

<http://www.ukclimbing.com/news/item.php?id=49981>

## Physiology

- **Define:** The study of how living organisms function
- Structure & function are important to understand function
- Why study: Curiosity. Better understand how humans function under normal conditions. Thus, modifications of pathological states back to a 'normal' state might be possible.
- Many of the physiological process are described by chemical and physical properties
- It is important to integrate these concepts with biology.

- The physiology of an animal is well suited to the environment in which it has evolved.

This is explained by the process of **ADAPTATION**- gradual change over many generations.

- **Acclimatization** is a change of an individual over its lifetime of biochemical or anatomical alterations
- **Acclimation** is like acclimatization but induced by experimentation.

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- Homeostasis- The tendency of an organism to regulate and maintain relative internal stability.

- Claude Bernard coined this term 1800's

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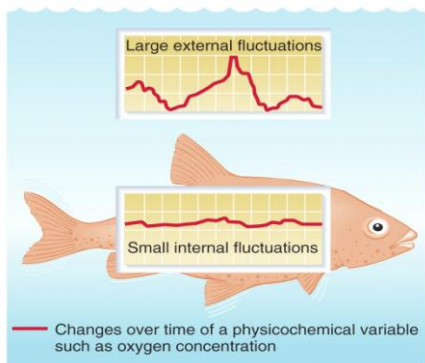
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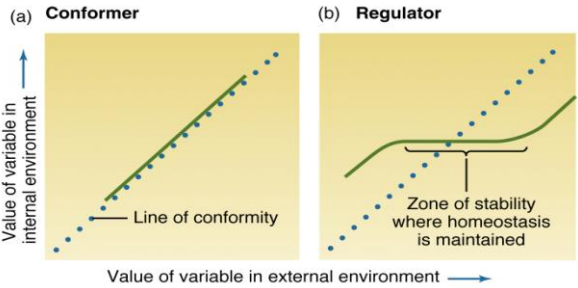
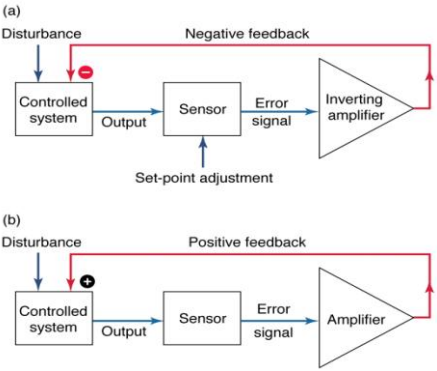
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This mostly works by a feed back control.

Such as by a negative feedback.

Examples - Temp, pH, salinity within the body



## Chapter 2

- Read over it for your own benefit
  - Know August Krogh principle.
- that there is an animal optimally suited to yield an answer of a physiological problem to be addressed

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## Chapter 3

- Understanding of molecules and their function for physiological processes.
- Water- it is polar can be used to interact with other ions. It lets  $\text{Na}^+\text{Cl}^-$  dissolve in body fluids. This is needed for rapid transport of ions.

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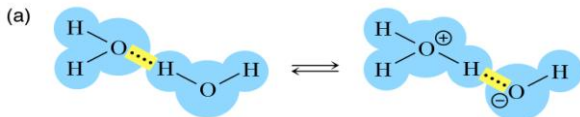
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- Acid: any substance that can donate a proton.
- Base: any substance that combines with a proton.

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(b)

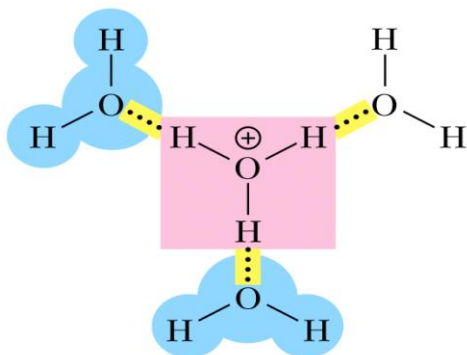
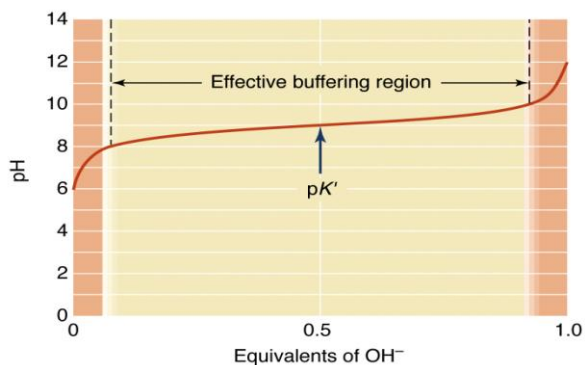


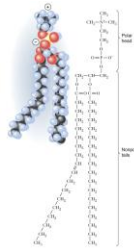
Table 3-2 The pH scale

|                         | pH | $[\text{H}^+]$ (mol · L <sup>-1</sup> ) | $[\text{OH}^-]$ (mol · L <sup>-1</sup> ) | Examples                |
|-------------------------|----|---|--|-------------------------|
|                         | 0  | 10 <sup>0</sup>                         | 10 <sup>-14</sup>                        |                         |
| ↑ Increasing acidity    | 1  | 10 <sup>-1</sup>                        | 10 <sup>-13</sup>                        | Human gastric fluids    |
|                         | 2  | 10 <sup>-2</sup>                        | 10 <sup>-12</sup>                        |                         |
|                         | 3  | 10 <sup>-3</sup>                        | 10 <sup>-11</sup>                        | Household vinegar       |
|                         | 4  | 10 <sup>-4</sup>                        | 10 <sup>-10</sup>                        |                         |
|                         | 5  | 10 <sup>-5</sup>                        | 10 <sup>-9</sup>                         | Interior of lysosomes   |
|                         | 6  | 10 <sup>-6</sup>                        | 10 <sup>-8</sup>                         | Cytoplasm of muscle     |
| Neutral                 | 7  | 10 <sup>-7</sup>                        | 10 <sup>-7</sup>                         | Pure water at 25°C      |
|                         | 8  | 10 <sup>-8</sup>                        | 10 <sup>-6</sup>                         | Seawater                |
|                         | 9  | 10 <sup>-9</sup>                        | 10 <sup>-5</sup>                         |                         |
| ↓ Increasing alkalinity | 10 | 10 <sup>-10</sup>                       | 10 <sup>-4</sup>                         | Alkaline lakes          |
|                         | 11 | 10 <sup>-11</sup>                       | 10 <sup>-3</sup>                         | Household ammonia       |
|                         | 12 | 10 <sup>-12</sup>                       | 10 <sup>-2</sup>                         | Saturated lime solution |
|                         | 13 | 10 <sup>-13</sup>                       | 10 <sup>-1</sup>                         |                         |
|                         | 14 | 10 <sup>-14</sup>                       | 10 <sup>0</sup>                          |                         |



FATS:

1. Lipids
2. Phospholipids for membranes
3. Waxes- water proof the surface of insects




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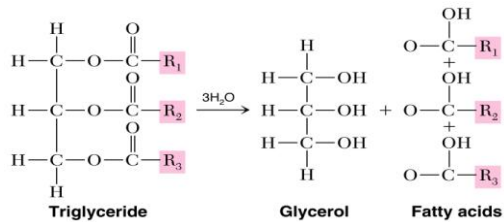
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## Biological Molecules

### LIPIDS

Fats - triglyceride




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### 2. Phospholipids for membranes

Saturated- each carbon has a single bond and is saturated with 'H'

Unsaturated- some carbons have a double bond and thus are not saturated with 'H'.

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|                 |              |   |
|-----------------|--------------|---|
|                 | <b>Unsat</b> | <b>Sat</b>  |
| 1. Melting Pt.: | Decreased    | increased   |
| 2. Metab:       | -            | easily converted to sterols<br>(ie, Cholesterol). |

Need for cholesterol for steroid based hormone production.

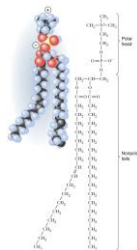
Fats are good as a energy store. 1. Little water needed. 2. Stored in adipose tissue. 3. High “H & C” with little “O” so one gets a good energy yield. (1 gram of fat=2 grams of carbohydrates)

**Table 3-3** The energy content of the three major categories of foodstuffs

| Substrate     | Energy content (kcal · g <sup>-1</sup> ) |
|---------------|--|
| Carbohydrates | 4.0                                      |
| Proteins      | 4.5                                      |
| Fats          | 9.5                                      |

FATS:

1. Lipids
2. Phospholipids for membranes
3. Waxes- water proof the surface of insects



# Carbohydrates

- I. Monosaccharides
- II. Disaccharides

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- I. Monosaccharides
  - Six carbon-- hexoses
  - five carbon-- pentoses

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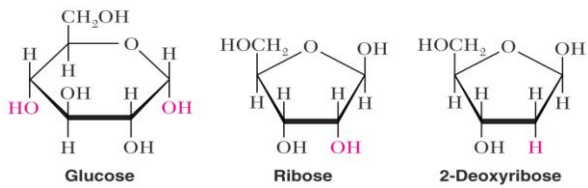
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## (a) Monosaccharide sugars



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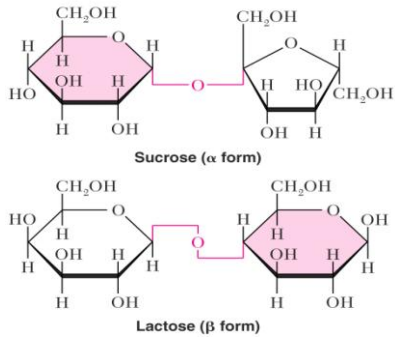
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## (b) Disaccharide sugars



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The disaccharides are units of monosaccharides

- A. Polymers     – Plants—starch  
                  -- Animals— glycogen

Little water needed for storage. So good energy  
for plants and animals.

B. Cellulose

C. Chitin (exoskeleton in insects/crustaceans)

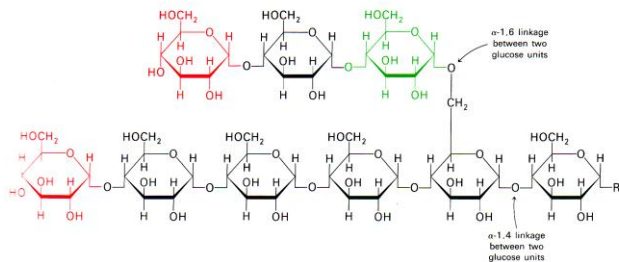
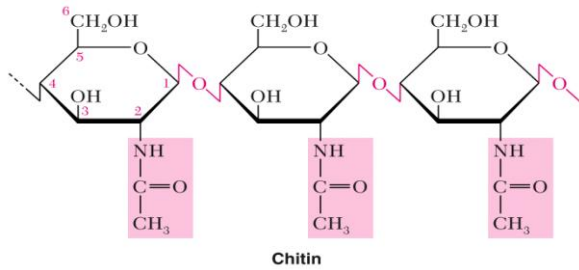


Table 16.1  
Glycogen storage diseases

| Type                      | Defective enzyme  | Organ affected   | Glycogen in the affected organ                 | Clinical features   |
|---------------------------|---|------------------|--|---|
| I<br>VON GIERKE'S DISEASE | Glucose 6-phosphatase   | Liver and kidney | Increased amount; normal structure.            | Massive enlargement of the liver. Failure to thrive. Severe hypoglycemia, ketosis, hyperuricemia, hypocalcemia.                 |
| II<br>POMPE'S DISEASE     | $\alpha$ -1,4-Glucosidase (lysosomal)                         | All organs       | Massive increase in amount; normal structure.  | Cardiorespiratory failure causes death, usually before age 2.   |
| III<br>CORI'S DISEASE     | Amylo-1,6-glucosidase (debranching enzyme)                    | Muscle and liver | Increased amount; short outer branches.        | Like Type I, but milder course.   |
| IV<br>ANDERSEN'S DISEASE  | Branching enzyme ( $\alpha$ -1,4 $\rightarrow$ $\alpha$ -1,6) | Liver and spleen | Normal amount; very long outer branches.       | Progressive cirrhosis of the liver. Liver failure causes death usually before age 2.  |
| V<br>MCKARDLE'S DISEASE   | Phosphorylase   | Muscle           | Moderately increased amount; normal structure. | Limited ability to perform strenuous exercise because of painful muscle cramps. Otherwise patient is normal and well developed. |
| VI<br>HERS DISEASE        | Phosphorylase   | Liver            | Increased amount.                              | Like Type I, but milder course.   |
| VII                       | Phosphofructokinase   | Muscle           | Increased amount; normal structure.            | Like Type V.  |
| VIII                      | Phosphorylase kinase  | Liver            | Increased amount; normal structure.            | Mild liver enlargement. Mild hypoglycemia.  |

Note: Types I through VII are inherited as autosomal recessives. Type VIII is sex-linked.




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## Proteins

- A lot in cells. ½ of the dry mass.
- Various structures.
- Primary, secondary, tertiary, and quaternary

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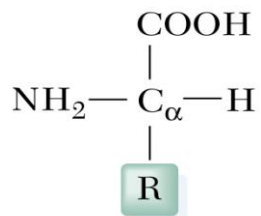
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(a) General structure of alpha-amino acids




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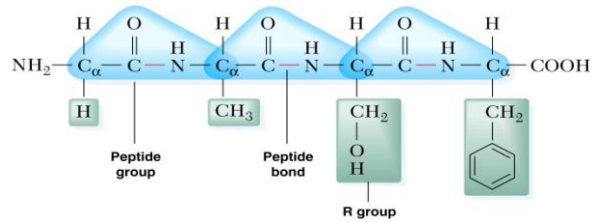
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(b) Structure of a tetrapeptide




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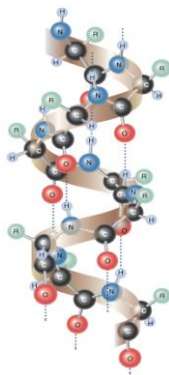
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Secondary  
Alpha helix




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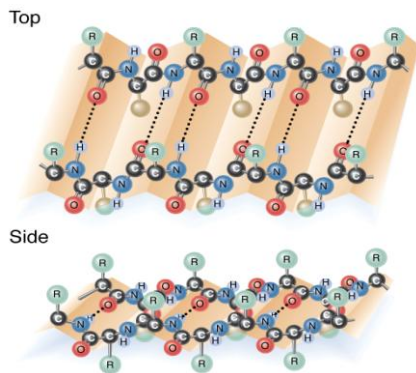
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Secondary  
Beta- sheet




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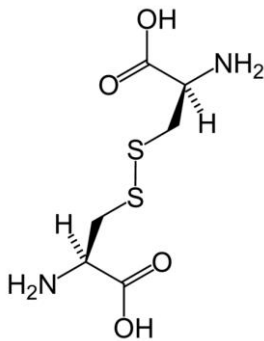
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## Secondary types

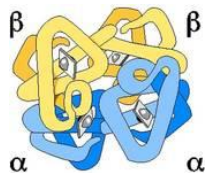
- alpha helix: alpha-Keratins for hair and wool
- Beta sheets: (Harder) beta- Keratins for reptile scales and turtle shells

Tertiary



Quaternary- a couple of subunits coming together like Heme units.

ie., Hemoglobin



## Molecular chaperones

- Fold other proteins. Maintain their structure.
- To protect proteins from degradation and preserve the overall integrity of intracellular protein pool.
- Example HSP or stress proteins.

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## DNA & RNA

- DNA isolated in 1869 by Friedrich Miescher.  
White blood cells and fish sperm.
- DNA- genes
- RNA translating the coded DNA message.

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- DNA with mRNA--- Transcription
  - mRNA to protein --- Translation
- A-T and C-G in DNA  
But A-U in RNA

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## Energy in Living cells

- Chemical Rx's taking place (types of tissues)
- Where within the cells? (organelles)

Animals are like chemical machines.

Different forms of Energy- ie., thermal and mechanical.

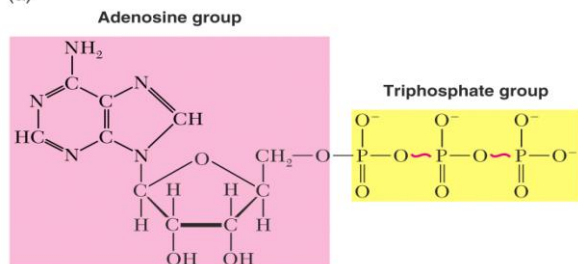
- A measure of a systems energy that it can provide.

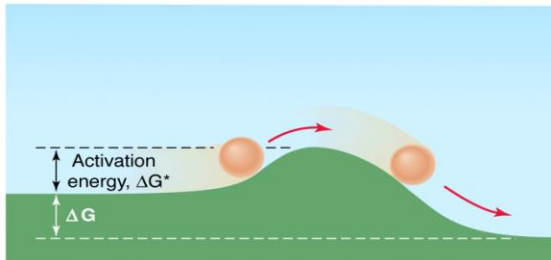
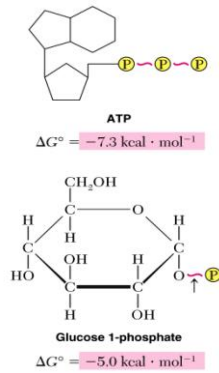
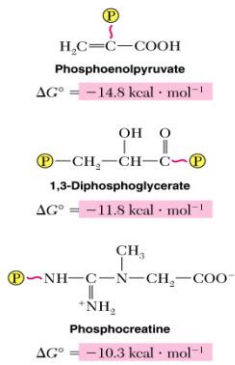
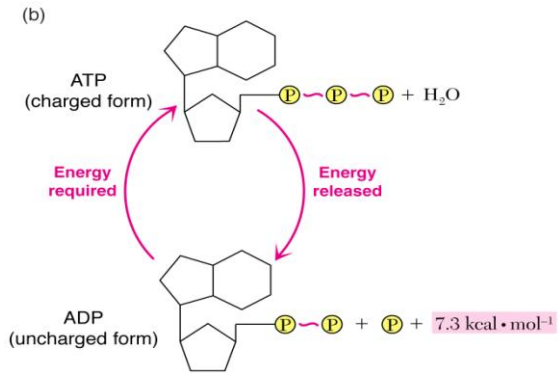
$\Delta G = \Delta H - T\Delta S$  Is Gibbs free Energy

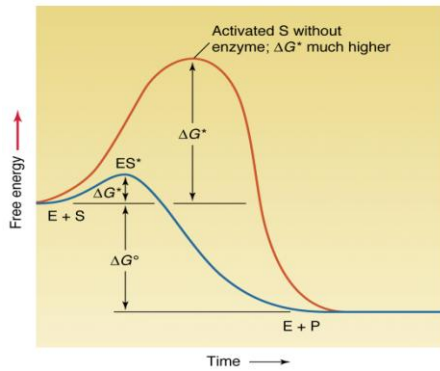
$\Delta H$  is heat;  $\Delta S$  is Entropy

If  $\Delta G$  is neg then the Rx is spontaneous. Also hold for biochemical Rx's.

(a)








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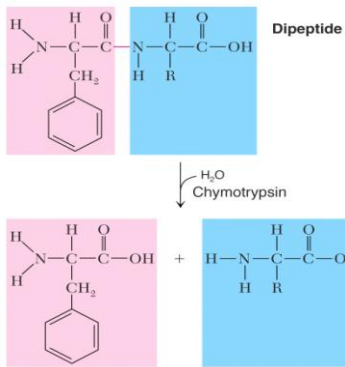
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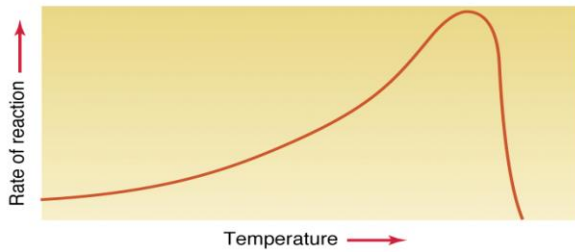
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(a) Enzyme activity versus temperature




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(b) Enzyme activity versus pH

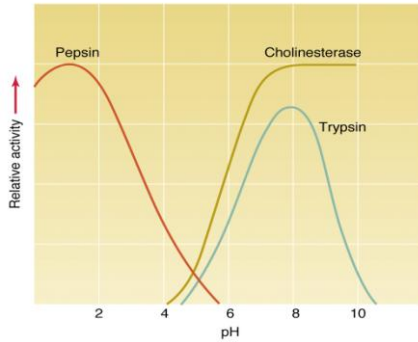
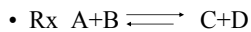


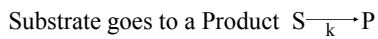
Table 3-6 Metal ions functioning as cofactors

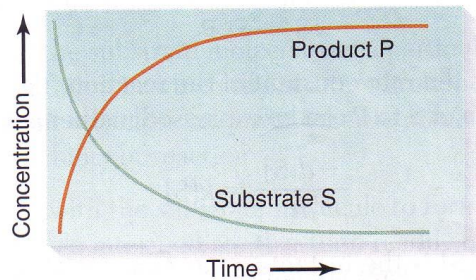
| Metal ion                            | Some enzymes requiring this cofactor                                      |
|--------------------------------------|---|
| $\text{Ca}^{2+}$                     | Phosphodiesterase<br>Protein kinase C                                     |
| $\text{Cu}^{2+}$ ( $\text{Cu}^+$ )   | Cytochrome oxidase<br>Tyrosinase  |
| $\text{Fe}^{2+}$ or $\text{Fe}^{3+}$ | Catalase<br>Cytochromes<br>Ferredoxin<br>Peroxidase                       |
| $\text{K}^+$                         | Pyruvate phosphokinase (also requires $\text{Mg}^{2+}$ )                  |
| $\text{Mg}^{2+}$                     | Phosphohydrolases<br>Phosphotransferases                                  |
| $\text{Mn}^{2+}$                     | Arginase<br>Phosphotransferases   |
| $\text{Na}^+$                        | Plasma membrane ATPase (also requires $\text{K}^+$ and $\text{Mg}^{2+}$ ) |
| $\text{Zn}^{2+}$                     | Alcohol dehydrogenase<br>Carbonic anhydrase<br>Carboxypeptidase           |

Source: Adapted from Nelson and Cox, 2000.



Reactants to Products






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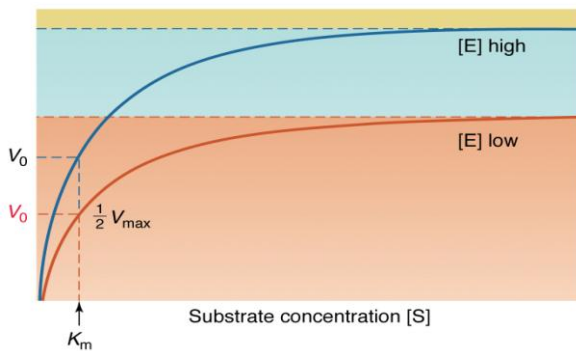
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- Then the Michaelis-Menton equation for a single enzyme  $V_0$  of initial rate varies but can be used for determining other factors.

Key is when  $V_0 = \frac{1}{2} V_{\max}$

Then;  $\frac{V_{\max}}{2} = \frac{V_{\max} [S]}{K_m + [S]}$  will be  $K_m = [S]$

So  $K_m$  can be determined by a plot. In this kind of plot many values are needed for the curve. A linear plot is easier and more accurate.

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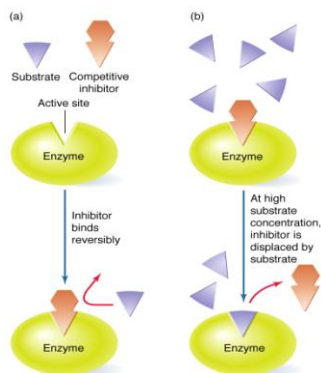
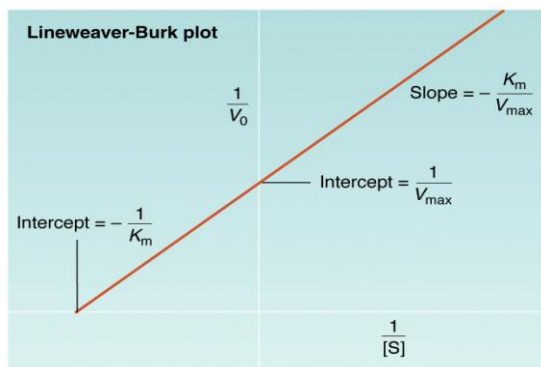
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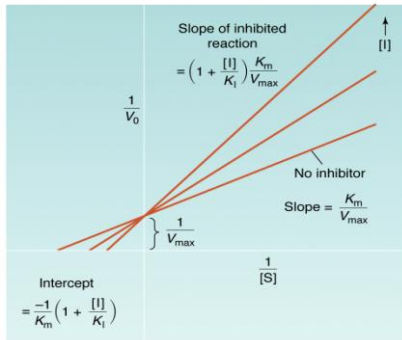
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$$\frac{1}{V_o} = \frac{K_m}{V_{max}[S]} + \frac{1}{V_{max}}$$



(a) Competitive inhibition




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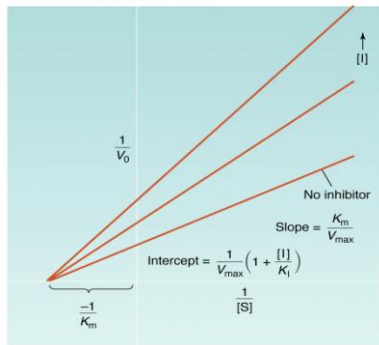
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(b) Noncompetitive inhibition




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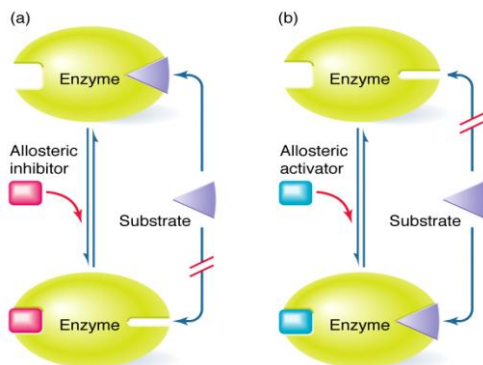
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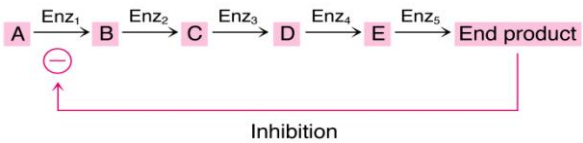
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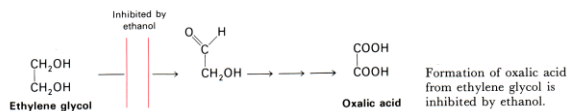
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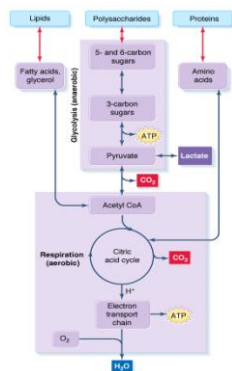
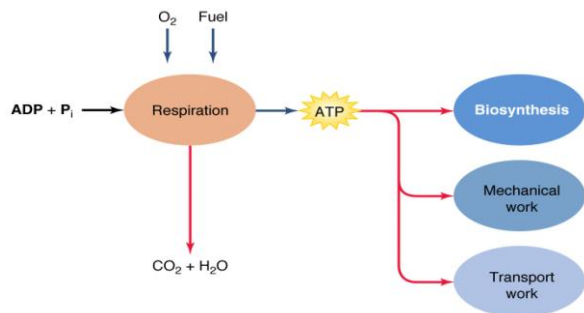
Metabolic Rx in the body must be regulated.  
Not all or none Rx for process like ATP (energy)  
Many Rx regulated by enzymes. They are proteins;  
they are made and broken down.  
These are amino acids so Temp, pH, and salts will  
have an effect on their function.  
Also the cell does not want too much excess protein  
around so make Enzymes as needed.



Ex: The neurotransmitter/hormone: Norepinephrine can inhibit  
tyrosine hydroxylase which is an enzyme that makes Norepi.



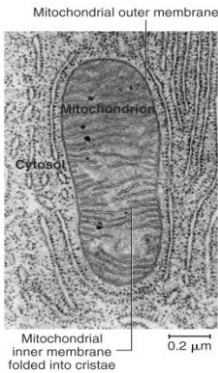
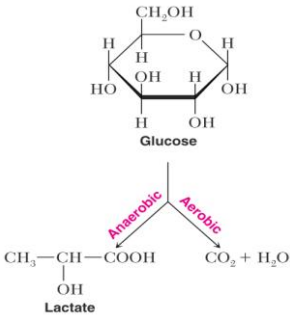
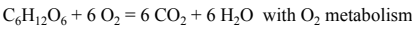
Energy within the cells used to drive the tissue- to organ- to system.



2 ATP

36 ATP (?)

Total 38 ATP (?)



Inner membrane ETC and most ATP is produced

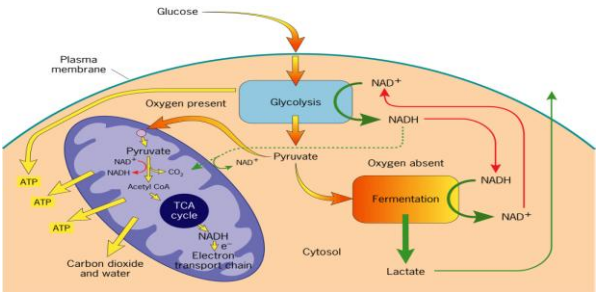


Figure 5.5

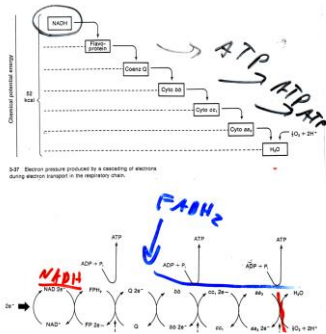
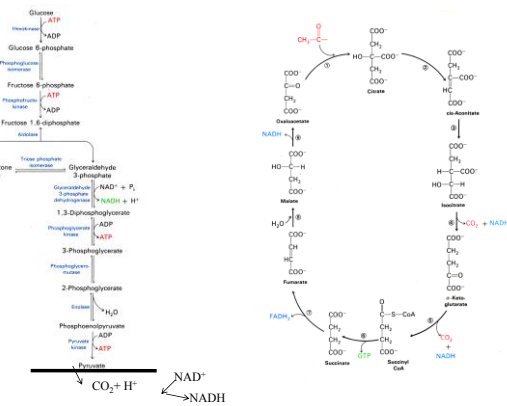


Table 14-4  
ATP yield from the complete oxidation of glucose

| Reaction sequence   | ATP yield per glucose |
|---|-----------------------|
| <b>Glycolysis: glucose into pyruvate (in the cytosol)</b>   |                       |
| Phosphorylation of glucose  | -1                    |
| Phosphorylation of fructose 6-phosphate   | -1                    |
| Dephosphorylation of 2 molecules of 1,3-DPG   | +2                    |
| Dephosphorylation of 2 molecules of phosphoenolpyruvate   | +2                    |
| 2 NADH are formed in the oxidation of 2 molecules of glyceraldehyde 3-phosphate                               |                       |
| <b>Conversion of pyruvate into acetyl CoA (inside mitochondria)</b>   |                       |
| 2 NADH are formed   |                       |
| <b>Citric acid cycle (inside mitochondria)</b>  |                       |
| 2 molecules of guanosine triphosphate are formed from 2 molecules of succinyl CoA                             | +2                    |
| 6 NADH are formed in the oxidation of 2 molecules each of succinate, α-ketoglutarate, and malate              |                       |
| 2 FADH <sub>2</sub> are formed in the oxidation of 2 molecules of succinate                                   |                       |
| <b>Oxidative phosphorylation (inside mitochondria)</b>  |                       |
| 2 NADH formed in glycolysis, each yields 2 ATP (assuming transport of NADH by the glycerol phosphate shuttle) | +4                    |
| 2 NADH formed in the oxidative decarboxylation of pyruvate; each yields 3 ATP                                 | +6                    |
| 2 FADH <sub>2</sub> formed in the citric acid cycle; each yields 2 ATP  | +4                    |
| 6 NADH formed in the citric acid cycle; each yields 3 ATP   | +18                   |
| <b>NET YIELD PER GLUCOSE</b>  | <b>+38</b>            |

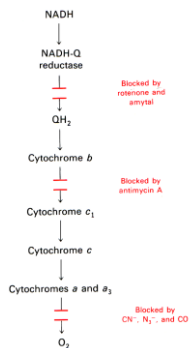
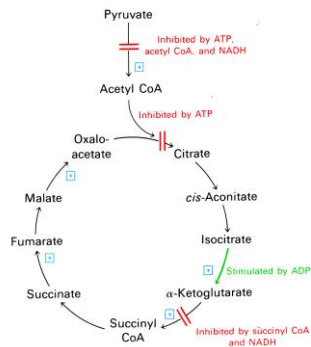


### TCA and ETC:

| Molecule   | ATP worth | Total |
|--|-----------|-------|
| 4 NADH   | 3         | 12    |
| 1 FADH <sub>2</sub>                                      | 2         | 2     |
| (Above, I wrote 2 FADH <sub>2</sub> in class by mistake) |           |       |
| 1 GTP  | 1         | 1     |
|  |           | 15    |
| X2 TCA cycles =30 ATP                                    |           |       |

### Glycolysis:

|  |                           |
|--|---------------------------|
| Gly Net ATP                              | 2                         |
| 2 NADH in Gly (3 ATP/each)               | 32 Total                  |
|  | 6                         |
|  | 38 Total                  |
| But use 2 ATP for the transport into Mit | -2 (1 ATP for each NADH)  |
|  | <b>36 ATP Grand total</b> |



Endosymbiotic theory  
Lynn Margulis

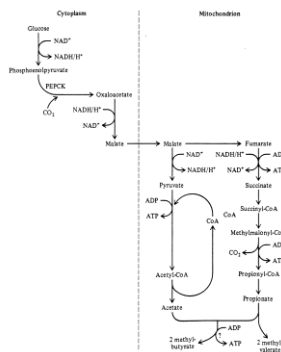
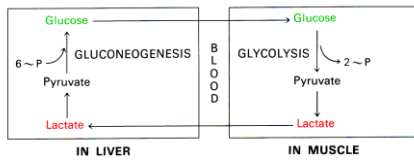
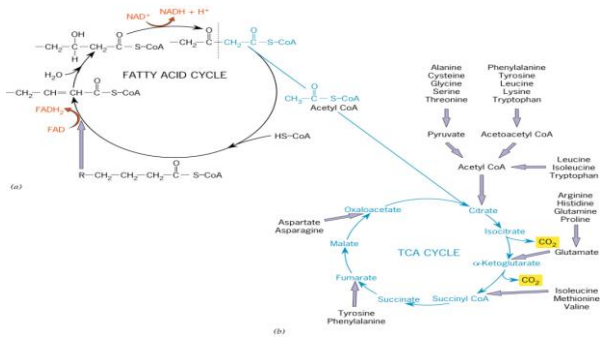


FIGURE 3-18 Linkage of glycolysis with citric acid cycle pathways in an anaerobic metabolic pathway that provides additional ATP formation. These pathways are found, for example, in many platyhelminth worms.

Anaerobic- Bacteria, some yeasts, some invertebrates can live without  $O_2$ .

Ex. Clostridium botulinum can not grow in  $O_2$ .

Aerobic- require a supply of  $O_2$ . Some tissues like muscle can function anaerobically and build up an “ $O_2$  debt” but pay back occurs.

With  $O_2$  the cells are 20 times more efficient to produce ATP.

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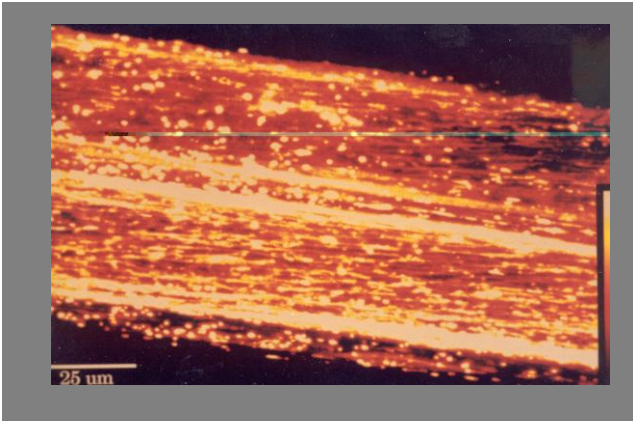
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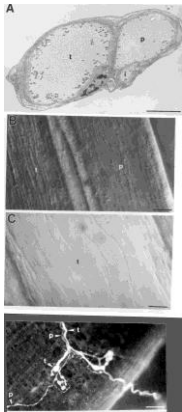
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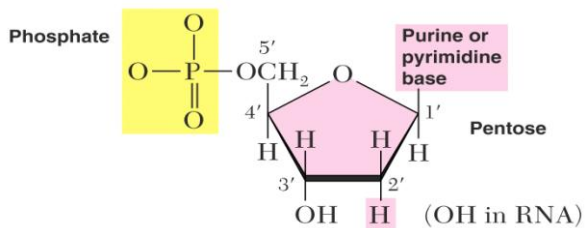
## Homework problems

### Competitive

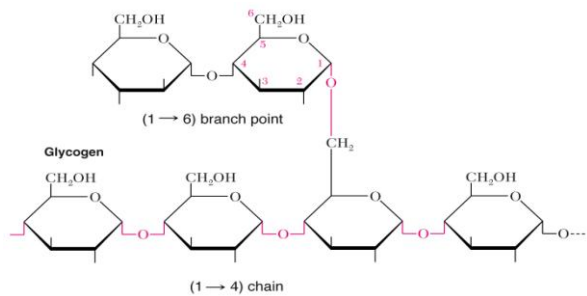
1. Binds at active site
2. Increase [S] removes inhibition
3. Increase [I] more block till saturated  
(alter rate and Increase  $K_m$ , without a change on  $V_{max}$ )
4. Slope changes

### Non-Competitive

1. Dose not binds at active site
2. Increase [S] no effect
3. Increase [I] more block till saturated  
(no change on  $K_m$ , but decrease  $V_{max}$ )
4. Slope changes








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