Membranes 1

Steven E. Massey, Ph.D. Assistant Professor Department of Biology University of Puerto Rico – Río Piedras

Office & Lab: NCN#343B Tel: 787-764-0000 ext. 7798 E-mail: stevenemassey@gmail.com



Membranes form cells and compartmentalize the components of the cell in eukaryotes. They are composed of lipids and form a bilayer

This bilayer is self assembling

They are selectively permeable and flexible, allowing growth and movement, fusion and fission

Membranes are impermeable to polar compounds and permeable to nonpolar compounds

Transport across the membrane occurs using protein transporters



When phospholipids come into contact with water they will self assemble into various assemblages, depending on the conditions:



There is a commonly accepted idea that the first 'cells' emerged from the primordial soup and were self – assembling

This idea proposes that life is an emergent property

Emergence is defined as where the whole is greater than the sum of the parts or order out of disorder

Other emergent properties in biology include flocks of birds, schools of fish, human consciousness, bee hives, human society.....



Membranes have different compositions, depending on tissue type and subcellular organelle eg. the mitochondrial membrane is different from the cell membrane

Subcellular components may be isolated by differential centrifugation, otherwise known as subcellular fractionation

Membranes also differ in their protein composition

TABLE 11–1 Major Components of Plasma Membranes in Various Organisms									
Components (% by weight)									
	Protein	Phospholipid	Sterol	Sterol type	Other lipids				
Human myelin sheath	30	30	19	Cholesterol	Galactolipids, plasmalogens				
Mouse liver	45	27	25	Cholesterol	—				
Maize leaf	47	26	7	Sitosterol	Galactolipids				
Yeast	52	7	4	Ergosterol	Triacylglycerols, steryl esters				
Paramecium (ciliated protist)	56	40	4	Stigmasterol	—				
E. coli	75	25	0	—	—				

Note: Values do not add up to 100% in every case, because there are components other than protein, phospholipids, and sterol; plants, for example, have high levels of glycolipids.

The fluid mosaic model implies that the proteins move laterally

They are embedded in the lipid bilayer and are stabilized by hydrophobic interactions



Membrane proteins may be integral, peripheral or amphitropic

Integral membrane proteins have a hydrophobic component or have a covalent bond

Peripheral membrane proteins form electrostatic interactions and H bonds with the polar groups of membrane lipids

Amphitropic proteins are found in the cytosol and associated with membranes - may have a noncovalent interaction with a membrane protein or lipid, or lipids may be attached to the protein

The reversible association with the membrane may be regulated by phosphorylation or ligand binding

Integral, peripheral and amphitropic proteins



Many integral membrane proteins are transmembrane, going through the membrane

Glycophorin for example has a single transmembrane domain

This region is made of hydrophobic residues

On the outer surface are oligosaccharides (red and blue hexagons on figure)



<u>The different classes of integral</u> <u>membrane proteins</u>

Type V and VI are held by covalent linkages to lipids



Bacteriorhodopsin is one of the best studied membrane proteins

It is a light driven proton pump found in archaea

It has seven transmembrane alpha helices



Sequences can be used to prediction protein function – this is bioinformatics

A hydropathy plot can be used to predict the regions of the protein sequence that are transmembrane regions

The graph plots the hydrophobicity of an amino acid versus the amino acid number



Charged residues tend to be on the surface of the membrane protein that is exposed to the aqueous environment

Positively charged residues tend to be on the cytoplasmic face of the membrane

This is the positive-inside rule

Distribution of residues in a range of membrane proteins



Transmembrane domains can also be composed of beta barrels



Proteins may be covalently attached to the membrane by covalently bonding to lipids

A glycosyl phosphatidylinositol (GPI) anchor is phosphatidylinositol attached to an oligosaccharide attached to phosphoethanolamine. The phosphoethanolamine is attached to the peptide chain

Man = mannose GlcNAc = acetyl glucosamine



Heat can alter the state of the membrane

At low temperatures it will form a gel

At physiological temperatures it is in a fluid state

(a) Paracrystalline state (gel)

Heat produces thermal motion of side chains (gel \rightarrow fluid transition)



As a consequence, cells regulate their lipid composition according to temperature

TABLE 11–2	Fatty Acid Composition of <i>E. coli</i> Cells Cultured at Different Temperatures						
		Percentage of total fatty acids*					
		10 °C	20 °C	30 °C	40 °C		
Myristic acid (14:0)		4	4	4	8		
Palmitic acid (16:0)		18	25	29	48		
Palmitoleic acid (16:1)		26	24	23	9		
Oleic acid (18:1)		38	34	30	12		
Hydroxymyristic acid		13	10	10	8		
Ratio of unsaturated to saturated [†]		2.9	2.0	1.6	0.38		

Movement of a phospholipid through a membrane can be slow

It may be catalyzed by enzymes called flippases, floppases and scramblases

Flippases: out \rightarrow in Floppases: in \rightarrow out Scramblases: both directions

(a) Uncatalyzed transbilayer ("flip-flop") diffusion



(b) Uncatalyzed lateral diffusion





(c) Catalyzed transbilayer translocations



Flippase (P-type ATPase) moves PE and PS from outer to cytosolic leaflet

ATP

Floppase (ABC transporter) moves phospholipids from cytosolic to outer leaflet Scramblase moves lipids in either direction, toward equilibrium

Outside

Inside

Fluorescence Recovery After Photobleaching

(FRAP)

is a method that is used to measure the lateral diffusion of lipids



The membrane is made up of distinct domains, and lipids are restricted in each domain, and then jump by a process called 'hop diffusion'



Hop diffusion

Atomic force microscopy (AFM) can be used to visualize membranes

A microscopic probe will detect peaks and valleys on the membrane surface





bacteriorhodopsin



E.coli aquaporin

Chloroplast ATP synthase



Membrane curvature and fusion are important in many biological processes

There are three mechanisms for inducing curvature in membranes

(as follows.....)

A protein with intrinsic curvature on its surface interacts strongly with a curved membrane surface, allowing both membrane and protein to achieve their lowest energy.

If a membrane region spontaneously curves, monomeric subunits of certain proteins can polymerize into a superstructure that favors and maintains the curvature.

A protein with one or more amphipathic helices inserted into one leaflet of the bilayer crowds the lipids in that leaflet, forcing the membrane to bend. Fusion proteins mediate the fusion of membranes

Membrane fusion occurs at synapses when neurotransmitters are released from vesicles

This is mediated by v-SNAREs and t-SNAREs

