

BioChem 330 - Course Outline

October 4-6, 2011

- **Bio-molecular Structure/Function (I cont'd)**

- NUCLEIC ACID

- DNA sequence and structure
- Protein/nucleic acid interactions

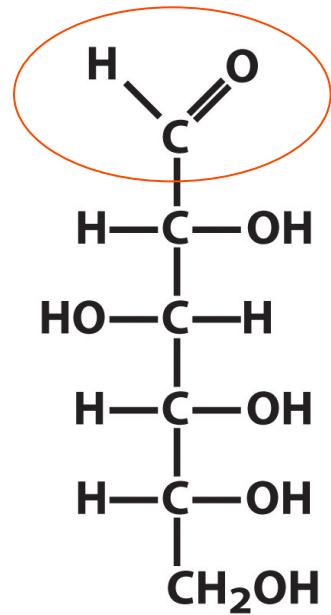
- CARBOHYDRATES

- Sugars - mono and disaccharides
- Polysaccharides
- Glycerides and glycerol

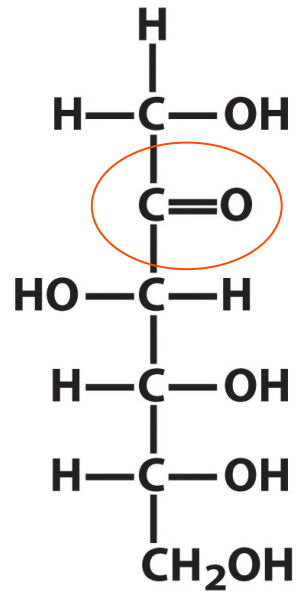
- FATS AND LIPIDS

- Chemistry and nomenclature for fatty acids
- Saturated and unsaturated fatty acids
- Fluid mosaic model of membrane structure

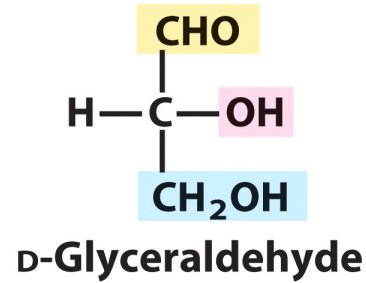
sugar chemistry



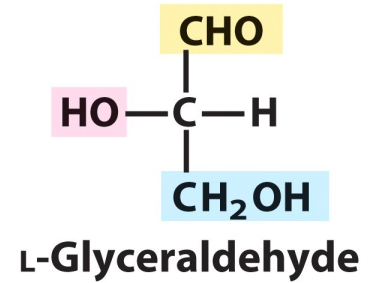
**D-Glucose,
an aldohexose**



**D-Fructose,
a ketohexose**

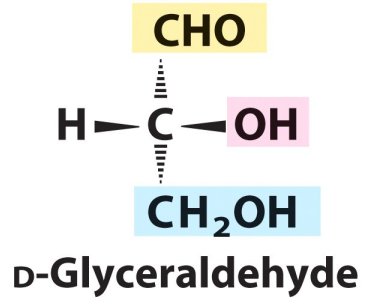


D-Glyceraldehyde

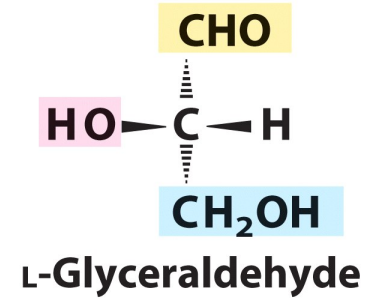


L-Glyceraldehyde

Fischer projection formulas



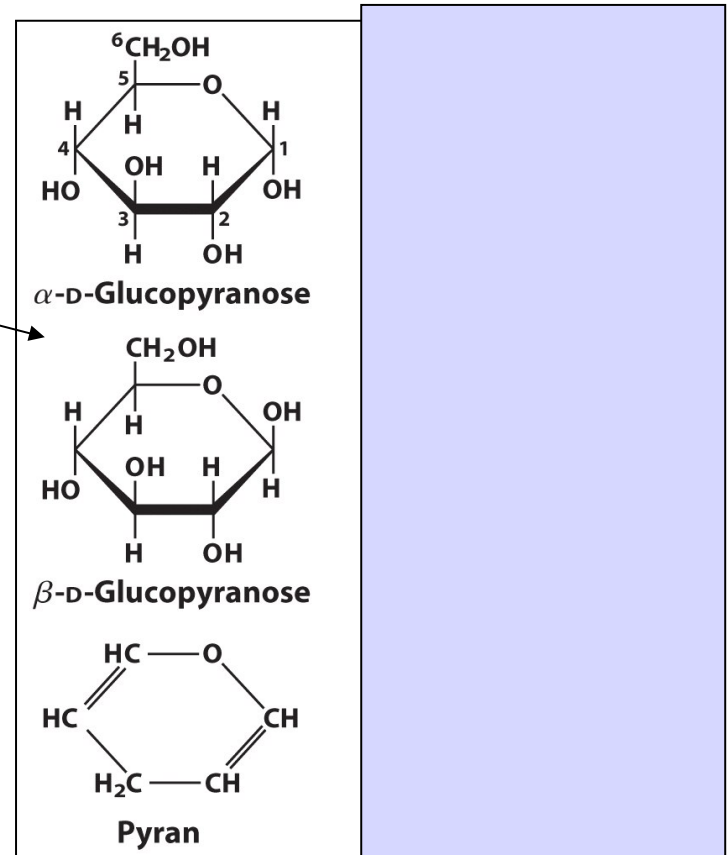
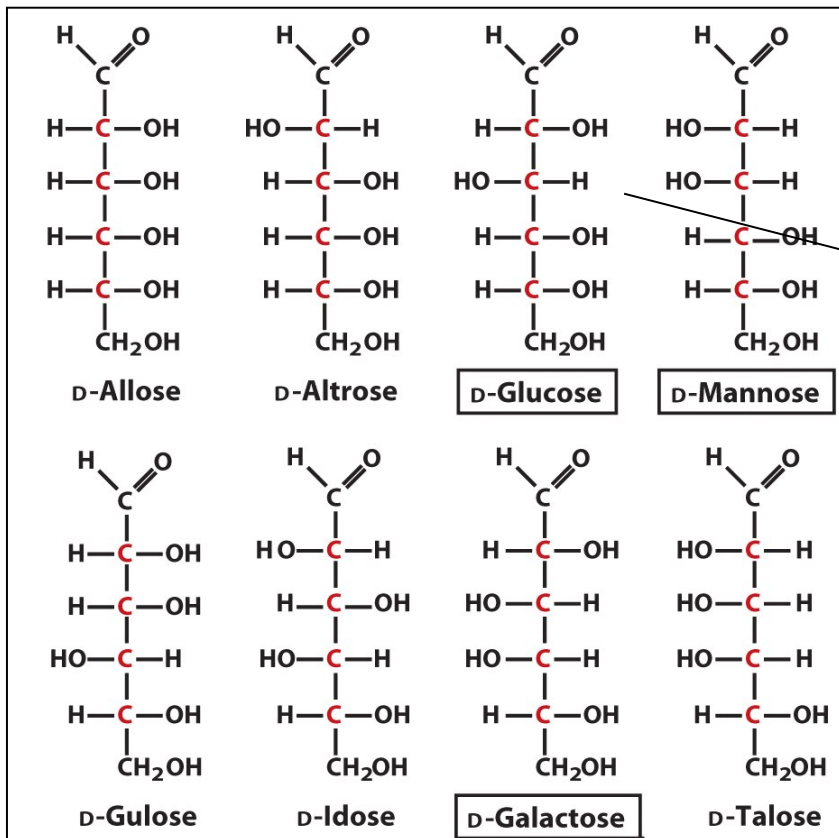
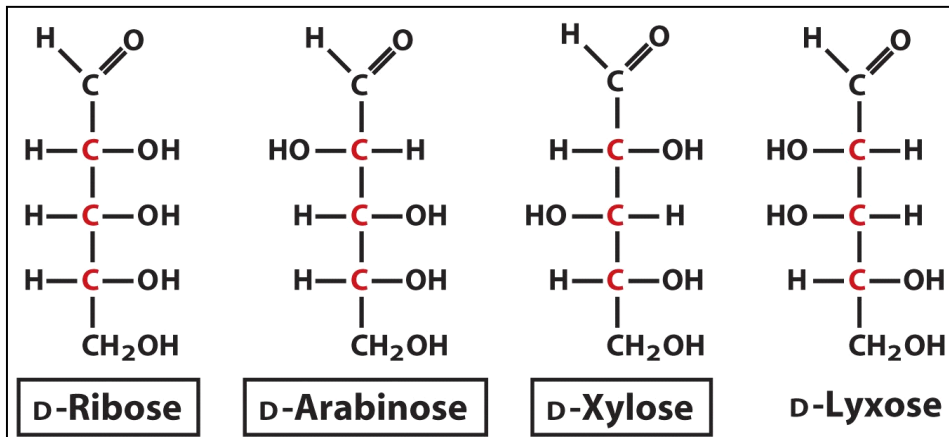
D-Glyceraldehyde



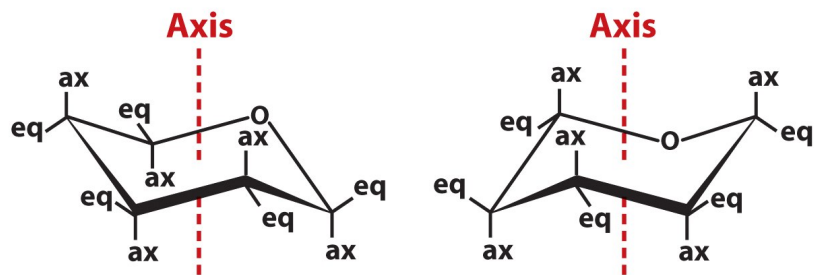
L-Glyceraldehyde

Perspective formulas

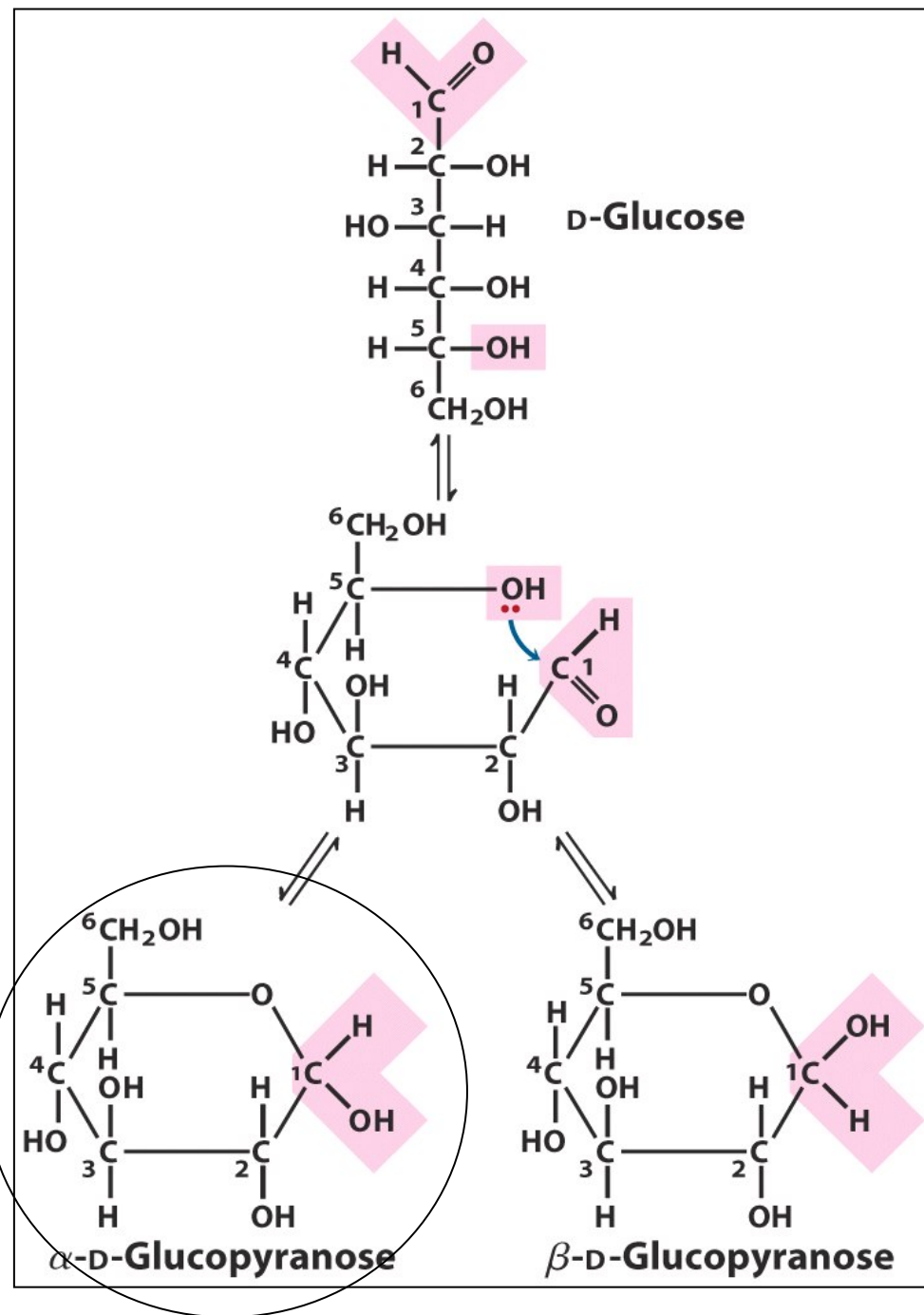
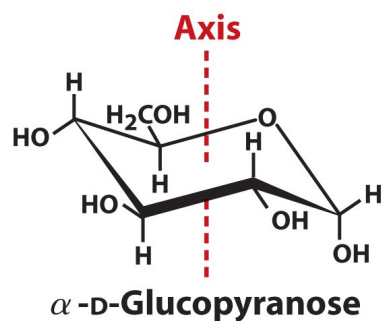
Five and Six Carbon Aldose Sugars important in Biology

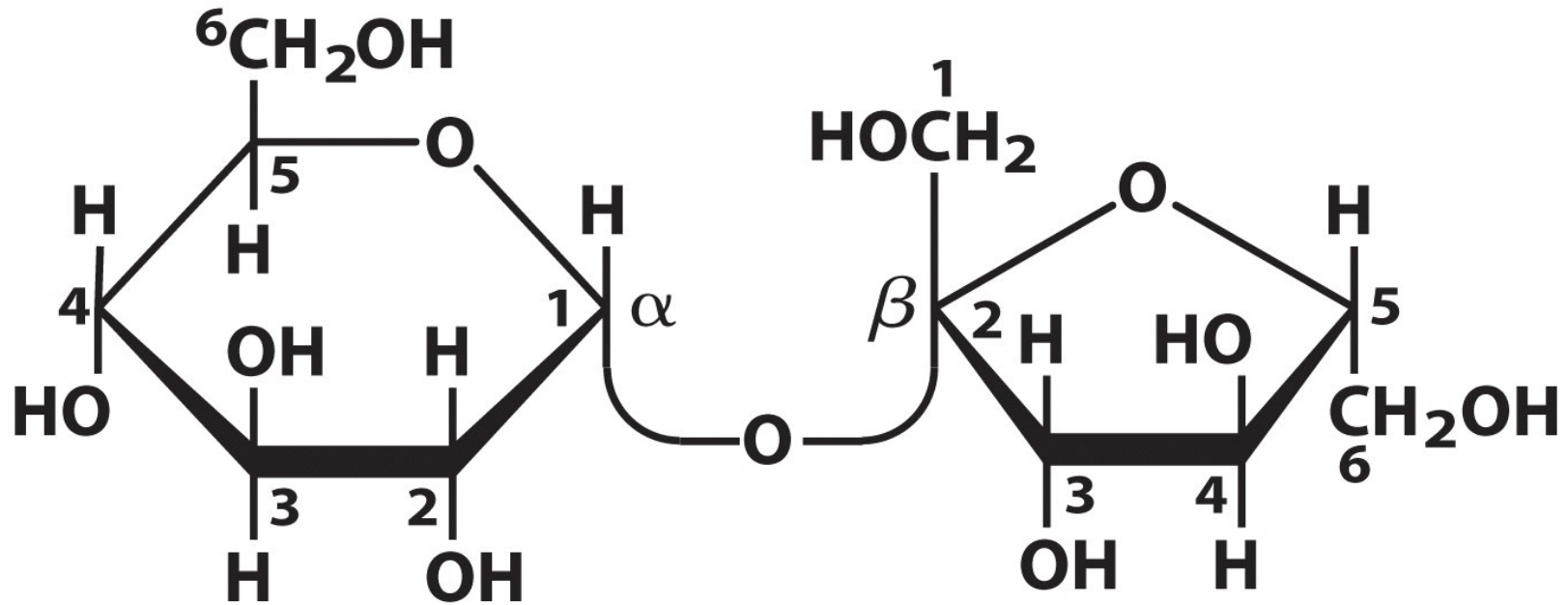


Cyclization of six carbon sugar to form α or β pyranose



Two possible chair forms





Sucrose

α -D-glucopyranosyl β -D-fructofuranoside
Glc(α 1 \leftrightarrow 2 β)Fru

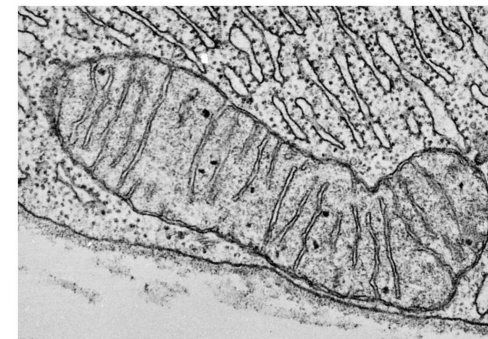
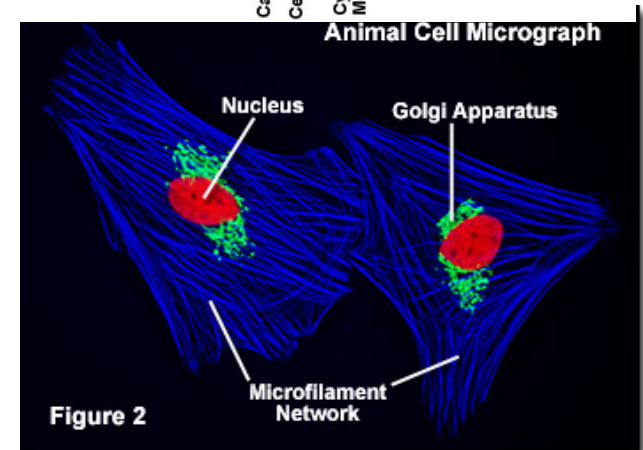
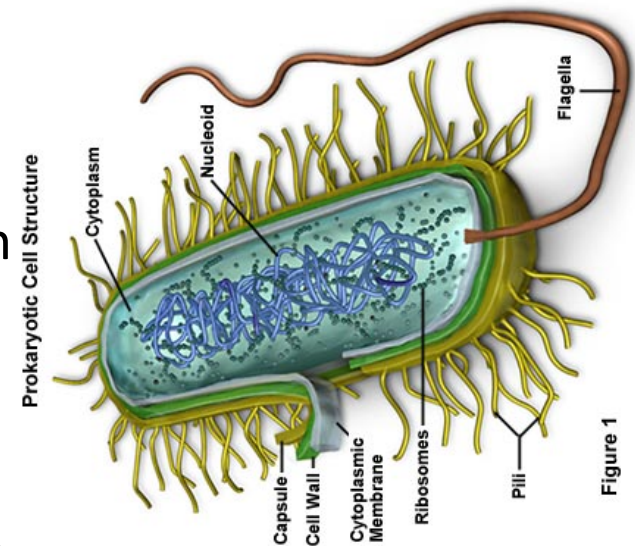
Sugars are frequently packaged as complex carbohydrates, here shown as the disaccharide sucrose, but also as polysaccharides such as glycogen (D-glucose α 1 \rightarrow 4 linkages plus occasional branches α 1 \rightarrow 6) or cellulose (D-glucose β 1 \rightarrow 4)

Job Description for a Biological Membrane

- **Control the ionic composition** of the cytoplasm by highly specific transporters (act as a barrier to diffusion)
- **Form a dynamic matrix** for enzymatic reactions, receptor processes and immunological recognition (act as a surface)
- **Guarantee the integrity** of the cell and influence its shape and movement as well as the displacement of organelles (act as a mechanical structure)
- **Contain a mosaic of various passive and active electrical devices**, control the potential at and near the membrane, control electrodynamic conditions (act as an electrically isolating leaflet)

Barriers to the world outside

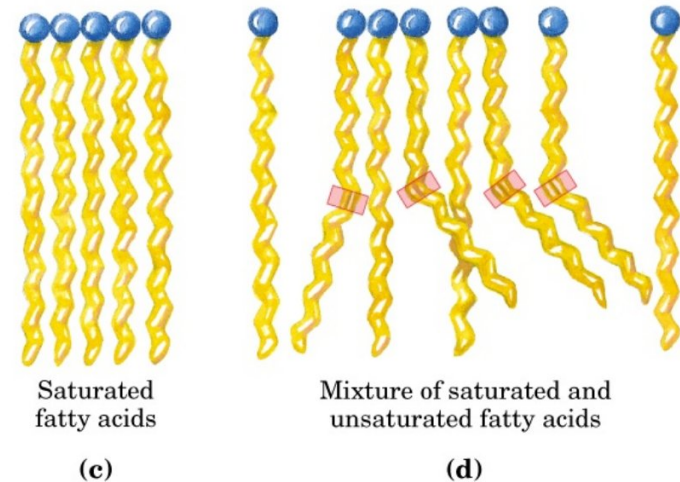
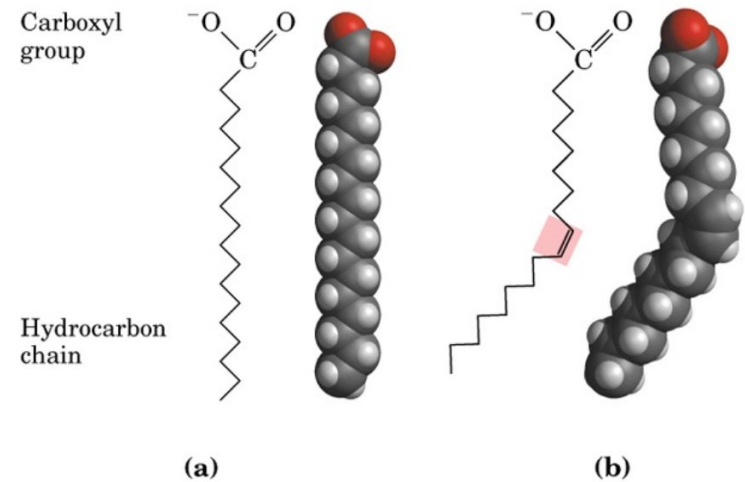
- Prokaryotes lack inner organelles, but have much more complicated outer walls, with capsules, cell walls that can be several layers thick and a cytoplasmic membrane (1 micron)
- Eukaryotic cells can be larger (10 microns-0.1 m) and the lipid bilayer is the plasma membrane that separates inside from outside. Inside are organelles many of which such as the nucleus, also have their own lipid bilayers. Plant cells have chloroplasts and an outer wall
- Mitochondrion organelles within the eukaryotic cells have a double wall. These inner and outer membranes, vestigial remnant of their former prokaryotic origin (electron micrograph at right)



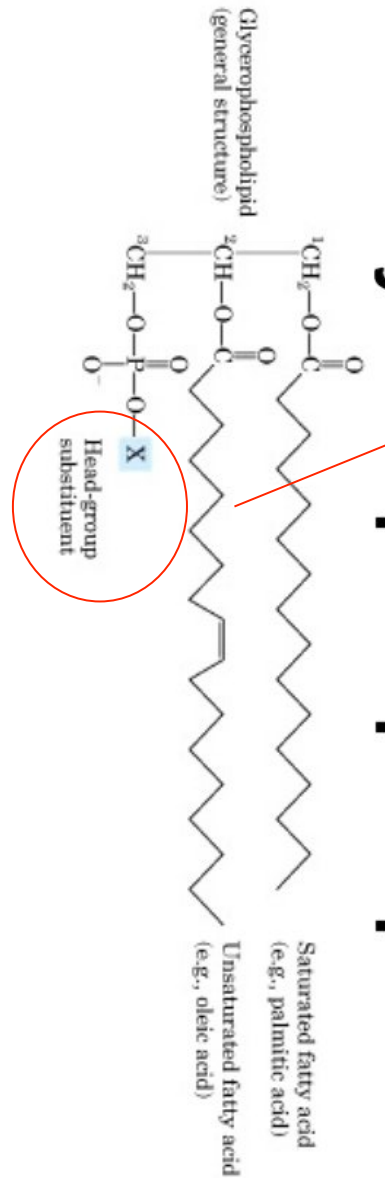
<http://www.molecularexpressions.com/cells>

Fatty acids self assemble into bilayers and micelles

- Polar head group (acid component) orients towards aqueous phase
- Non-polar tail (long hydrocarbon chain) orients towards inside (micelle) or towards other layer in a bilayer (shown)



Biological Membranes are made up of Phospholipids



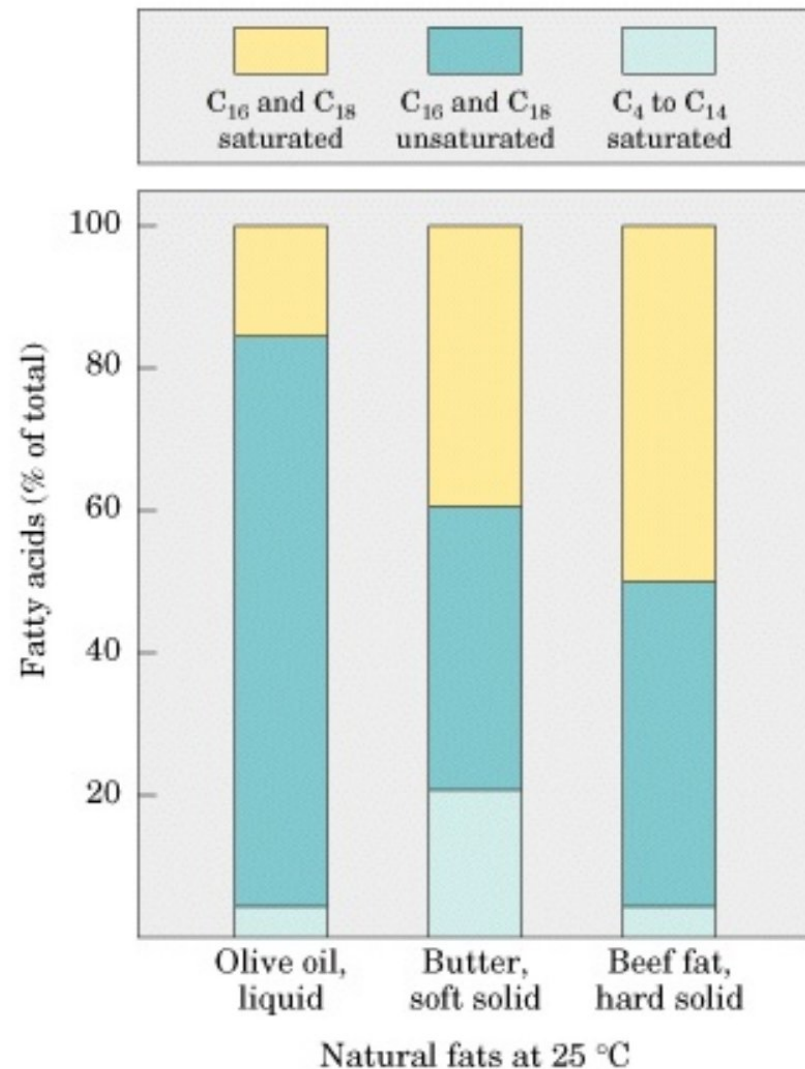
Name of glycerophospholipid	Name of X	Formula of X	Net charge (at pH 7)
Phosphatidic acid	—	— H	-1
Phosphatidylethanolamine	Ethanolamine	— CH ₂ —CH ₂ —NH ₃ ⁺	0
Phosphatidylcholine	Choline	— CH ₂ —CH ₂ —N ⁺ (CH ₃) ₃	0
Phosphatidylserine	Serine	— CH ₂ —CH—NH ₃ ⁺ COO ⁻	-1
Phosphatidylglycerol	Glycerol	— CH ₂ —CH—CH ₂ —OH OH	-1
Phosphatidylinositol 4,5-bisphosphate	<i>myo</i> -Inositol 4,5-bisphosphate		-4
Cardiolipin	Phosphatidyl-glycerol	$ \begin{array}{c} \text{— CH}_2 \\ \\ \text{CHOH} \\ \\ \text{CH}_2\text{—O—P—O—CH}_2 \\ \quad \\ \text{O} \quad \text{O} \\ \quad \quad \\ \quad \quad \text{CH—O—C—R}^1 \\ \quad \quad \\ \quad \quad \text{O} \\ \quad \quad \\ \quad \quad \text{CH}_2\text{—O—C—R}^2 \\ \quad \quad \\ \quad \quad \text{O} \end{array} $	-2

Common fatty acid components of biological membranes

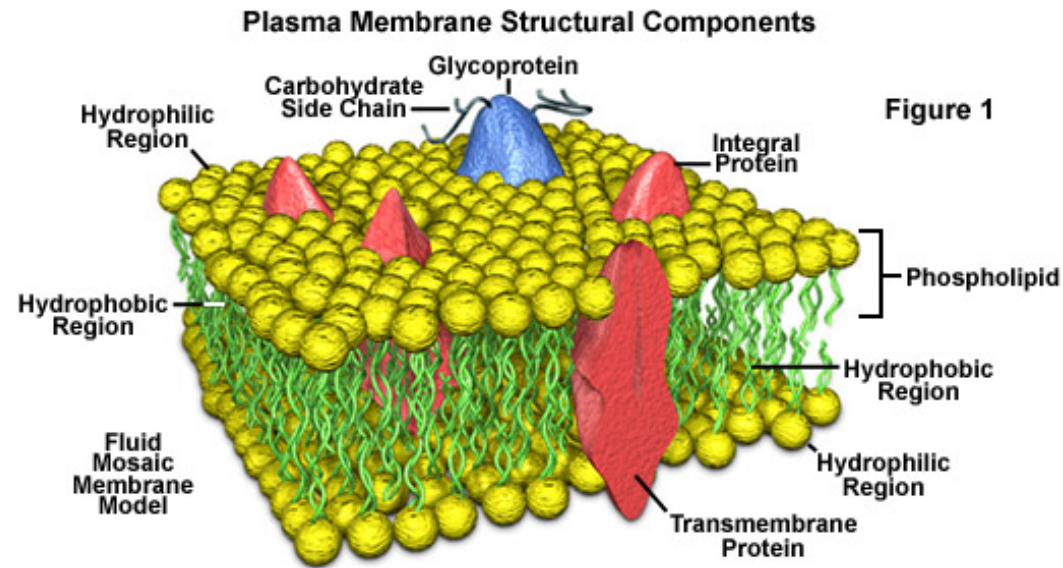
Common Name	Systematic Name	Abbreviation	Structure	Melting Point (°C)
Saturated Fatty Acids				
Capric	<i>n</i> -Decanoic	10:0	CH ₃ (CH ₂) ₈ COOH	31.6
Lauric	<i>n</i> -Dodecanoic	12:0	CH ₃ (CH ₂) ₁₀ COOH	44.2
Myristic	<i>n</i> -Tetradecanoic	14:0	CH ₃ (CH ₂) ₁₂ COOH	53.9
Palmitic	<i>n</i> -Hexadecanoic	16:0	CH ₃ (CH ₂) ₁₄ COOH	63.1
Stearic	<i>n</i> -Octadecanoic	18:0	CH ₃ (CH ₂) ₁₆ COOH	69.6
Arachidic	<i>n</i> -Eicosanoic	20:0	CH ₃ (CH ₂) ₁₈ COOH	76.5
Behenic	<i>n</i> -Docosanoic	22:0	CH ₃ (CH ₂) ₂₀ COOH	81.5
Lignoceric	<i>n</i> -Tetracosanoic	24:0	CH ₃ (CH ₂) ₂₂ COOH	86.0
Cerotic	<i>n</i> -Hexacosanoic	26:0	CH ₃ (CH ₂) ₂₄ COOH	88.5
Unsaturated Fatty Acids				
Palmitoleic	<i>cis</i> -9-Hexadecenoic	16:1cΔ9	CH ₃ (CH ₂) ₅ CH=CH(CH ₂) ₇ COOH	0
Oleic	<i>cis</i> -9-Octadecenoic	18:1cΔ9	CH ₃ (CH ₂) ₇ CH=CH(CH ₂) ₇ COOH	16
Linoleic	<i>cis,cis</i> -9,12-Octadecadienoic	18:2cΔ9,12	CH ₃ (CH ₂) ₄ CH=CHCH ₂ CH=CH(CH ₂) ₇ COOH	5
Linolenic	all- <i>cis</i> -9,12,15-Octadecatrienoic	18:3cΔ9,12,15	CH ₃ CH ₂ CH=CHCH ₂ CH=CHCH ₂ CH=CH(CH ₂) ₇ COOH	-11
Arachidonic	all- <i>cis</i> -5,8,11,14-Eicosatetraenoic	20:4cΔ5,8,11,14	CH ₃ (CH ₂) ₄ CH=CHCH ₂ CH=CHCH ₂ CH=CHCH ₂ CH=CH(CH ₂) ₃ COOH	-50
Branched and Cyclic Acids				
Tuberculostearic	<i>I</i> -D-10-Methyloctadecanoic		$\begin{array}{c} \text{CH}_3 \\ \\ \text{CH}_3(\text{CH}_2)_7\text{CH}(\text{CH}_2)_8\text{COOH} \end{array}$	13.2
Lactobacillic	ω -(2- <i>n</i> -Octylcyclopropyl)-octadecanoic		$\begin{array}{c} \text{CH}_2 \\ \diagdown \\ \text{CH}_3(\text{CH}_2)_5\text{CH}-\text{CH}(\text{CH}_2)_9\text{COOH} \end{array}$	29

Fatty acid compositions of fats

- Saturated (no double bonds)
 - Tend to form more rigid structures because hydrocarbon chains can be closely packed
 - Example animal fats (crisco, butter)
- Unsaturated (one or more double bonds)
 - Tend to form less rigid structures (higher T_m) because kinks due to double bonds resist packing
 - Examples plant fats (olive oil)
 - Can be cis or trans fats, **cis** fats can be more easily metabolized by fatty acid machinery in cell



Membranes: fluid mosaic model



- Fluid mosaic model –

- two dimensional lipid bilayer with random motion within layers, minimal diffusion between layers http://www.youtube.com/watch?v=Qqsf_UJcBc (Singer and Nicholson 1972)
- Embedded proteins can be integral or transmembrane, and many of them are glycosylated (that is they have complex carbohydrate groups added to the protein side chains).
- Hydrophobic substances such as cholesterol further add rigidity to the membrane

Membrane Deformations

- A. Translation and Rotation
- B. Planar extension
- C. Shear deformation
- D. Translocation of membrane proteins by spectrin network
- E. Mending of membrane by asymmetric lipids or proteins
- F. Extrusion of membrane components with lower flexibility from bended areas

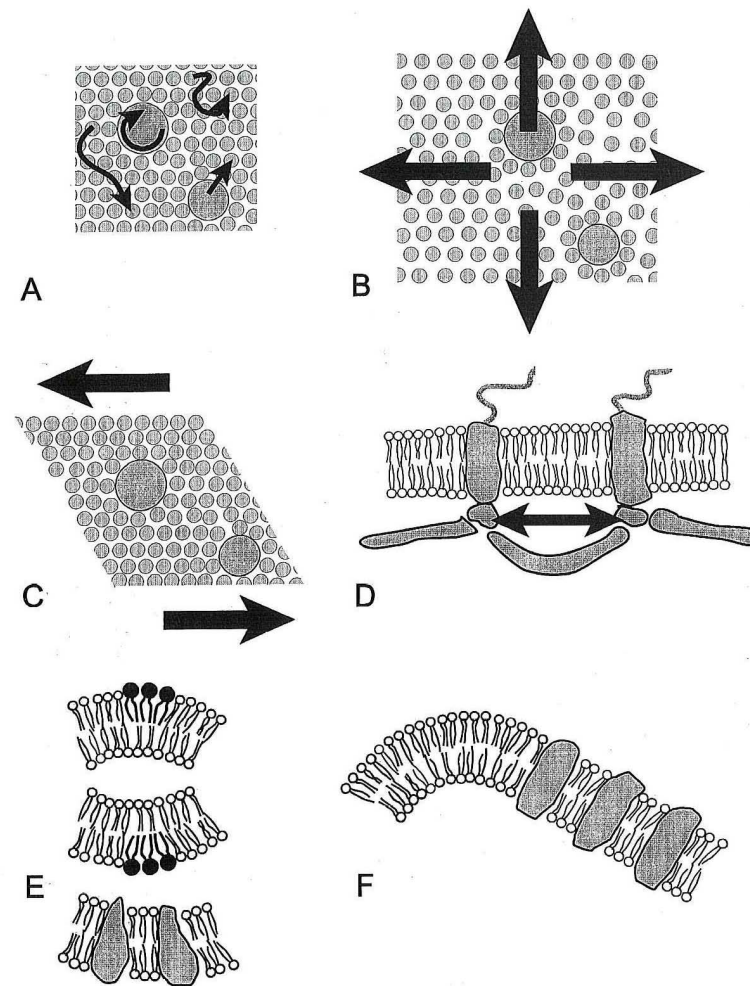


Figure 2.41 Glaser p 89

Membrane Deformations

- Rapid lateral (within surface) diffusion for lipids (us to ms)
- Proteins (attached to cytoskeletal components) travel slowly or not at all
- Slower transverse diffusion between surfaces (min to hours) but can be assisted by lipid exchange systems

Glaser p 89

Easy deformation by sheering interactions (area is conserved)

Nearly impossible to extend the area of bilayer

Elasticity measured by property Y (Young's modulus)

Rubber: $Y = 1$ MPa

Erythrocytes: $Y = 56$ MPa

Bone: $Y = 10,000$ MPa

1-2 % stretching breaks cell, cell lysis by swelling common first step in protein purification

Review

Dynamics in the plasma membrane: how to combine fluidity and order

Didier Marguet^{1,2,3,*}, Pierre-François
Lenne^{4,5}, Hervé Rigneault^{4,5} and
Hai-Tao He^{1,2,3}

¹Centre d'Immunologie de Marseille-Luminy, Université de la Méditerranée, Marseille, France, ²INSERM, UMR 631, Marseille, France, ³CNRS, UMR 6102, Marseille, France, ⁴Institut Fresnel, MOSAIC group, Université Paul Cézanne, Marseille, France and ⁵CNRS UMR 6133, Marseille, France

Cell membranes are fascinating supramolecular aggregates that not only form a barrier between compartments but also harbor many chemical reactions essential to the existence and functioning of a cell. Here, it is proposed to review the molecular dynamics and mosaic organization of the plasma membrane, which are thought to have important functional implications. We will first summarize the basic concepts of Brownian diffusion and lipid domain formation in model membranes and then track the development of ideas and tools in this field, outlining key results obtained on the dynamic processes at work in membrane structure and assembly. We will focus in particular on findings made using fluorescent labeling and imaging procedures to record these dynamic processes. We will also discuss a few examples showing the impact of lateral diffusion on cell signal transduction, and outline some future methodological challenges which must be met before we can answer some of the questions arising in this field of research.

The EMBO Journal (2006) 25, 3446–3457. doi:10.1038/sj.emboj.7601204; Published online 22 June 2006

Subject Categories: membranes & transport

Keywords: actin meshwork; cell membrane; fluorescence microscopy; lateral diffusion; membrane microdomain
