



## CHAPTER 6

# An Introduction to Metabolism

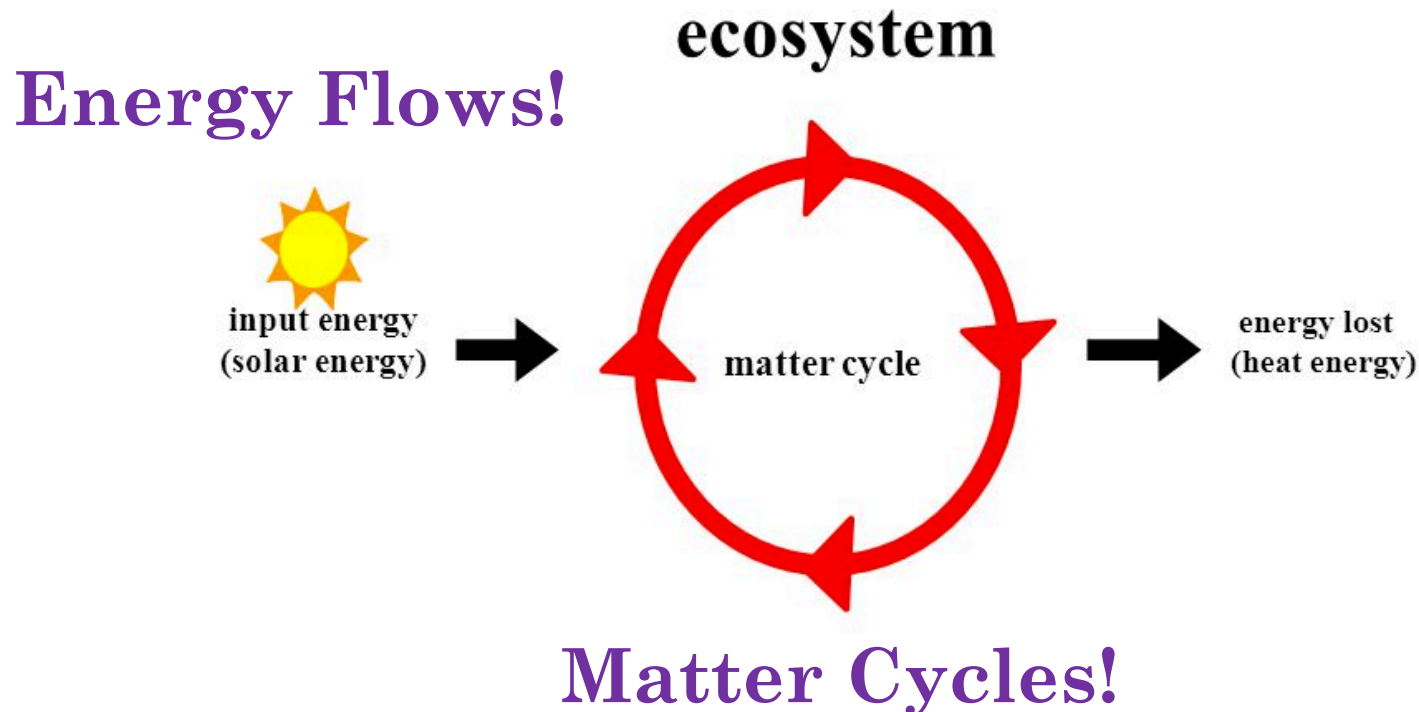
# WHAT YOU NEED TO KNOW:

- Exergonic reactions release free energy ( $\Delta G$  is negative); endergonic reactions store free energy ( $\Delta G$  is positive).
- ATP powers cellular work by coupling exergonic reactions to endergonic reactions.
- Enzymes work by lowering the energy of activation.
- The catalytic cycle of an enzyme that results in the production of a final product.
- Enzymes are specific in the reactions they catalyze because of the molecular shape of their active site.
- Factors that change the shape of the active site of enzymes and how they influence enzyme activity.
- How feedback inhibition is used to maintain appropriate levels of enzymes and enzyme products in a pathway.

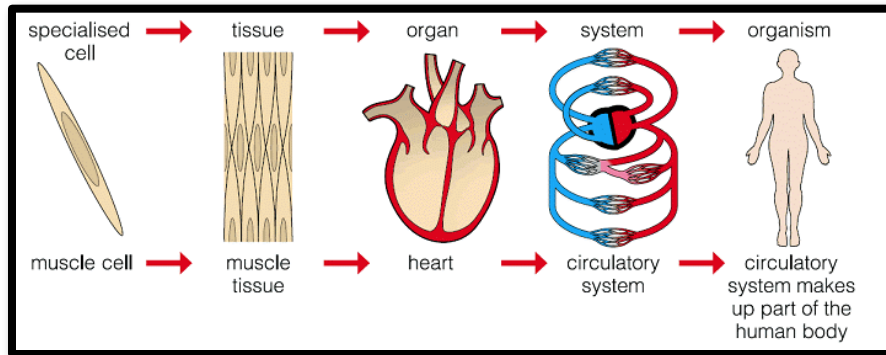


# ENERGY DYNAMICS

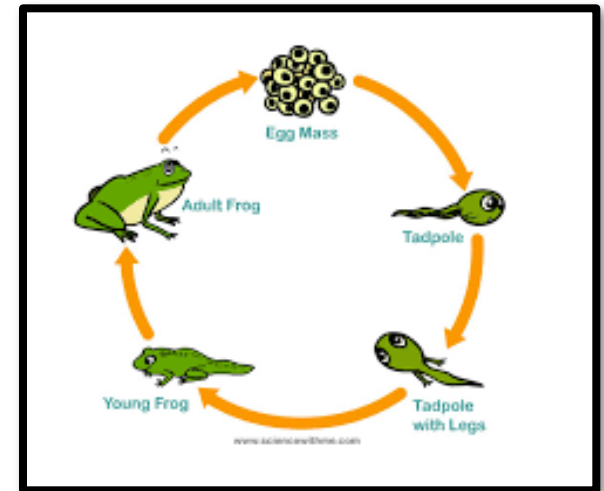
The highly complex organization of living systems requires constant input of energy and the exchange of macromolecules.



# ORGANISMS USE **ENERGY** TO:



Maintain Organization

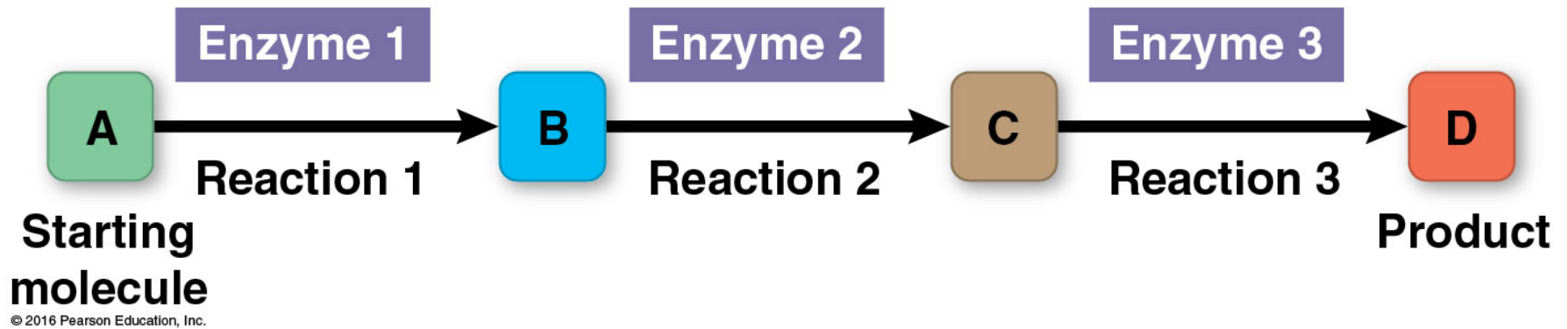


Reproduce



Grow





**Metabolism** is the totality of an organism's chemical reactions

- Manage the materials and energy resources of a cell
- **Metabolic rate** = total amount of energy an animal uses in a unit of time
- In general, the *smaller* the organism, the *higher* the metabolic rate



- **Catabolic pathways** release energy by breaking down complex molecules into simpler compounds
  - Eg. digestive enzymes break down food → release energy
- **Anabolic pathways** consume energy to build complex molecules from simpler ones
  - Eg. amino acids link to form muscle protein



# ENERGY = CAPACITY TO DO WORK

- Kinetic energy (KE): energy associated with motion
  - *Heat* (thermal energy) is KE associated with random movement of atoms or molecules
- Potential energy (PE): stored energy as a result of its position or structure
  - *Chemical energy* is PE available for release in a chemical reaction
- Energy can be **converted** from one form to another
  - Eg. chemical → mechanical → electrical





**A diver has more potential energy on the platform.**

**Diving converts potential energy to kinetic energy.**



**Climbing up converts the kinetic energy of muscle movement to potential energy.**

**A diver has less potential energy in the water.**





Thermodynamics is the study of energy transformations that occur in matter

- **Closed** system: isolated from its surroundings (eg. liquid in a thermos)
- **Open** system: energy and matter can be transferred between the system and its surroundings
- **Organisms = Open Systems**
  - A **net gain in energy** results in **energy storage** or the **growth** of an organism
  - A **net loss of energy** results in **loss of mass**, and/or **death** of an organism



# *THE FIRST LAW OF THERMODYNAMICS (CONSERVATION OF ENERGY)*

- The energy of the universe is constant
  - Energy can be transferred and transformed
  - Energy cannot be created or destroyed



**(a) First law of thermodynamics**

# THE SECOND LAW OF THERMODYNAMICS

- Every energy transfer or transformation **increases the entropy** (disorder) of the universe
- During every energy transfer or transformation, some energy is *unusable*, often lost as **heat**



(b) Second law of thermodynamics

# THERMOREGULATION

- Maintain an internal temperature within a tolerable range
- Endothermic animals use thermal energy generated by metabolism to maintain homeostatic body temperatures (birds and mammals)
- Ectothermic animals gain heat from external sources (invertebrates, fishes, amphibians, and nonavian reptiles)





**(a) A walrus, an endotherm**

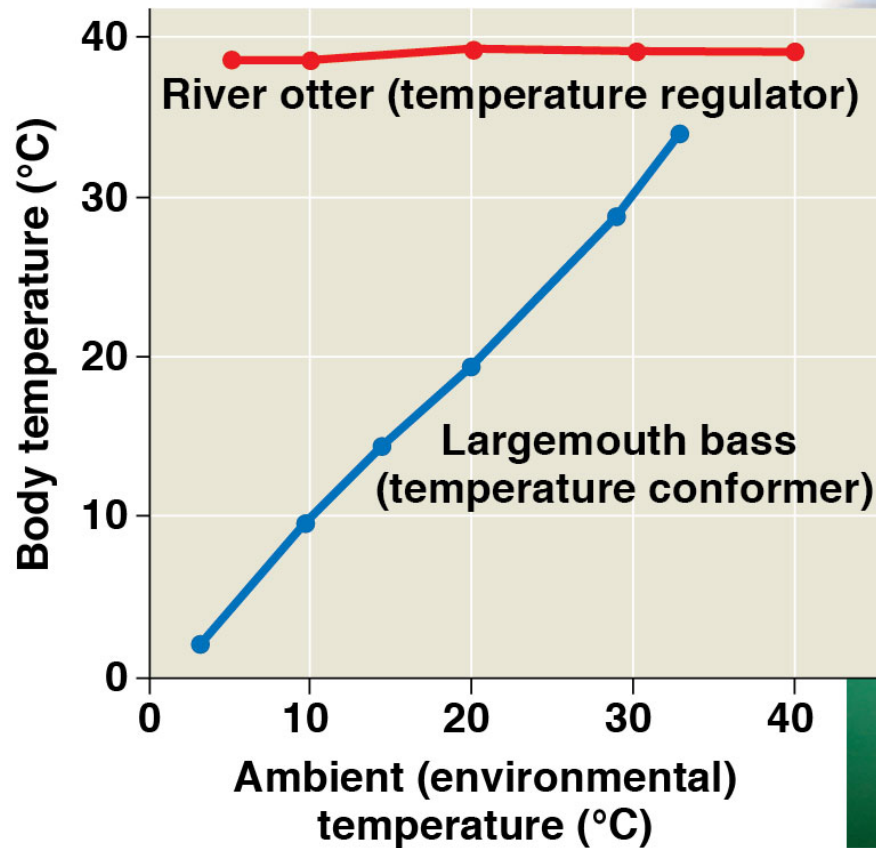


**(b) A lizard, an ectotherm**





# ENDOTHERMS VS. ECTOTHERMS





- **Free energy**: part of a system's energy available to perform work
  - $\Delta G$  = change in free energy
- **Exergonic reaction**: energy is released
  - Spontaneous reaction
  - $\Delta G < 0$
- **Endergonic reaction**: energy is required
  - Absorb free energy
  - $\Delta G > 0$

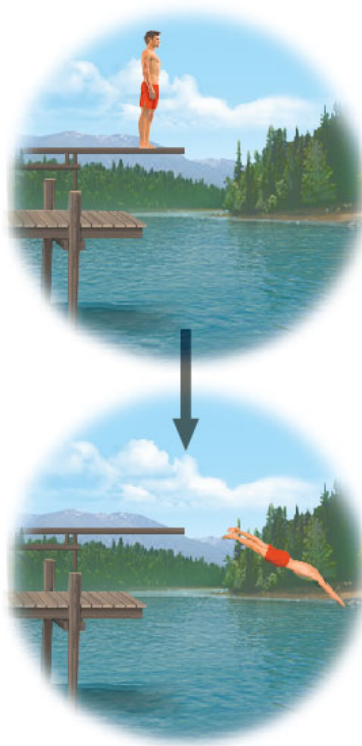


- More free energy (higher  $G$ )
- Less stable
- Greater work capacity

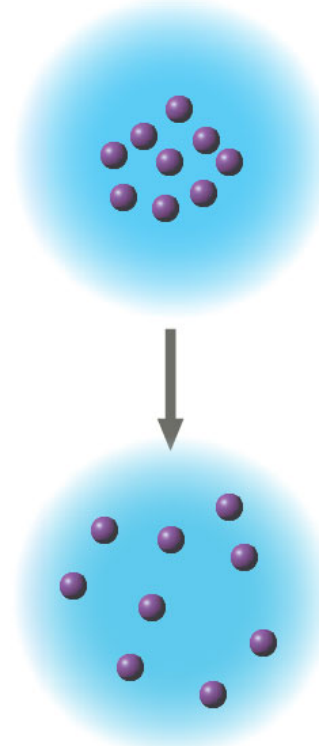
In a spontaneous change

- The free energy of the system decreases ( $\Delta G < 0$ )
- The system becomes more stable
- The released free energy can be harnessed to do work

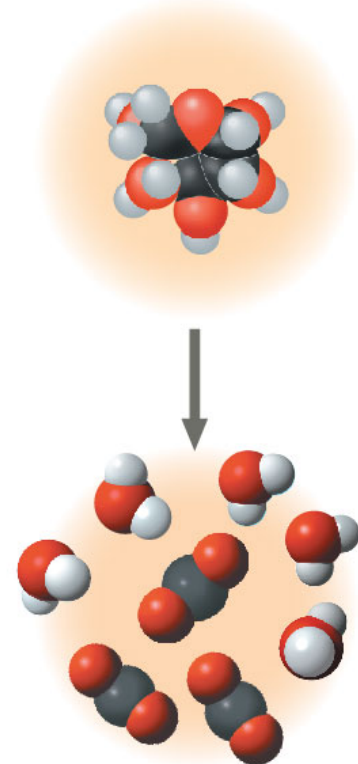
- Less free energy (lower  $G$ )
- More stable
- Less work capacity



(a) Gravitational motion

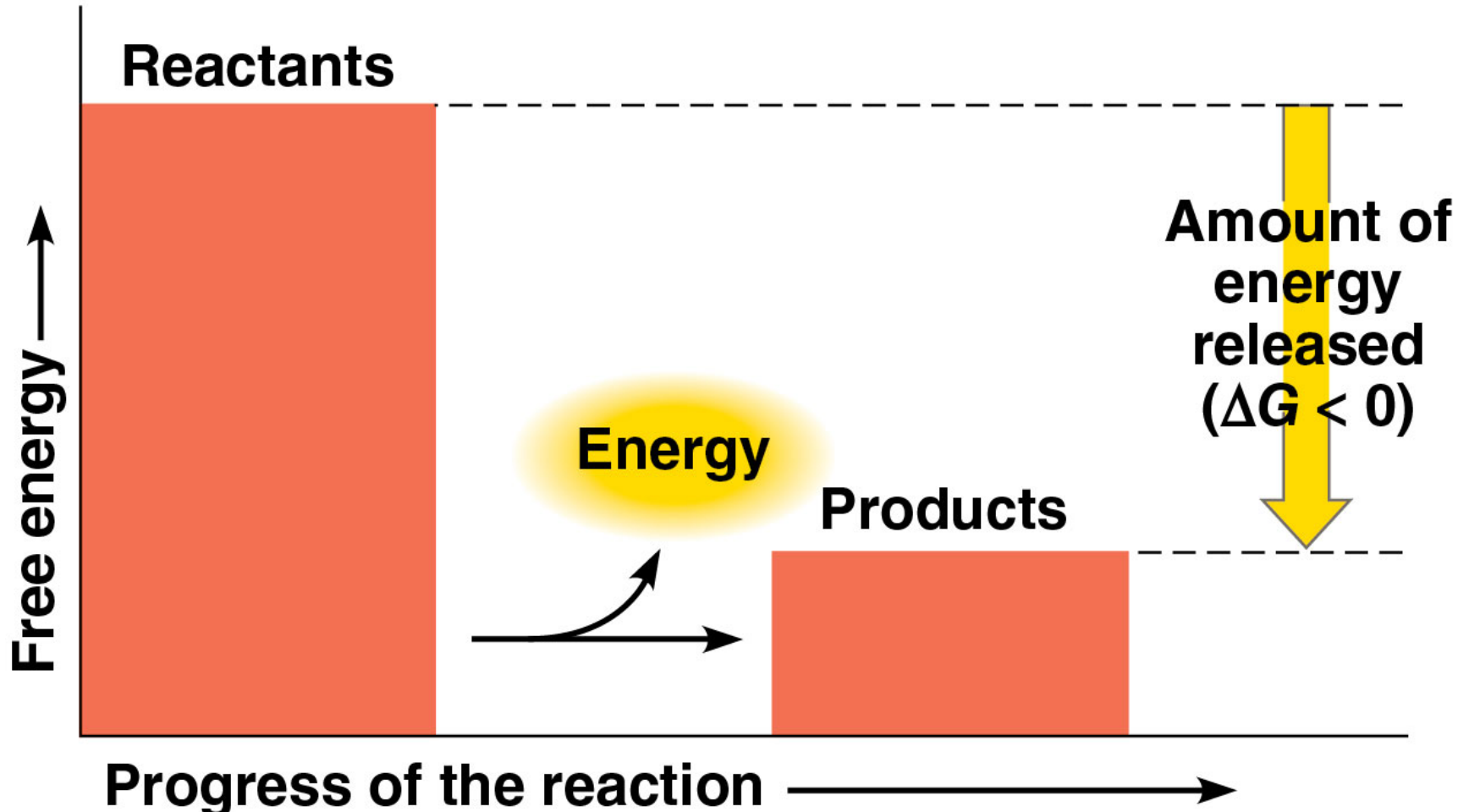


(b) Diffusion

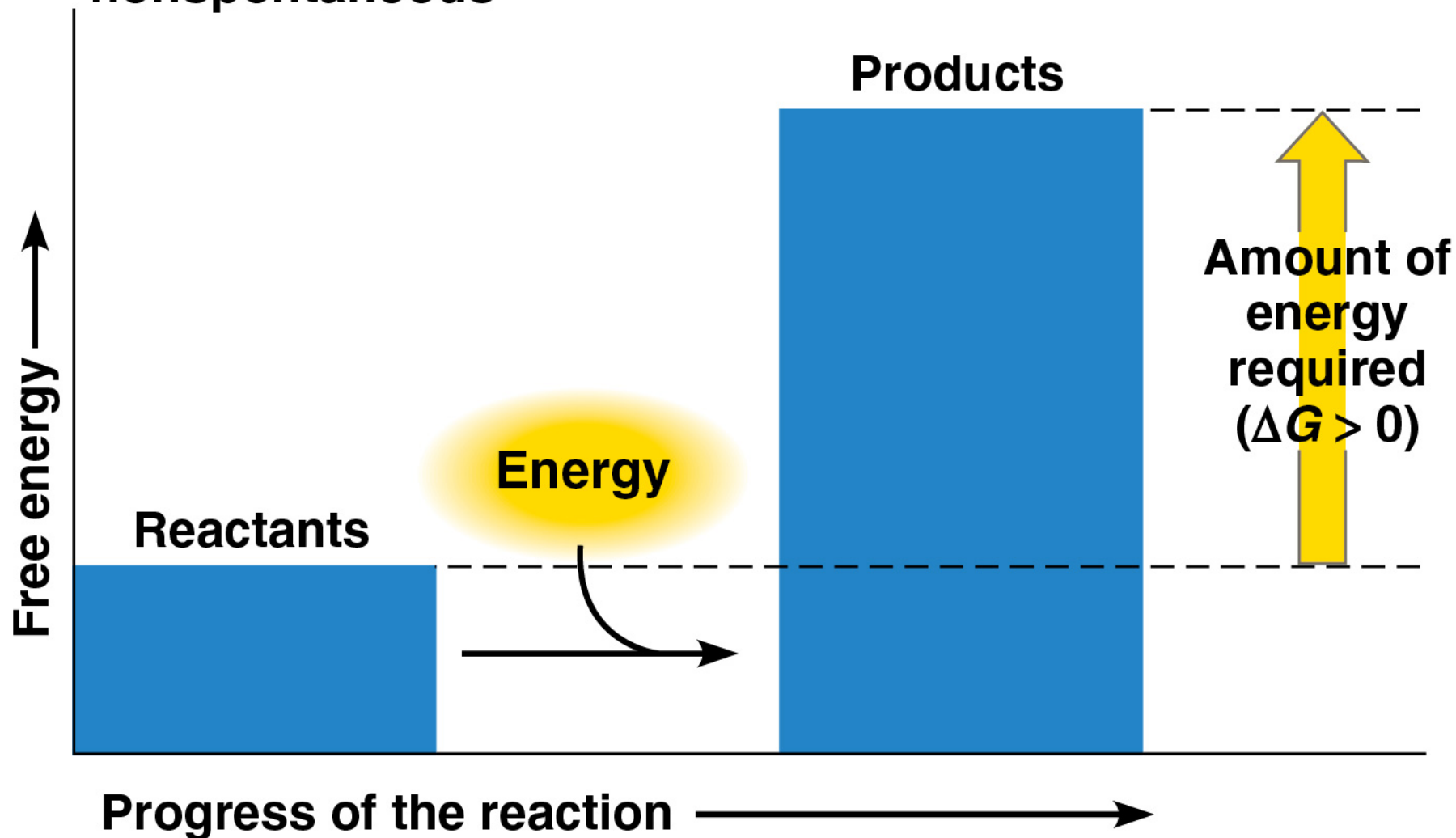


(c) Chemical reaction

**(a) Exergonic reaction: energy released, spontaneous**



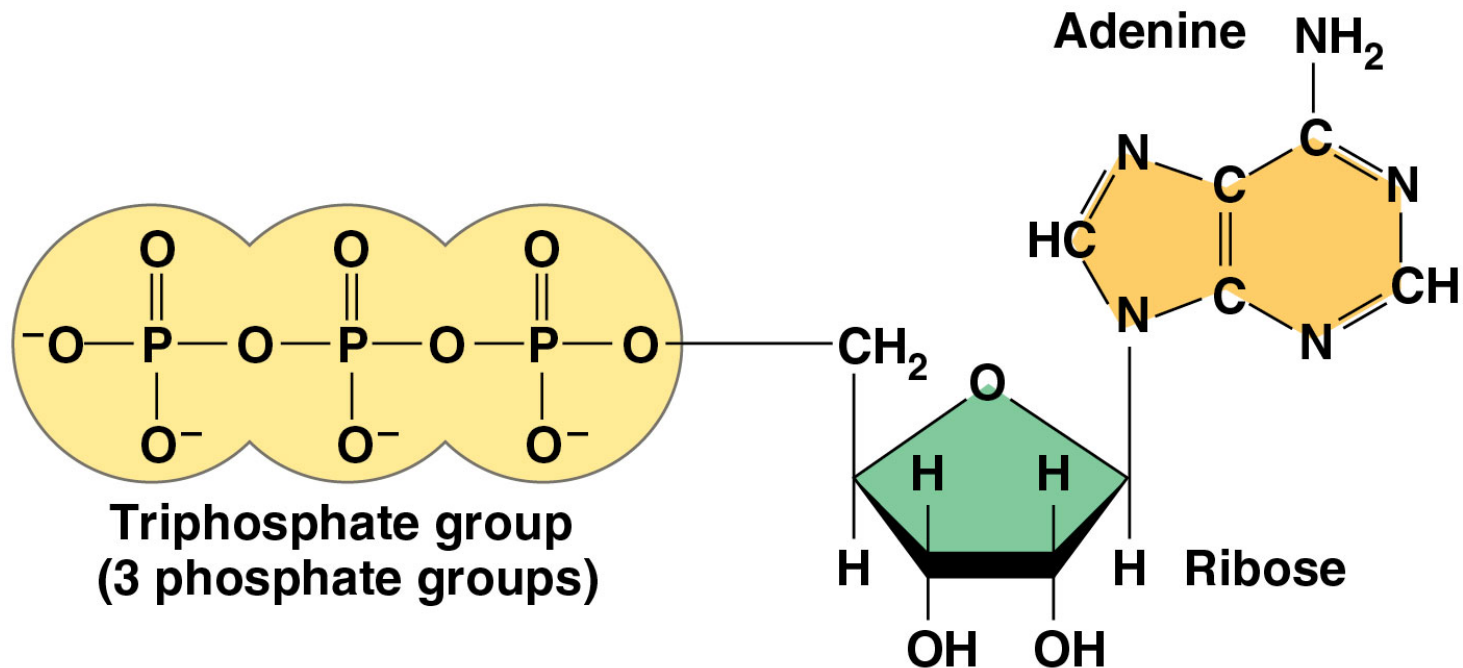
**(b) Endergonic reaction: energy required, nonspontaneous**



- A living cell is NOT at equilibrium
  - Constant flow of materials in/out of cell
- A cell does three main kinds of work:
  1. Mechanical
  2. Transport
  3. Chemical
- Cells manage energy resources to do work by energy coupling: using an **exergonic** process to drive an **endergonic** one



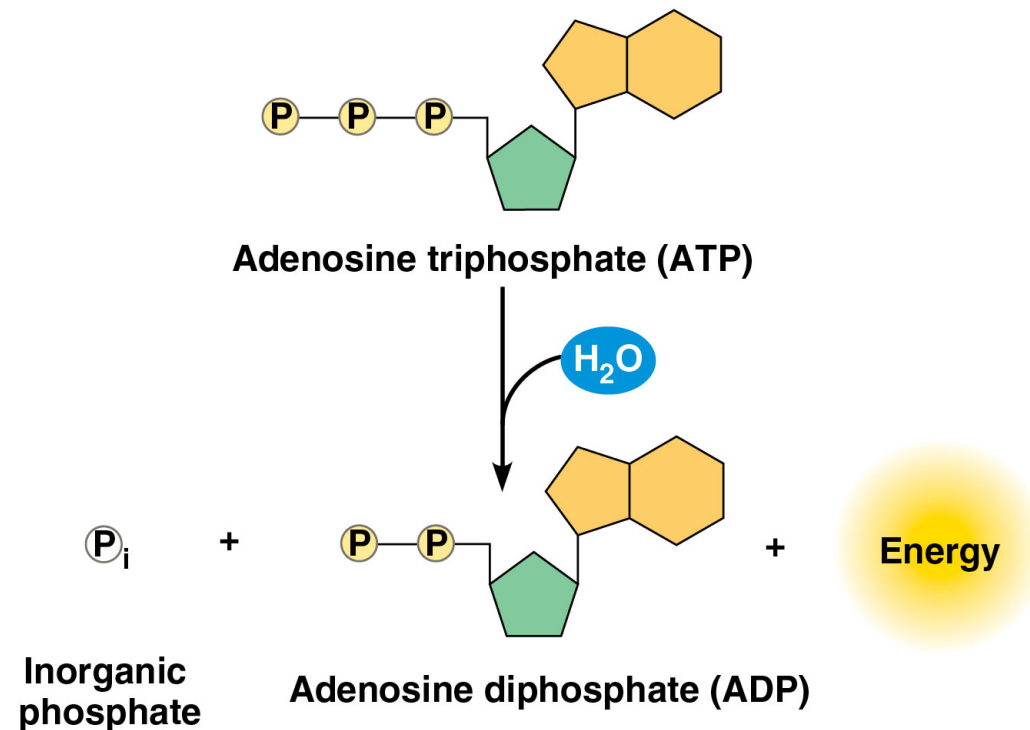
- **ATP** (adenosine triphosphate) is the cell's main energy source in energy coupling
- ATP = adenine + ribose + 3 phosphates



(a) The structure of ATP



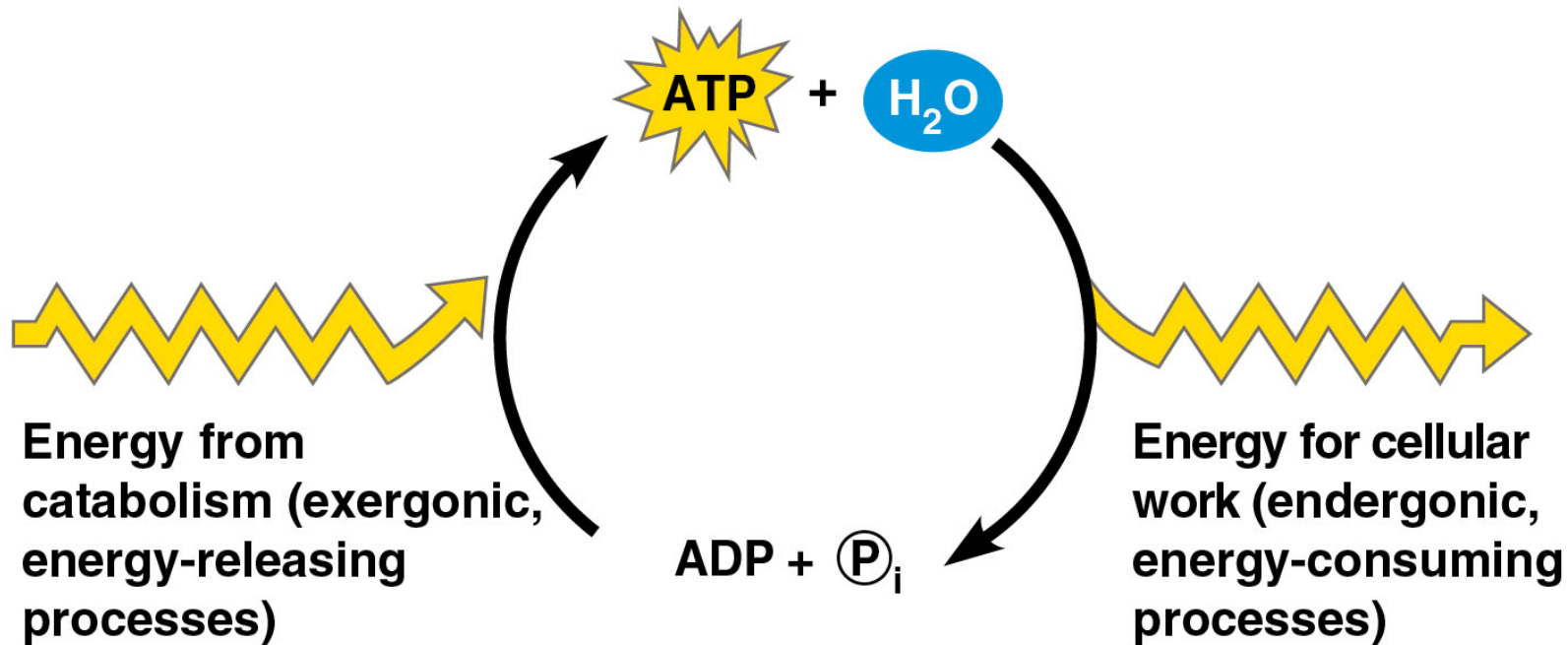
- When the bonds between the phosphate groups are broken by **hydrolysis** → ***Energy is released***
- This release of energy comes from the **chemical change to a state of lower free energy**, not in the phosphate bonds themselves

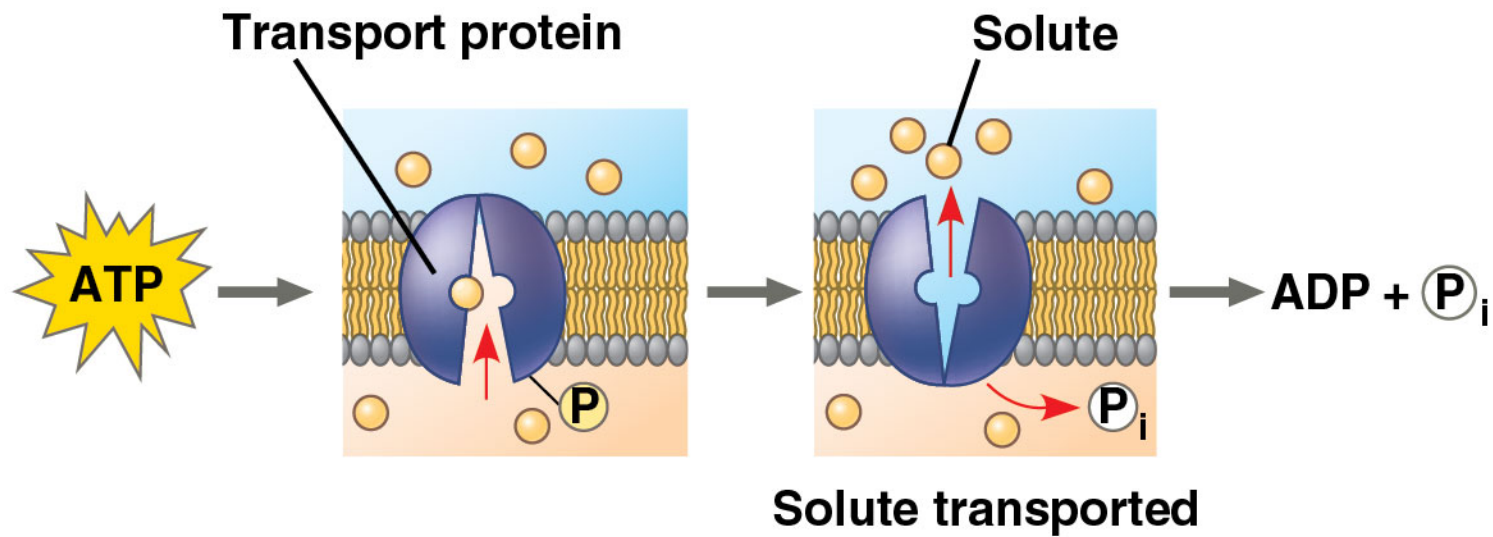


(b) The hydrolysis of ATP

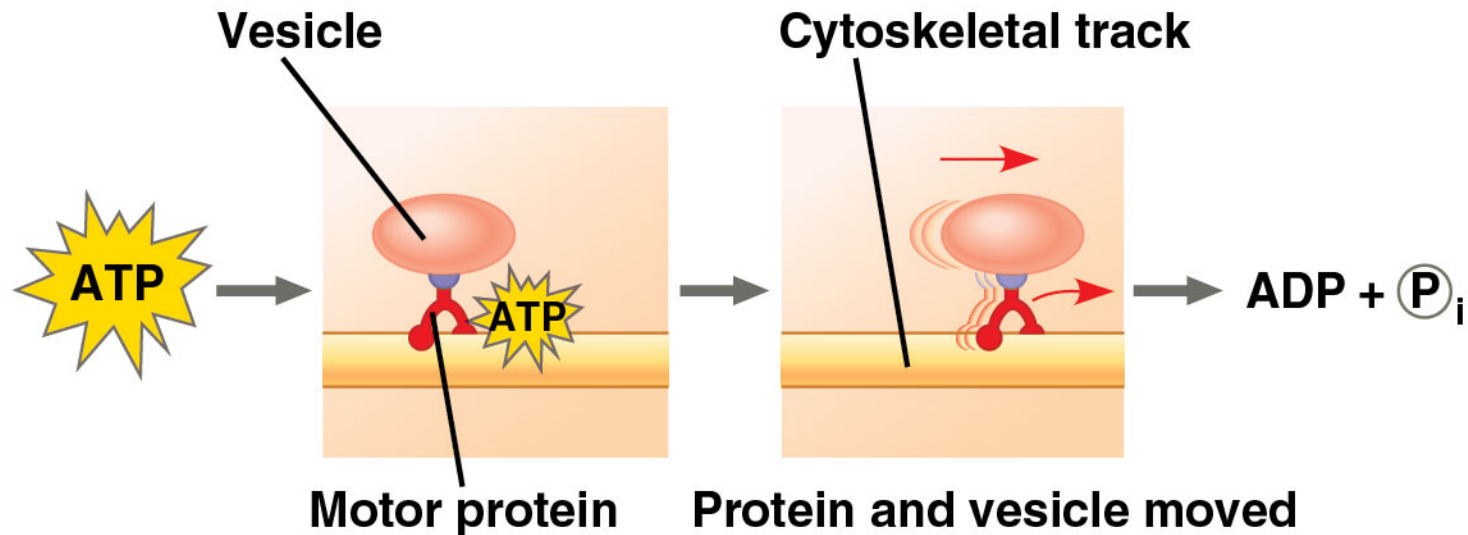
# HOW ATP PERFORMS WORK

- *Exergonic* release of  $P_i$  is used to do the *endergonic* work of cell
- When ATP is hydrolyzed, it becomes ADP (adenosine diphosphate)



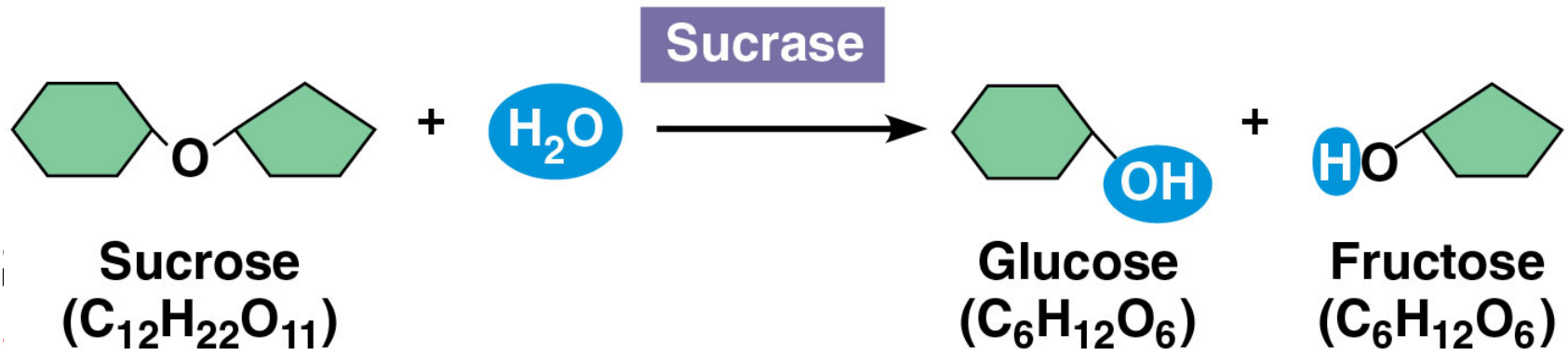


**(a) Transport work: ATP phosphorylates transport proteins.**



**(b) Mechanical work: ATP binds noncovalently to motor proteins and then is hydrolyzed.**

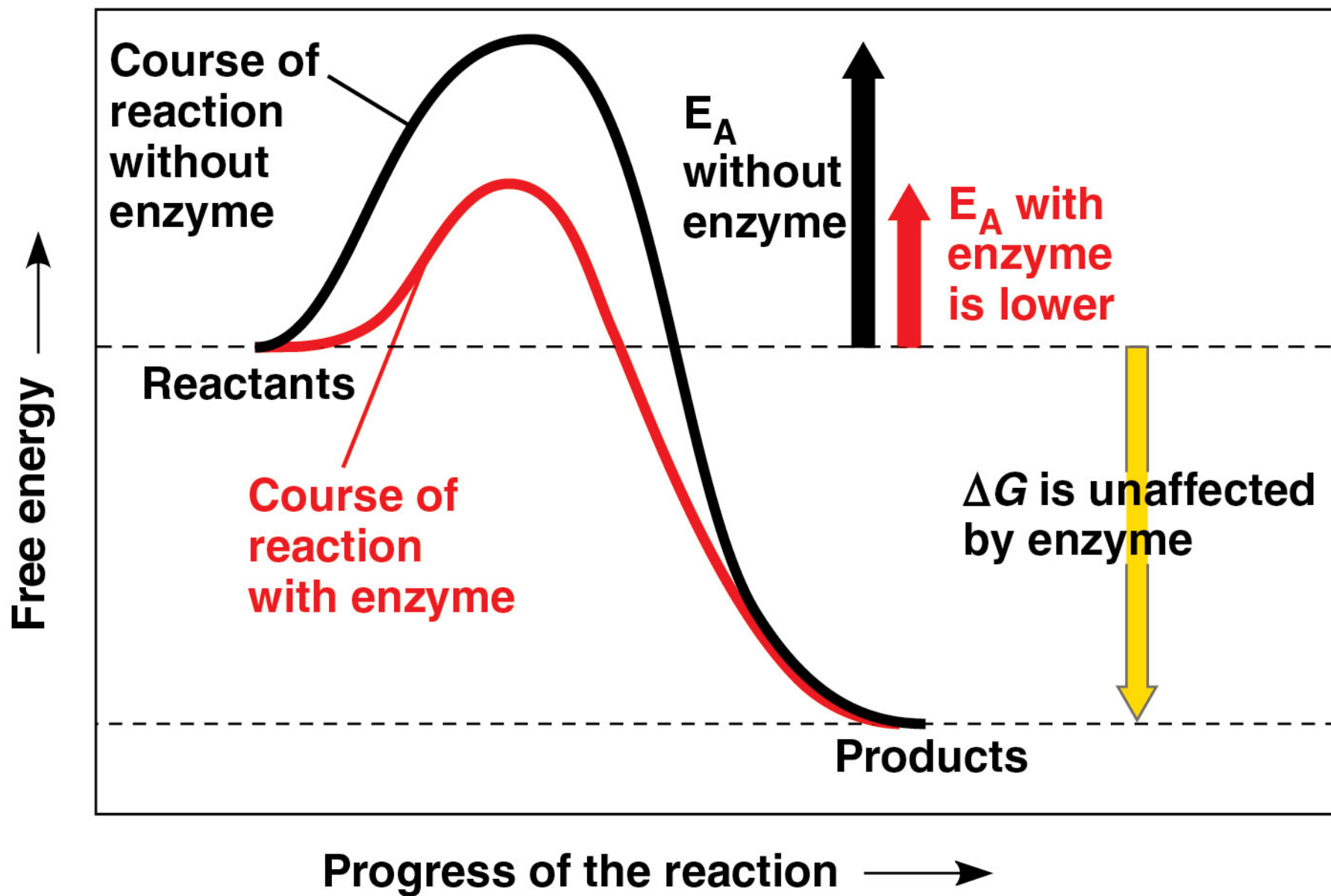
- **Catalyst**: substance that can change the rate of a reaction without being altered in the process
- **Enzyme** = biological catalyst



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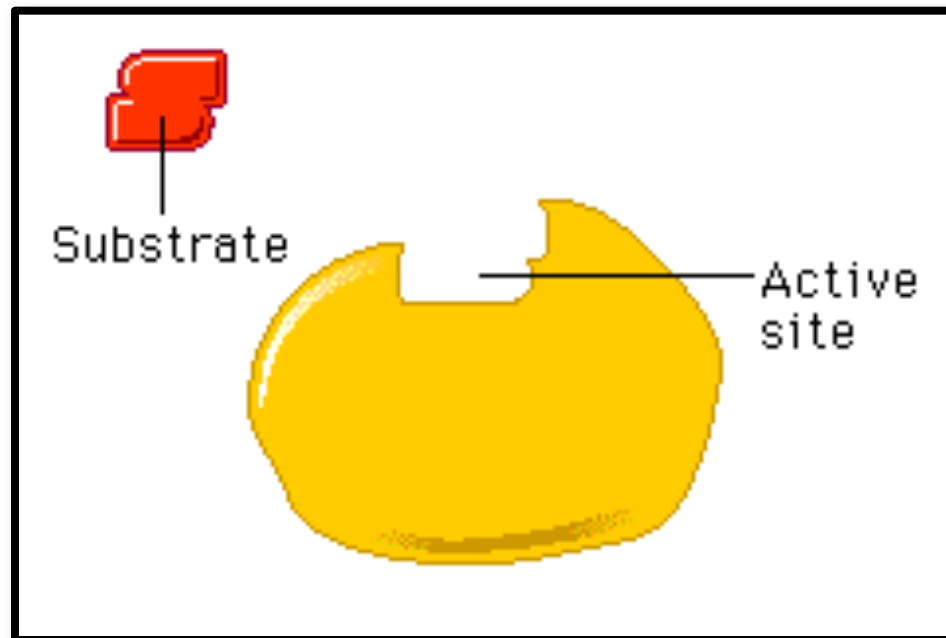
breaking bonds)





# SUBSTRATE SPECIFICITY OF ENZYMES

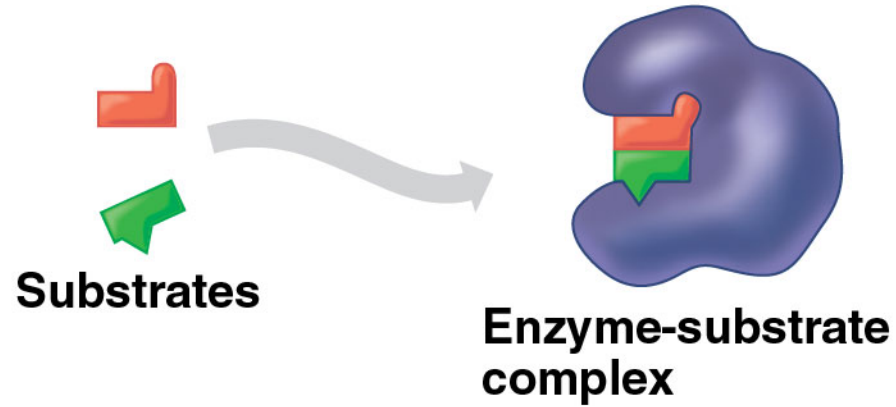
- The reactant that an enzyme acts on is called the enzyme's **substrate**
- The enzyme binds to its substrate, forming an **enzyme-substrate complex**
- The **active site** is the region on the enzyme where the substrate binds





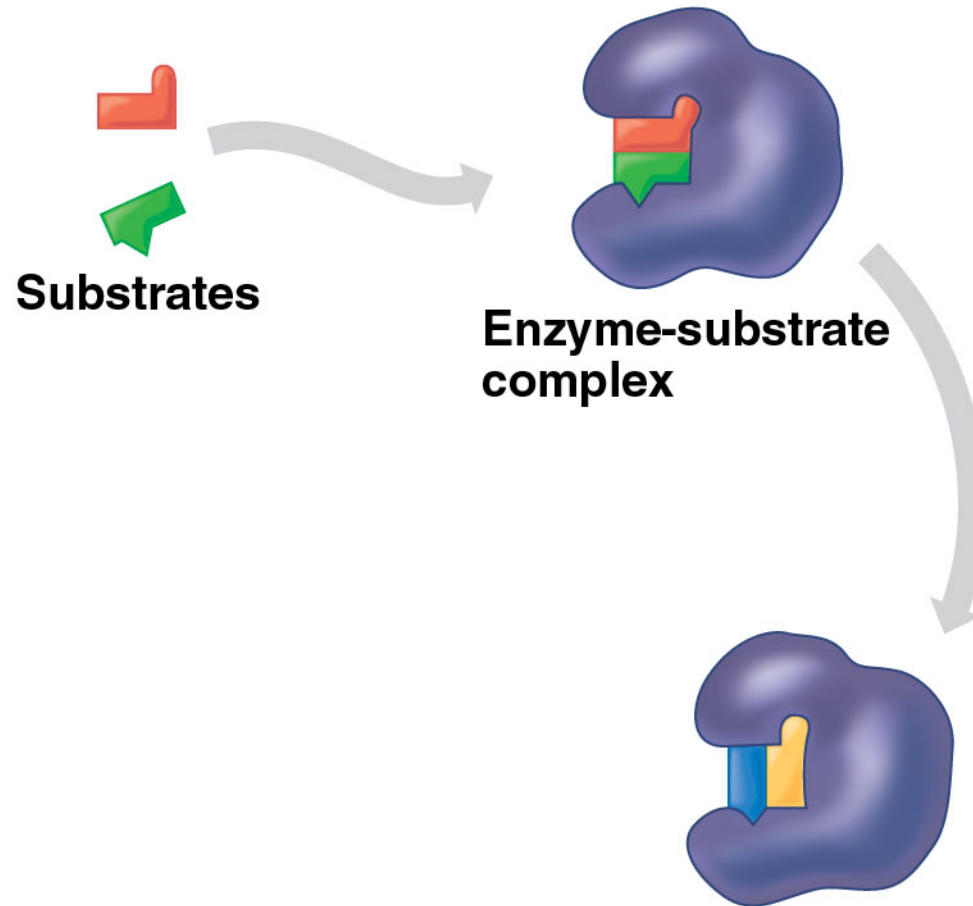
**1** Substrates enter active site.

**2** Substrates are held in active site by weak interactions.



**1** Substrates enter active site.

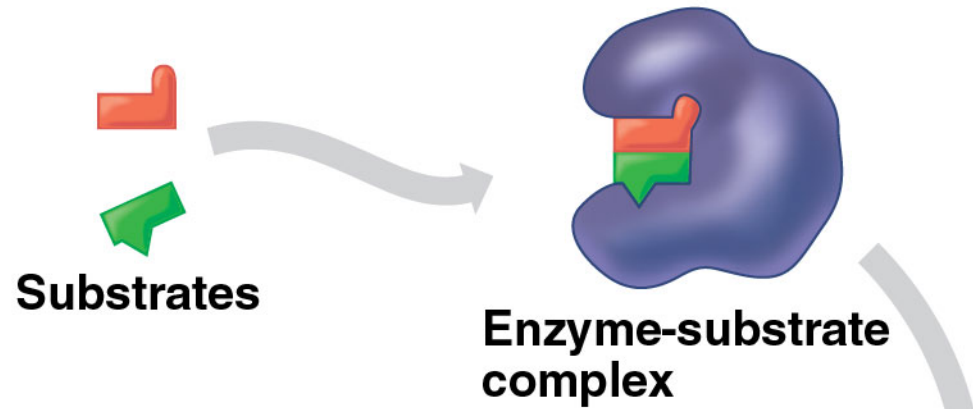
**2** Substrates are held in active site by weak interactions.



**3** Substrates are converted to products.

**1** Substrates enter active site.

**2** Substrates are held in active site by weak interactions.



**4** Products are released.



**3** Substrates are converted to products.



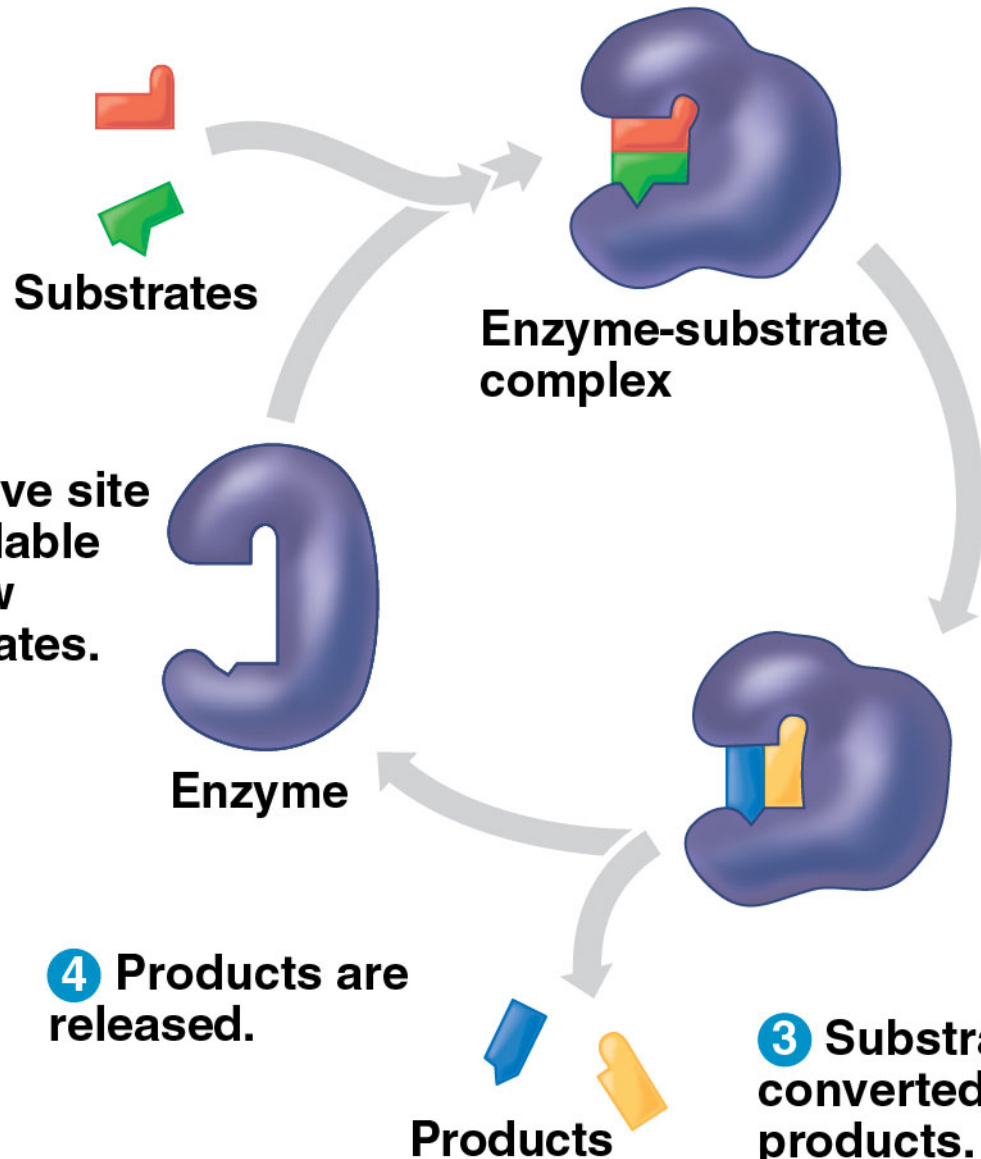
**1** Substrates enter active site.

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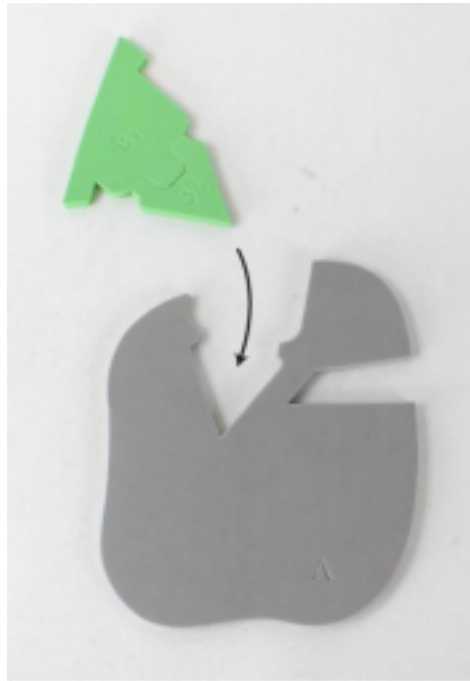
**5** Active site is available for new substrates.

**4** Products are released.

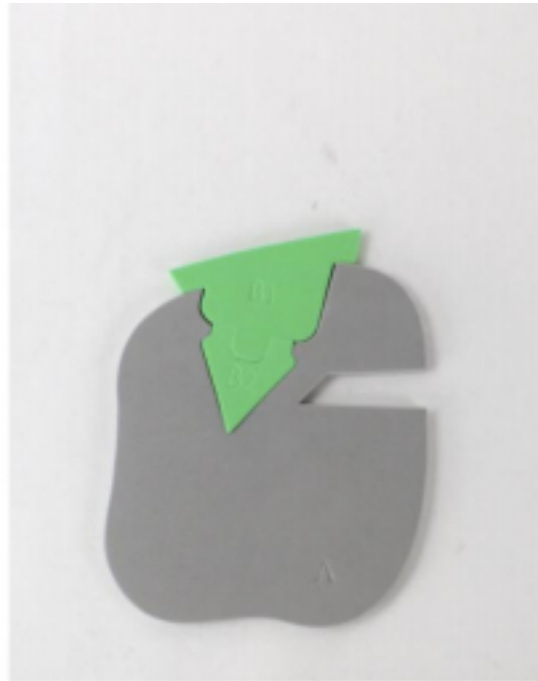
**3** Substrates are converted to products.



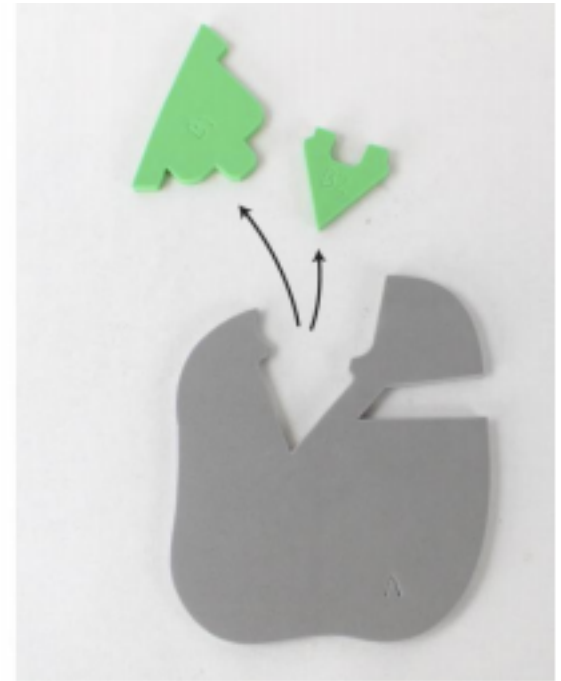
# ENZYME ACTION: CATABOLISM



**Step 1**



**Step 2**



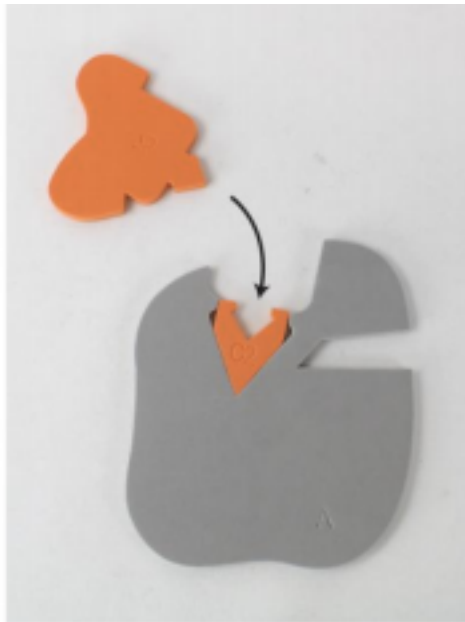
**Step 3**



# ENZYME ACTION: ANABOLISM



**Step 1**



**Step 2**



**Step 3**

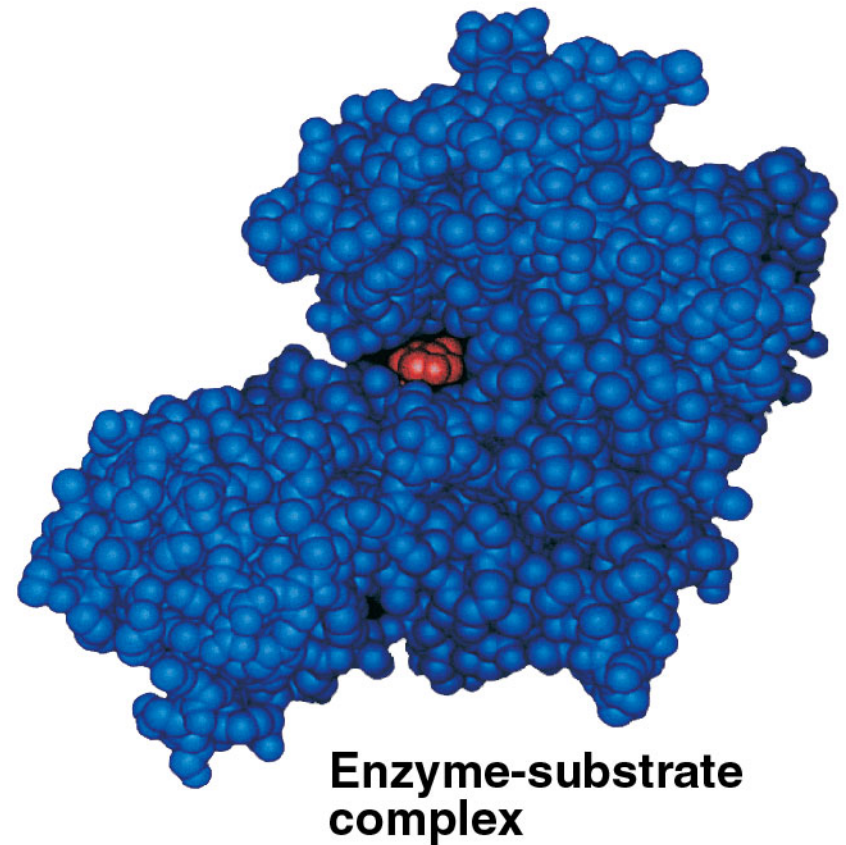
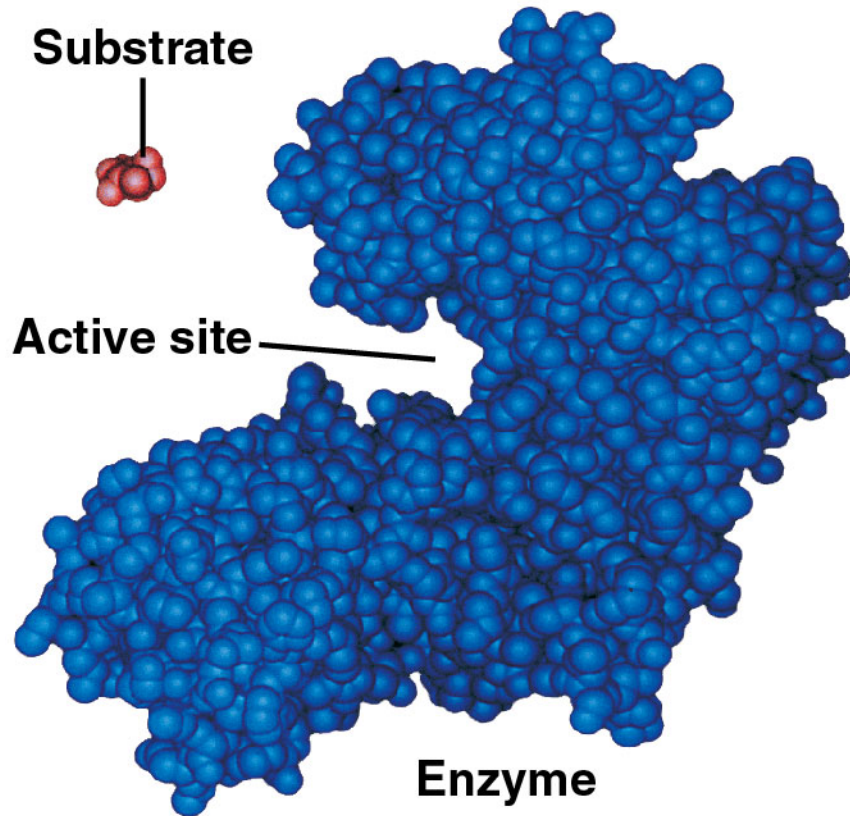


**Step 4**



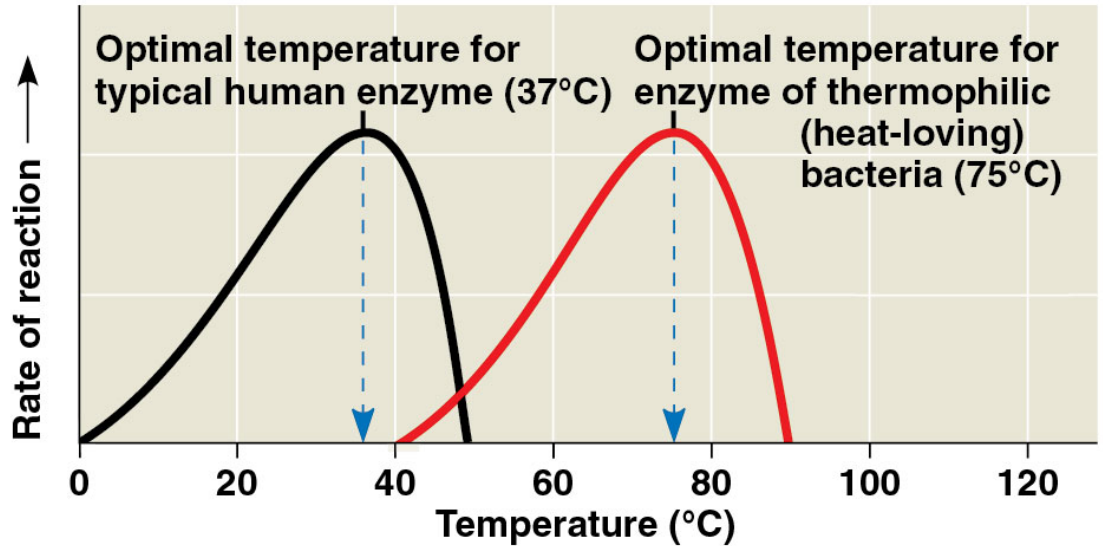


**INDUCED FIT:** ENZYME FITS SNUGLY AROUND  
SUBSTRATE -- “CLASPING HANDSHAKE”

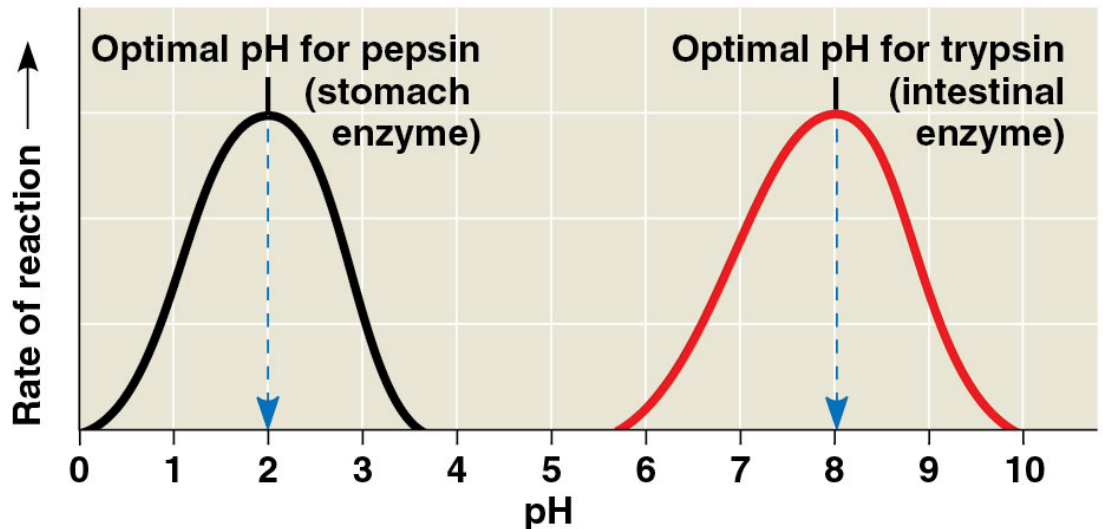


An enzyme's activity can be affected by:

- Temperature
- pH
- Chemicals



(a) Optimal temperature for two enzymes



(b) Optimal pH for two enzymes

# ENZYME STRUCTURE & FUNCTION

- Change to the **molecular structure** of a component in an enzymatic system may result in a change of **function** or **efficiency** of the system
- **Denaturation**: disrupt protein structure  
→ reduce enzymatic activity
- **Environmental pH**: alter efficiency of enzyme activity; disruption of H-bonds
- In some cases, enzyme denaturation is *reversible* → enzyme regains activity



# COFACTORS

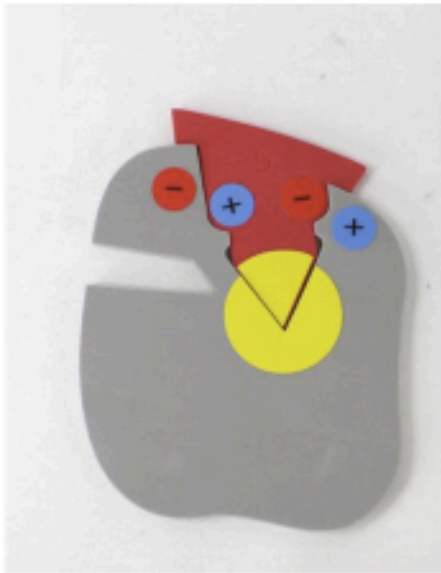
- Cofactors: nonprotein enzyme helpers such as minerals (eg. Zn, Fe, Cu)
- Coenzymes: organic cofactors (eg. vitamins)

## Enzyme Inhibitors

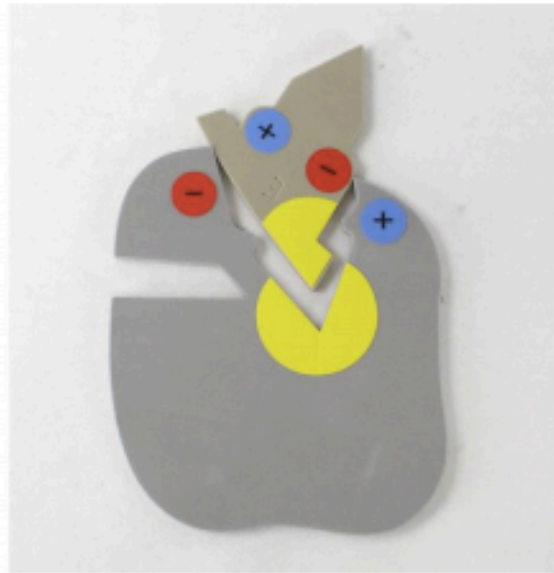
- Competitive inhibitor: binds to the *active site* of an enzyme, competes with substrate
- Noncompetitive inhibitor: binds to *another part* of an enzyme → enzyme changes shape → active site is *nonfunctional*



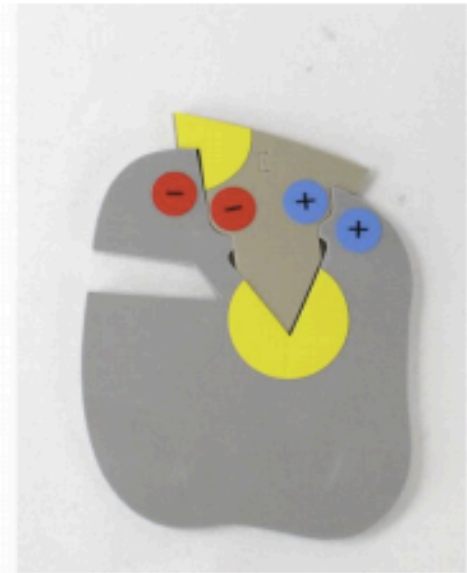
# ENZYME SPECIFICITY



**Figure 1:** Enzyme-substrate complex



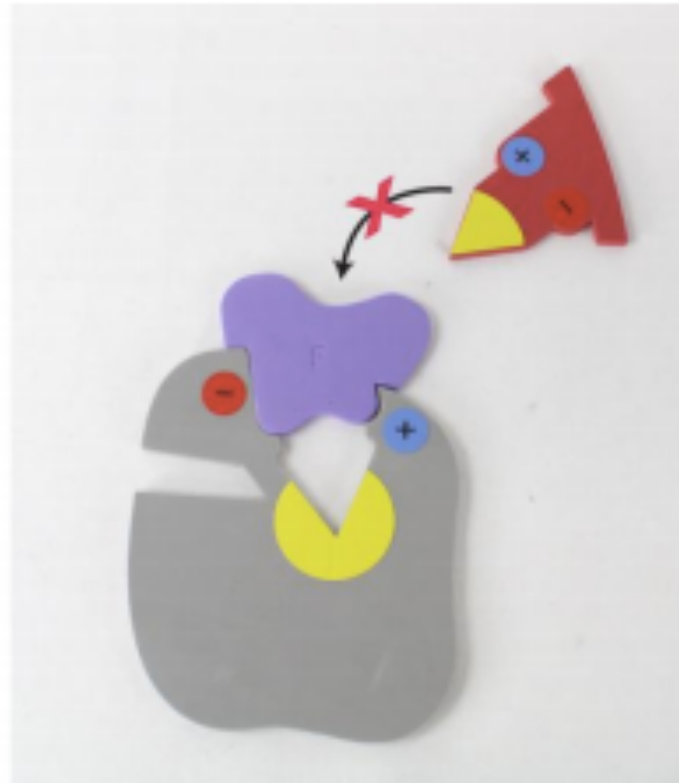
**Figure 2:** The charges align between the enzyme and the substrate; however, the enzyme's shape will not "fit".



**Figure 3:** The shape of the substrate appears to fit but the charges do not align in the active site of the enzyme.

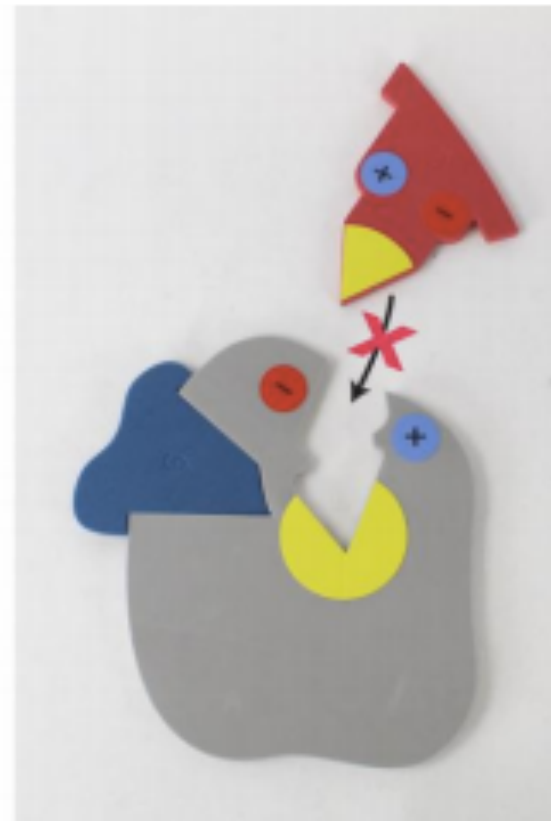


# COMPETITIVE INHIBITION



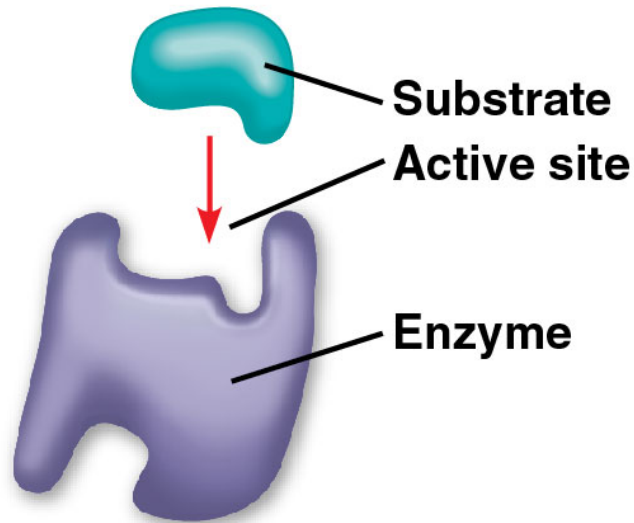


# NONCOMPETITIVE INHIBITION

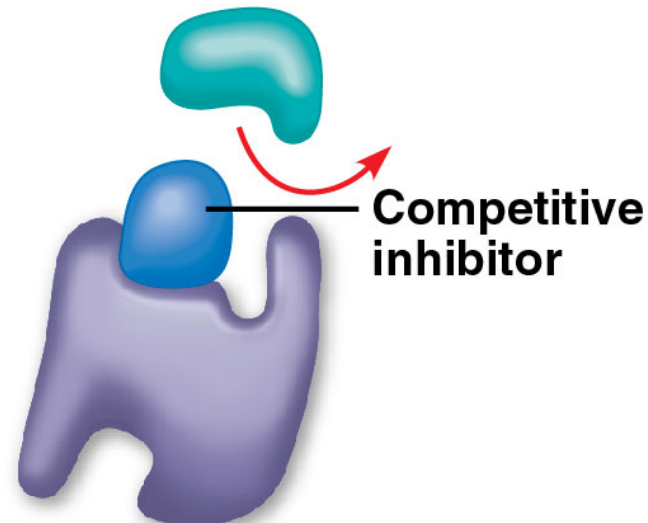


# INHIBITION OF ENZYME ACTIVITY

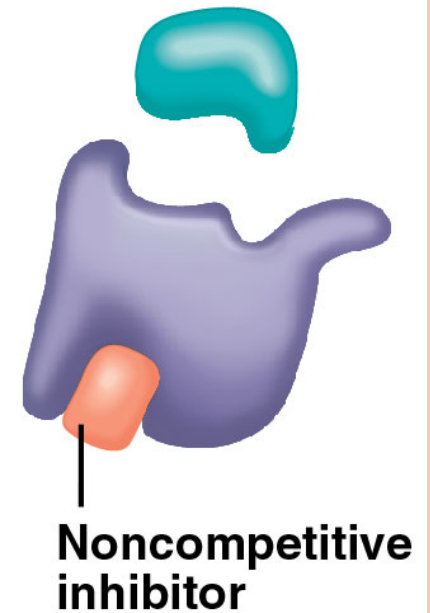
(a) Normal binding



(b) Competitive inhibition



(c) Noncompetitive inhibition

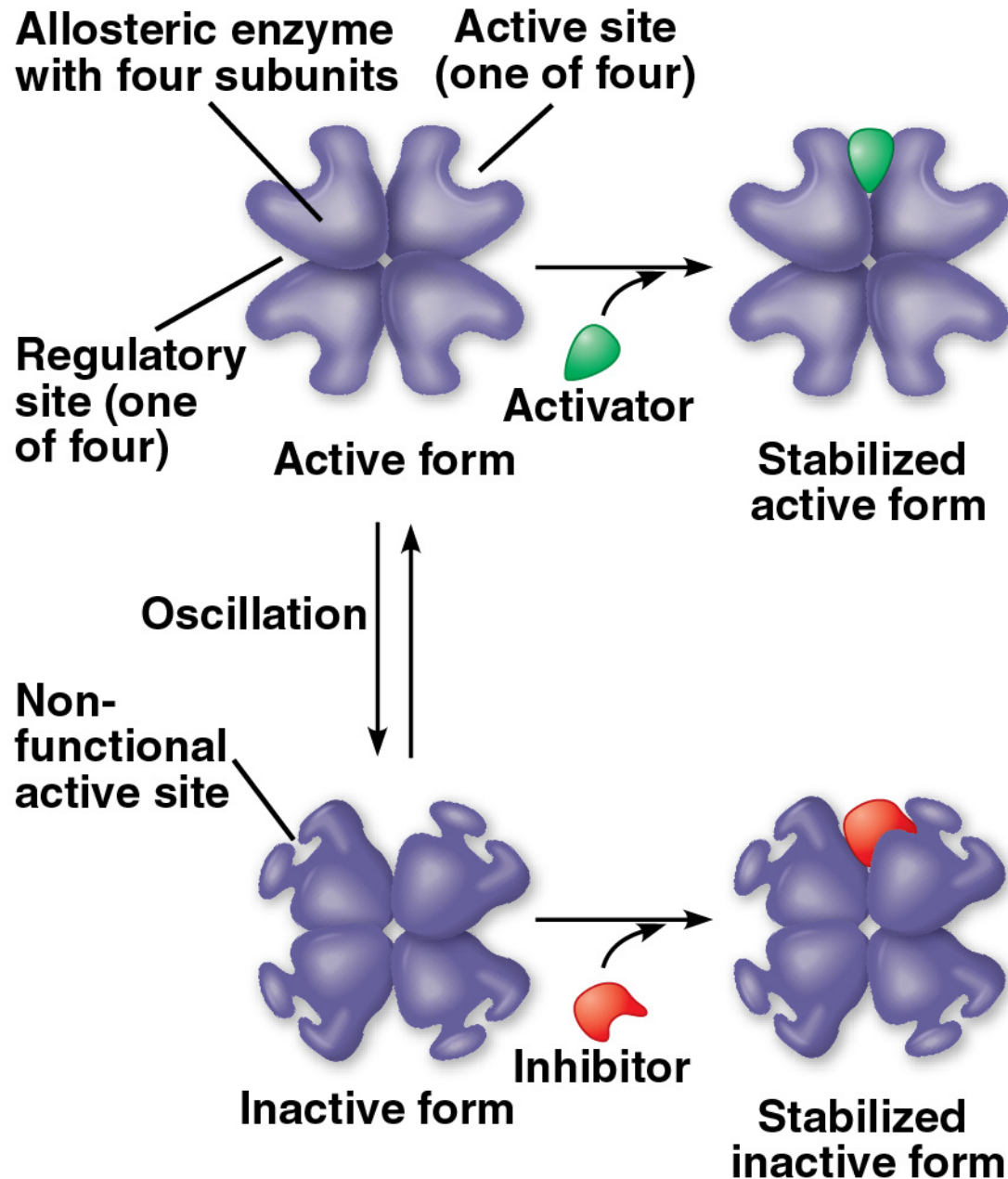


# REGULATION OF ENZYME ACTIVITY

- To regulate metabolic pathways, the cell switches on/off the genes that encode specific enzymes
- **Allosteric regulation**: protein's function at one site is affected by binding of a **regulatory molecule** to a separate site (allosteric site)
  - **Activator** – stabilizes active site
  - **Inhibitor** – stabilizes inactive form
  - **Cooperativity** – one substrate triggers shape change in other active sites → increase catalytic activity

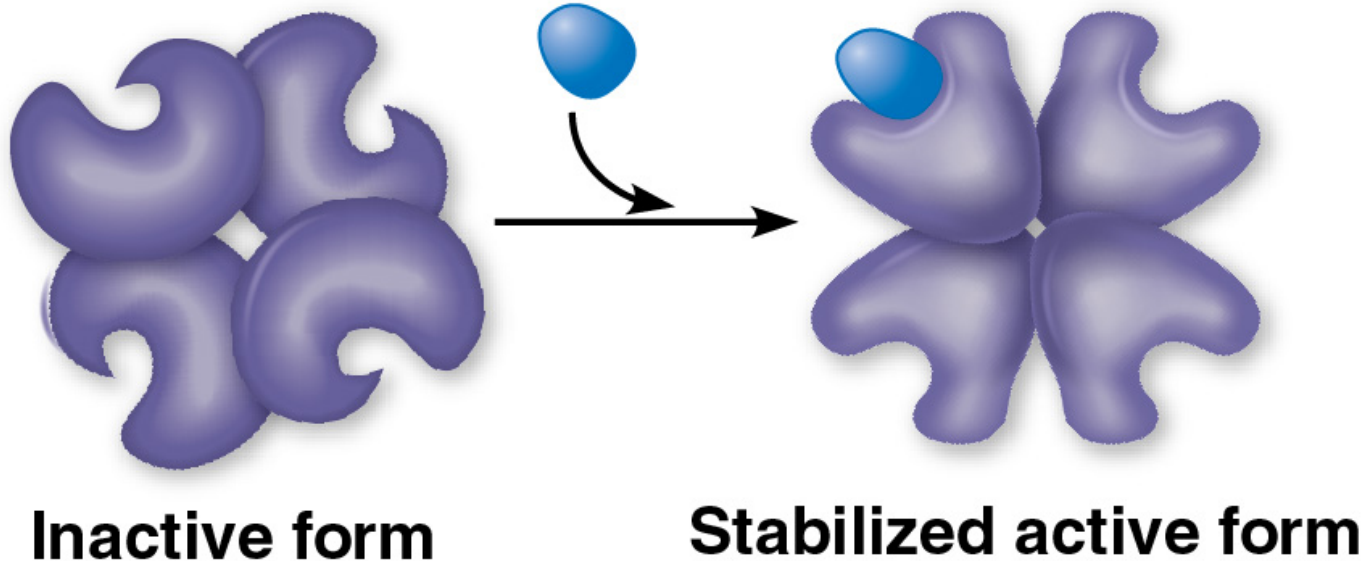


## (a) Allosteric activators and inhibitors



**(b) Cooperativity: another type of allosteric activation**

**Substrate**



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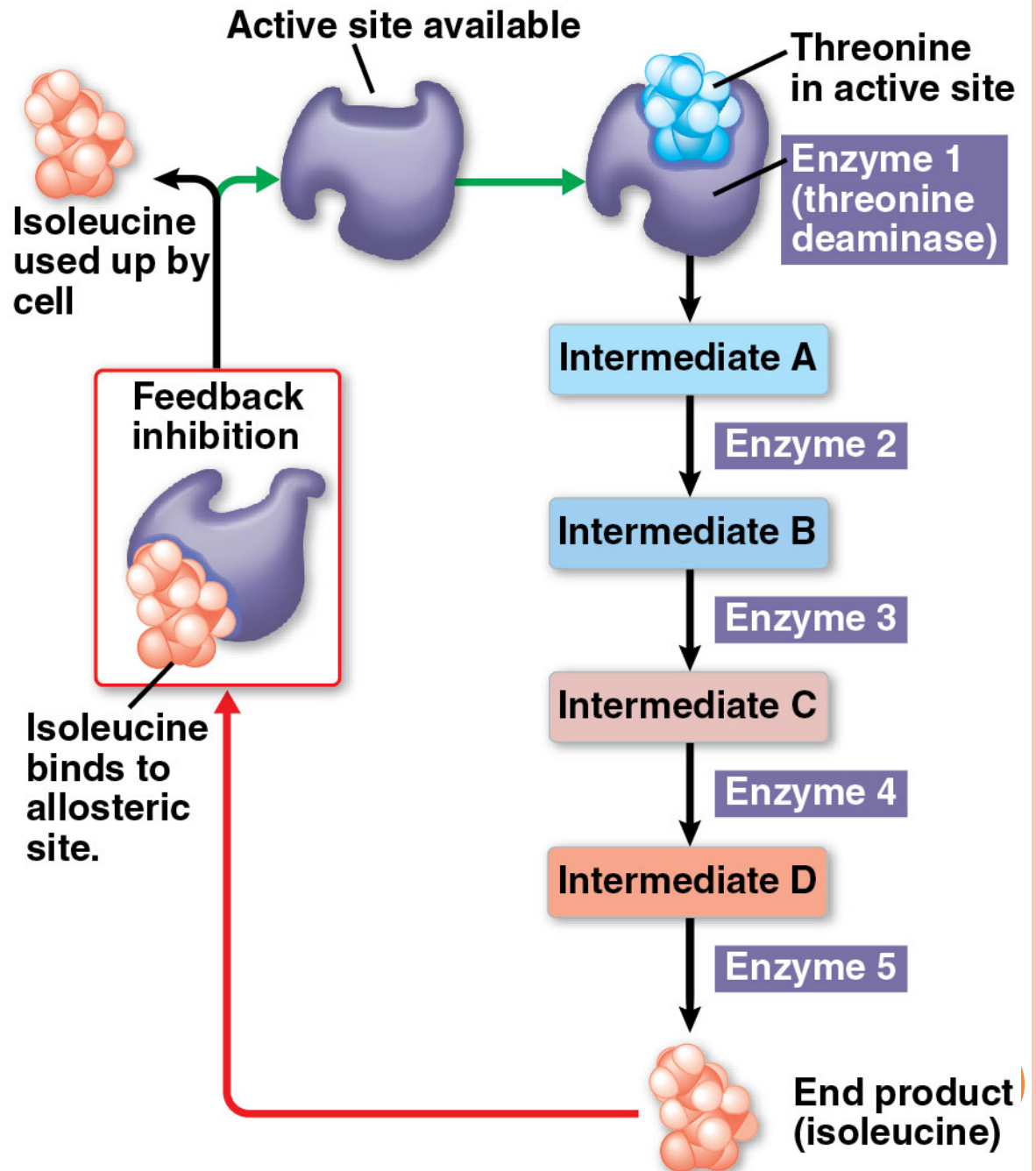
# *FEEDBACK INHIBITION*

- End product of a metabolic pathway shuts down pathway by binding to the allosteric site of an enzyme
- Prevent wasting chemical resources, increase efficiency of cell

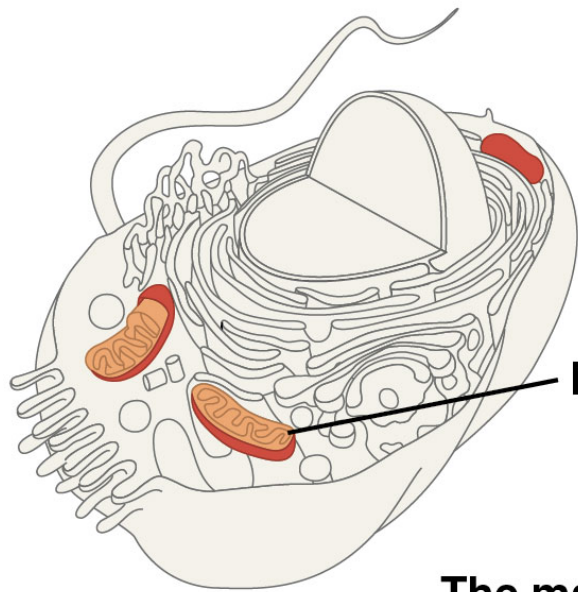




# FEEDBACK INHIBITION



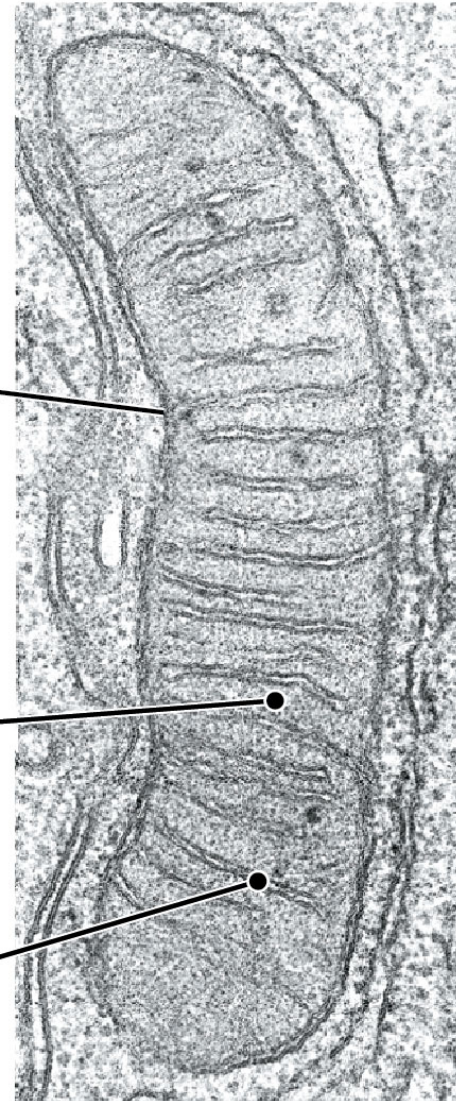
# ORGANIZATION OF ENZYMES WITHIN A CELL



Mitochondrion

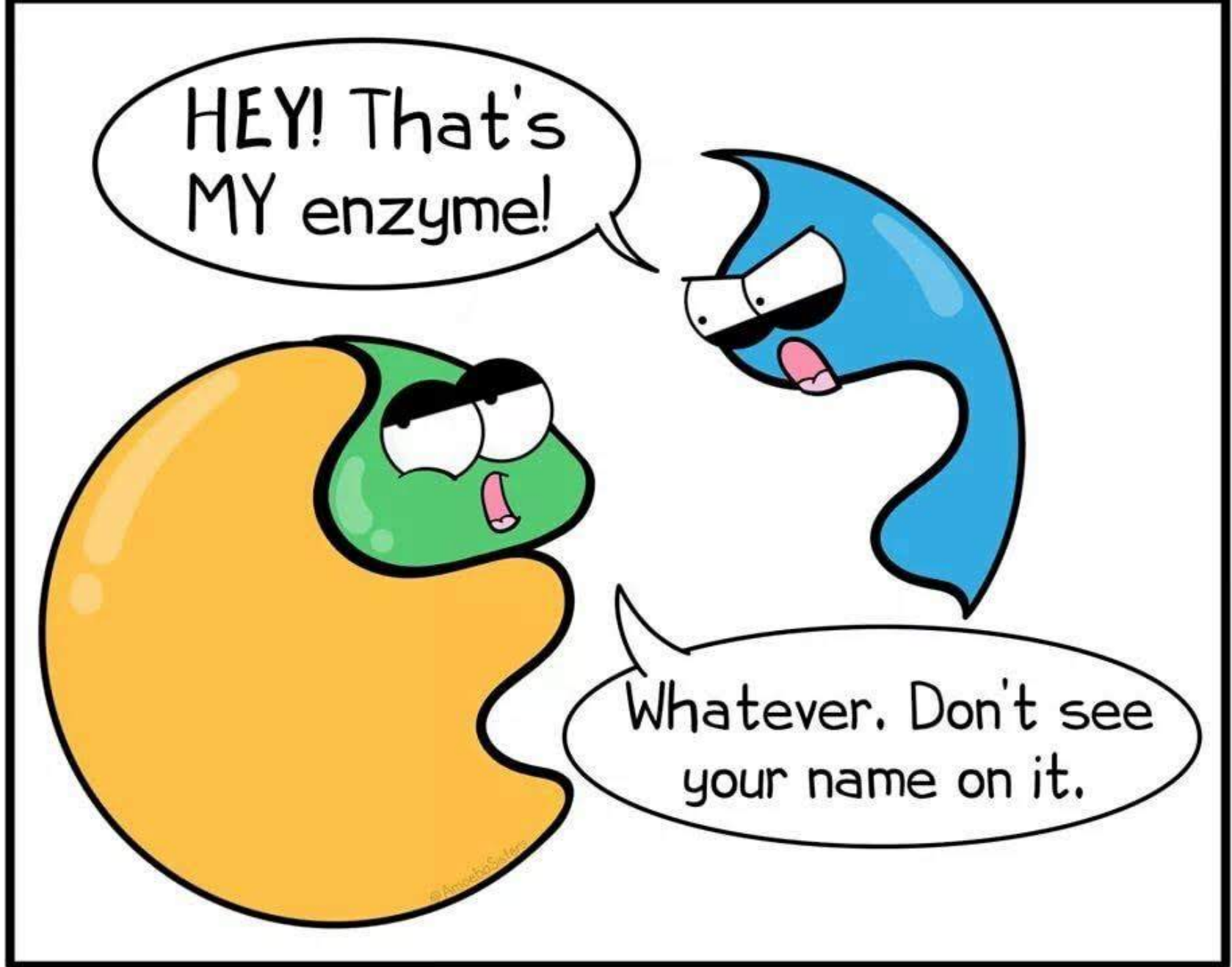
**The matrix contains enzymes in solution that are involved in one stage of cellular respiration.**

**Enzymes for another stage of cellular respiration are embedded in the inner membrane.**



1  $\mu\text{m}$





**Competitive Inhibitors:** If it fits, it sits.