ME 411 / ME 511

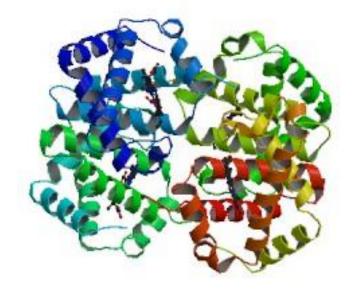
Biological Frameworks for Engineers





Class Organization

- HW2 due on Friday
- Lab 1 Protein Structure
 - Bring your laptops on Wed
 - Handouts provided





ME 411 / ME 511

Proteins





Protein Functions

- Different shapes and sizes mediate a diverse array of activities
- Function based on proteins binding to themselves, other proteins, small molecules, or ions
- Life is nothing without the function of proteins...

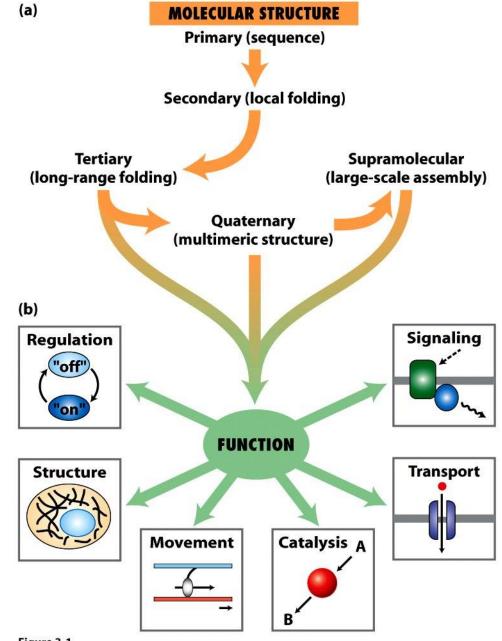


Figure 3-1

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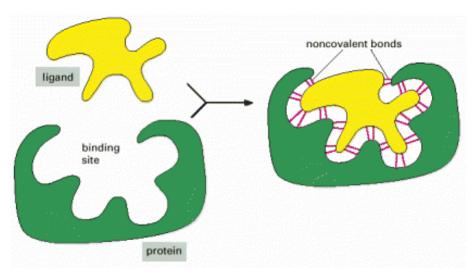
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Binding

 Specific binding of a target molecule regulates protein function

Specificity

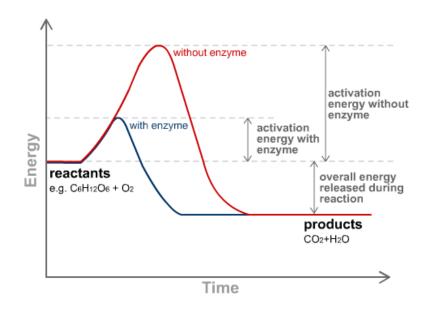






Enzymatic Function

- Enzyme catalyze the rate of reactions inside a cell
- Substrate target for enzymes that become the products of the reaction



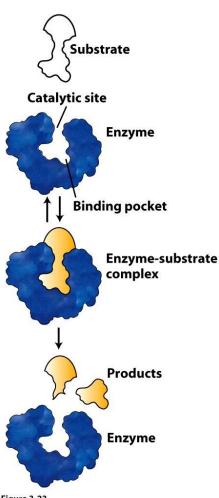
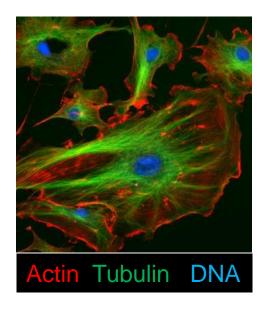


Figure 3-23

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Structure

- Cytoskeleton actin, microtubules, intermediate filaments, cadherins, integrins, and others
- Extracellular matrix collagen, elastin, laminin, fibronectin, and others



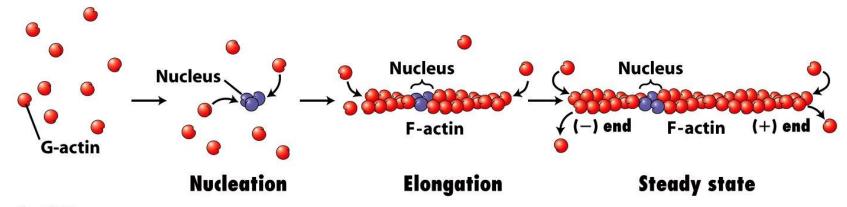


Figure 17-7a

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Signaling

Signaling Proteins – molecules and receptors

(a) Residues essential to (b) (c) tight binding with receptor

Growth hormone

Residues essential to (b) (c)

Residues essential to (b) (c)

Residues essential to (b) (c)

Electrical

Figure 15-3

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Co-receptor (blue)

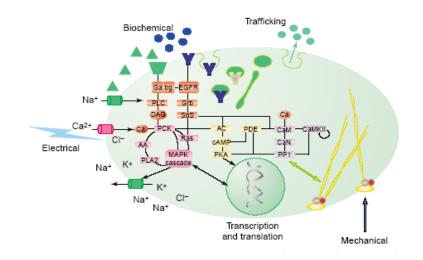
Transcription and translation

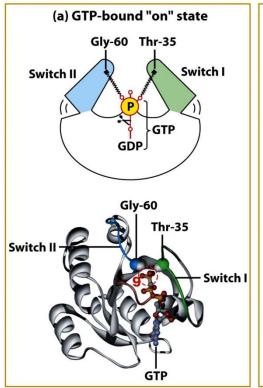
Mechanical



Regulation

- Regulatory Proteins –
 kinases, phosphatases,
 GTPases, etc. interpret a
 receptor signal for gene
 expression or cell
 function
- RasGTP has allosteric change in conformation
- Dissociation of GTP to GDP is an "egg timer"





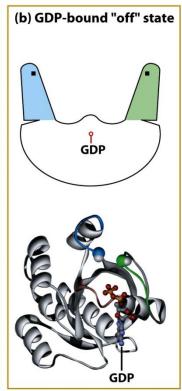
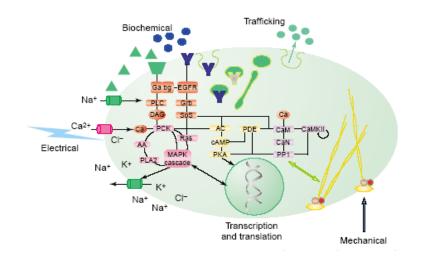


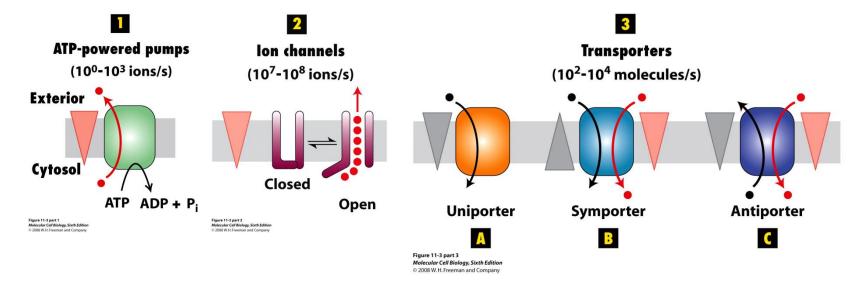
Figure 15-8

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Transport

 Membrane transport proteins – control the transport of ions and small molecules across membranes



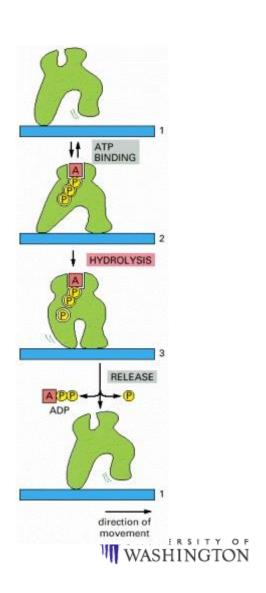






Motor Proteins

- Allosteric motor protein
- Transition between three conformations allows stepping motion
- Regulated by
 - ATP binding
 - Hydrolysis of ATP to ADP
 - ADP unbinding



Dissociation Constant

<u>Affinity</u>

Binding Reaction:

$$P + L \leftrightarrow PL$$

Dissociation Constant:

$$K_d = [P][L] / [PL]$$

where [] is concentration e.g. molarity (mol/L)

Comparisons

Weak:

$$K_d \ge 10^{-3} M$$

Moderate

$$K_d \approx 10^{-6} M$$

• Tight:

$$K_{cl} \le 10^{-9} M$$

• Biotin-Avidin:

$$K_d \ge 10^{-15} M$$



- Consider a cell having
 - 100 copies of protein P
 - 100 copies of ligand L

- 10⁻¹⁵ L volume
- Reaction at equilibrium



- Consider a cell having
 - 100 copies of protein P
 - 100 copies of ligand L

- 10⁻¹⁵ L volume
- Reaction at equilibrium

$$\frac{[P][L]}{[PL]} = 10^{-6} \text{ mol/L}$$



- Consider a cell having
 - 100 copies of protein P
 - 100 copies of ligand L

- 10⁻¹⁵ L volume
- Reaction at equilibrium

$$\frac{\left(\frac{(100-PL)}{N_A V}\right)\left(\frac{(100-PL)}{N_A V}\right)}{\frac{(PL)}{N_A V}} = 10^{-6} \text{ mol/L}$$



- Consider a cell having
 - 100 copies of protein P
 - 100 copies of ligand L

- 10⁻¹⁵ L volume
- Reaction at equilibrium

$$\frac{\frac{\binom{(100-PL)}{N_A V}\binom{(100-PL)}{N_A V}}{\frac{(PL)}{N_A V}} = 10^{-6} \text{ mol/L}$$

$$10^4 + (-200 - N_A \times V \times 10^{-6})PL + PL^2 = 0$$



- Consider a cell having
 - 100 copies of protein P
 - 100 copies of ligand L

- 10⁻¹⁵ L volume
- Reaction at equilibrium

• If $K_d = 10^{-6} M$, then

$$\frac{\left(\frac{(100-PL)}{N_A V}\right)\left(\frac{(100-PL)}{N_A V}\right)}{\frac{(PL)}{N_A V}} = 10^{-6} \text{ mol/L}$$

$$10^4 + (-200 - N_A \times V \times 10^{-6})PL + PL^2 = 0$$

88 copies of unbound P

88 copies of unbound L

12 copies of PL (P bound to L)



- Consider a cell having
 - 100 copies of protein P
 - 100 copies of ligand L

- 10⁻¹⁵ L volume
- Reaction at equilibrium





- Consider a cell having
 - 100 copies of protein P
 - 100 copies of ligand L

- 10⁻¹⁵ L volume
- Reaction at equilibrium

- If $K_d = 10^{-9} M$, then
 - 8 copies of unbound P
 - 8 copies of unbound L
 - 92 copies of PL (P bound to L)



Questions?

