- 8. Others:
 - Loss of libido.(Loss of sexual desire)
 - Menstrual cycle disturbance

<u>Based on the Investigations</u>

	T3,T4	TSH
primary hypothyroidism	low	High
secondary hypothyroidism	low	low

Treatment:

- L-Thyroxine (Is a synthetic form of thyroid hormone)
 - Starting dose is 25-50 µg
 - Increase to 200 µg
 - At 2-4 weeks period
 - ☑ The first response seen is the weight loss





	Hyperthyroidism	Hypothyroidism
Symptoms	Increased basal metabolic rate	Decreased basal metabolic rate
	Weight loss	Weight gain
	Negative nitrogen balance	Positive nitrogen balance
	Increased heat production	Decreased heat production
	Sweating	Cold sensitivity
	Increased cardiac output	Decreased cardiac output
	Dyspnea (shortness of breath)	Hypoventilation
	Tremor, muscle weakness	Lethargy, mental slowness
	Exophthalmos	Drooping eyelids
	Goiter	Myxedema
		Growth retardation
		Mental retardation (perinatal)
		Goiter
Causes	Graves' disease (increased thyroid-stimulating	Thyroiditis (autoimmune or Hashimoto's thyroid
	immunoglobulins)	Surgery for hyperthyroidism
	Thyroid neoplasm	I ⁻ deficiency
	Excess TSH secretion	Congenital (cretinism)
	Exogenous T ₃ or T ₄ (factitious)	Decreased TRH or TSH
TSH Levels Decreased (feed anterior lobe) Increased (if dec	Decreased (feedback inhibition of T ₃ on the anterior lobe)	Increased (by negative feedback if primary defec thyroid gland)
	Increased (if defect is in anterior pituitary)	Decreased (if defect is in hypothalamus or anteri pituitary)
Treatment	Propylthiouracil (inhibits peroxidase enzyme and thyroid hormone synthesis)	Thyroid hormone replacement therapy
	Thyroidectomy	
	¹³¹ I ⁻ (destroys thyroid)	
	B-Adrenergic blocking agents (adjunct therapy)	

Cretinism

- Extreme hypothyroidism during infancy and childhood (failure of growth)

Causes:

- 1- Congenital lake of thyroid gland (congenital cretinism)
- 2- Genetic deficiency leading to failure to produce hormone
- 3- Iodine lake in the diet (endemic cretinism)

Symptoms:

- 1- Infant is normal at birth but abnormality appears within weeks
- 2- Protruding tongue
- 3- Dwarf with short limbs
- 4- Mental retardation
- 5- Often umbilical hernia
- 6- Teeth

Treatment:

- Changes are irreversible unless treatment is given early.

Q1: A 30-year-old female complains of palpitations, fatigue, and insomnia On physical exam, her extremities are warm and she is tachycardic. There is diffuse thyroid gland enlargement and proptosis. There is a thickening

of the skin in the pretibial area Which of the following lab values would you expect in this patient?

- a. Increased TSH, total thyroxine, total T3
- b. Decreased TSH, increased total thyroxine
- c. Increased T3 uptake, decreased T3
- d. Decreased TSH, normal T4

Q2: The cause of this patient's thyrotoxicosis is (related to 1st question):

- a. Autoimmune disease
- b. Benign tumor
- c. Malignancy
- d. Viral infection of the thyroid

Q3: A 30-year-old female complains of fatigue, constipation, and weight gain. There is no prior history of neck surgery or radiation. Her voice is hoarse and her skin is dry. Serum TSH is elevated and T4 is low. The most likely cause of these findings is

- a. Autoimmune disease
- b. Postablative hypothyroidism
- c. Pituitary hypofunction
- d. Thyroid carcinoma

answers:	1 (B).
	2 (A).
	3 (A).