

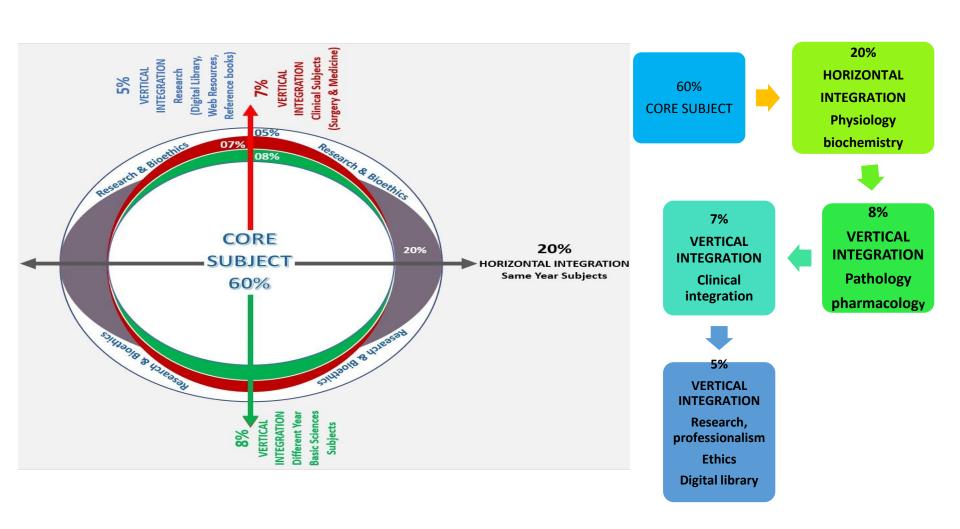
# **Motto** Vision; The Dream/Tomorrow



- To impart evidence based research oriented medical education
- To provide best possible patient care
- To inculcate the values of mutual respect and ethical practice of medicine



#### **Professor Umar Model of Integrated Lecture**





# Endocrinology Module 2<sup>nd</sup> Year MBBS (CBL) - Biochemistry Thyrotoxicosis

H.O.D

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**Deptt of Biochemistry** 

**RMU** 

Date: 11-03-25

#### **Learning Objectives**

At the end of this session student should be ab

- 1. Discuss the Synthesis of Thyroid Hormone
- 2. Understand the Regulation of Plasma level of T3, T4
- 3. Describe the Mechanism of Action of Thyroxin
- 4. Explain the Functions of Thyroid Hormone in the body
- 5. Discuss Causes and Clinical Features of Hyperthyroidism
- 6. Practice the principles of bioethics & apply strategic use of A.I in the related clinical condition.
- 7. Read relevant research articles related to the Core Knowledge.

#### Clinical Scenario - Case

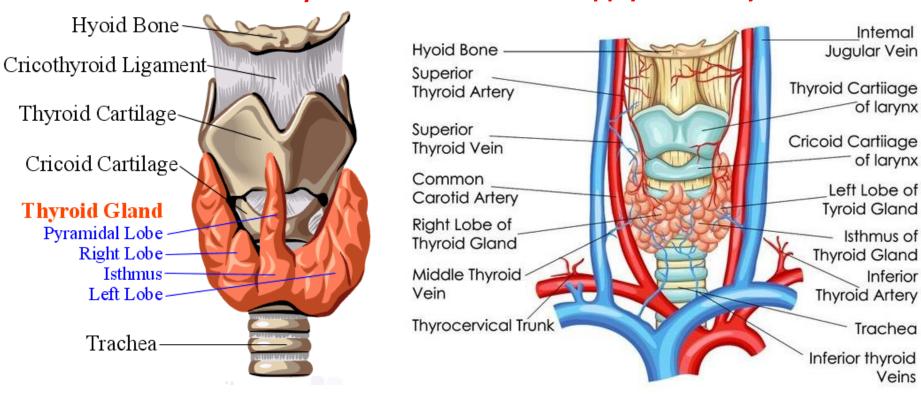
#### **Clinical Scenario (Hyperthyroidism)**

• A 20 year old unmarried female (previously healthy) presented to the OPD of DHQ Hospital Rawalpindi with a 2 months history of increased appetite, hair & weight loss, fatigue, difficulty in sleep, excessive sweating and altered menstrual cycle. Examination showed rapid pulse, warm hands and slightly enlarged thyroid gland. Serum TSH level was very low and free T3, T4 levels were raised. She was provisionally diagnosed as case of hyperthyroidism.

# **Anatomical Aspects**

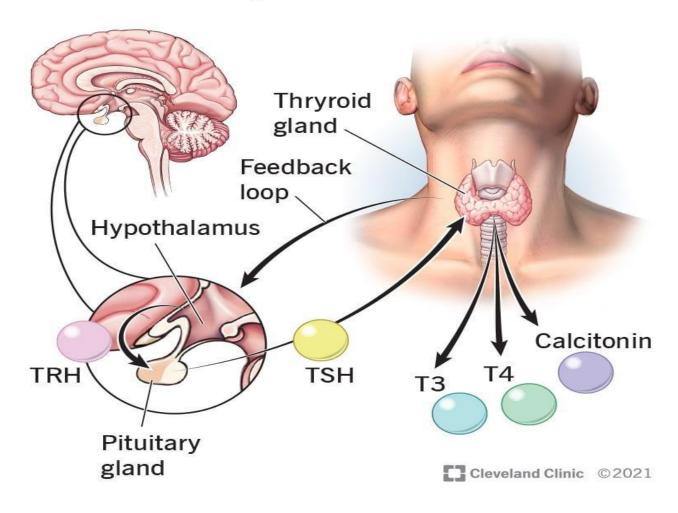
#### Parts & Relations of the Thyroid Gland

#### **Blood Supply of the Thyroid Gland**



# PHYSIOLOGICAL ASPECTS

#### **Thyroid Hormones**



# **Thyroid Hormones**

- The thyroid hormones are α-amino acid derivatives of tyrosine
- Thyroxine & Triiodothyronine are exceptionally rich in iodine, which comprises more than half of their molecular weight
- Thyroxine contains four atoms of iodine and is abbreviated as T4
- Triiodothyronine, which has three atoms of iodine, is abbreviated as

Reverse T3 (rT3)

Amount

Site of synthesis

Site of stores

Structure

Thyroid gland hormones الكمية عكس الاكتفتي يعني كلما زادت الكمية قل نشاط الهرمون (أقلهم كمية أكثرهم نشاطاً)				
	T3	T4 Thyroxine (Tetraiodiothyronine)	Calcitonia Harmana	Reverse T3 (not important)

Contains 4 iodine

When it reaches the tissue

its majority will be

converted to T3

90%

In colloid

Calcitonin Hormone

(Humoral stimulation) #L1 !

(For Ca++ metabolism)

Parafollicular cells

C -cell

Biologically inactive,

but it appears in

chemical reactions

Mirrored image of T3 having 3 lod, but the

problem is the mispositioning of iod, making it inactive.

Hormones

(Triiodothyronine)

Contains 3 iodine

Most potent

10% Less but has

stronger action than T4

Apical and basal membrane in follicular cells

# Transport of thyroid hormones in blood

#### Thyroid hormones are transported in blood in two forms

#### 1-Bound forms:

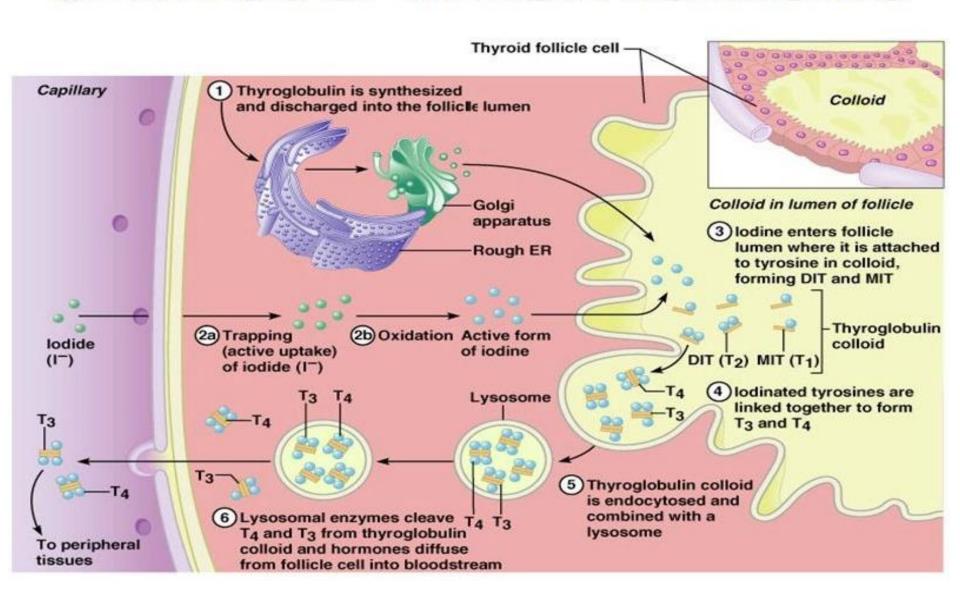
- Thyroid hormones are transported in plasma bound to plasma proteins:
  - Thyroid Binding Globulin (TBG)
     binds 70% of plasma T4 & 80% of plasma T3
  - Others: Pre-albumin & Albumin

#### 2- Free form:

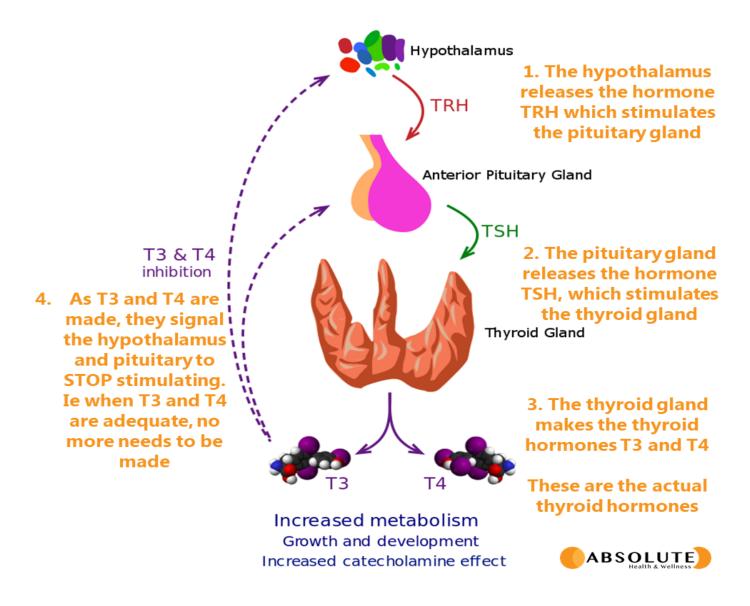
0.05% of plasma T4 & 0.02% of plasma T3 are free (not bound to plasma proteins)

Only the <u>free</u> fraction can cross the cell membrane & affect intracellular metabolism

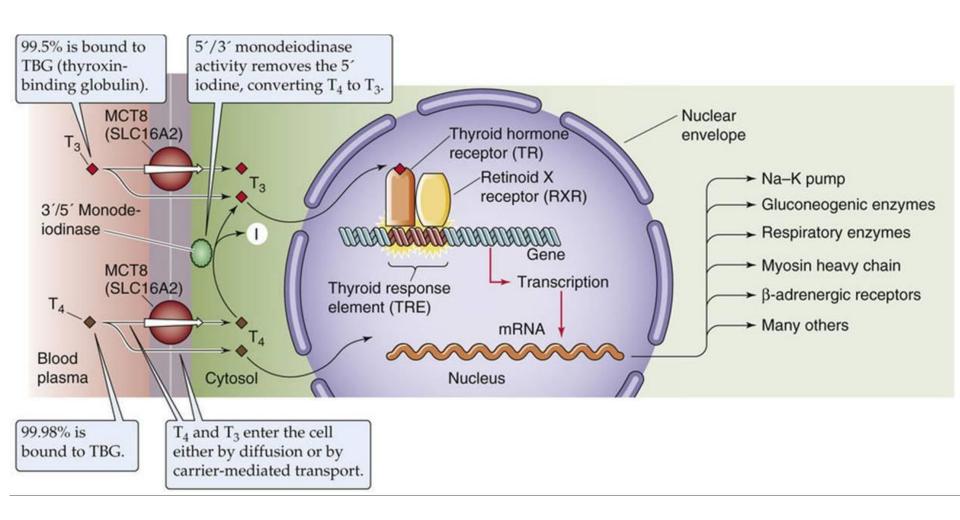
#### SYNTHESIS OF THYROID HORMONES



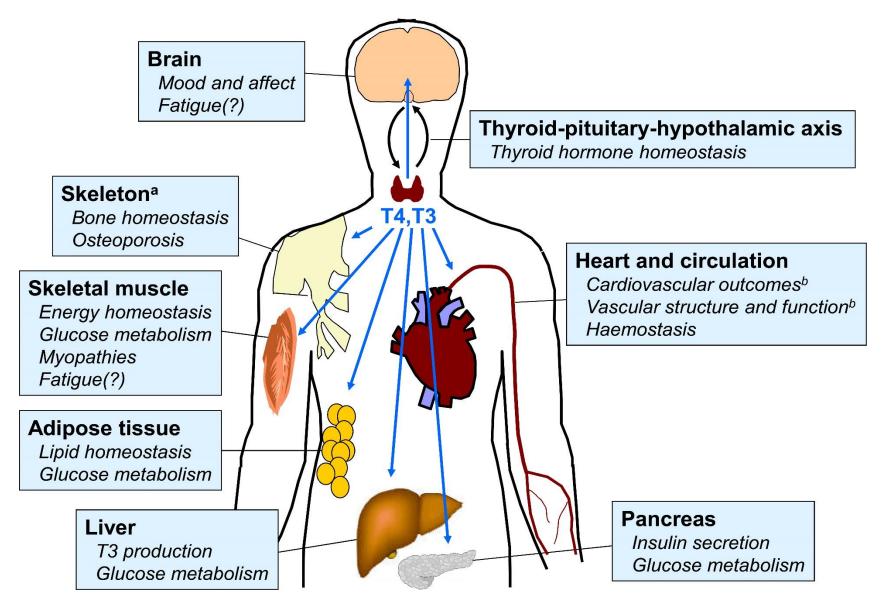
#### Regulation of Plasma level of T3, T4



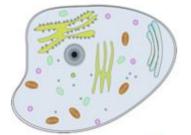
#### Mechanism of Action of Thyroxin



#### Functions of Thyroid Hormone in the body



# Thyroid Effects on the Body

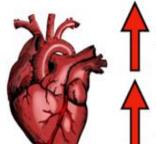




Protein Synthesis and Metabolic Activity in Cells

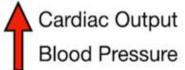


Body Metabolism Body Heat



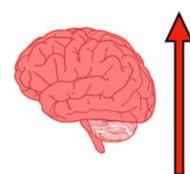
Beta-1 Receptors

Heart Rate Stroke Volume





GI Motility
GI Secretions



Neurogenesis
Myelination
Synaptogenesis
Dendrite Formation
Sympathetic Activity



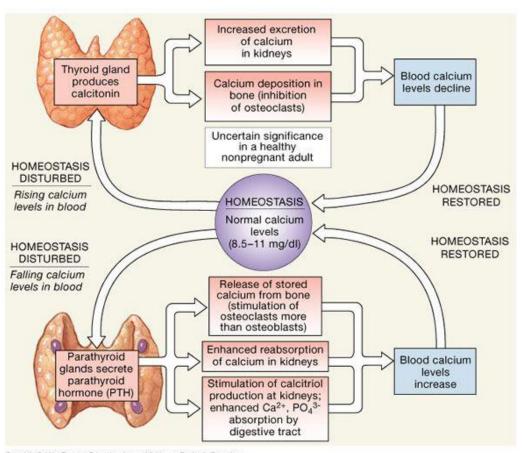
Proliferation of Skin Cells Sweat & Skin Secretions Hair Growth Nail Growth

#### **Actions of thyroid hormones:**

 $T_3$  and  $T_4$  have the following effects:

- 1- General metabolism: Thyroid hormones increase the metabolic rate and O<sub>2</sub> consumption of all tissues of the body except the adult brain, lungs, lymphoid tissues, retina and anterior pituitary gland.
- 2- Protein metabolism: Normal levels of thyroid hormones stimulate protein synthesis (anabolic effect) whilst excess Thyroid hormones secretions cause protein catabolism (breakdown of protein molecules).
- 3- Carbohydrate metabolism: Thyroid hormones stimulate glucose uptake and utilization by tissues, insulin hormone secretion, liver glycogenolysis and intestinal absorption of glucose.
- 4- Lipid metabolism: Thyroid hormones decrease the level of lipid and cholesterol in the blood (i.e. increased cholesterol catabolism than lipogenesis)

# **Calcitonin**



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#### **THYROTOXICOSIS**

#### DEFINITION

 Thyrotoxicosis is any syndrome caused by excess thyroid hormone and can be related to excess hormone production (hyperthyroidism).



#### Causes of thyrotoxicosis

- Graves' disease
- Toxic multinodular goitre
- Toxic adenoma
- Thyroiditis
- Drugs (e.g. amiodarone, lithium)
- Gestational thyrotoxicosis
- Choriocarcinoma
- Hydatidiform mole
- Adenoma secreting thyroid-stimulating hormone
- Others (e.g. neuroendocrine tumour, thyroid cancer)

#### **CAUSES OF THYROTOXICOSIS**

# Causes of Hyperthyroidism





The causes of hyperthyroidism are mainly categorised into:

#### Primary Hyperthyroidism

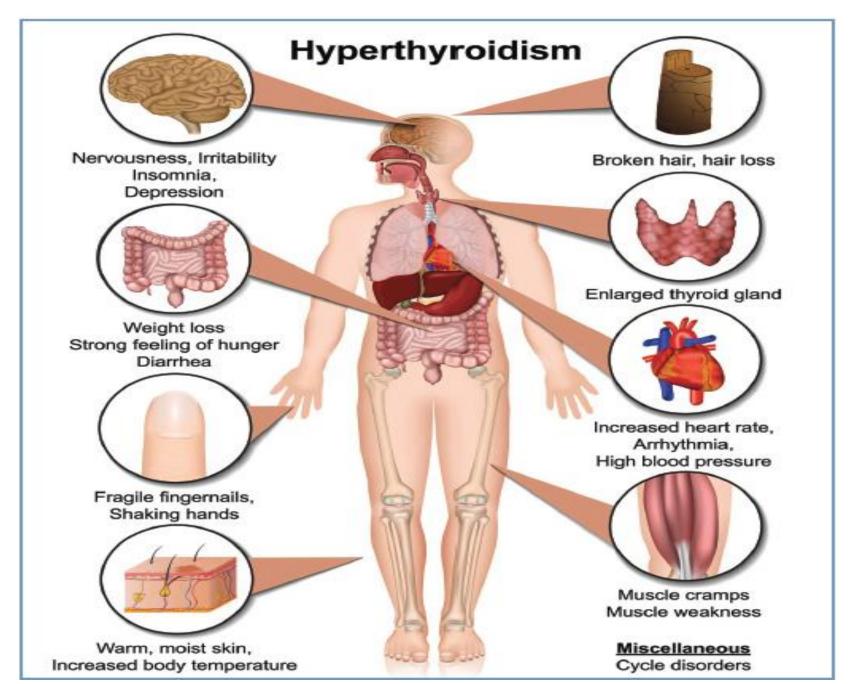
- · Graves' disease
- Toxic adenoma, toxic multinodular goiter or Plummer's disease
- Functioning thyroid carcinoma metastases
- Activating mutation of the TSH receptor
- Activating mutation of GSα (McCune-Albright syndrome)
- Struma ovarii
- Drugs: iodine excess (Jod-Basedow phenomenon)

#### Secondary Hyperthyroidism

- TSH-secreting pituitary adenoma
- Thyroid hormone resistance syndrome
- Chorionic gonadotropin-secreting tumours
- Gestational thyrotoxicosis







# Clinical Features of Hyperthyroidism

# HYPERTHYROIDISM SIGNS & SYMPTOMS

**Mnemonic: THYROIDISM** 

- T Tremor
- H Heart rate up
- Y Yawning (Fatigue)
- R Restlessness
  - O Oligomenorrhoea
  - I Intolerance to heat
- D Diarrhoea
- I Irritability
- S Sweating
- M Muscle wasting

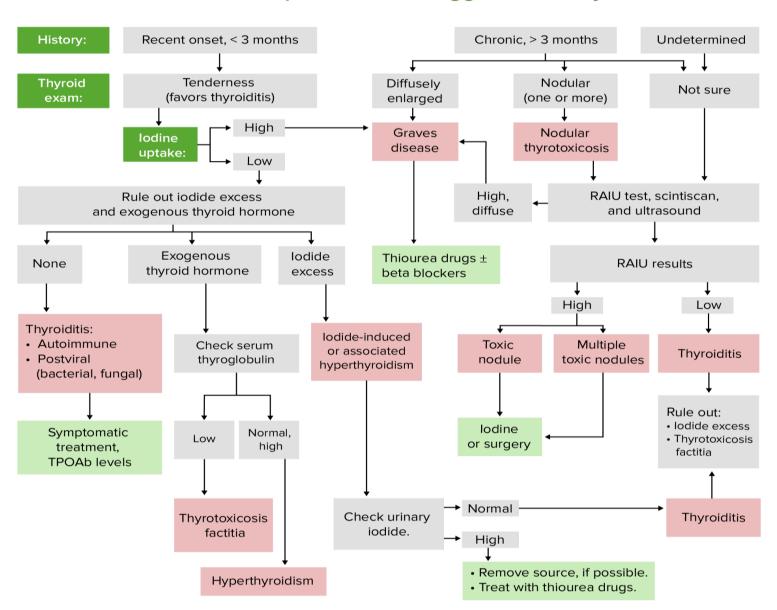
#### **HYPERTHYROIDISM**





#### Management

Patient with clinical presentation suggestive of thyrotoxicosis



#### **Family Medicine**

# Management of

# Role of AI in Management of

- Al can potentially aid in enhancing diagnostic accuracy and efficiency.
- Al-powered decision support systems can also help clinicians in selecting appropriate treatment modalities

 Al-driven predictive models may help anticipate the risk of complications of the Disease in susceptible populations

### **Ethical Considerations**

- From an ethical standpoint, the scenario raises considerations regarding patient autonomy, informed consent, and confidentiality
- The physician must ensure that patient fully understands her diagnosis, treatment options, and potential implications
- Discuss the necessity of a healthy lifestyle & treatment plan.
   This requires clear communication and understanding of risks and benefits.
- Additionally, the physician must respect patient's privacy and confidentiality throughout the diagnostic and treatment process

# **Suggested Research Article**

- https://www.lecturio.com/conc epts/thyrotoxicosis-andhyperthyroidism/
- https://www.researchgate.net/fi gure/Production-and-action-ofthyroid-hormone-The-keycomponents-required-forthyroidhormone fig1 264711792

#### **CBL** --- **QUESTIONS**

- 1. What is the role of thyroxin on protein metabolism?
- 2. Which of the two is more active, T3 or T4?
- 3. What is the chemical nature of thyroid hormone?
- 4. What is the effect of TSH on plasma T3 level?
- 5. How is thyroid hormone transported in plasma?

# **Learning Resources**

- Lippincott Illustrated Reviews BIOCHEMISTRY, Eighth Edition, chapter 29, pages 452 455.
- Harper's Illustrated Biochemistry 32<sup>nd</sup> Edition
- Google Scholar
- Google Images

# **How To Access Digital Library**

- 1. Steps to Access HEC Digital Library
- 2.Go to the website of HEC National Digital Library.
- 3.On Home Page, click on the INSTITUTES.
- 4.A page will appear showing the universities from Public and Private Sector and other Institutes which have access to HEC National Digital Library HNDL.
- 5. Select your desired Institute.
- 6. A page will appear showing the resources of the institution
- 7. Journals and Researches will appear
- 8. You can find a Journal by clicking on JOURNALS AND DATABASE and enter a keyword to search for your desired journal.

Thank You!