# ENDOCRINOLOGY (INTRODUCTION)

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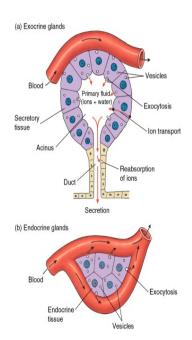
- Endocrine vs exocrine gland
- Chemical messengers
- Hormone
  - Definition
  - Chemical structure
  - Paracrine, autocrine, endocrine, neuroendocrine
  - Transport and clearance
- Mechanism of action
  - Receptors, down-regulation and up-regulation
  - Intracellular signaling
  - Second messenger (cAMP, IP3)

#### A. Exocrine gland

- Ducts
- Secrets enzymes
- Lumen and surfaces

#### B. Endocrine gland

- No ducts
- Secrets Chemical messengers
- Blood stream



## CHEMICAL MESSENGERS

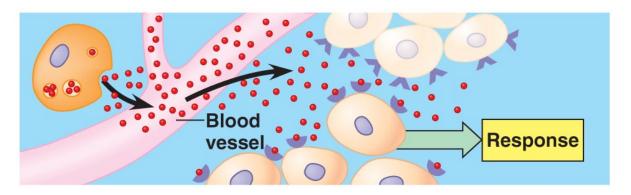
- The activities of cells, tissues and organs are coordinated by chemical messengers
  - Neurotransmitters
  - Endocrine hormones
  - Neuroendocrine hormones
  - Paracrines
  - Autocrines
  - Cytokines

# SMALL GROUP ACTIVITY

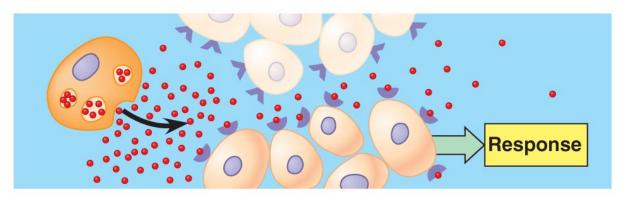
- Divide into 6 groups
- Explain
  - Neurotransmitter
  - Neuroendocrine
  - Endocrine
  - Paracrine
  - Autocrine
  - Cytokines



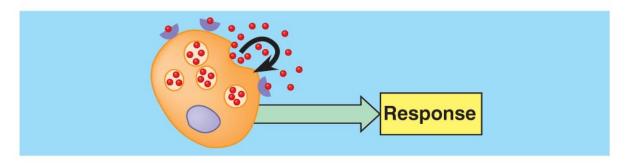




#### (a) Endocrine signaling

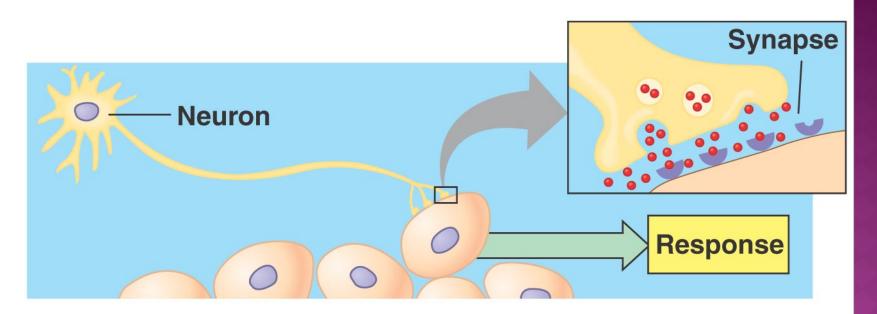


#### (b) Paracrine signaling

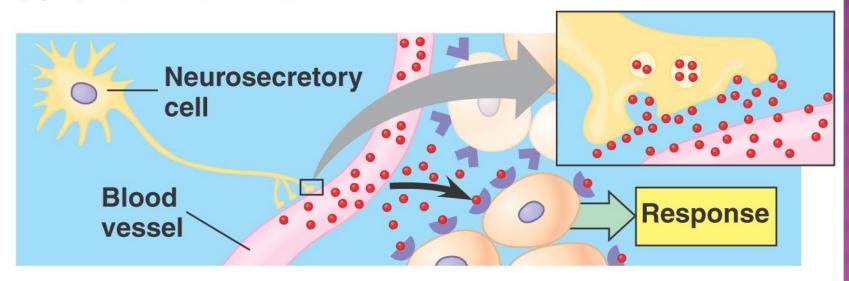


#### (c) Autocrine signaling

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#### (d) Synaptic signaling

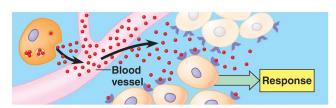


#### (e) Neuroendocrine signaling

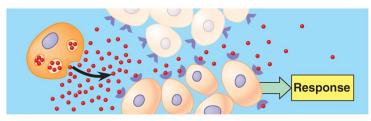
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## CYTOKINES

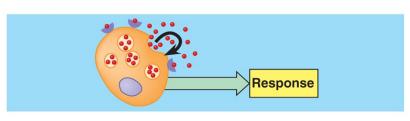
- Peptides (interleukins, lymphokines, adipokines)
- Secreted by cells into extracellular fluid.
- Can function as



(a) Endocrine signaling



(b) Paracrine signaling



(c) Autocrine signaling

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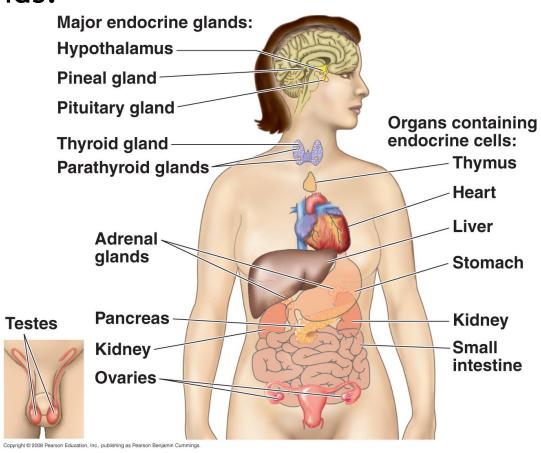
## LARGE GROUP ACTIVITY

List the endocrine glands.

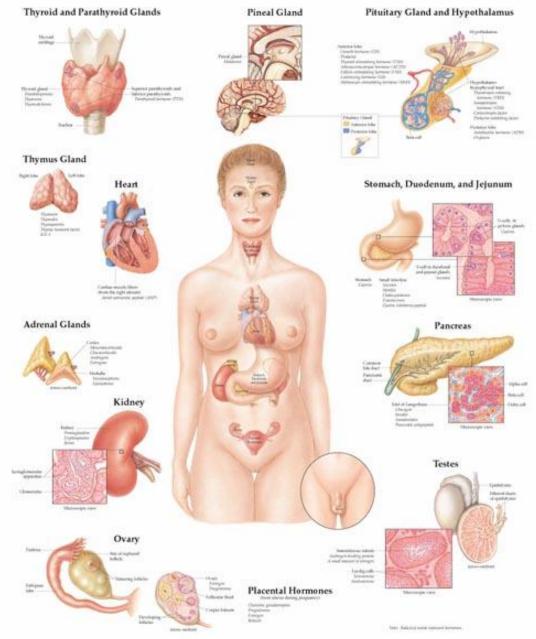


### • Endocrine glands:

- Pituitary
- Thyroid
- Parathyroid
- Adrenal
- Pancreas
- Ovaries
- Testes



#### THE ENDOCRINE SYSTEM



- The multiple hormone systems play a key role in regulating almost all body functions:
  - Metabolism
  - Growth and development
  - Water and electrolyte balance
  - Reproduction
  - Behavior

#### • Definition :

 Hormone is a chemical substance released by group of cells to control the function of other type of cells.

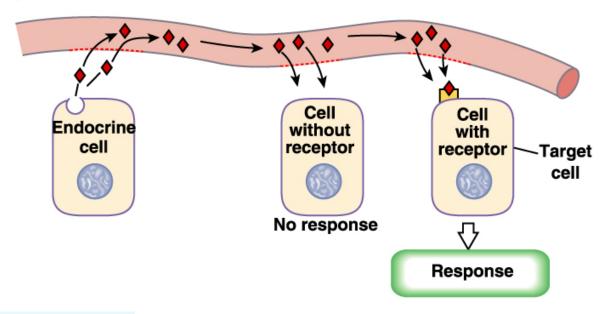
### Types of hormones

- Affect many different types of cells (eg. GH and Thyroxin)
- Affect only specific target cells (eg. ACTH and estrogen)

• What are target cells?

Target cells refer to cells that contain specific receptors (binding sites) for a particular hormone.

#### Hormone

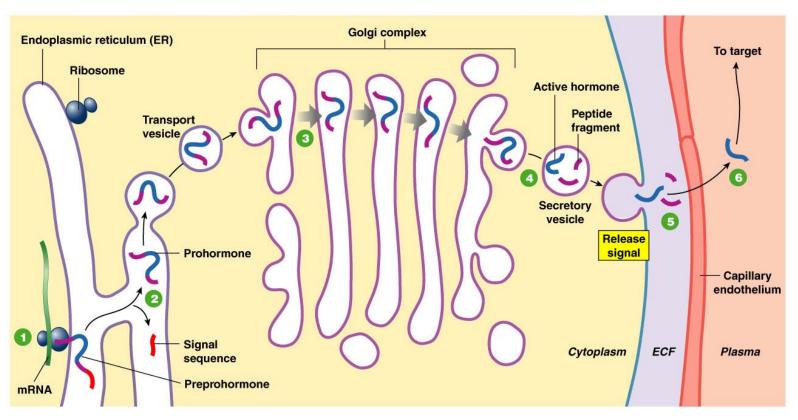


Silverthorn, Human Physiology, 3rd edition Figure 6-1&2

- Chemical structure of hormones
  - Three general classes of hormones:
    - Proteins and polypeptides (anterior and posterior pituitary, pancreas and parathyroid hormones) stored in vesicles until needed
    - Steroids (adrenal cortex, ovarian and testicular hormones) diffuse across the cell membrane
    - Derivatives of amino acid tyrosine (thyroid hormones and catecholamines)

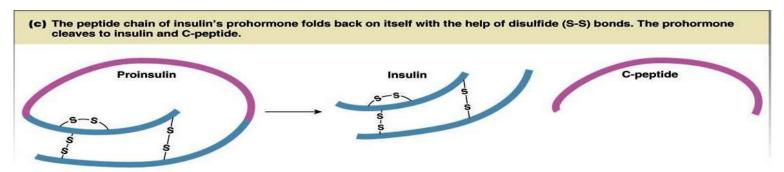
## PEPTIDE (PROTEIN) HORMONES

 Synthesized as preprohormone → posttranslational modification to prohormone → then hormone



# PEPTIDE (PROTEIN) HORMONES

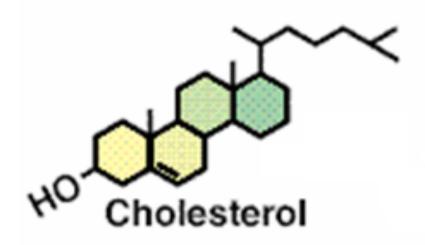
- Example of protein hormone
  - Insulin



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## STEROID HORMONES

- Secreted by gonads, adrenals, placenta
- Derived from cholesterol (lipophilic)
  - Cross membranes (no storage)
- On-demand synthesis (SER)
- Usually Bound to Carrier proteins



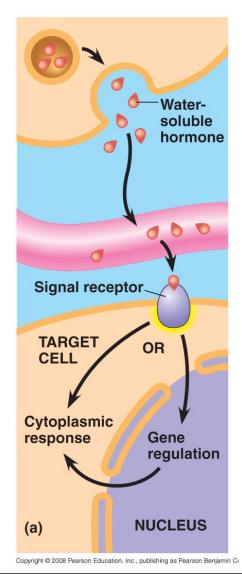
### AMINE HORMONES

- Derived from tyrosine or tryptophan
- 3 groups
  - Tryptophan ⇒ Melatonin
  - Tyrosine ⇒ Catecholamines
     behave like peptide hormones
  - Tyrosine ⇒ Thyroid hormones
     behave like steroid hormones

## TRANSPORT OF HORMONES

- Water soluble hormones- hydrophilic (peptides & catecholamines) dissolved in plasma
- Fat soluble hormones hydrophobic Steroids and thyroid hormones transported bound to plasma proteins (90%),
   binding to proteins helps to
  - Provide reservoirs
  - Slow hormones clearance

# DIFFERENCES BETWEEN WATER AND FAT SOLUBLE HORMONES



Fat-soluble hormone **Transport** protein Signal receptor Cytoplasmic Gene response regulation (b)

ummings.





## HORMONES & RECEPTORS

### • Receptors:

- Hormonal receptors are large proteins
- 2000-100,000 receptors/cell
- Receptors are highly specific for a single hormone

### • Receptor's Location:

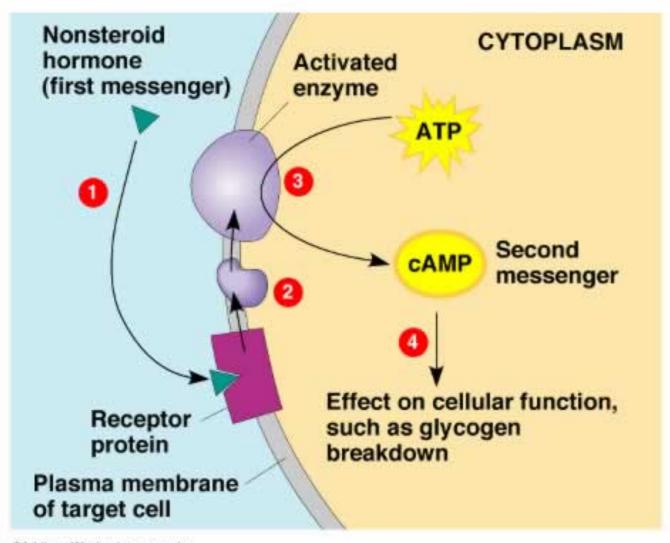
- On the surface of cell membrane (proteins, peptides and catecholamines)
- In the cell cytoplasm (Steroids)
- In the cell nucleus (thyroid hormones)

## MECHANISM OF ACTION OF HORMONES

#### • Mechanism of action :

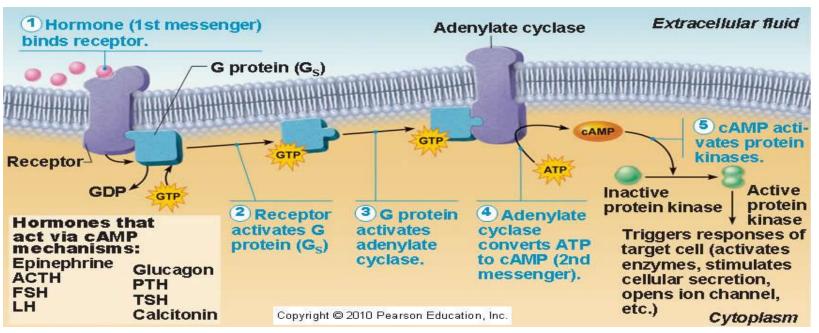
- 1. Hormone-receptor interaction (1st messenger)
- 2. Enzyme activation
- 3. Release of the second messenger
- 4. Effects on cellular function

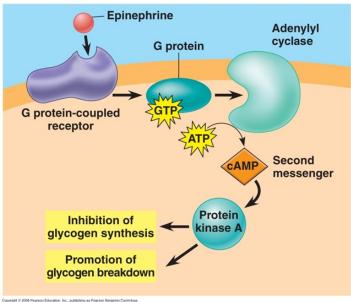
# MECHANISM OF ACTION (PEPTIDES AND PROTEIN HORMONES)



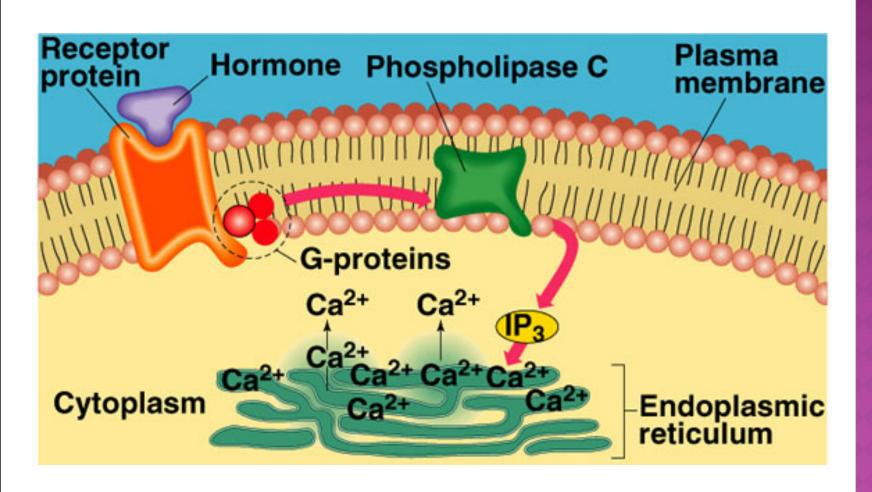
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## SECOND MESSENGER (ADYNYLATE CYCLASE-CAMP)

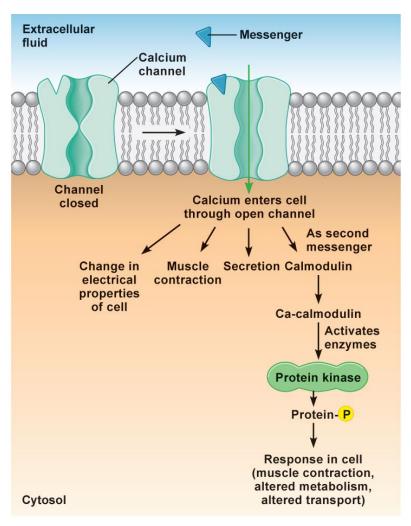




# SECOND MESSENGER (PHOSPHOLIPASE C-IP3)

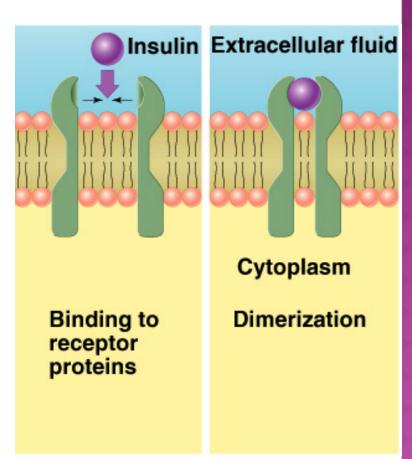


# SECOND MESSENGER (CALCIUM-CALMODULIN COMPLEX)



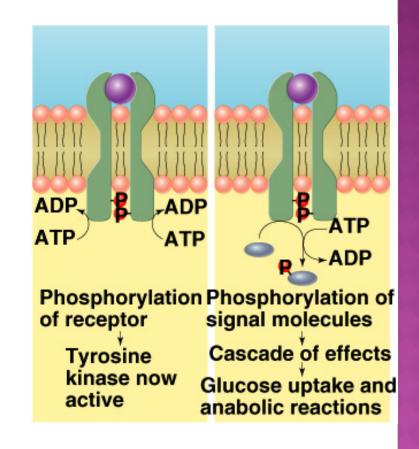
# SECOND MESSENGER (TYROSINE KINASE SYSTEM)

- Is used by insulin & many growth factors to cause cellular effects
- Surface receptor is tyrosine kinase
  - Consists of 2 units that form active dimer when insulin binds

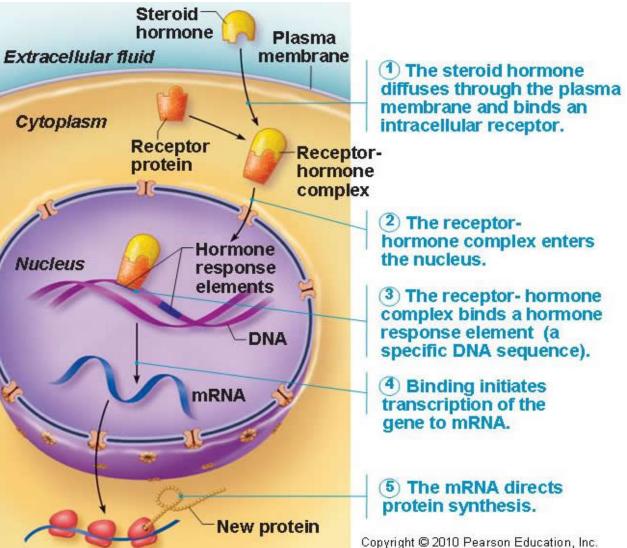


# SECOND MESSENGER (TYROSINE KINASE SYSTEM)

- Activated tyrosine kinase phosphorylates signaling molecules
- Induction of hormone/growth factor effects



# MECHANISM OF ACTION (STEROID HORMONES)



### REGULATION OF HORMONAL RECEPTORS

- Receptors does not remain constant
  - Inactivated or destroyed
  - Reactivated or manufactured
- Downregulation
  - Increase hormone concentration leads to decrease in the number of active receptors
  - Most peptide hormones have <u>pulsatile</u> <u>secretion</u> which prevents downregulation
- Upregulation
  - The hormone induces greater than normal formation of a receptor or intracellular signaling proteins

## CLEARANCE OF HORMONES

- Two factors control the concentration of a hormone in the blood:
  - The rate of its secretion
  - The rate of its removal (metabolic clearance)
- Hormones are cleared by:
  - Metabolic destruction by tissues
  - Binding with tissues
  - Excretion by the liver into bile
  - Excretion by the kidney into urine
- Clearance of protein bound hormones is slower than clearance of peptide hormones

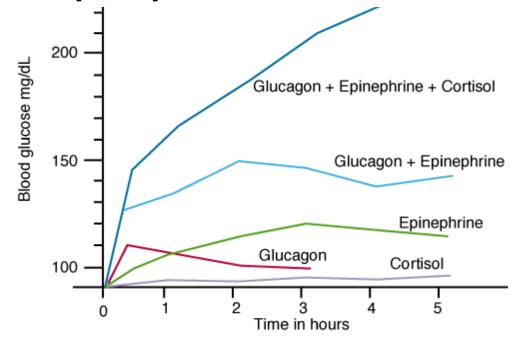
## HORMONE INTERACTIONS

- Multiple hormones can affect a single target simultaneously
- Three types of hormone interactions:
  - 1. Synergism
  - 2. Permissiveness
  - 3. Antagonism

What is Synergism?
What is permissiveness?
What is antagonism?

## SYNERGISM

- Combined action of hormones is more than just additive!
- Example: Blood glucose levels & synergistic effects of glucagon, cortisol and epinephrine



## PERMISSIVENESS

- One hormone allows another hormone to have its full effect
  - Especially during growth
- Example
  - Thyroid hormone have permissive effect on growth hormone action
  - Deficiency of thyroid hormone in infants leads to dwarfism.

## ANTAGONISM

- Antagonistic hormones have opposing physiological actions -
  - Hormone B diminishes the effect of hormone A
- Example
  - Glucagon antagonizes the action of insulin
  - Can you mention another example?

