Second Messenger Systems: G Protein Coupled Receptors

◆ G protein-coupled receptors (GPCRs): class of membrane-bound receptors that initiate signaling		
• Signaling cascade: chemical messe	ngers linked in Cyclic AMP () is a common example.
1. Hormone binds to: transmits signal across membrane.	4. Adenylate cyclase converts ATP to	5. cAMP binds to the enzyme 6. Kinase phosphorylates proteins; triggers response.
2. Receptor (GPCR)	3. G proteins diffuses along	
activates:	membrane, binds to, and activates	
GTP replaces GDP on G protein.	the enzyme	
Camp Kinase	Holding Really Great Activities at	t cAMP K inase.

EXAMPLE

The steps of the cascade using cAMP are given below. Write the letter for the appropriate step in the blanks below to put the pathway in order.

a. Protein kinase is activated.

c. G protein is activated.

b. ATP is converted to cAMP.

d. Adenylate cyclase is activated.

Hormone binds extracellular receptor → _____ → ___ → Cellular Response.

PRACTICE

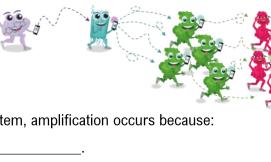
What is the role of G proteins in GPCR signaling?

- a) Catalyzing DNA synthesis.
- b) Initiating apoptosis.
- c) Transcription of mRNA.
- d) Activating downstream effectors.

Amplification

◆ Recall: signaling cascade: series of linked _____ messengers.

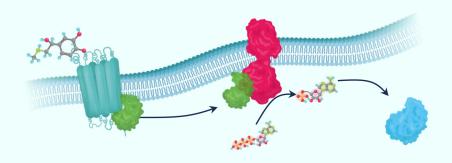
◆ Amplification: when one molecule _____ the signal to multiple molecules - ___ signal.



- ◆ In cAMP secondary messenger system, amplification occurs because:
 - 1. GPCR will activate multiple ______.
 - 2. Adenylate cyclase will produce many _____ molecules.
 - 3. Protein kinase will phosphorylate many _____.

EXAMPLE

The image below shows a signaling cascade using cAMP as a second messenger. Circle the places on the image where amplification can happen.



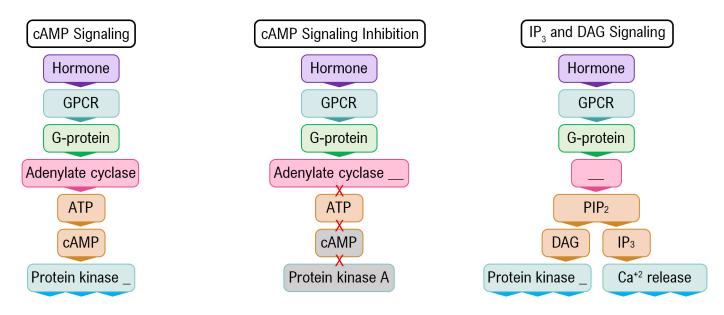
PRACTICE

Hormones can bring about substantial physiological changes at very low concentrations. How does this relate to the concept of second messenger systems?

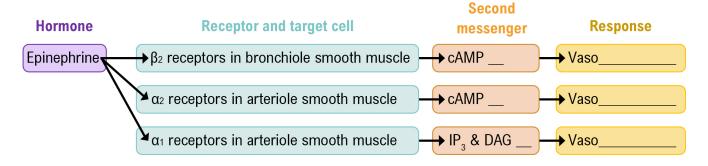
- a) Second messenger systems act more quickly, creating a larger cellular response.
- b) Second messenger systems modify the DNA directly, allowing the hormone to make an impact even at low concentrations.
- c) Second messenger systems prevent hormones from being degraded so they can exert their effect for a longer duration.
- d) Second messenger systems amplify the original hormone signal, allowing it to work at low concentrations.

Secondary Messenger Systems: cAMP, DAG, & IP,

- ◆ Hormones may induce different signaling cascades depending on:
 - The presence of a specific _____. ► The secondary messenger involved. ► The activity of the _____.



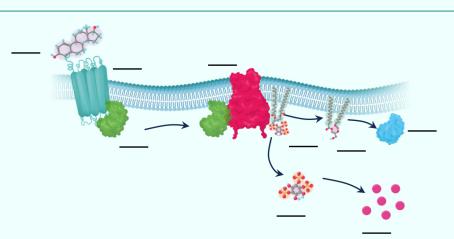
◆ One hormone; many responses:



EXAMPLE

The image below shows parts of a signaling cascade that uses IP₃ and DAG as secondary messengers. Label each component and circle the places on the image that are different from a signaling cascade that uses cAMP as a secondary messenger.

- a. Phospholipase C
- b. G protein
- c. PIP₂
- d. Kinase
- e. Hormone
- f. Receptor
- g. DAG & IP₃
- h. Calcium Ions



PRACTICE

You are studying how a hormone affects a cell and find that when oxytocin binds to the receptor, the intracellular Ca⁺² concentration increases. Based on this information, what could you logically conclude about this cell and hormone?

- a) The cell uses adenylate cyclase as a second messenger.
- b)Oxytocin directly interacts with the DNA of the cell.
- c) The cell uses $\ensuremath{\text{IP}_{\scriptscriptstyle 3}}$ and DAG as secondary messengers.
- d) Both A & B are correct.

PRACTICE

cAMP, IP₃, and DAG are all molecules that are used as secondary messengers. Which statement below correctly identifies a difference between the pathways in which they are found?

- a) cAMP secondary messenger systems result in the activation of a kinase, while systems that use IP₃ and DAG do not.
- b) cAMP and IP₃ are part of the same signaling cascade, while DAG is found in different cascades.
- c) Adenylate cyclase produces cAMP while phospholipase C produces $\ensuremath{\text{IP}_{\scriptscriptstyle 3}}$ and DAG.
- d) Both A & C are correct.