





- GENERAL ASPECTS OF THYROID GLAND
 - Anatomy: weight range from 12 to 30g
 - Located in the neck, anterior to the traquea
 - Produces: T4 & T3 (active hormone)
 - Regulation: "negative Feed-back" axis



THYROID GLAND REGULATION
 "negative Feed-back" axis

(negative effect)

- Hypothalamus

(TRH positive effect)

Pituitary gland

(TSH, positive effect)

Thyroid gland

T3 & T4



Thyroid hormones:

 T4: (Thyroxine) is made exclusively in thyroid gland

Ratio of T4 to T3;
 5::1

Potency of T4 to T3; 1::10

 T4 is the most important source of T3 by peripheral tissue deiodination "T4 to T3"



- Thyroid hormones:
 - T3: (Triiodothyronine) main source is peripheral deiodination:
 - Ratio of T3 to T4;
 1::5
 - Potency of T3 to T4; 10::1
 - T3 is the most important because more than 90% of the thyroid hormones physiological effects are due to the binding of T3 to Thyroid receptors in peripheral tissues.



PHYSIOLOGY EFFECTS OF THYROID HORMONES

THEY ARE NOT ESSENTIAL
FOR LIFE, BUT ARE
EXTREMELY HELPFUL



- THYROID HORMONE EFFECTS:
 - Affects every single cell in the body
 - Modulates:
 - Oxygen consumption
 - Growth rate
 - Maturation and cell differentiation
 - Turnover of Vitamins, Hormones, Proteins, Fat, CHO



MECHANISMS OF THYROID HORMONE ACTION

- Act by binding to Nuclear receptors, termed Thyroid Hormone Receptors (TRs), Increasing synthesis of proteins
- At mitochondrial level increases number and activity to increasing ATP production
- At Cell membrane increases ions and substrates transmembrane flux



- CALORIGENESIS
- GROWTH & MATURATION RATE
- C.N.S. DEVELOPMENT & FUNCTION
- CHO, FAT & PROTEIN METABOLISM
- MUSCLE METABOLISM
- ELECTROLYTE BALANCE
- VITAMIN METABOLISM
- CARDIOVASCULAR SYSTEM
- HEMATOPOIETIC SYSTEM
- GASTROINTESTINAL SYSTEM
- ENDOCRINE SYSTEM
- PREGNANCY



- CALORIGENESIS
 - Controls the Basal Metabolic Rate (BMR)
- CHO METABOLISM
 - Increases:
 - Glucose absorption of the GI tract
 - Glucose consumption by peripheral tissues
 - Glucose uptake by the cells
 - Glycolysis
 - Gluconeogenesis
 - Insulin secretion



- GROWTH & MATURATION RATE
- C.N.S. DEVELOPMENT & FUNTION
 - "ESSENTIAL" in the newborn to prevent development of "CRETINISMS" & to a normal "IQ"
 - Modulation of brain cerebration
 - Mood modulation



- FAT & PROTEIN METABOLISM
 - Increase lipolysis and lipid mobilization with:
 - Cholesterol
 - Triglicerides
 - Free fatty acids
 - MUSCLE METABOLISM
 - Modulates;
 - Strength & velocity of contraction



- ELECTROLYTE BALANCE
 - Low Thyroid hormones could induce hyponatremia
- VITAMIN METABOLISM
 - Modulates vitamin consumption
- HEMATOPOIETIC SYSTEM
 - Could induce anemia



THYROID HORMONE EFFECTS

- CARDIOVASCULAR SYSTEM

- Hyperthyroidism, increases:
 - Heart rate & myocardial strenght
 - Cardiac output
 - Peripheral resistances (Vasodilatation)
 - Oxygen consumption
 - Arterial pressure
- Hypothyroidism, reduces:
 - Heart rate & myocardial strenght
 - Cardiac output
 - Peripheral resistances (Vasodilatation)
 - Oxygen consumption
 - Arterial pressure



- GASTROINTESTINAL SYSTEM
 - Modulate bowel movements and absorption
- ENDOCRINE SYSTEM
 - Modulates pituitary axis, affecting GH, ACTH, FSH, LH, so-on
- PREGNANCY
 - Modulates growth rate and affects lactation



- DIVIDED INTO:
 - THYROTOXICOSIS (Hyperthyroidism)
 - Overproduction of thyroid hormones
 - HYPOTHYROIDISM (Gland destruction)
 - Underproduction of thyroid hormones
 - **NEOPLASTIC PROCESSES**
 - Beningn
 - Malignant



THYROID GLAND DISORDERS LABORATORY EVALUATION

TSH normal, practically excludes abnormality

- If TSH is abnormal, next step: Total & Free T4 & T3
- TSI (Thyroid Stimulating Ig)
- TPO (Thyroid Peroxidase Ab)
- Antimitochondrial Ab
- Serum Tg (Thyroglobulin)
- Radioiodine uptake & Thyroid scaning
- FNA, Fine-needle aspiration
- Thyroid ultrasound



- TSH High usually means Hypothyroidism
 - Rare causes:
 - TSH-secreting pituitary tumor
 - Thyroid hormone resistance
 - Assay artifact
- TSH low usually indicates Thyrotoxicosis
 - Other causes
 - First trimester of pregnancy
 - After treatment of hyperthyroidism
 - Some medications (Esteroids-dopamine)



THYROTOXICOSIS:

- is defined as the state of thyroid hormone excesss

HYPERTHYROIDISM:

- is the result of excessive thyroid gland function



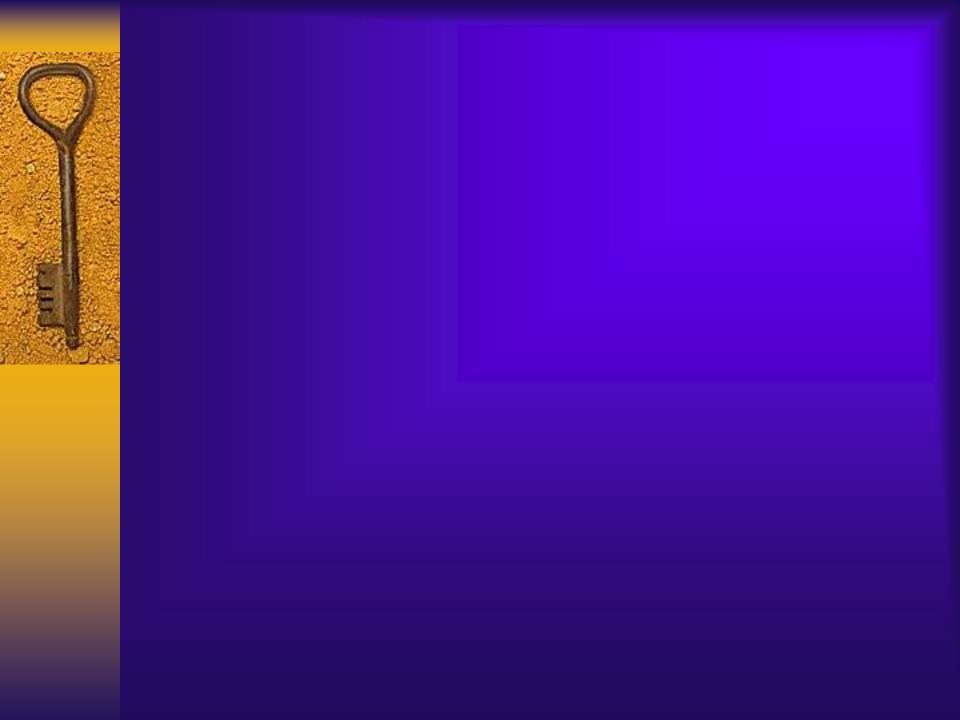
- Abnormalities of Thyroid Hormones
 - Thyrotoxicosis
 - Primary
 - Secondary
 - Without Hyperthyroidism
 - Exogenous or factitious
 - Hypothyroidism
 - Primary
 - Secondary
 - Peripheral



- Causes of Thyrotoxicosis:
 - Primary Hyperthyroidism
 - Grave's disease
 - Toxic Multinodular Goiter
 - Toxic adenoma
 - Functioning thyroid carcinoma metastases
 - Activating mutation of TSH receptor
 - Struma ovary
 - Drugs: Iodine excess



- Causes of Thyrotoxicosis:
 - Thyrotoxicosis without hyperthyroidism
 - Subacute thyroiditis
 - Silent thyroiditis
 - Other causes of thyroid destruction:
 - Amiodarone, radiation, infarction of an adenoma
 - Exogenous/Factitia
 - Secondary Hyperthyroidism
 - TSH-secreting pituitary adenoma
 - Thyroid hormone resistance syndrome
 - Chorionic Gonadotropin-secreting tumor
 - Gestational thyrotoxicosis





THYROTOXICOSIS

Symptoms:

- Hyperactivity
- Irritability
- Dysphoria
- Heat intolerance & sweating
- Palpitations
- Fatigue & weakness
- Weight loss with increased appetite
- Diarrhea
- Polyuria
- Sexual dysfunction

Signs:

- Tachycardia
- Atrial fibrillation
- Tremor
- Goiter
- Warm, moist skin
- Muscle weakness, myopathy
- Lid retraction or lag
- Gynecomastia
- * Exophtalmus
- * Pretibial myxedema



- Differential diagnosis:
 - Panic attacks
 - Psychosis
 - Mania
 - Pheochromocytoma
 - Hypoglycemia
 - Occult malignancy



Treatment:

- Reducing thyroid hormone synthesis:
 - Antithyroid drugs (Methimazole, Propylthyouracil)
 - Radioiodine (¹³¹I)
 - Subtotal thyroidectomy
- Reducing Thyroid hormone effects:
 - Propranolol
 - Glucocorticoids
 - Benzodiazepines
- Reducing peripheral conversion of T4 to T3
 - Propylthyouracil
 - Glucocorticoids
 - Iodide (Large oral or IV dosage) (Wolf-Chaikoff effect)



- Treatment: Special considerations:
 - Thyrotoxic crisis or Thyroid storm:
 - It's a life-threatening exacervation of thyrotoxicosis, acompanied by fever, delirium, seizures, coma, vomiting, diarrhea, jaundice.
 - Mortality rate reachs 30% even with treatment
 - It's usually precipitated by acute illness, such as:
 - Stroke, infection,trauma, diabeic ketoacidosis, surgery, radioiodine treatment
 - Propylthyouracil IV or Nasogastric tube
 - Radioiodine (¹³¹I)
 - Propranolol
 - Glucocorticoids
 - Benzodiazepines
 - Iodide (Large oral or IV dosage) (Wolf-Chaikoff effect)



- HYPOTHYROIDISM
 - Primary
 - Autoimmune (Hashimoto's)
 - latrogenic Surgery or ¹³¹I
 - Drugs: amiodarone, lithium
 - Congenital (1 in 3000 to 4000)
 - Iodine defficiency
 - Infiltrative disorders



 Hashimoto's Thyroiditis or Goitrous thyroiditis

- Mean anual incidence:
 - Women 4:1000 Men 1:1000
 - Risk factors; TPO antibodies (90%)
 Japanese, previous history, high I intake
 - Average age: 60
 - Frequently associated to other autoimmune disorders such as: AR, SLE, Sjogren's so-on.
 - Treatment: Levothyroxine



- CONGENITAL HYPOTHYROIDISM
- Prevalence: 1 in 3000 to 4000 newborns
 - Cause: Dysgenesis 85%
 - Dx: Blood screning (TSH &/or T4)
- Treatment:
 - Supplemental Tx. With Levothyroxine is "essential" for a normal C.N.S. Development and prevention of mental retardation



HYPOTHYROIDISM

- Secondary
 - Pituitary gland destruction
 - Isolated TSH deficiency
 - Bexarotene treatment
 - Hypothalamic disorders
- Peripheral:
 - Rare, familial tendency



HYPOTHYROIDISM

- Symptoms:
 - Tiredness
 - Weakness
 - Dry skin Sexual dysfunction
 - Dry skin
 - Hair loss
 - Difficulty concentrating

Signs:

- Bradycardia
- Dry coarse skin
- Puffy face, hands and feet
- Diffuse alopecia
- Peripheral edema
- Delayed tendon reflex relaxation
- Carpal tunel syndrome
- Serous cavity effusions.



- SPECIAL TREATMENT CONSIDERATIONS
- Myxedema coma
 - Reduced level of consciousness, seizures
 - Hypotension/shock
 - Hypothermia
 - Hyponatremia
- Usually in elderly hypothyroid pts.
- Usually precipitated by intercurrent illnesses that impairs ventilation
- It's an Emergency with a high mortality rate
- Treatment: Lyotironine(T3) or T4, Hydrocortisone, external warming, IV fluids



- SPECIAL TREATMENT CONSIDERATIONS
- Elderly patients
- Coronary Artery Disease
- Poor adrenal gland reserve
- Childrens
- Pregnancy
- Emergency surgery (Non thyroid related)



THYROID GLAND NEOPLASIAS

Out of the focus of this lecture