

HUMAN PHYSIOLOGY

AN INTEGRATED APPROACH

The Endocrine System

glands ↓ Hormones ↓ Homeostasis ↓ Control

we have no bad
Hormons
an **increase** or **decrease**
in them cause illness

التوازن

Endocrine system
+
Nervous system
Complete each
other

→ What is the Function?!

Regulation / growth + development / Reproduction / metabolism

Dr. Howard D. Booth, Professor of Biology, Eastern Michigan University

Endocrine glands versus Exocrine glands

glands (Secretory cells / organs)

(Are not Related to the endocrine system)

• **Exocrine glands:** these glands have ducts + they empty their content into cavity / Lumen.

Produce massive amount

➤ produce non-hormonal substances and have ducts to transport them to their final destination

➤ Sweat and Saliva Lumen, Cavity, Surface of the body

the endocrine system.

• **Endocrine glands:** One which produce (hormones) / they don't have ducts

chemicals

Release their secretions directly into blood

➤ produce hormones and are lack ducts. They release their hormones into blood → then to the rest parts of the body

➤ **Insulin**, adrenalin

From pancreas

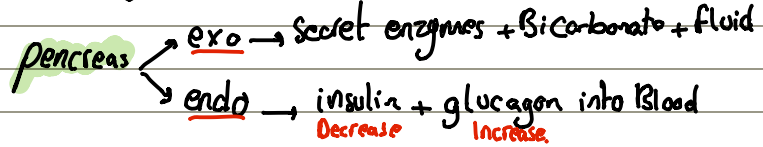
from blood to the rest of the body

Examples about exocrine glands:

Salivary glands → Release saliva $\xrightarrow{\text{through duct}}$ to release it into the Oral cavity

Pancreas → duct that empties into duodenum + gastro intestinal gland.

Sweat glands → duct which release content on the body's surface.



We have chemicals that are produced + not secretive through ducts + not into blood → they are similar to hormones.

a) Autocrine:

A chemical compound released from a cell, effect the same cell. (have it's own receptors)

b) Paracrine:

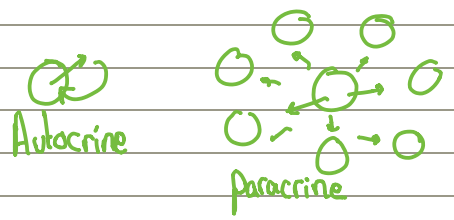
Chemical released from one single cell → effect the neighboring cells.

c) Phermons:

Chemical secreted by 1 individual of species to effect another individual from the same species.

They use it for perfums → Female perfums: Period

Men perfums: tobacco, sweat



Hormones

Target most body cells producing diverse effects and their control include:

- ***Reproduction***
- ***Growth and Development***
- ***Maintenance of Electrolyte, Water and Nutrient
Balance of the Blood***
- ***Regulation of Cellular metabolism and energy balance***
- ***Mobilization of body defence***

Autocrines Paracrines Phermones

Characteristics of Hormones

✓ Made in gland(s) or cells (Secretory Cells)

Any chemical compound that is not secreted by gland or into the blood is not a hormone

✓ Transported by blood

✓ Distant target tissue receptors: does not effect tissues in place but effect tissues at a distance

✓ Exert their effect at very low conc. In Picomoles (directly to the receptors) (their Receptor is too sensitive to low conc.)

✓ Activates physiological response Have physiological effects.

enzymes → High Conc.

✓ Have receptors → to perform their Response

To be called a hormone it must has all these characteristics

✓ Must be terminated → neurotransmitter
→ Protein

Chemistry of Hormones

We have 2 main Categories of hormones

- Amino acid based (Hydrophilic)

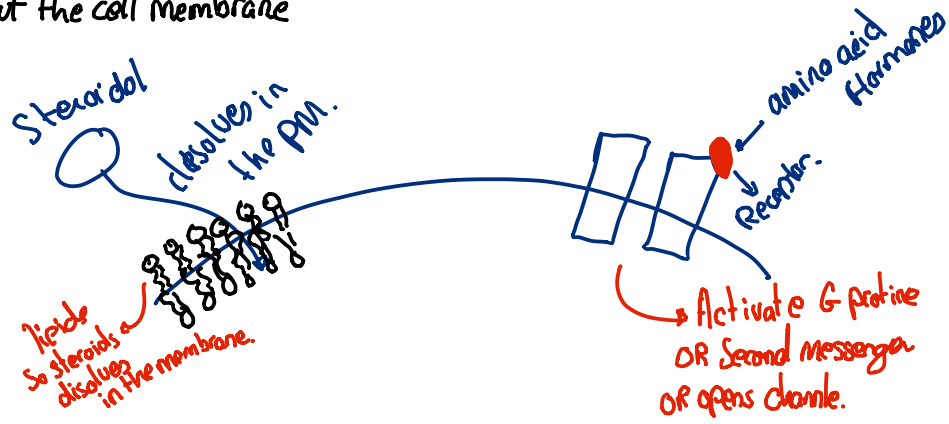
Vary from simple amino acid derivatives (thyroxine) to peptides to proteins

- Steroids (steroidal based) (Hydrophobic)

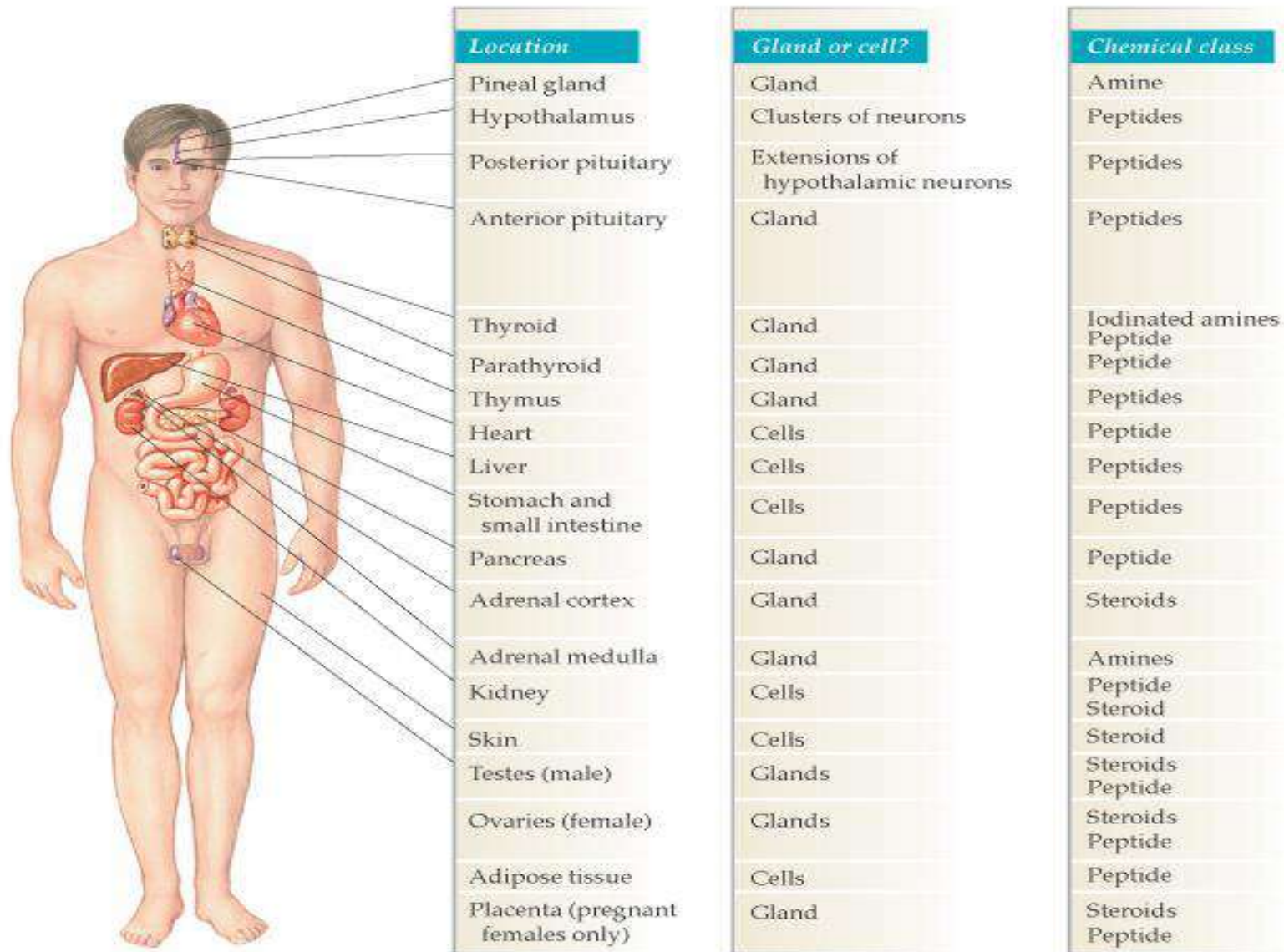
Synthesized from cholesterol → We can't stop it from going into the cell. Produced Upon need.

→ What is the difference between amino acid based and Steroids?

- 1) Receptors for steroidal hormones must be located inside the cell whereas amino acid based hormones → receptors are located at the cell membrane
- 2) Steroidal hormone cannot be stored in the cell while amino acids can be stored in vesicles.



Summary of the Endocrine System



Summary of the Endocrine System Cont.

Hormone	Target	Main Effect
Melatonin	Unclear in humans	Circadian rhythms. Other effects uncertain
Trophic hormones (see Fig. 7-13) See posterior pituitary	Anterior pituitary	Release or inhibit pituitary hormones
Oxytocin (OT) Hypothalamus	Breast and uterus	Milk ejection; labor and delivery; behavior
Vasopressin (ADH) → Hypothalamus ✓ Prolactin (PRL) Growth hormone (GH, somatotropin) Corticotropin (ACTH) Thyrotropin (TSH) Follicle stimulating hormone (FSH) Luteinizing hormone (LH)	Kidney Breast Many tissues Adrenal cortex Thyroid gland Gonads Gonads	Water reabsorption Milk production Growth and metabolism Cortisol release Thyroid hormone synthesis and release Egg or sperm production; sex hormone production Sex hormone production; egg or sperm production
Triiodothyronine and thyroxine (T ₃ , T ₄) Calcitonin (CT)	Many tissues Bone	Metabolism, growth and development Plasma calcium levels (minimal effect in humans)
Parathyroid hormone (PTH)	Bone, kidney	Regulate plasma calcium and phosphate levels
Thymosin, thymopoietin	Lymphocytes	Lymphocyte development
Atrial natriuretic peptide (ANP)	Kidneys	Increase sodium excretion
Angiotensinogen Insulin-like growth factors (IGF)	Adrenal cortex, blood vessels, brain Many tissues	Aldosterone secretion, increase blood pressure Growth

Summary of the Endocrine System Cont.

<i>Hormone</i>	<i>Target</i>	<i>Main Effect</i>
Gastrin, cholecystokinin (CCK), secretin, and others	GI tract and pancreas	Assist digestion and absorption of nutrients
Insulin, glucagon, somatostatin (SS), pancreatic polypeptide	Many tissues	Metabolism of glucose and other nutrients
Aldosterone Cortisol Androgens	Kidney Many tissues Many tissues	Na ⁺ and K ⁺ homeostasis Stress response Sex drive in females
Epinephrine, norepinephrine	Many tissues	Fight-or-flight response
Erythropoietin (EPO) 1,25 Dihydroxy-vitamin D ₃ (calciferol)	Bone marrow Intestine	Red blood cell production Increase calcium absorption
Vitamin D ₃	Intermediate form of hormone	Precursor of 1,25 dihydroxy -vitamin D ₃
Androgen Inhibin	Many tissues Anterior pituitary	Sperm production, secondary sex characteristics Inhibit FSH secretion
Estrogens and progesterone Ovarian inhibin Relaxin (pregnancy)	Many tissues Anterior pituitary Uterine muscle	Egg production, secondary sex characteristics Inhibit FSH secretion Relaxes muscle
Leptin	Hypothalamus, other tissues	Food intake, metabolism, reproduction
Estrogens and progesterone (P) Chorionic somatomammotropin (CS) Chorionic gonadotropin (CG)	Many tissues Many tissues Corpus luteum of ovary	Fetal and maternal development Metabolism Hormone secretion

1: PRIMARY ENDOCRINE ORGANS

Major endocrine organs:

- Hypothalamus and pituitary gland ✓
- Pineal gland
- Thyroid gland and parathyroid glands
- Thymus
- Adrenal glands
- Pancreas
- Gonads

When we talk about the endocrine organs are the ones which release hormones → Tropic hormones: effect the release of other hormones.

① Hypothalamus and Pituitary Gland

- Hypothalamus in brain
- Pituitary gland
 - Anterior lobe
 - Posterior lobe
- Infundibulum → the connection between hypothalamus and pituitary gland.

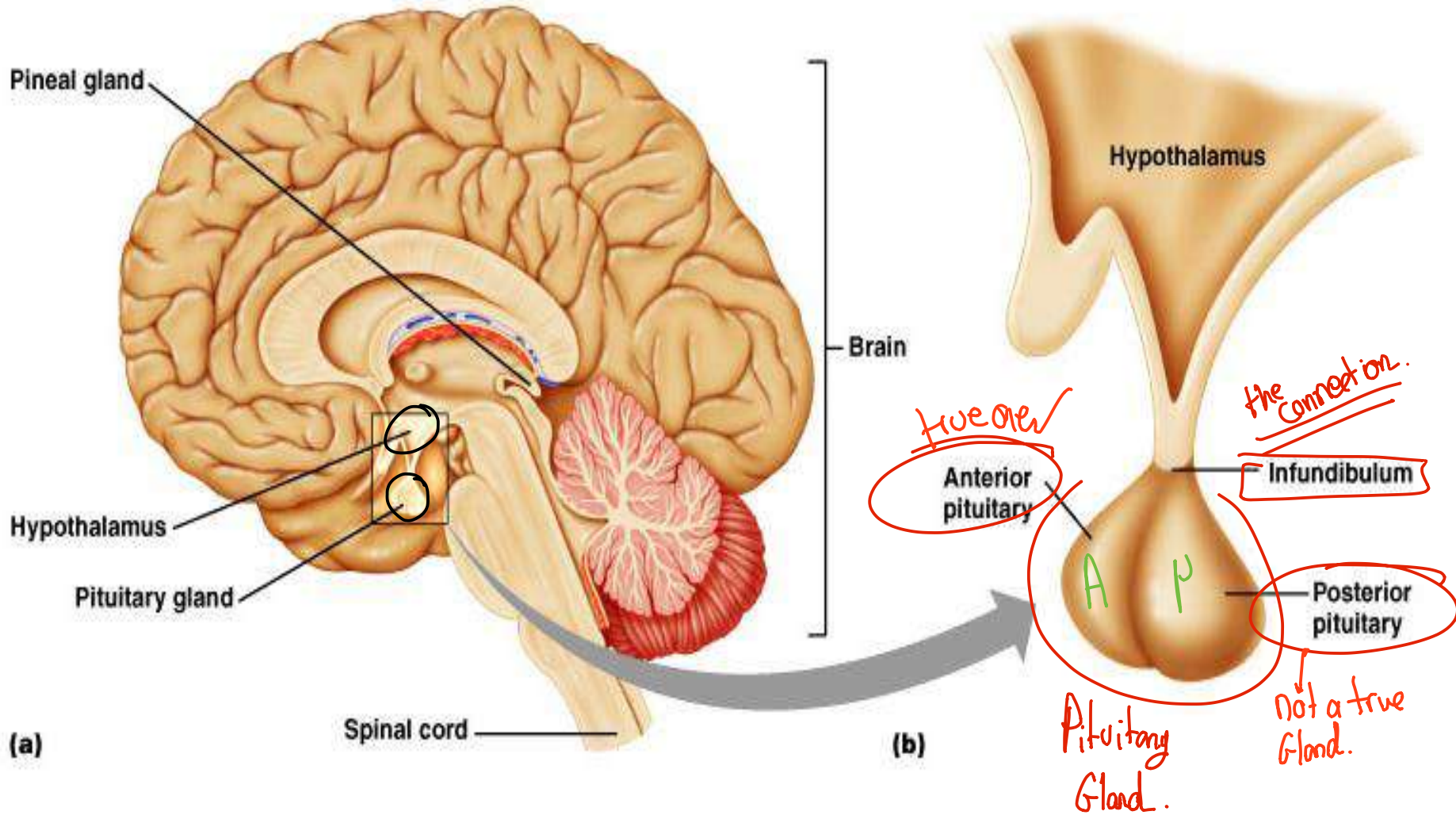
Primary endocrine organs:

• Hypothalamus and pituitary gland: Hypothalamus is directly connected to the pituitary gland

↳ the connection between hypothalamus and pituitary gland
↳ Infundibulum

Anterior lobe
Posterior lobe

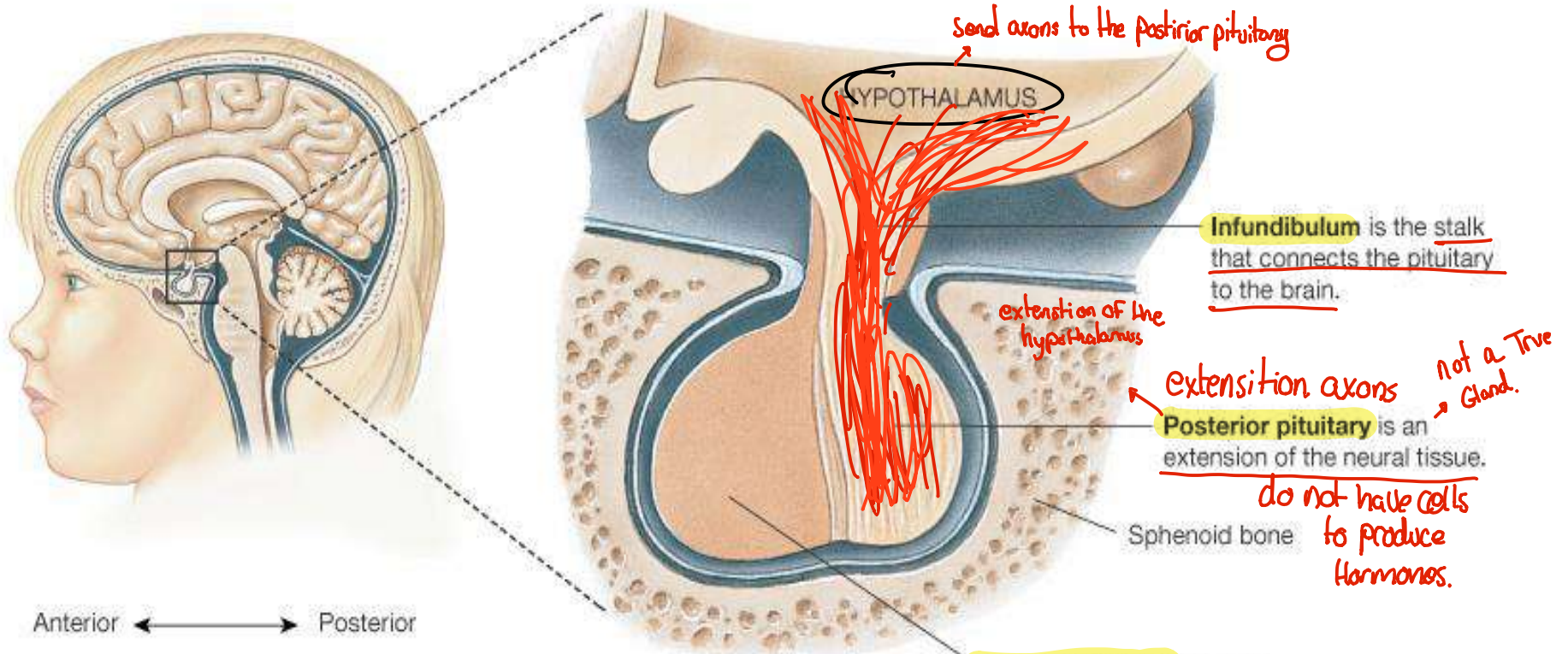
Hypothalamus & Pituitary Gland



Pituitary gland anatomy

Posterior Pituitary is not a true gland → Extension of hypothalamus

Send axons to the posterior pituitary



what we understand → neurosecretory cells that Release Hormones → located in Hypothalamus

Release their content into blood but not in the hypothalamus Area in the posterior pituitary

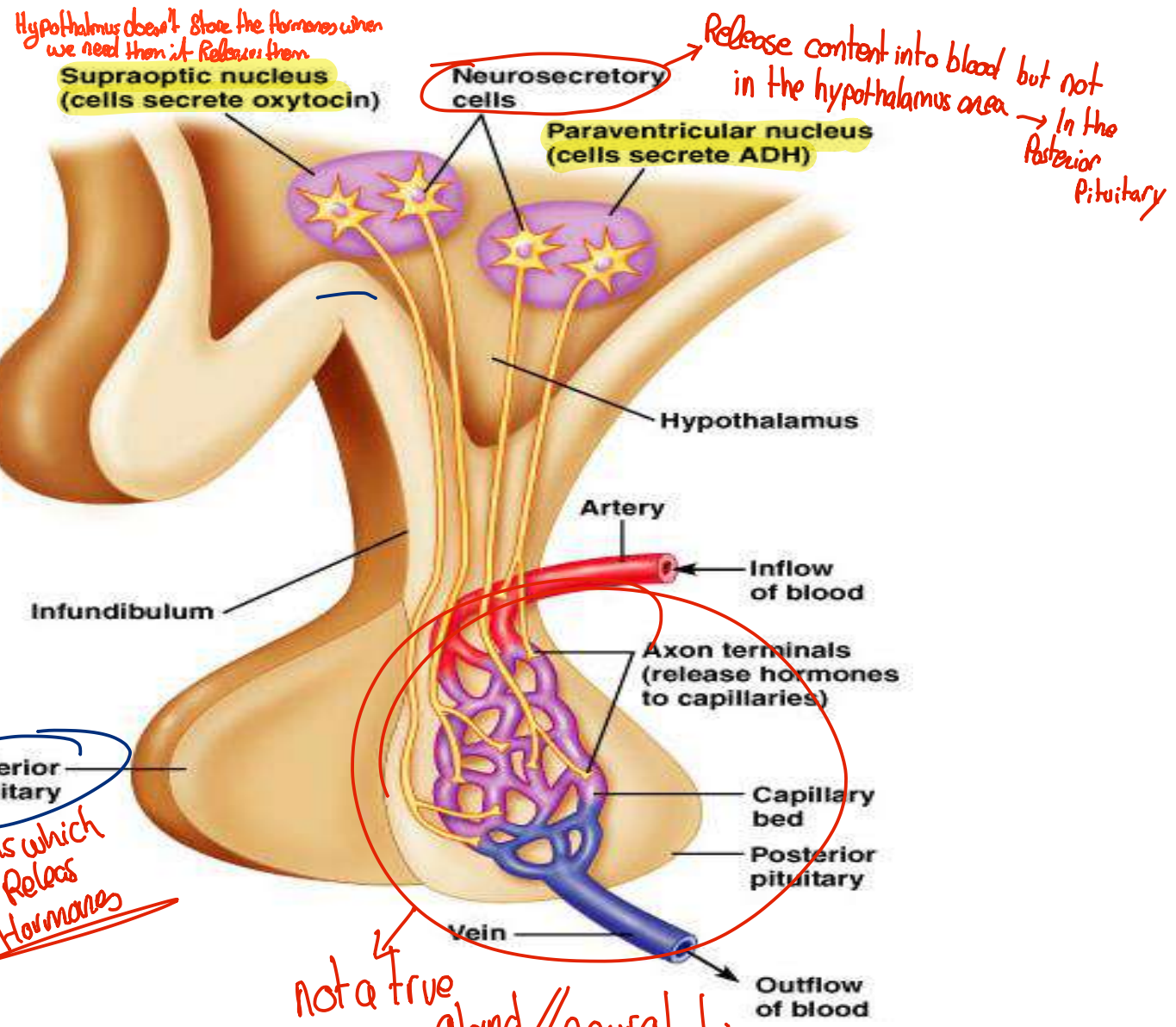
Neural Connection to Posterior Pituitary

Hormones by Hypothalamus:
Oxytocin + ADH Hormone.

Oxytocin → Targets Breast and uterus.
→ Milk ejection + labor and delivery.

ADH Hormone → Target Kidney
→ Water absorption.

Supraoptic Nucleus: Cells secrete oxytocin
Paraventricular nucleus: Cells secrete ADH Hormone



Hormones of Posterior Pituitary

- **Antidiuretic Hormone (ADH or vasopressin)** *→ kidneys*

- Paraventricular nucleus
- Water balance and osmolarity

- **Oxytocin**

- Supraoptic nucleus
- milk ejection + *labor and delivery.*

Blood Connection to Anterior Pituitary

Anterior pituitary has a complicated system
 ↳ it's hormones are released upon the release of other hormones from the Hypothalamus.

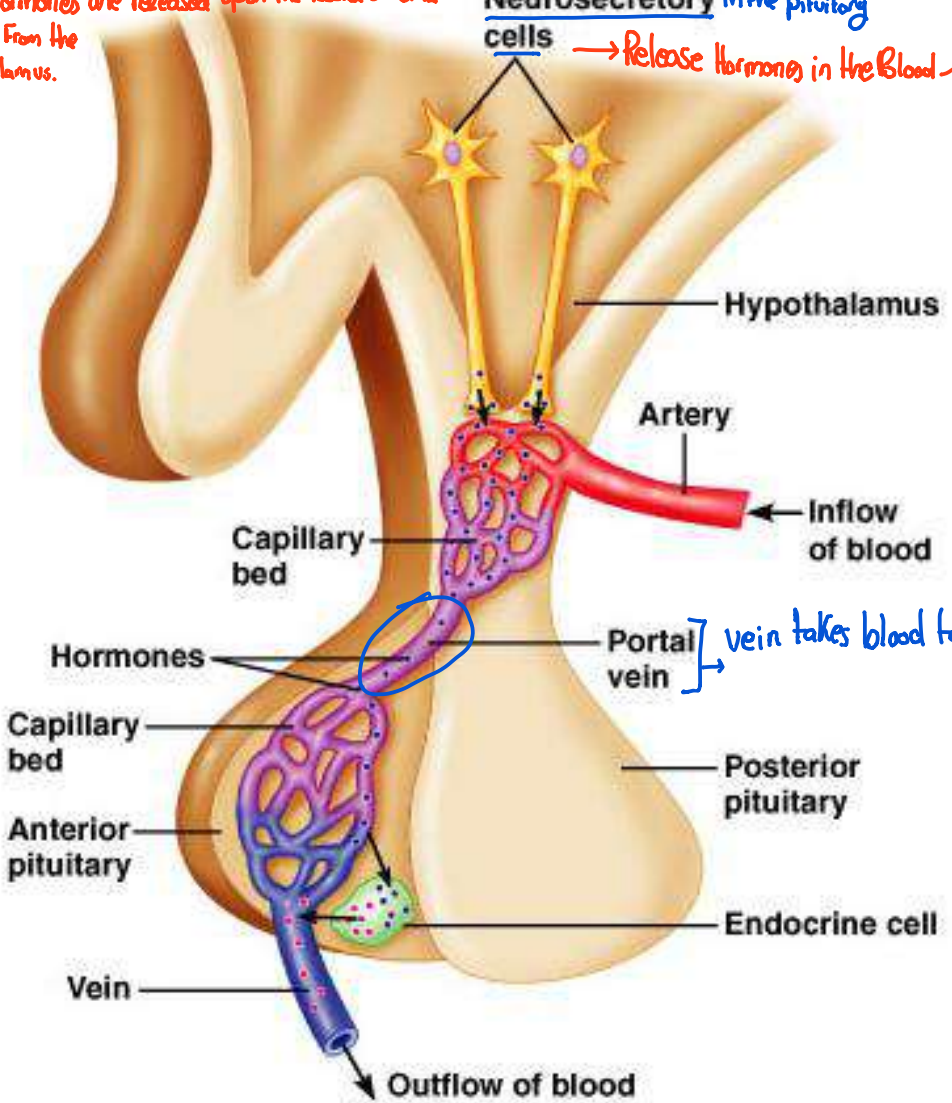
Releases in low conc. → effect only the hormones in the pituitary

So blood has to circulate until it gets in the anterior pituitary and then effecting cells in the anterior pituitary to release their content.

Neurosecretory cells → Release hormones in the blood

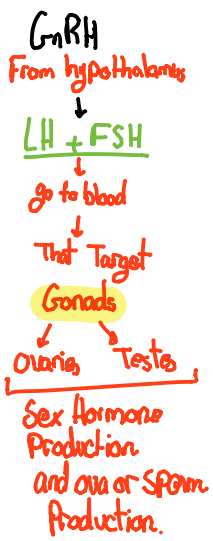
Portal vessels- link two capillary beds

↳ it's important because it moves blood from one section to another section before it goes to heart *why?* → Because we have a very low conc. of hormones released from hypothalamus.



vein takes blood to heart

In capillaries is where exchange occurs between blood and tissue



Tropic Hormones

↳ take action for the release of other hormones.

Tropic Hormones are Hormones Released From a gland effecting other glands to Release Hormones.

Affect release of another hormone

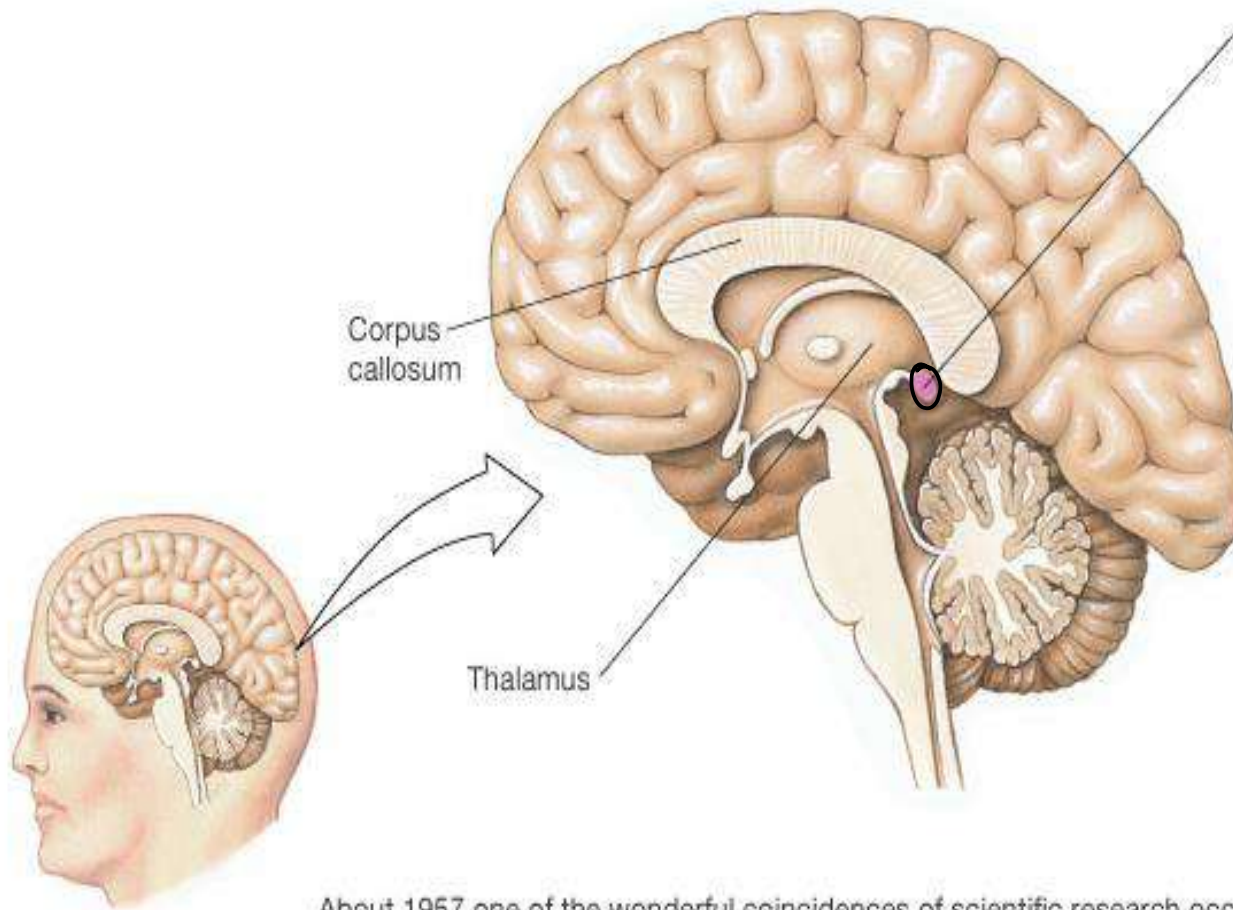
↳ From hypothalamus to anterior pituitary / From anterior pituitary to gonads.

- Releasing hormones
- Inhibiting hormones

GNRH Released From Hypothalamus → Affects the Anterior pituitary so it release LH + FSH Tropic Hormone → Affects Gonads → Egg/Sperm production by Releasing Sex Hormones.

Pineal Gland & Melatonin

THE PINEAL GLAND



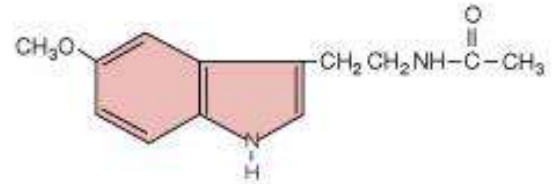
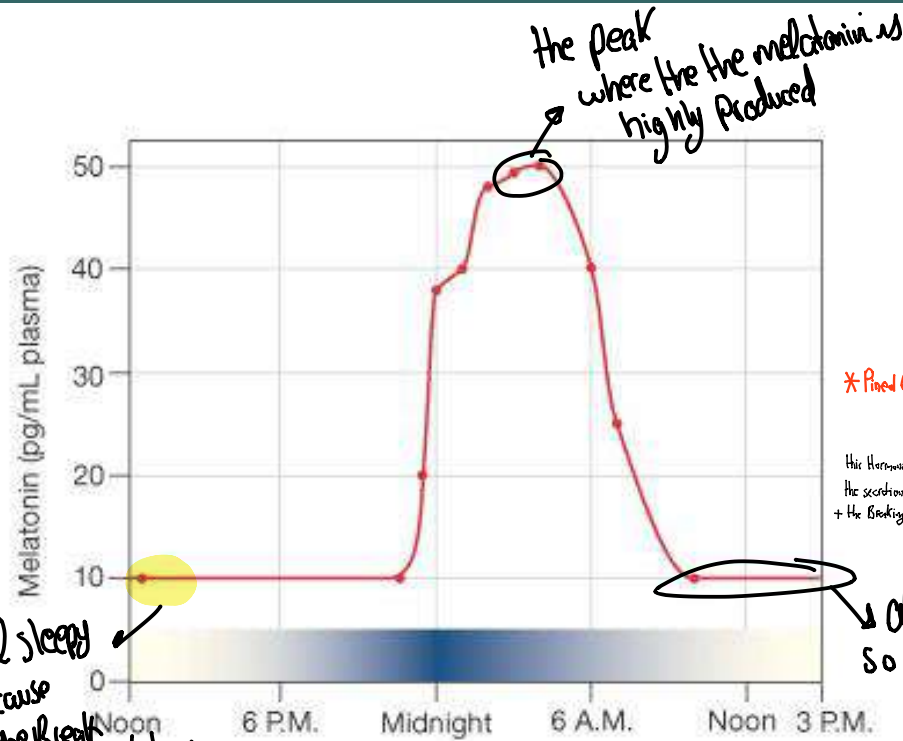
The pineal gland is a pea-sized structure buried deep in the brain of humans. Nearly 2000 years ago, this 'seat of the soul' was thought to act as a valve that regulated the flow of vital spirits and knowledge into the brain. By 1950, however, scientists had decided that it was an evolutionary remnant with no known function.

About 1957 one of the wonderful coincidences of scientific research occurred. An investigator heard about a factor in beef pineal glands that could lighten the skin of amphibians. Using the classical methodology of endocrinology, he obtained pineal glands from a slaughterhouse and started making extracts. His biological assay consisted of dropping pineal extracts into bowls of live tadpoles to see if their skin color blanched. Several years and hundreds of thousands of pineal glands later, he had isolated a small amount of melatonin.

Pineal Gland

- Glandular tissue in brain
- Secretes melatonin
 - Function unknown
 - May be involved in circadian rhythms body clock (A circadian rhythm is a roughly-24-hour cycle in the physiological processes of living beings)
- Influences the antioxidant activity
- Other roles need research: SAAD & sexual behavior

Pineal Gland and Melatonin



Melatonin is an amine hormone derived from the amino acid tryptophan.

* Pineal Gland: Produce melatonin . located in the brain

Hormone that later Regulate Biological Rhythms - sleep-wake cycle.

This Hormone is secreted in different amount during day and night:
the secretion of Melatonin: During night more or full sleep
+ the Breakdown of it wake you up

Melatonin → Amine
Hormone
From Tryptophan

all Breakdown
so you wake up → Biological Hour.

Melatonin is the "darkness hormone," secreted at night as we sleep. It is the chemical messenger that transmits information about light-dark cycles to the brain center that governs the body's biological clock.

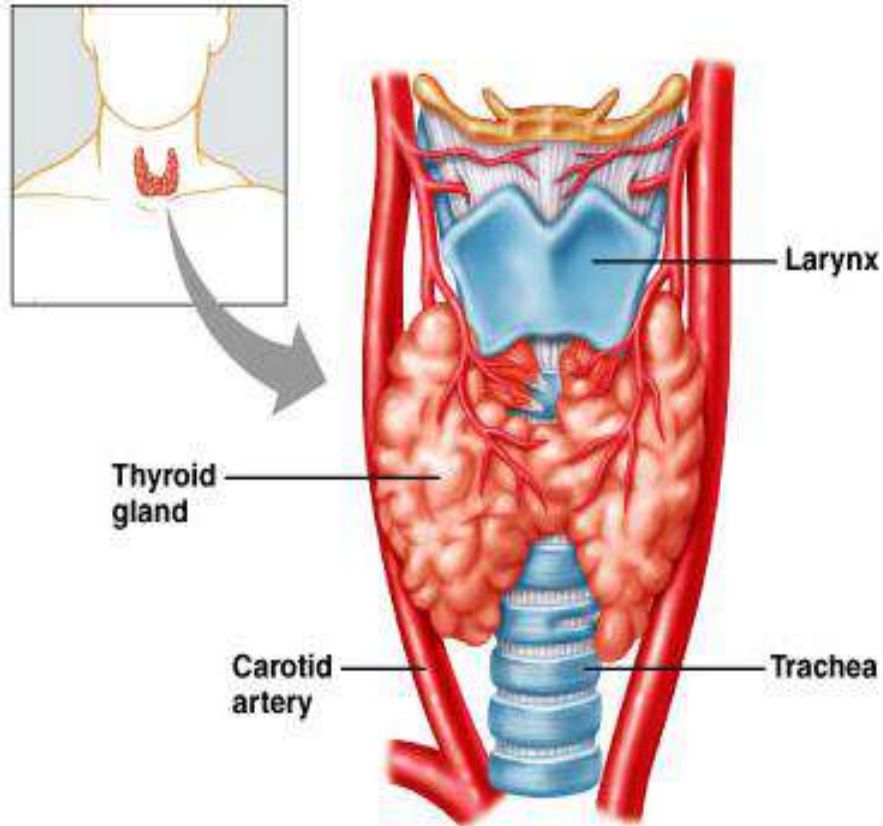
(Adapted from J. Arendt, *Clin. Endocrinol.* 29:205-229, 1988.)

Ten years ago scientists and the popular press were trying to link melatonin to sexual function, the onset of puberty, seasonal affective depressive disorder (SADD) in the darker winter months, and sleep-wake cycles. At that time, the only function supported with scientific evidence was the hormone's ability to help shift timing of the body's internal clock, making it useful in overcoming jet lag. Now there is evidence that melatonin is a powerful antioxidant that has the potential to protect the body from the damage caused by free radicals. For a recent review, see "Melatonin: lowering the high price of free radicals," *News in Physiological Sciences* 15: 246-250, Oct. 2000 (<http://nips.physiology.org>).

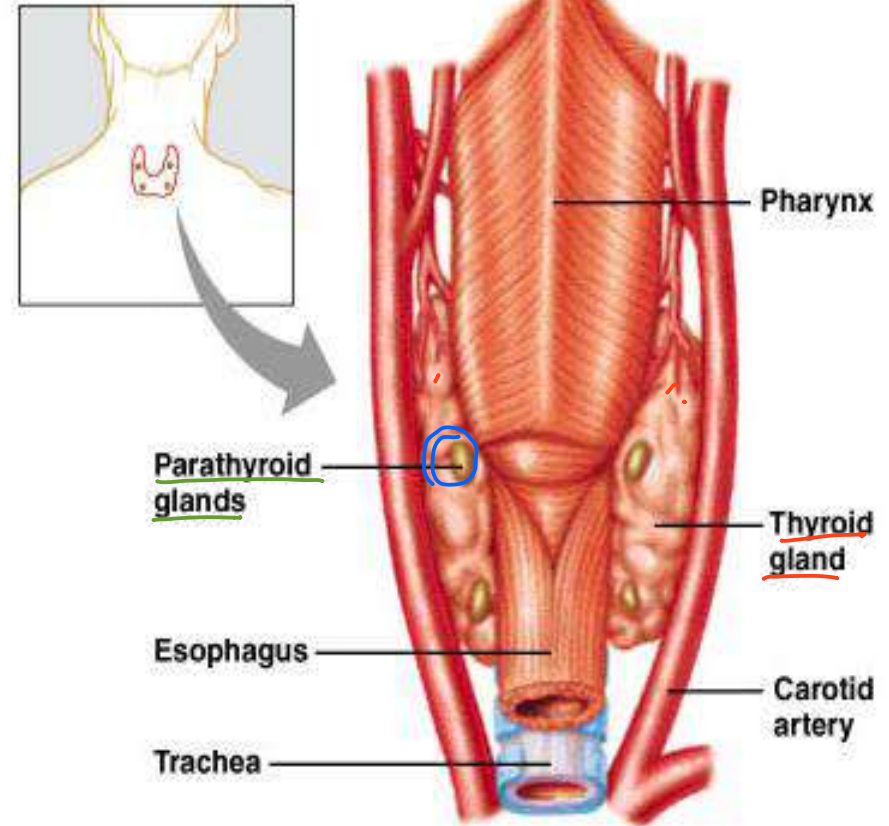
Thyroid and Parathyroid Glands

* Thyroid Gland is located at → Inferior to larynx + Anterior to trachea + Medial to Arms + Superior to lungs

* Parathyroid → Located on the posterior side of Thyroid gland.



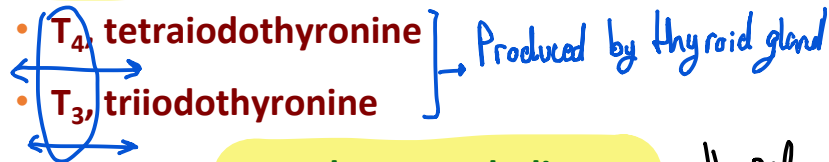
(a) Ventral view



(b) Dorsal view

Hormones of the Thyroid Gland

- Two thyroid hormones



- Regulate metabolism → the role of T_3 and T_4

- Calcitonin + Parathyroid hormone (PTH)

- Regulates calcium levels in blood

→ What effect would you expect to have if we have Hypothyroidism OR Hypoparathyroidism OR Hyperthyroidism OR Hyperparathyroidism?

1) Hypothyroidism: this is a situation where the thyroid doesn't release enough thyroid hormone to the blood stream which slows down metabolism → makes you feel tired OR gaining weight and unable to tolerate temperature

2) Hypoparathyroidism: low secretion OR decrease in activity of PTH → lead to the decrease of blood's calcium + increase in blood phosphorus → (irregular heart beats).

3) Hyperthyroidism: you may feel nervous, moody, weak OR tired → high metabolism + high calcium conc. → irregular heart beats.
↳ increase in the secretion of T_3 , T_4 and Calcitonin.

4) Hyperparathyroidism: increase in blood's calcium → non regulated heart beats → memory loss.

Parathyroid Hormone (PTH)

Regulates calcium levels in blood

Thymus

An organ located in the upper anterior portion of the chest cavity just behind the sternum

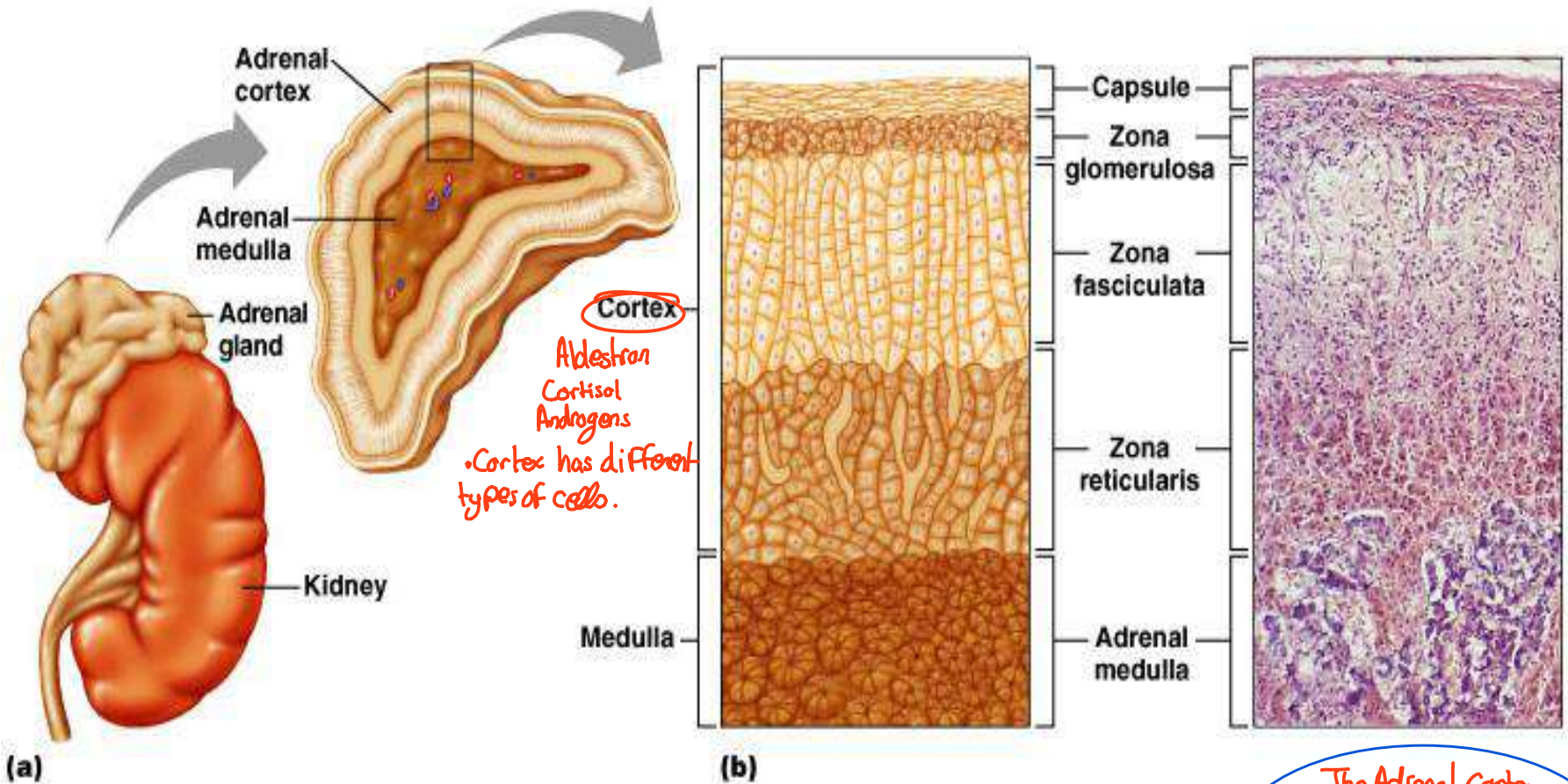
Posterior to

- Secretes thymosin
- Regulates T cell function

Thymus is more important for children.

↳ Immune cells.

Adrenal Glands



* Adrenal Gland is divided into Adrenal Cortex and Adrenal medulla

↳ localization: Below the liver, Posterior to inferior Vena cava, Anterior to diaphragm, Up of the Kidneys.

The Adrenal Cortex and the Adrenal medulla Release different types of Hormones.

Hormones of the Adrenal Gland: Adrenocorticoids

~> these hormones are secreted from **Cortical Area) Cortex**

1. Mineralocorticoids (aldosterone) →

- secreted from **zona glomerulosa**
- **regulates sodium and potassium levels**

Kidneys system: Ajiotinsin Renal Aldosterone

↳ group of hormones → work together → ↑ Pressure
↳ Help in the Reabsorption of Sodium which ↑ press.
in the case when a person loses a lot of liquids.

2. Glucocorticoids (cortisol) ~> Effects the Bones and lower the immunity if it was in Higher Conc.

- secreted from **zona fasciculata and reticularis**
- **regulates body's response to stress** ~> Slow down immune system → Gives pleasure.
- **regulates metabolism**

* increase the appetite which Release more glucose

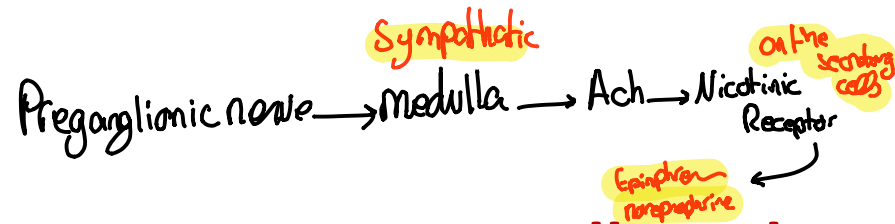
→ in moving Organs From one to other they use Cortisol

3. Sex hormones (androgens) ~> testosterone.

- secreted from **zona fasciculata and reticularis**
- **regulate reproductive function**

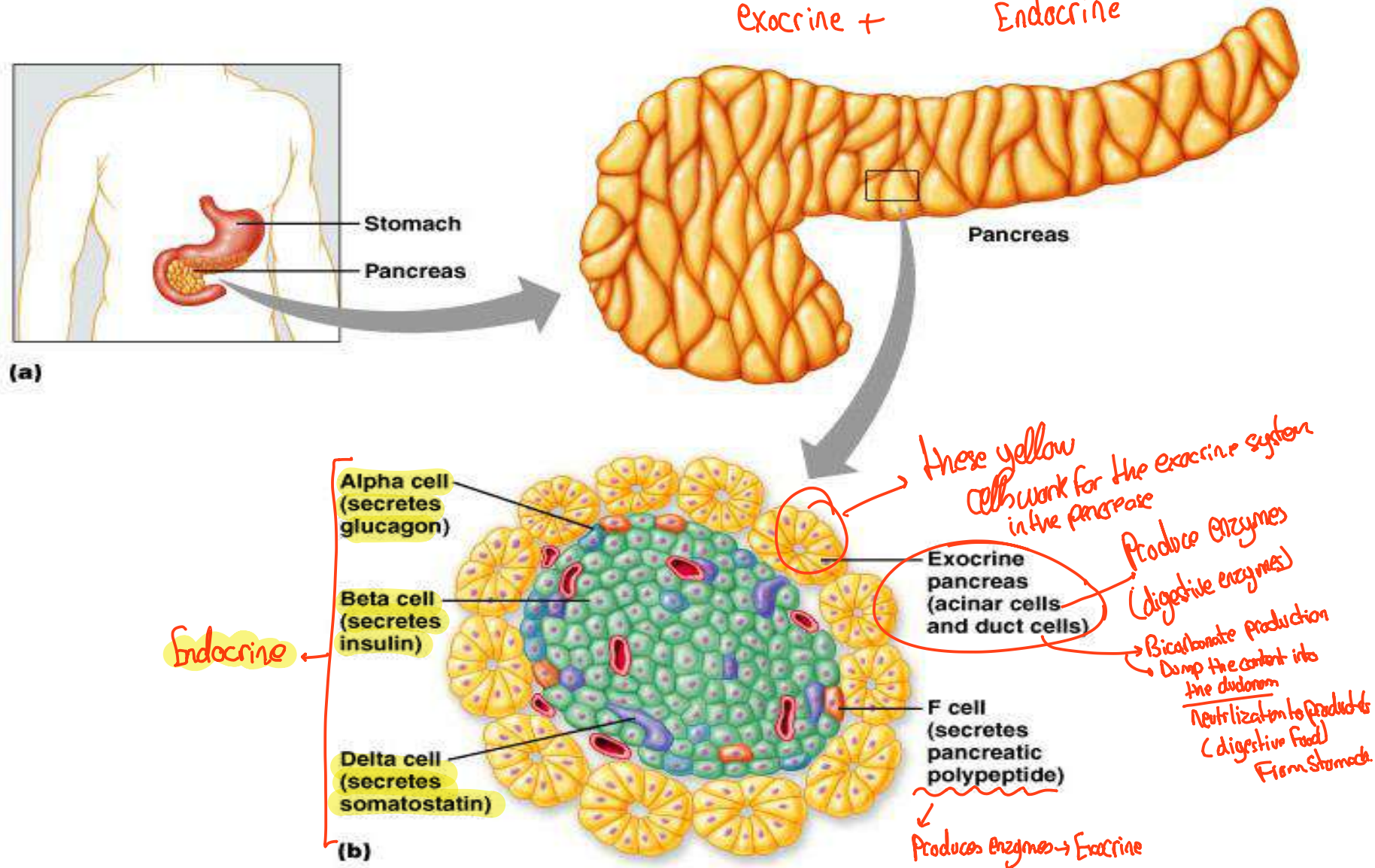
~> **Testosterone:** Produced For Female in low conc. → **Body Hair growth + Sex drive + growth and development of Sex organs.**

Adrenal Medulla



- **Secretory cells = Chromaffin cells**
 - 80% epinephrine
 - 20% norepinephrine
 - <1% dopamine
- **Under neural control**

Anatomy of the Pancreas



Exocrine Pancreas

**Acinar and duct cells secrete fluid
and enzymes into digestive tract**

Endocrine Pancreas

Islets of Langerhans

- **Alpha cells** - glucagon
- **Beta cells** - insulin
- **Delta cells** - somatostatin
- **F cells** - **pancreatic polypeptide** → we have some of them used for the endocrine system also.

Gonads

Male and Female Reproductive organs.

• Male - testes

We have certain cells → **Interstitial cells + Luteal cells** → Produce testosterone

* The conc. of test. is much higher in the Gonads than in the adrenal gland.

- Testosterone
- Androstenedione

• Female - ovaries

Non pregnant pregnant { • **Estradiol** Family of estrogens → Pregnancy Hormone (Maintain pregnancy)

• **Progesterone**

For pregnant women
• **Placenta of pregnant female**

→ estrogens and progesterone

• **Estrogen** → Reproductive development in female
Produced by placenta during pregnancy
the secretion of it is regulated by FSH
Formation sex characteristics
enlargement of uterus + breast during pregnancy.

→ **Pregnancy Hormone**

• **Progesterone**: 1. Prepares uterus for pregnancy
2. Ovulation → Secreted by Ovaries
3. Regulated LH
4. Stimulate the growth of mammary cells.

* Embryo Releases Hormone at the first weeks → if it was present in urine + blood → Pregnancy
↳ **HCG (Human chorionic gonadotropin)**
it is released to protect the embryo + more release of estrogens + progesterone

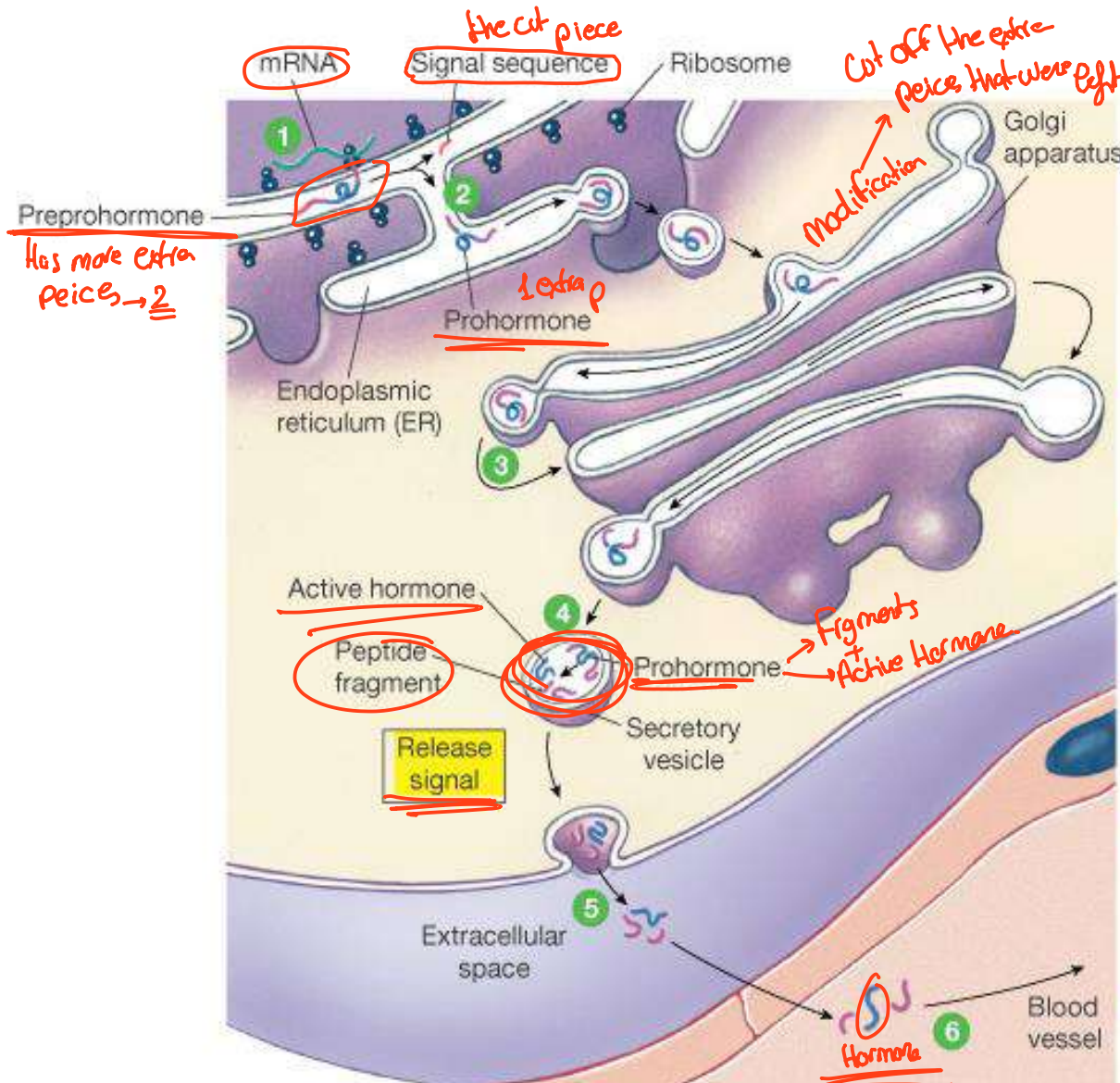
II. Secondary Endocrine Organs

- **Heart** - **atrial natriuretic peptide (ANP)** → targets Kidneys → to Increase sodium excretion.
- **Kidneys** - **Erythropoietin (EPO)** → Bone marrow → Red Blood cell Production
- **GI tract** - several
 - **Cholecystikin (CCK)** → Helps in digestion + Reduce Aptite.
 - **Secretin** → Regulates gastric Acid, Regulation of pancreatic Bicarbonate + Osmoregulation.
 - **Gastrin** → Enhancing gastric mucosal growth, Gastric motility + Secretion of HCl into the stomach.
- **Liver** - **insulin-like growth factors (somatomedins)** → Helps in growth + development.
- **Skin & Kidneys** - **calcitriol** → Prevent low levels of calcium + Bone Disease.

Assist
digestion and
Absorption of
nutrients

is used to treat Hypoparathyroidism
metabolic bone disease in people
who have chronic kidney failure + not
receiving dialysis.

PROTEIN AND POLYPEPTIDE HORMONES: SYNTHESIS AND RELEASE



- 1 Messenger RNA on the ribosomes of the ER binds amino acids into a peptide chain called a **preprohormone**. The chain is directed into the ER lumen by a **signal sequence** of amino acids.
- 2 Enzymes in the ER **chop off the signal sequence**, creating an **inactive prohormone**.
- 3 The prohormone passes from the ER through the **Golgi apparatus**.
- 4 **Secretory vesicles** containing enzymes and **prohormone** bud off the Golgi. The enzymes **chop the prohormone** into one or more **active peptides** plus **additional peptide fragments**.
- 5 The secretory vesicle releases its contents by **exocytosis** into the extracellular space.
- 6 The hormone moves into the circulation for transport to its target.

* How do Amino Acid Based Hormones Are produced?

↳ what is the main organelle do you expect to find in the cell?

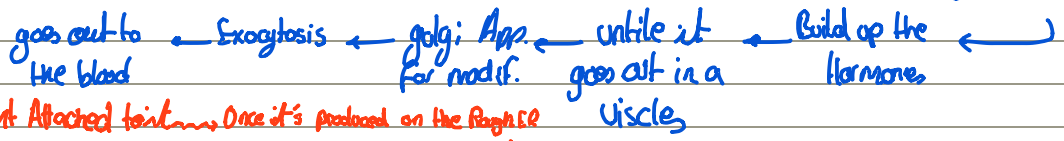
Ribosomes → On the Rough ER (Attached)

↳ what is the difference between attached and free Ribosomes? . Free Ribosomes produce proteins that are required inside the cell.

. Attached Ribosomes produce proteins that are required to be transported outside the cell.

. Modification of Amino Acid Based Hormones is done inside Golgi apparatus, So for these Hormones they need to be synthesised on the Attached Ribosomes in the Rough ER.

* Once we have a Hormone that is amino acid Based → to be synthesized → mRNA that comes from nucleus → Attached Ribosomes on the Rough ER



. PreProHormone → more than 1 part Attached part → Once it's produced on the Rough ER

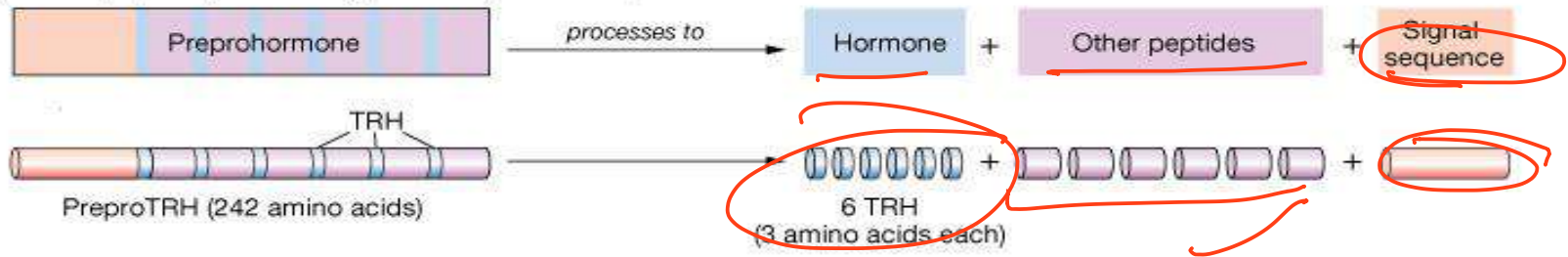
A small piece will be cut of → Signal Sequence →

. Diabetes → Insulin → Active → So they check for C Fragments → gives indication for the Active insulin. // PreProHormone Helps in giving indications in many diseases

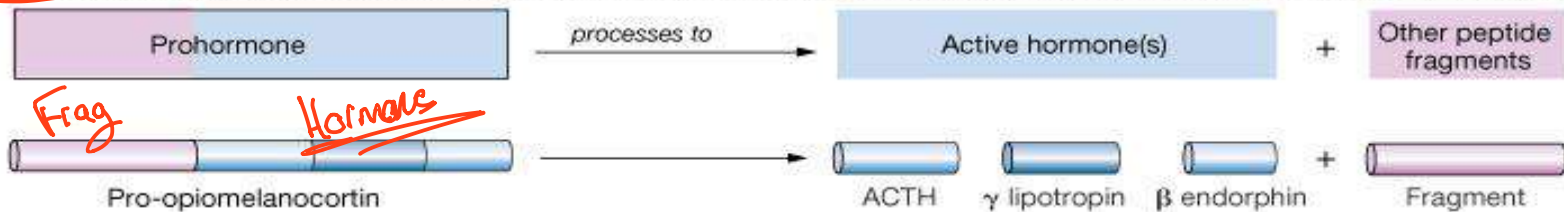
↳ not Active

Peptide hormone processing

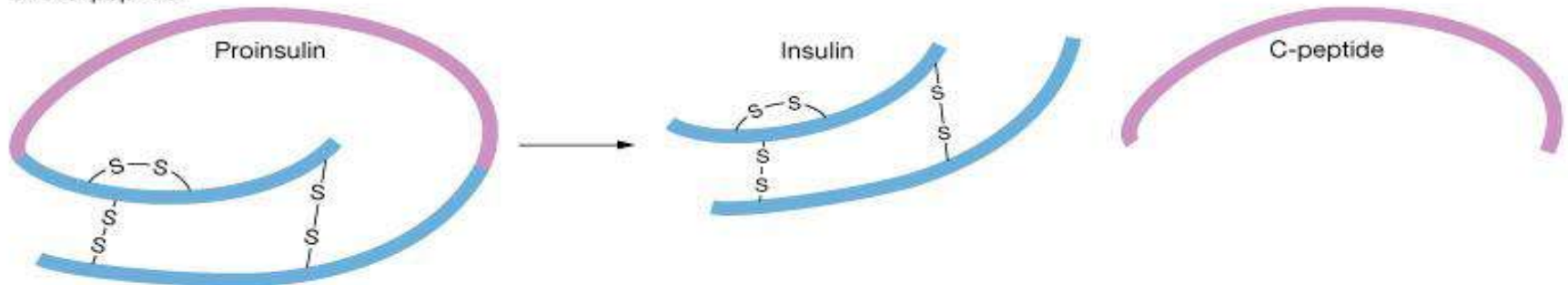
(a) **PreproTRH** (thyrotropin-releasing hormone) has six copies of the 3-amino acid hormone TRH.



(b) **Prohormones**, such as pro-opiomelanocortin, the prohormone for ACTH, may contain several peptide sequences with biological activity.



(c) The peptide chain of insulin's prohormone folds back on itself with the help of disulfide (S-S) bonds. The prohormone cleaves to insulin and C-peptide.



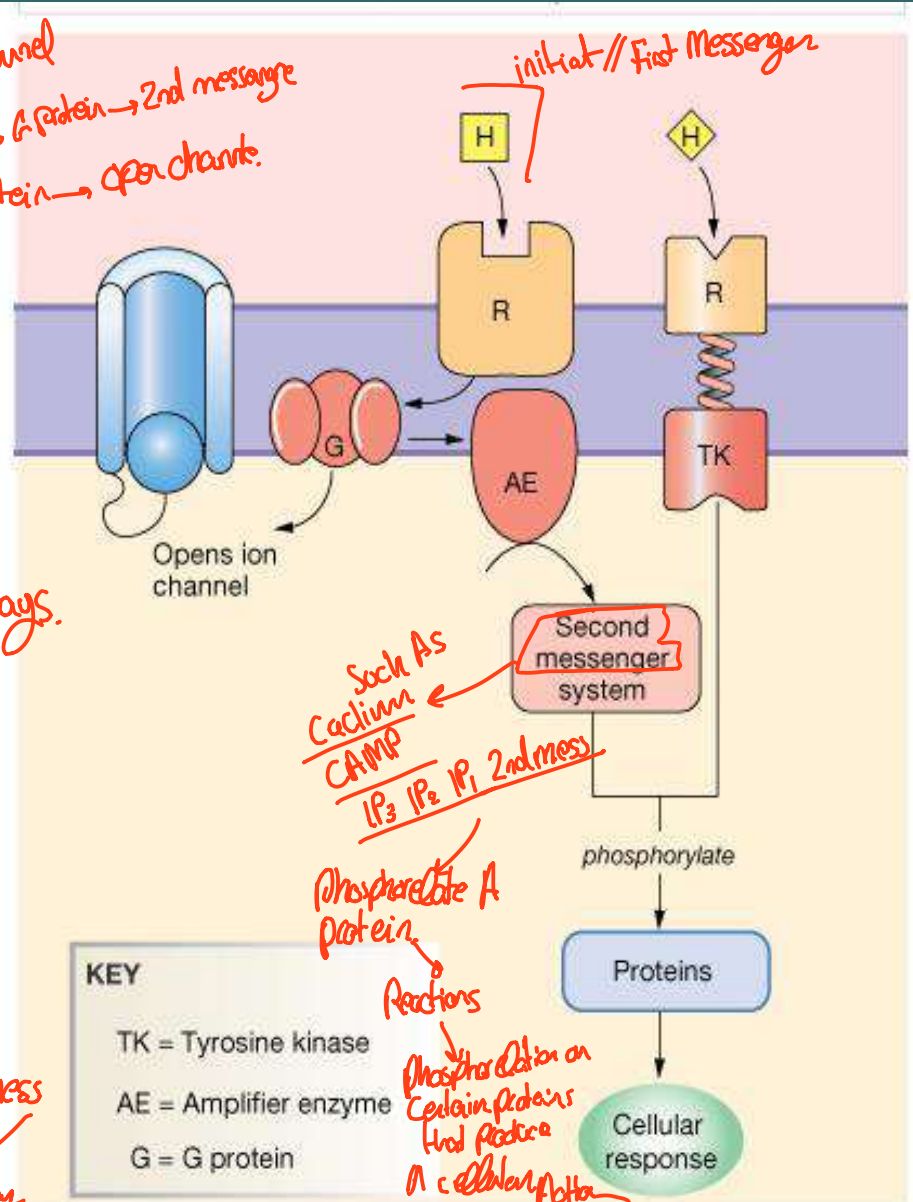
Protein and Polypeptide Hormone Receptors

Hormones come and attached on a Receptor on the cell membrane.

- **Surface receptor**

- Hormone binds
- Transduction
- Enzyme activation
- Open channels
- Second messenger systems
- Synthesis

Part of the channel
Activates G protein → 2nd message
G protein → open channel
initiat // First Messenger



2 ways.

Such As Calcium, cAMP, IP3, IP2, IP1 2nd mess

phosphorylate A protein

Reactions
phosphorylation on certain proteins had produce a cellular response

Surface Receptor → Activation → transduction / 2nd mess

cellular Response.

Steroid Hormones: Structure and Action

↳ Has different Actions

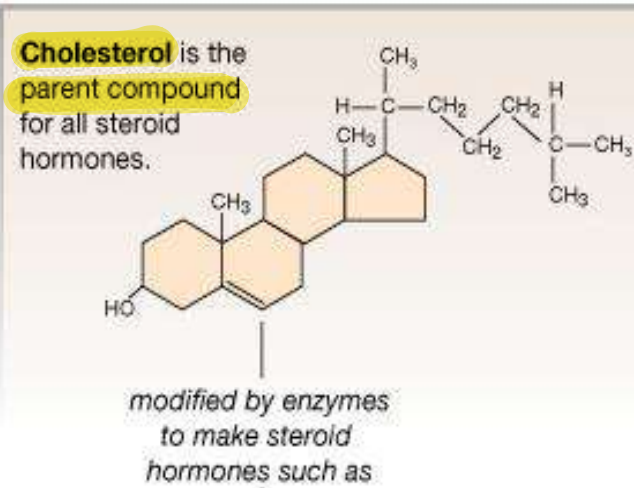
- From **cholesterol**, lipophilic, **enter target cell**, Directly
- **Cytoplasmic or nuclear receptors (mostly)**
↳ Receptor in Cytoplasm/nucleus → DNA to Produce Protein. (Have slower Action)
- **Activate DNA for protein synthesis**
- **Slower acting, longer half-life** ↳ Their action is slower than the action of Amino Acid Based Hormones. (Peptide Hormone) → Action/Cascade is longer.
- **Examples: cortisol, estrogen & testosterone**

Sex Hormones

Steroid hormones are derived from cholesterol

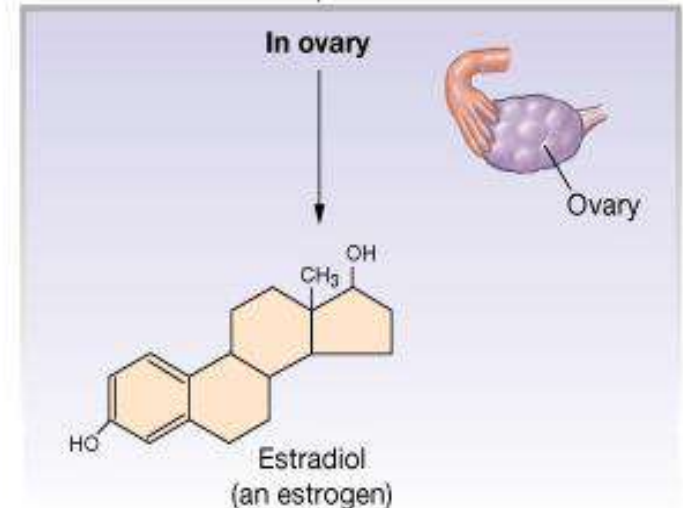
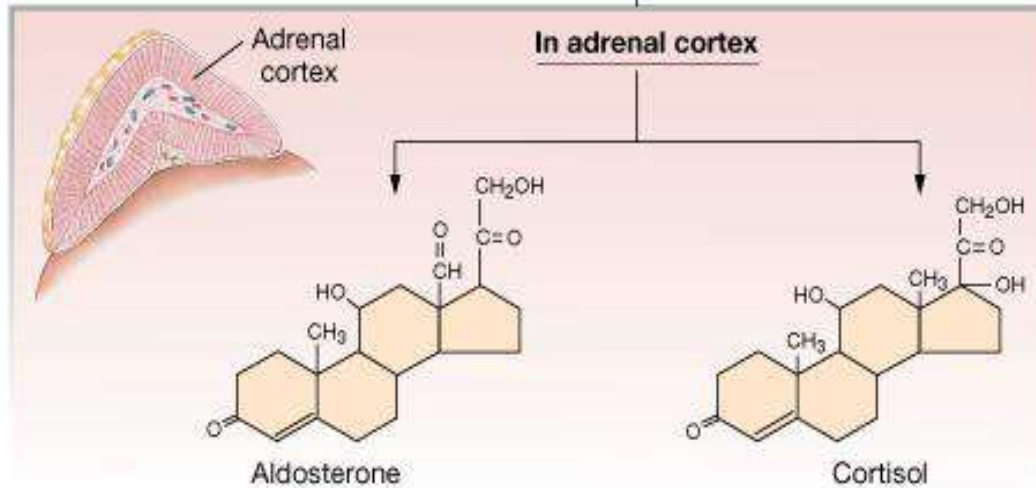
* Cholesterol is a main component of Steroid Hormones.

→ In Ovaries once Female Oocytes → what left from the cells → granulosa cells
 Change their function and start to store cholesterol as much as they can
 So it's ready to produce estrogen and progesterone

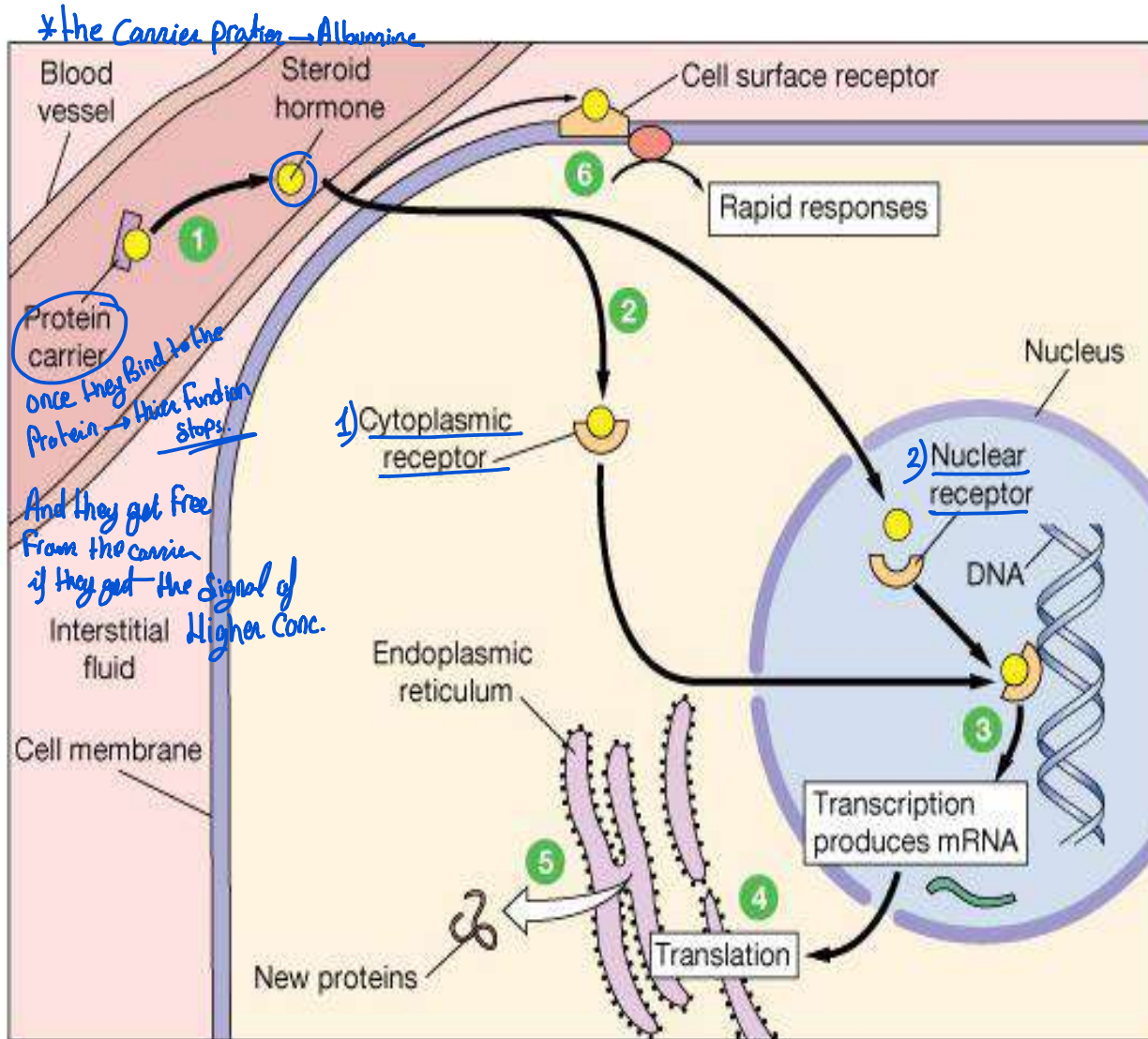


What kind of organelles you expect to find in these cells?

Smooth ER → which produce lipids + Hydrophobic Compounds



Steroid hormone action



- 1 Most hydrophobic steroids are bound to plasma protein carriers. Only unbound hormones can diffuse into the target cell.
- 2 Steroid hormone receptors are in the cytoplasm or nucleus.
- 3 The receptor-hormone complex binds to DNA and activates or represses one or more genes.
- 4 Activated genes create new mRNA that moves back to the cytoplasm.
- 5 Translation produces new proteins for cell processes.
- 6 Some steroid hormones also bind to membrane receptors that use second messenger systems to create rapid cellular responses.

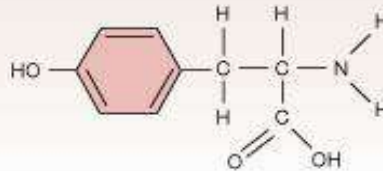
Amine Hormone Structures and Functions

- Made of 1-2 amino acids
- Receptors
 - Surface
 - Intracellular
- Small size, OH group
- Benzene ring
- Examples
 - Thyroxin
 - Epinephrine

* These types are connected together by Tyrosin

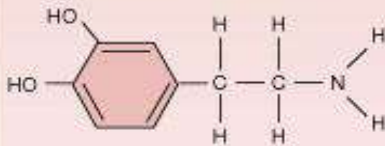
Amine Hormone Structures and Functions

Tyrosine
is the parent amino acid for catecholamines and thyroid hormones.

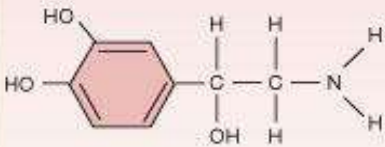


Catecholamines

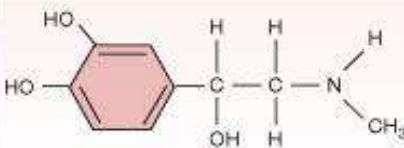
are made by modifying the side groups of tyrosine.



Dopamine



Norepinephrine



Epinephrine

Dopamine
norepinephrine
Epinephrine.

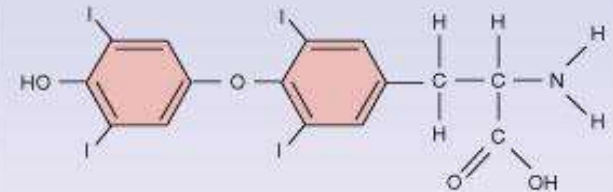
(their main thing is
Tyrosin)

Secreted from tyrosin

T₃+T₄

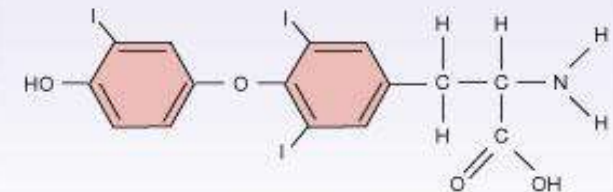
Thyroid hormones

are synthesized from two tyrosines and iodine (I) atoms.



Thyroxine (Tetraiodothyronine, T₄)

From Thyroid Gland



Triiodothyronine (T₃)

Comparison of Peptide, Steroid, and Amine Hormones

	PEPTIDE HORMONES	STEROID HORMONES	AMINES	
			CATECHOLAMINES	THYROID HORMONES
Synthesis and storage	Made in advance; stored in secretory vesicles	Synthesized on demand from precursors	Made in advance; stored in secretory vesicles	Made in advance; precursor stored in secretory vesicles
Release from parent cell	Exocytosis	Simple diffusion	Exocytosis	Simple diffusion
Transport in blood	Dissolved in plasma	Bound to carrier proteins	Dissolved in plasma	Bound to carrier proteins
Half-life	Short	Long	Short	Long
Location of receptor	On cell membrane	Cytoplasm or nucleus; some have membrane receptors also	On cell membrane	Nucleus
Response to receptor-ligand binding	Activation of second messenger systems. May activate genes	Activate genes for transcription and translation. May have nongenomic actions	Activation of second messenger systems	Activate genes for transcription and translation
General target response	Modification of existing proteins and induction of new protein synthesis	Induction of new protein synthesis	Modification of existing proteins	Induction of new protein synthesis
Examples	Insulin, parathyroid hormone	Estrogen, androgens, cortisol	Epinephrine, norepinephrine	Thyroxine (T ₄)

Endocrine Reflex Pathways: Overview

What's the pathway for the hormone? Any hormone to be activated we have to have a stimulus. → For insulin to be produced (Increase in Blood Sugar)
→ For Glucagon to be produced (Decrease in Blood Sugar)

- **Stimulus** (Increase in Blood Sugar)

- **Afferent signal**

- **Integration** (Integration Center → Pancreas)

- **Efferent signal (the hormone)** β cells

- **Physiological action** → Exocytosis of vesicles containing insulin

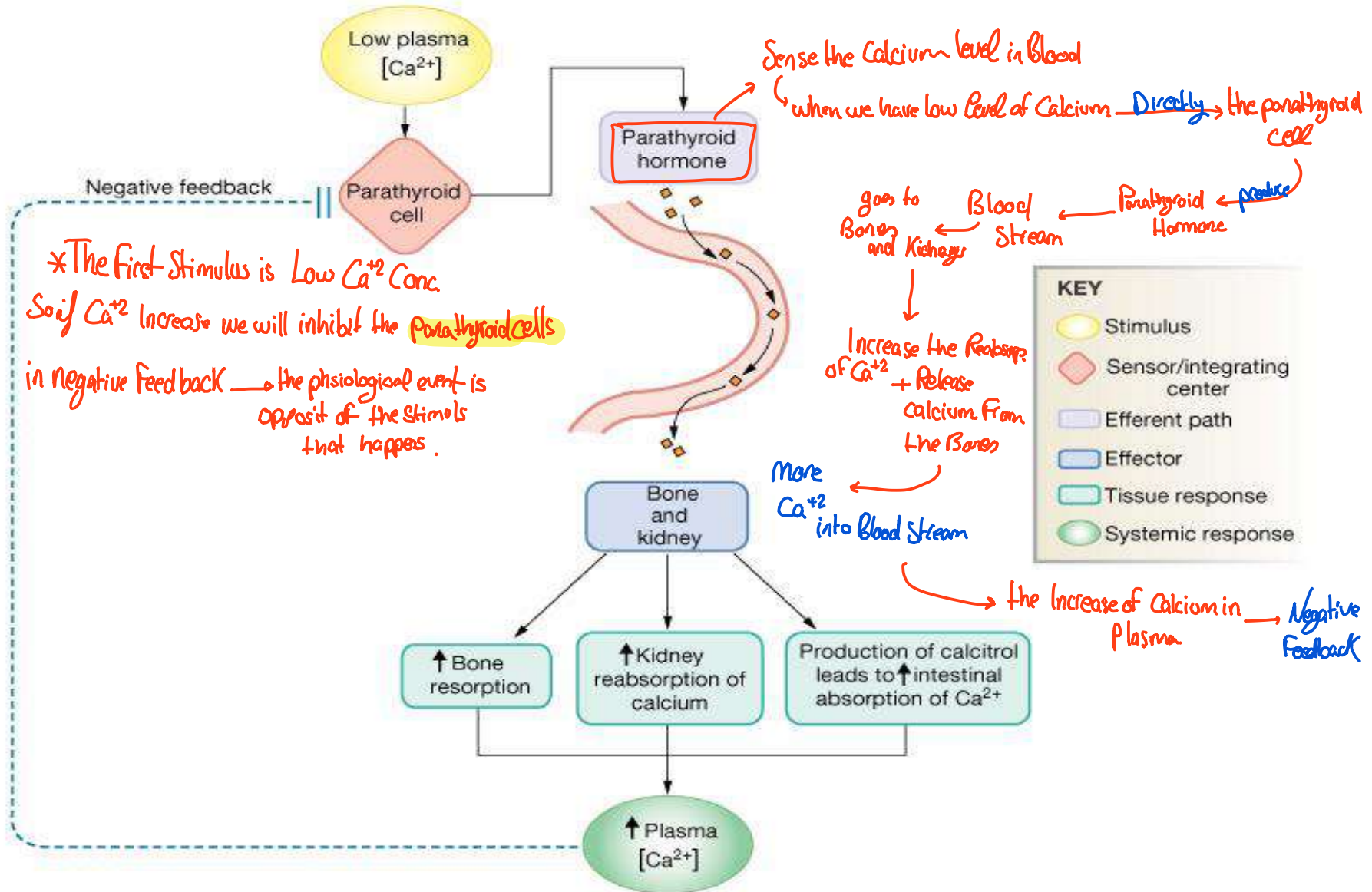
- **Negative feedback** → The Action.

* Insulin is a good example for negative feedback
discuss?

When Blood Sugar rises → Receptors in the Body sense the change → the Control center (Pancreas) secretes insulin into the Blood

Effectively lowering Blood Sugar level.

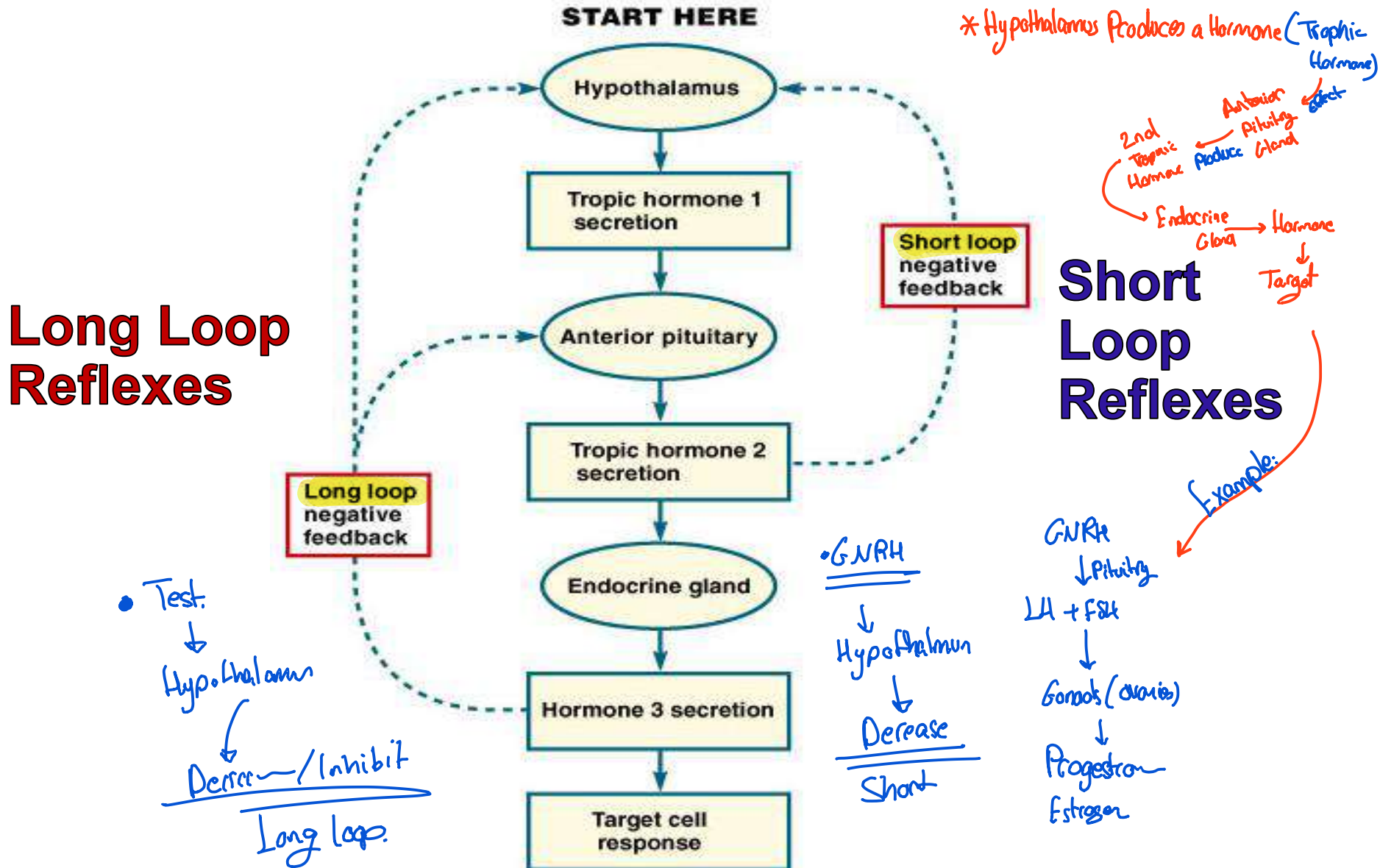
A simple endocrine reflex: Parathyroid hormone



Negative Feedback Loops

Long Loop Reflexes

Short Loop Reflexes



* An increase of insulin causes a decrease in Blood Sugar and the opposite.

↳ Short + Direct loop for negative feedback.

Hypothalamus ^{Tropic Hormone} sends → A Hormone (GnRH) ^{Tropic Hormone} sends → Pituitary gland (LH, FSH) → Glands Release Test, estrogen, progesterone.

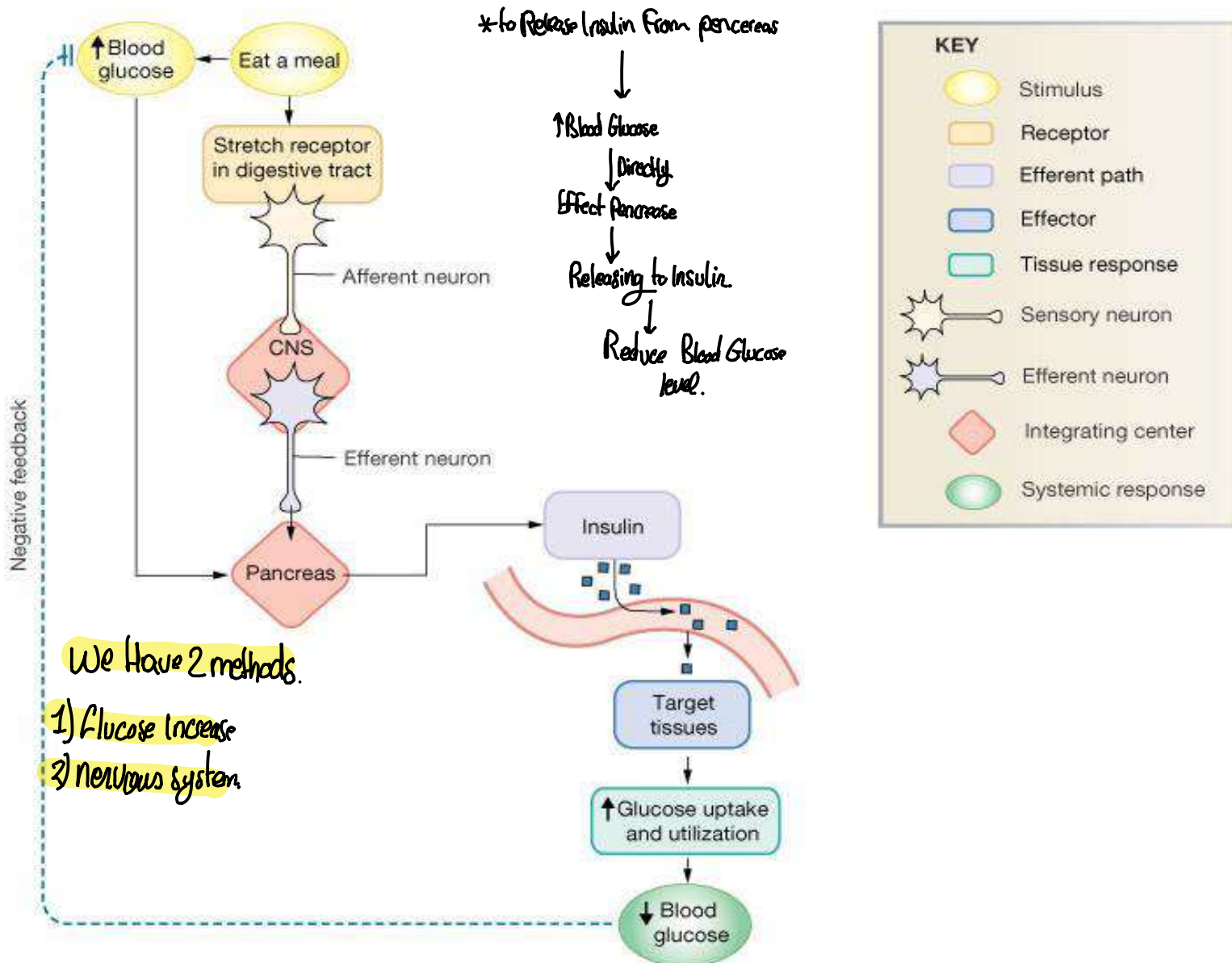
• FSH + LH → Can effect the Release of GnRH From the Hypothalamus
Short loop.

Increase in Test. → Anterior Pituitary to Reduce the Release of LH and FSH

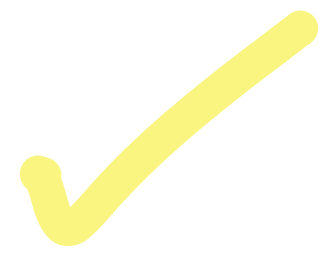
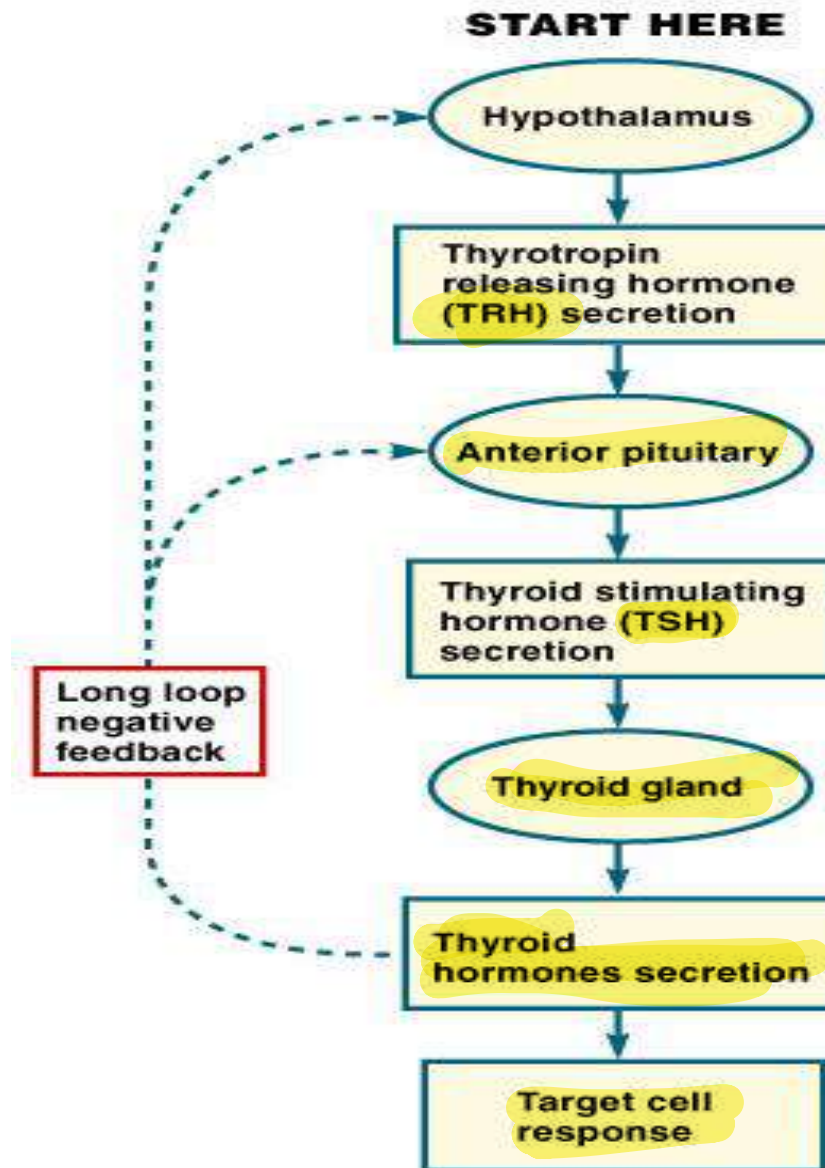
Long loop

Effect the Release of GnRH From the Hypothalamus. (*)

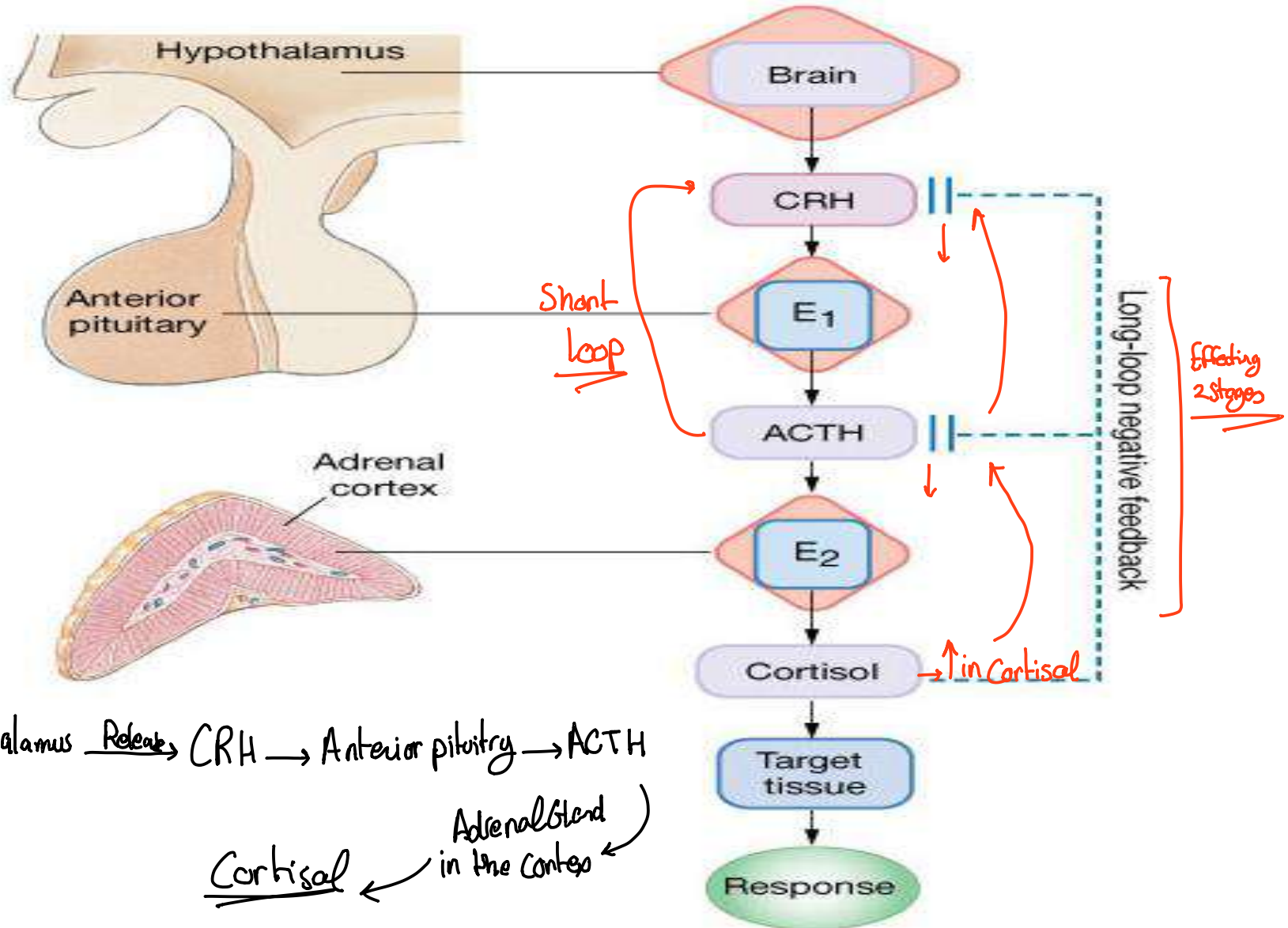
Endocrine Reflex Pathways: multiple stimuli



Example of Negative Feedback Control: Thyroid Hormones



Negative Feedback Controls Long & Short Loop Reflexes



✓ Tropic Hormones

Affect release of another hormone

- **Releasing hormones**
- **Inhibiting hormones**

Common Tropic Hormone Pathway

- 1. Hypothalamus secretes releasing or inhibiting hormone into capillary bed**
- 2. Blood with tropic hormones enters portal vein**
- 3. Hypothalamic tropic hormones access anterior pituitary secretory cells through capillary beds**
- 4. Alter release of anterior pituitary tropic hormones**
- 5. Anterior pituitary tropic hormones enter bloodstream in same capillary bed**
- 6. Travel to distant endocrine gland to trigger release of hormone**

Control of Hypothalamic Tropic Hormone Release

How to Control?

- **Neural input** → As for example the stretch in the stomach
- **Hormonal - negative feedback** → ↑Hormone → event → normal → negative feedback for the event.
- **Circadian rhythm** → sleep + wakeup cycle
 - **Suprachiasmatic Nucleus of Hypothalamus**

Neurohormones

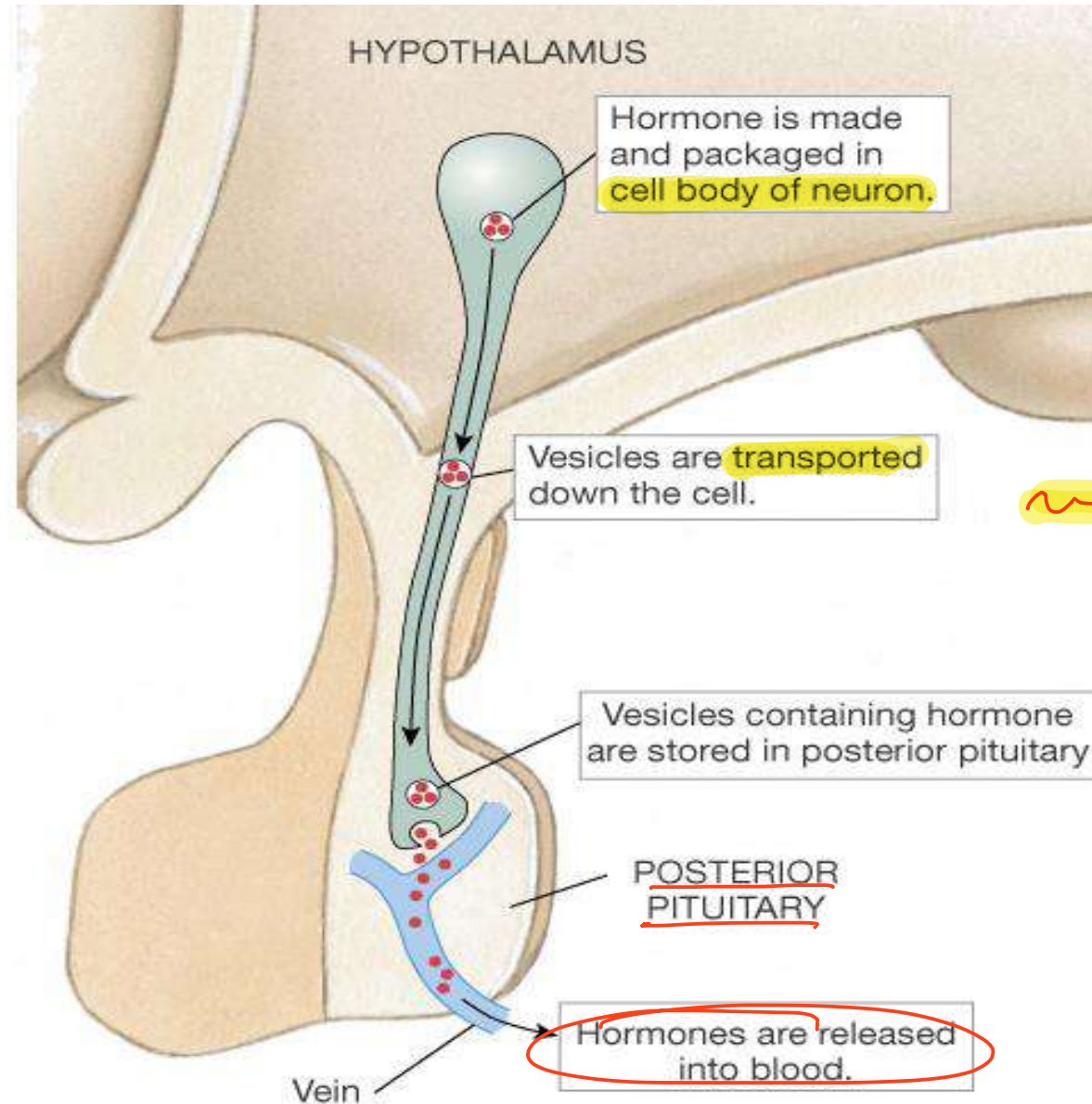
secreted into the Blood by Neurons

→ Another type of Hormones → Released to the blood but they are produced from nerve cells.

↓
Oxytocin
Vasopressin (ADH)
From the hypothalamus

- **Adrenal Medulla—catecholamines**
- **Hypothalamus to:**
 - **Anterior pituitary**
 - Tropic Hs ✓
 - Growth H. ✓
 - Prolactin ✓
 - **Posterior pituitary**
 - Vasopressin ✓
 - Oxytocin ✓

Neurohormones secreted into the Blood by Neurons

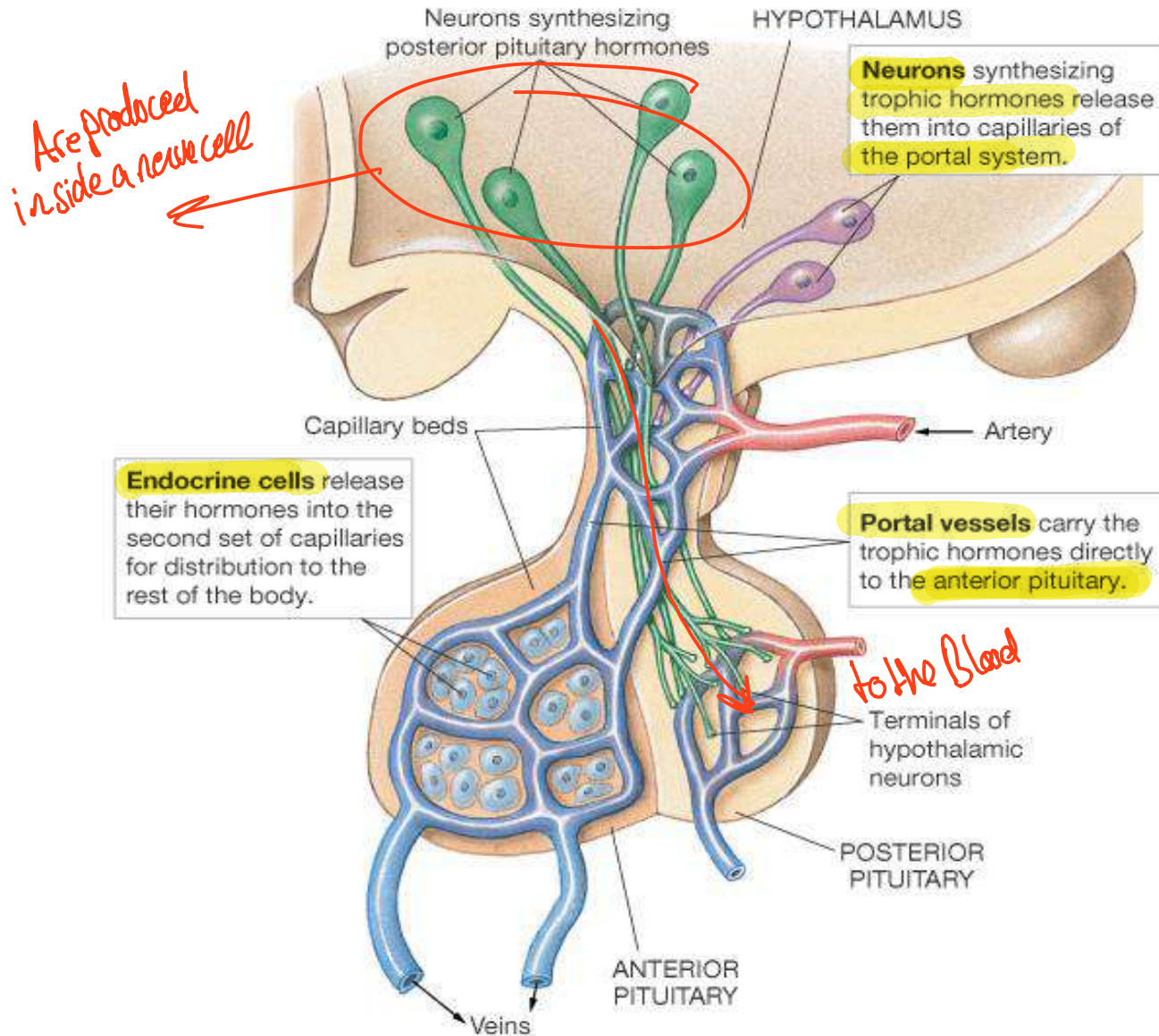


Posterior
Pituitary.

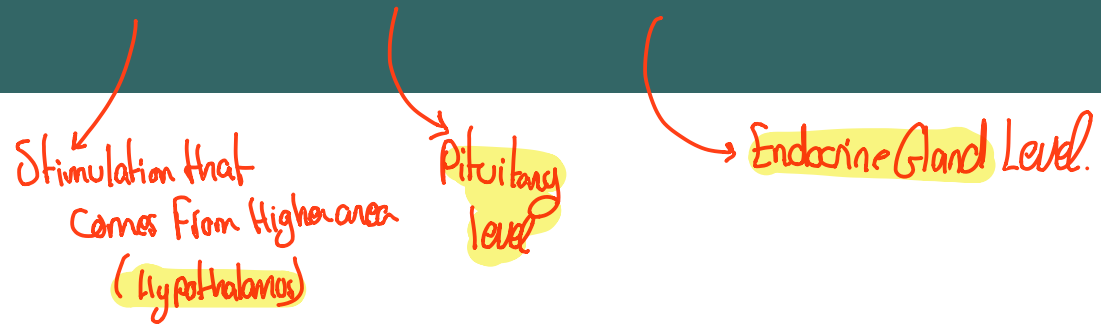
→ neurohormone

The hypothalamic-hypophyseal portal system

Anterior Pituitary

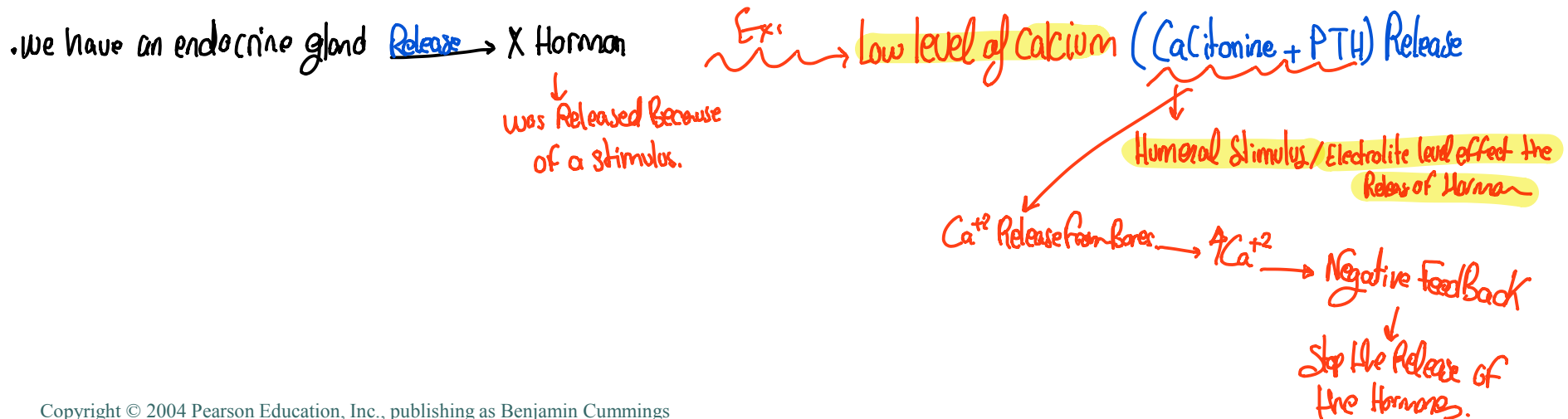


Endocrine Control



• Three Levels of Integration

- Hypothalamic stimulation—from CNS
- Pituitary stimulation—from hypothalamic trophic Hs
- Endocrine gland stimulation—from pituitary trophic Hs

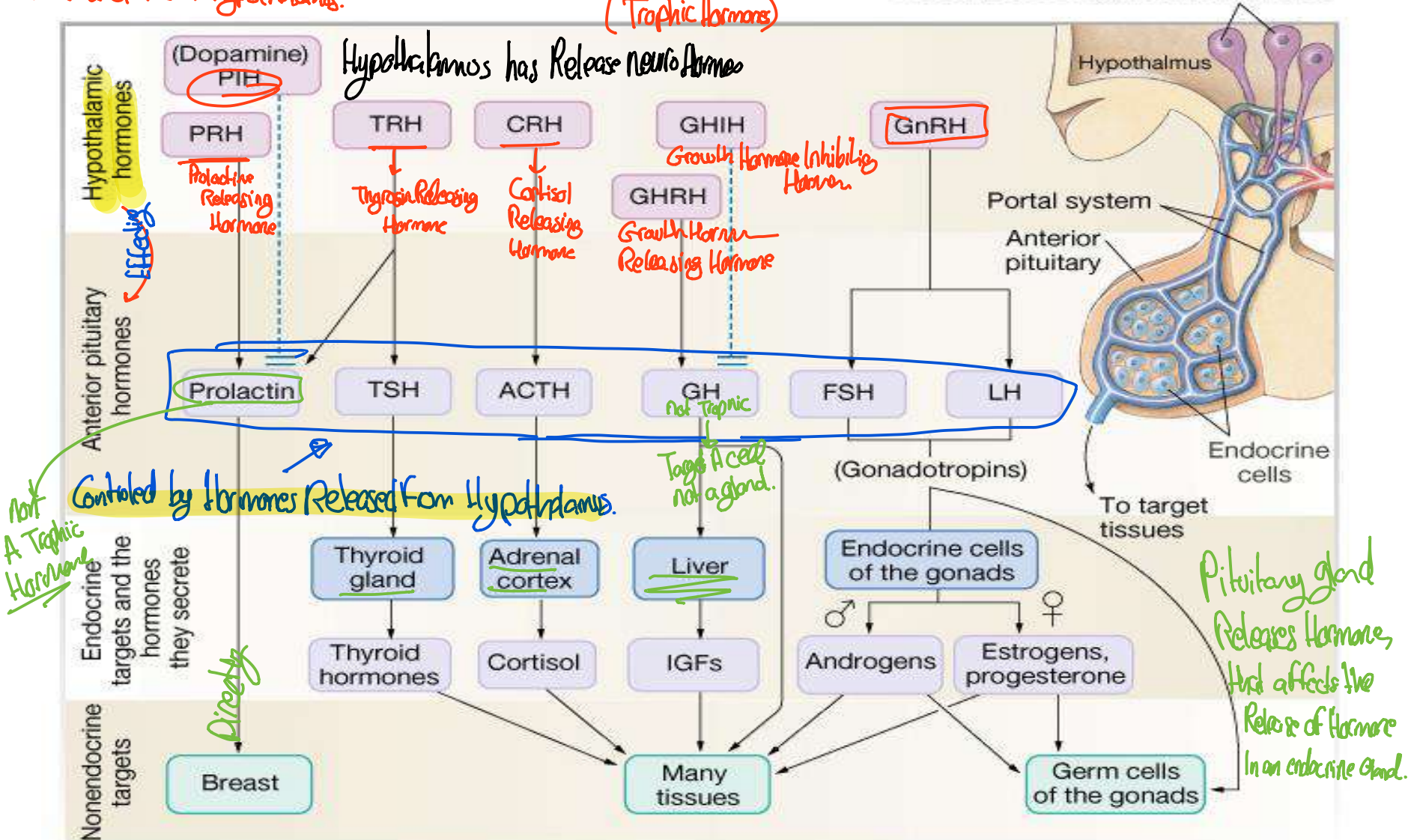


Endocrine Control: Three Levels of Integration

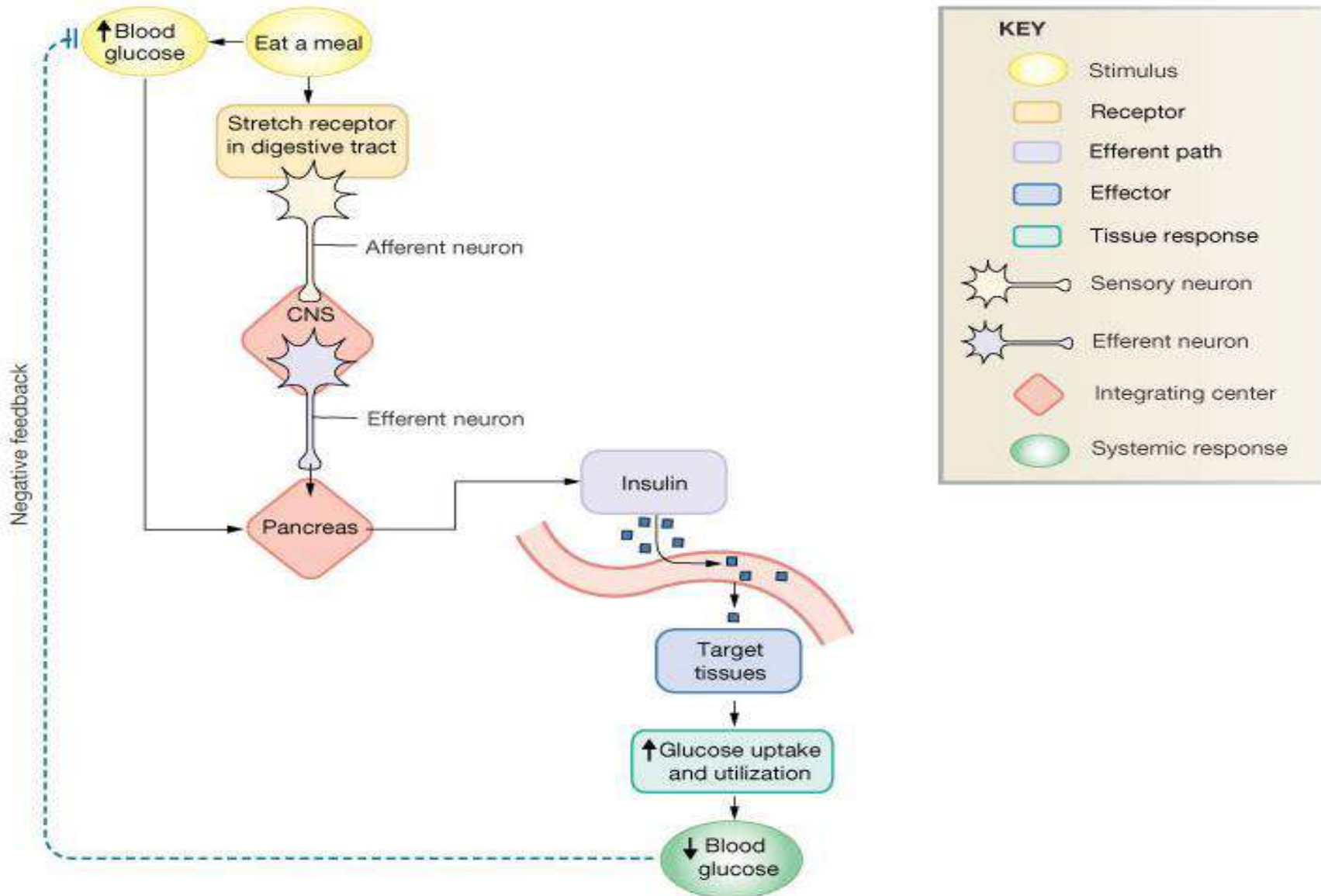
Released from hypothalamus.

(Trophic Hormones)

Neurons secreting trophic hormones

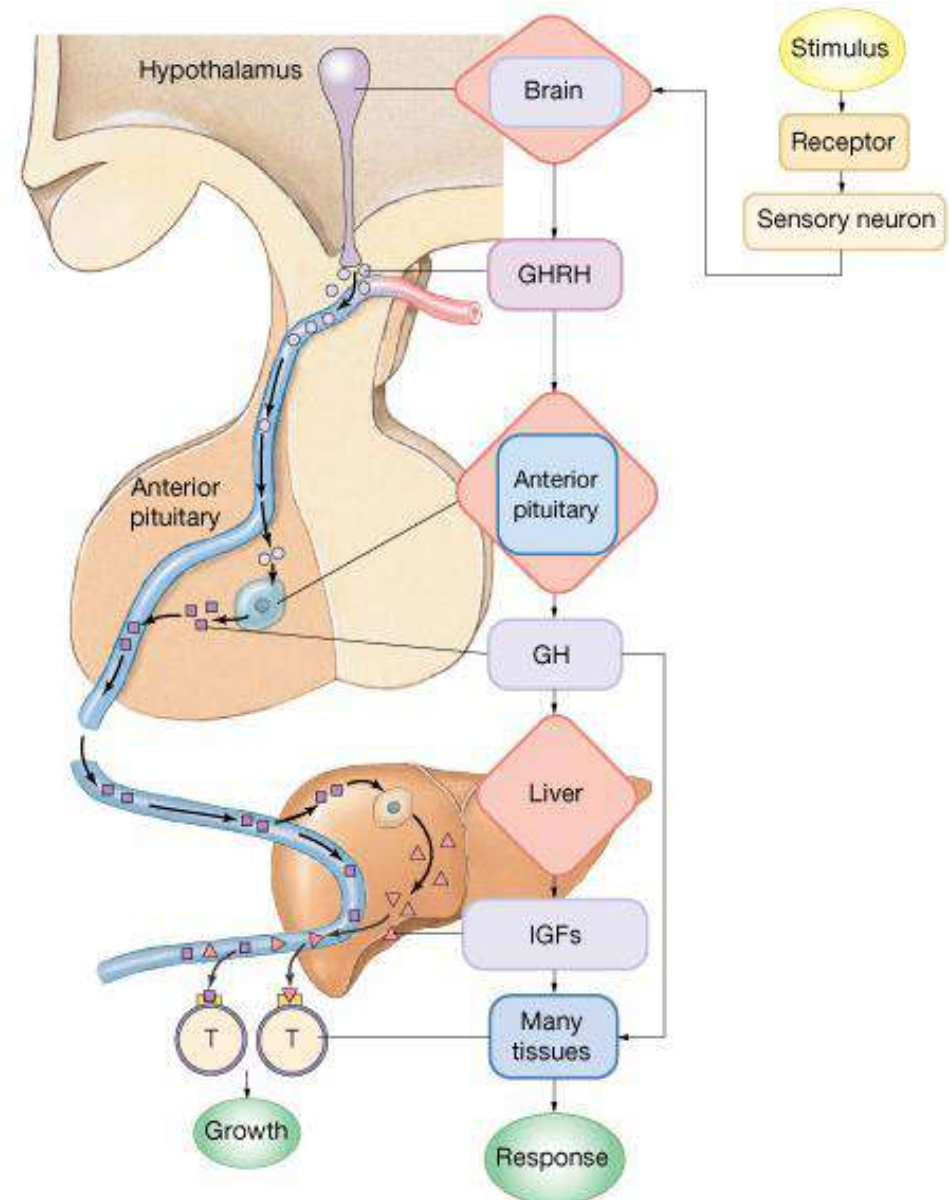


Multiple Stimuli for Hormone Release: Nervous & Endocrine



Multiple Hormones Can Target a Cell/Tissue

- **Growth H**
- **Somatomedins**
- **Thyroxin**
 - All have receptors on many tissues
 - Stimulate pathways for growth



Hormone Interactions

*Hormones do interact with each others → they do:

- **Antagonism**
- **Additive**
- **Synergism**
- **Permissiveness**

Antagonism

Effects of two hormones oppose each other

if one increase the other decrease

Example: **insulin** and **glucagon** on blood glucose

Lower Blood Sugar

Higher Blood Sugar Conc

2) Additive

We have more than one type of Growth Hormones → they add up together

Effects of two hormones favor each other and sum

Does the summation of effect.

Synergism

One Hormone Increases the Level of

effects of two hormones favor each other but
the net effect exceeds the sum of individual
effects ✓

Example: ⁷Glucagon, ⁸cortisol, and
epinephrine on blood glucose

4

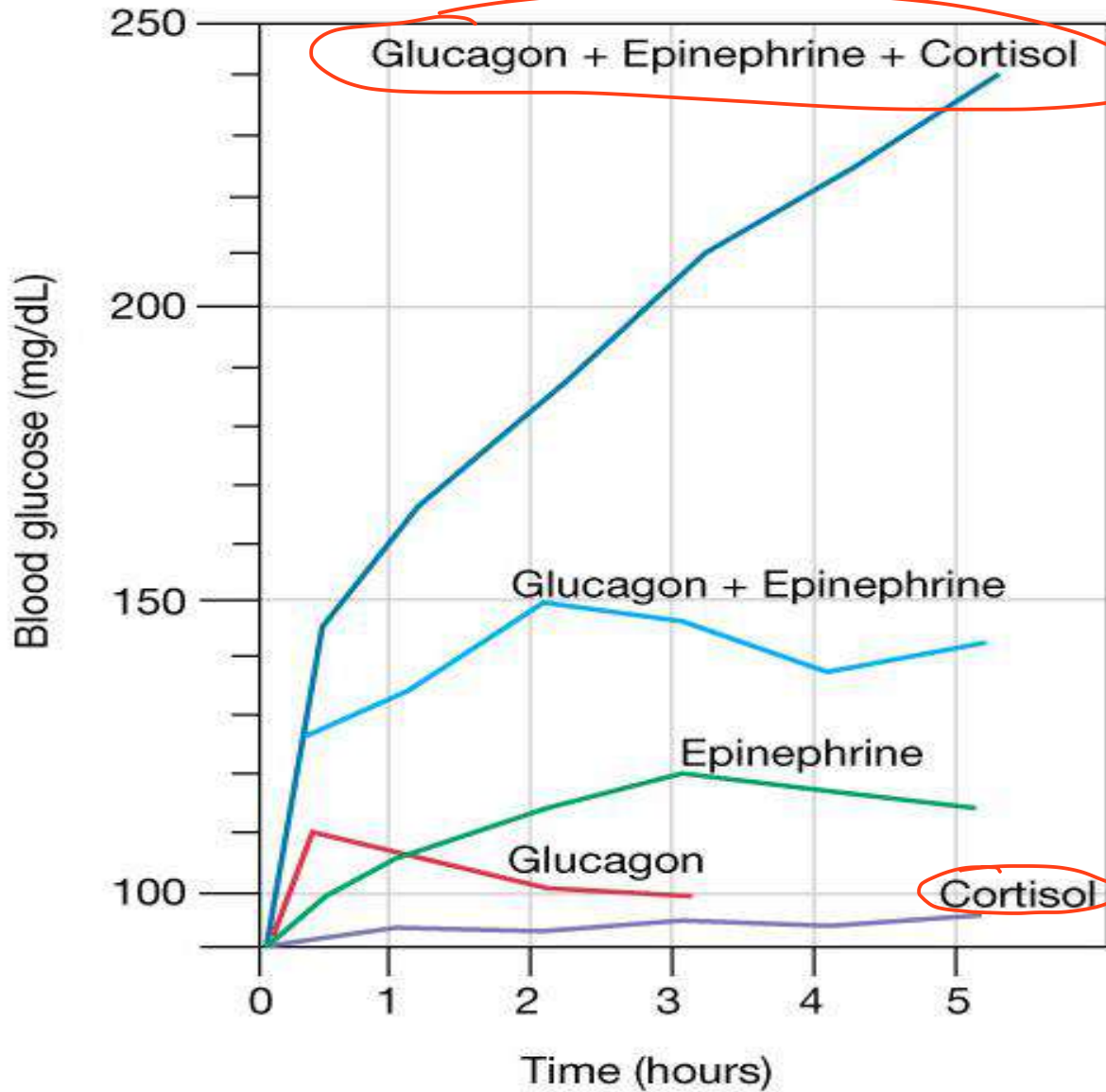
Increase

if I Release gl Cor Ep what will be the Blood Sugar?

→ 19: Additive α

→ One Inhibits the Release of another one.

Synergism



All together
↓
Massive Increase
که با هم بزرگ است
فعلی بزرگ

Permissiveness

The Release of Hormon effect the Release of other Hormones.

One hormone needed for another to exert its effects

Cause the production of Receptors.
→ where Adrenaline should Bind.

if Adrenaline was there
without Receptors
there will be no effects

Example 1: **Thyroid hormones cause expression of adrenergic receptors in bronchiolar smooth muscle**

Example 2: **Estrogen causes expression of progesterone receptors in uterus**

Pathologies

• "NO BAD HORMONES – JUST TOO MUCH OR TOO LITTLE"

- Exogenous medication

- Replaces & exceeds normal
- Cause atrophy of gland

↳ Adrenalitis
Hypersecretion OR Hyposecretion

- Hypersecretion: too much

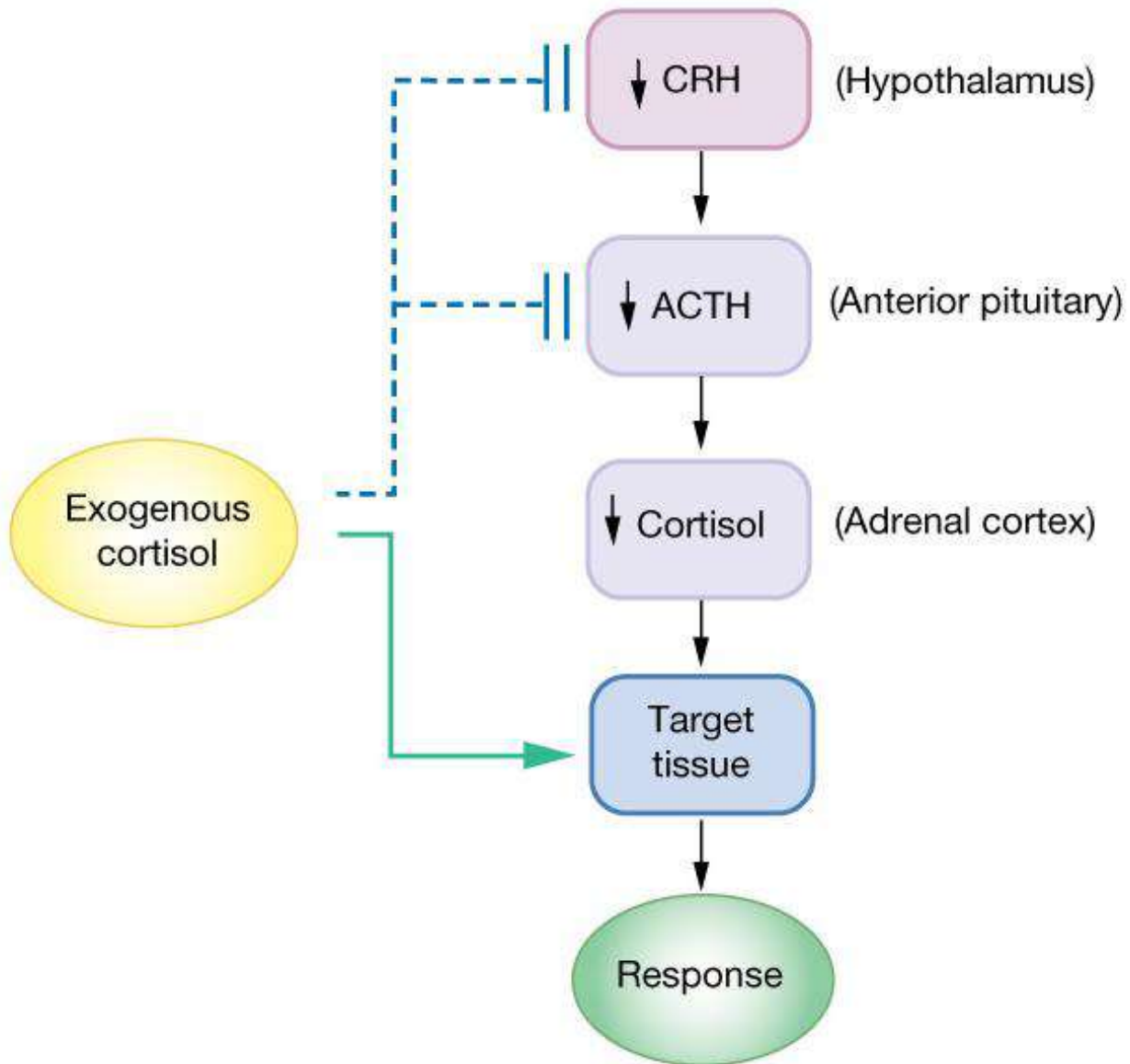
- ✓ Tumors or cancer
- ✓ Grave's disease- thyroxin (T4)

- Hyposecretion: too little

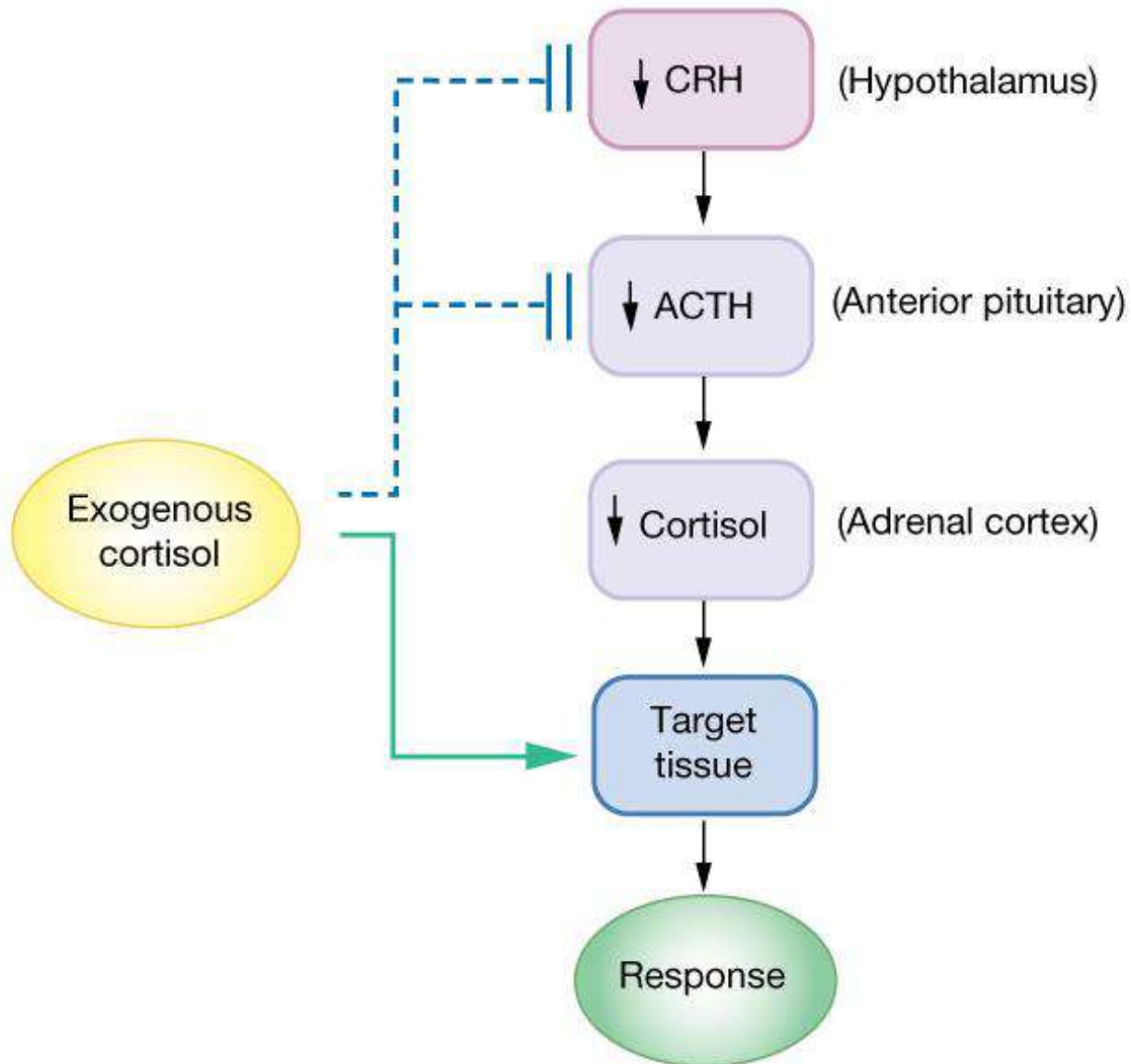
- Goiter – thyroxin (T4) ✓
- Diabetes – insulin ✓

↳ Dead cells → No insulin → Exogenously for insulin

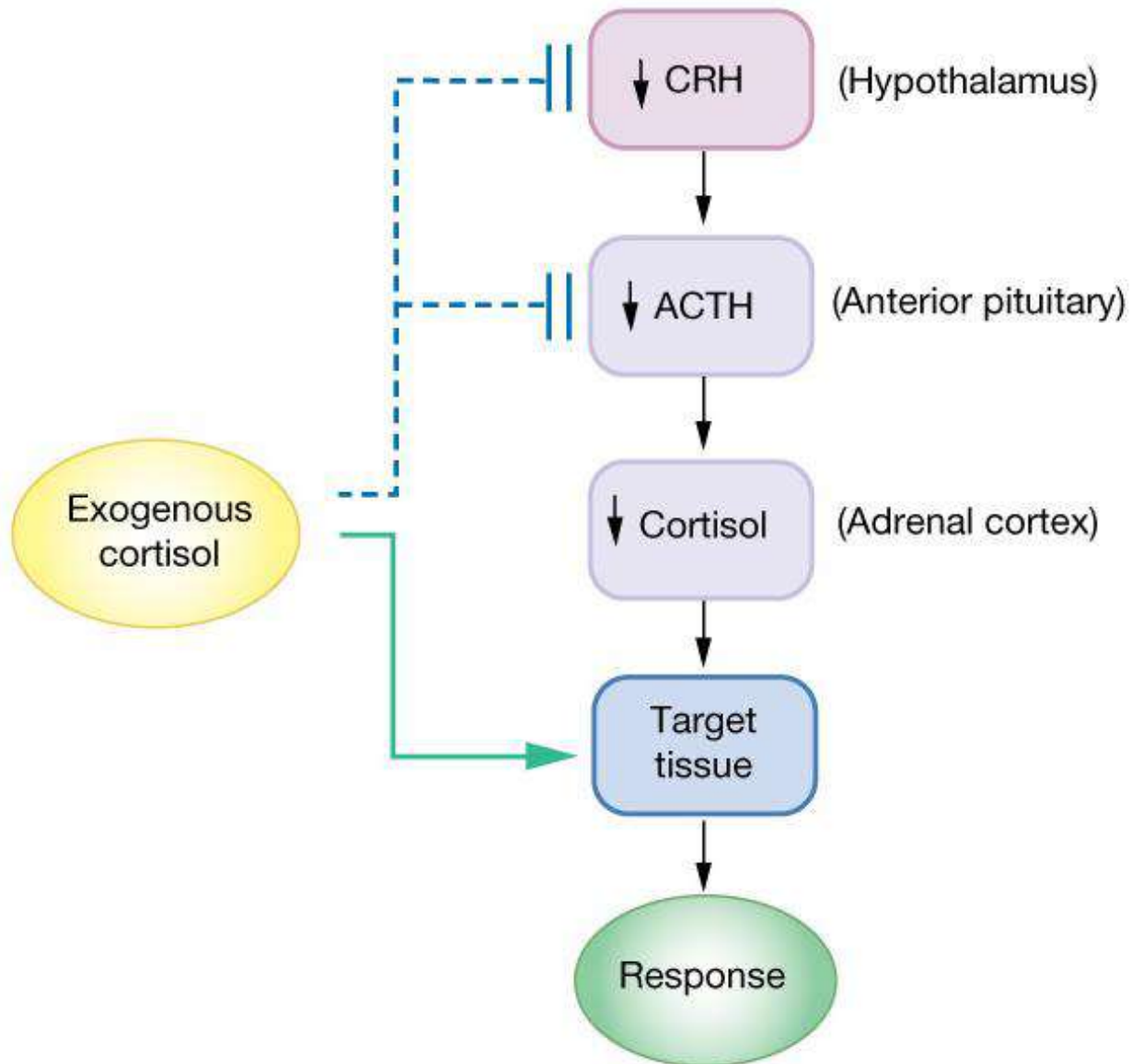
Pathologies: Over or Under Production



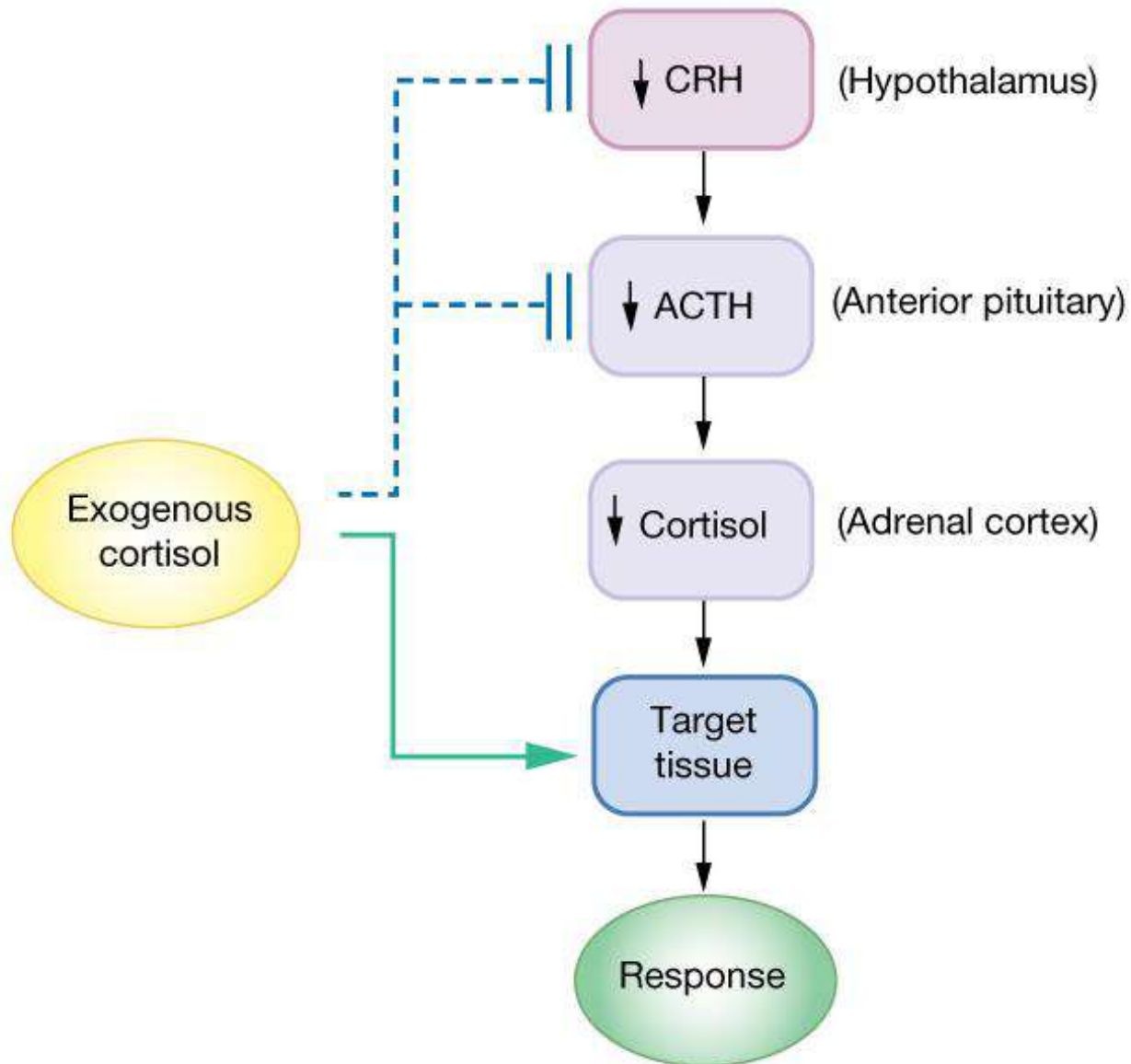
Pathologies: Over or Under Production



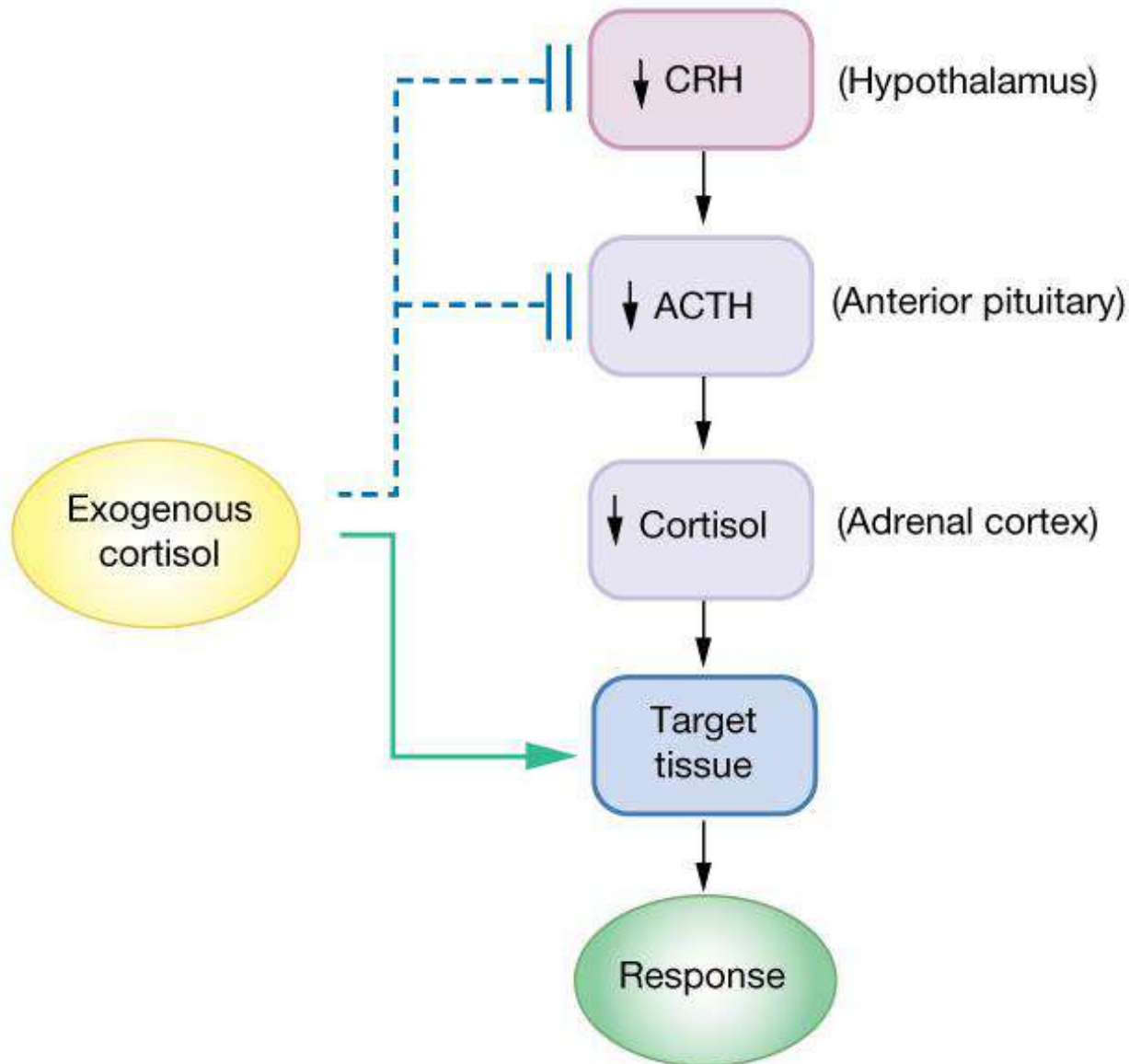
Pathologies: Over or Under Production



Pathologies: Over or Under Production



Pathologies: Over or Under Production



Pathologies: Over or Under Production

