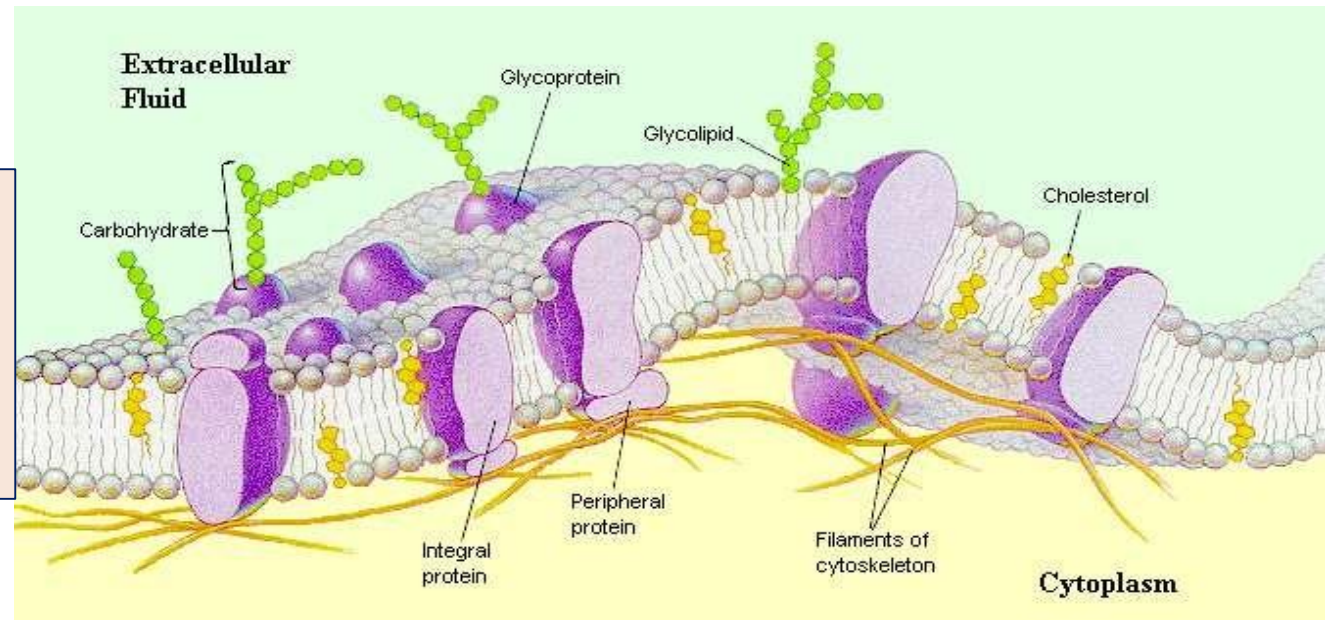


# Cell membranes

Plasma membrane consists of :  
cholesterol, phospholipid, sphingolipid and protein.

- The membrane is hypothesized in a model known as the fluid mosaic model. **Mainly on extra cellular surface**
- Components: 45% lipid, 45% protein and 10% carbohydrate
- They exist side by side without forming some other substance of intermediate nature.

This membrane is being in the membranes in the cell (structural perspective ) like in lysosome and nucleus . The difference is how this component is distributed ?? The inner mitochondrial membrane has much protein Than plasma membrane ,almost 75% protein and 25%lipid.





# Phospholipids

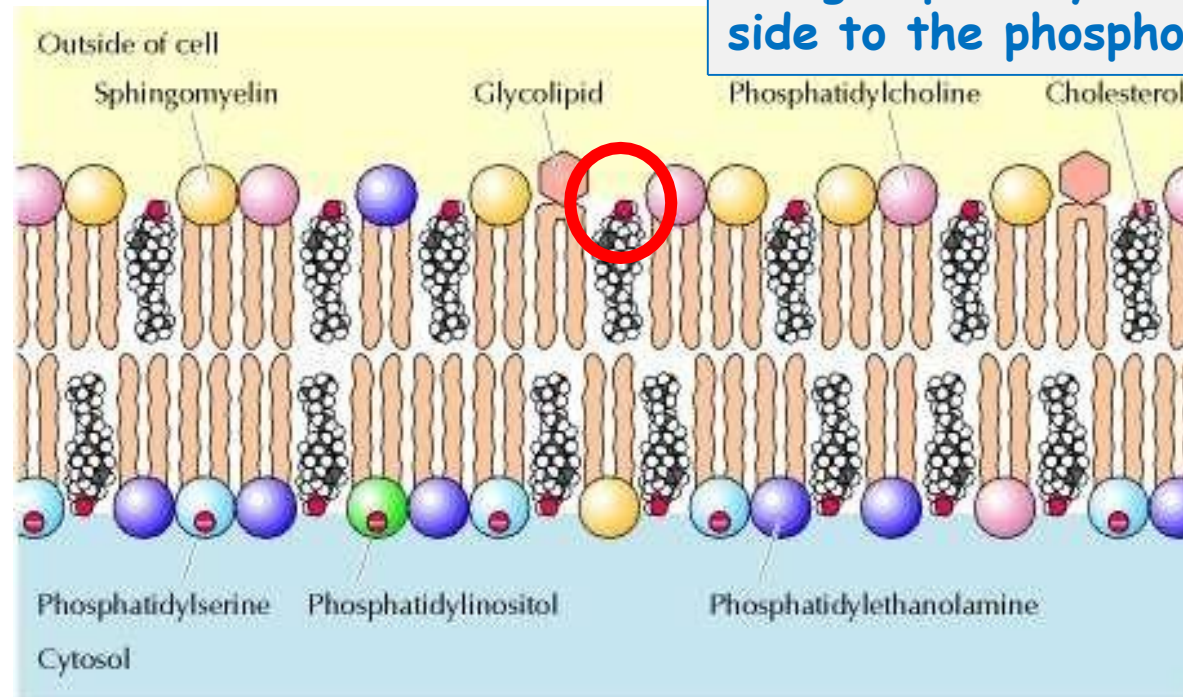
If there a lot of any component in the outer membrane that doesn't mean it not be in the inner membrane

- The outer: phosphatidylcholine, sphingomyelin, and glycolipids (cell recognition)
- The inner: phosphatidylethanolamine, phosphatidylserine, and phosphatidylinositol (signaling)

**Cholesterol is distributed in both leaflets**

**Animal cells vs. plant cells vs. prokaryotic cells**

OH group always exposed outside side by side to the phospholipid head





phosphatidylinositol is very important in signaling by producing (diacylglycerol and IP3 ) which transmit signals inside cell  
Cholesterol has equally distribution on both surfaces (sides) of the cell membrane .

Which molecules are abundant in the outer membrane ?  
Which molecules are abundant in the inner leaflet ?  
(in the previous slide )

The function of glycolipid (in the outer ) :cellular communication ,signaling and identification of the cells .

The most abundant type in the outer is :  
phosphatidylcholine



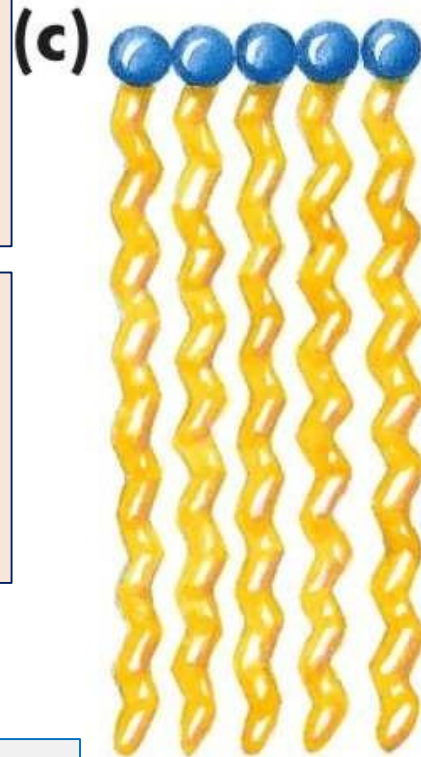
# Fatty acids and membrane fluidity



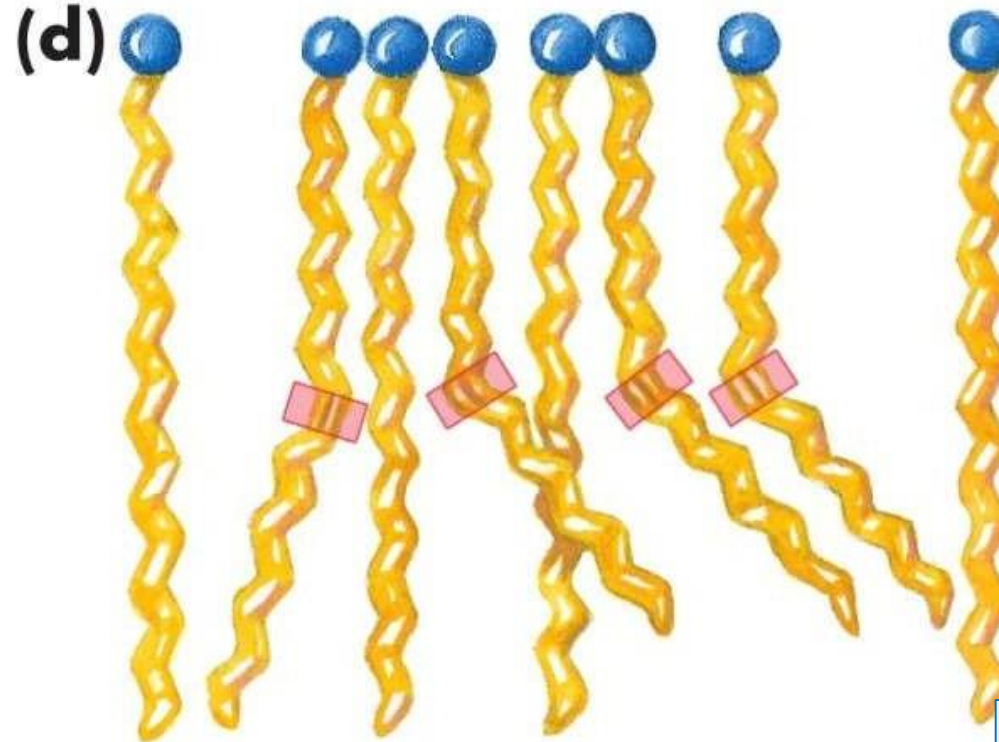
Sphingolipid is more Rigid than phospholipid

Saturated f.a = will be aligned, compacted and more rigidity of membrane

Rigid because it is saturated --> no double bonds present



**Saturated fatty acids**



**Mixture of saturated and unsaturated fatty acids**

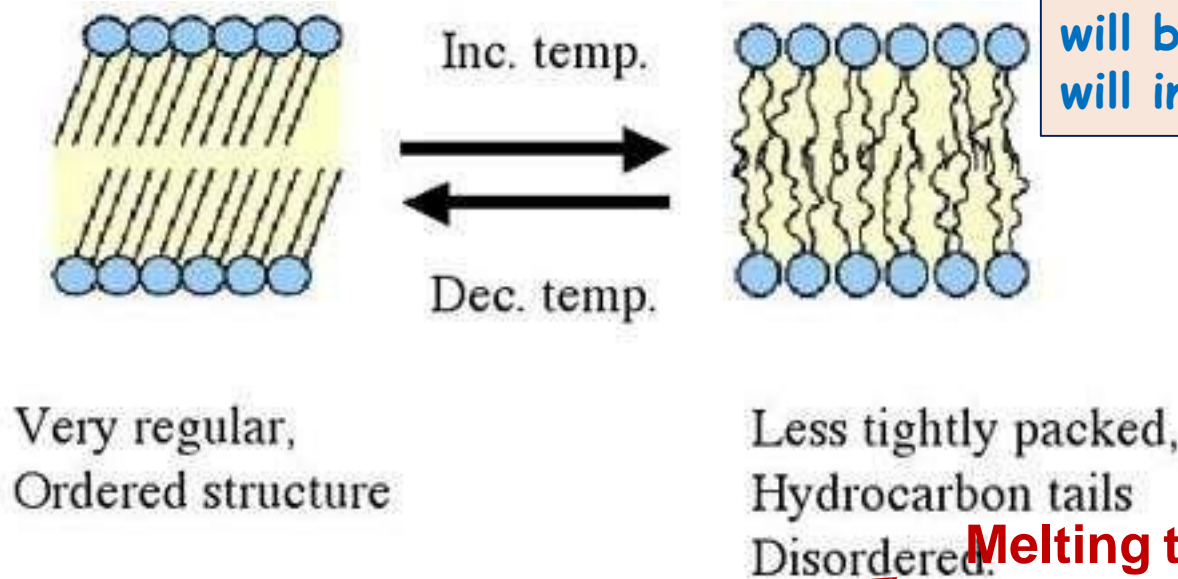
The more the dents, the more the more fluid.

More fluidic membrane because it has unsaturated fatty acids which is very important for signaling molecules

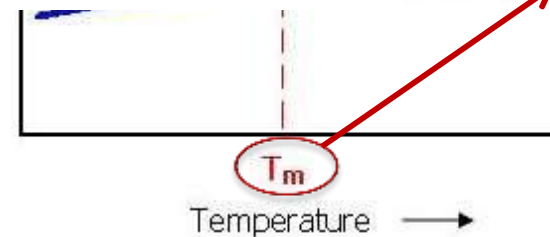
# Membrane fluidity and temperature



Low temperature --> more rigid membrane  
When we raise the temperature, the electrons will move in higher speed-->unstable membrane



Increase temperature will give energy, this energy will be converted to kinetic energy. As much as moving more, they will be more separated and this will increase the fluidity

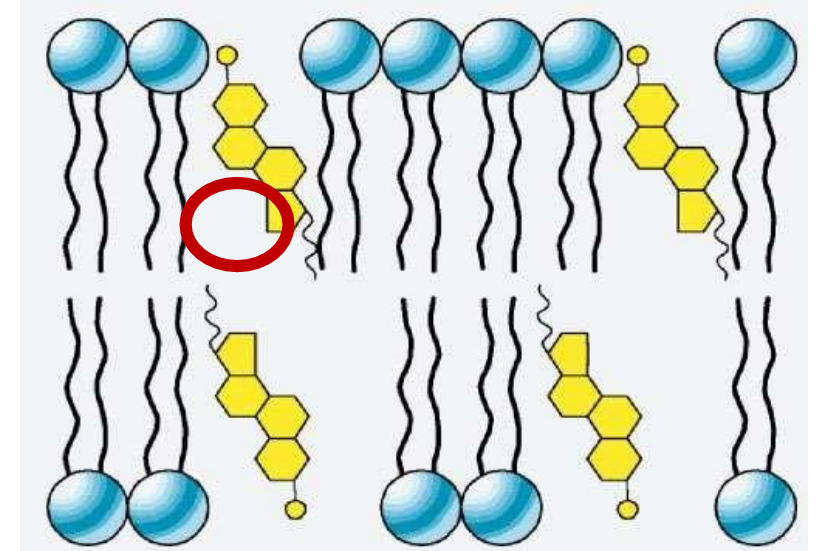


**Melting temperature  
(transition temperature)**

# Cholesterol and membrane fluidity



- The presence of cholesterol and the cis unsaturated fatty acids in the membrane prevent the hydrophobic chains from packing too closely together, allowing free membrane proteins and lipid molecules to move laterally in the plane of the leaflet making the membrane a dynamic environment.
- Cholesterol can also stabilize very fluid membranes by increasing interactions between the fatty acids of phospholipids through hydrophobic interactions with the cholesterol ring structure.



Increasing the concentration of cholesterol, the fluidity of membrane will decrease (the rigidity will increase).  
The most of place that cholesterol concent in it :(locked rafts )

Cholesterol helps regulate the fluidity of a cell membrane by limiting lateral movement of the phospholipids and adds stability on plasma membrane in low, high temperature. How?  
By making more spaces between phospholipids --> more liquid state

# Types of membrane proteins



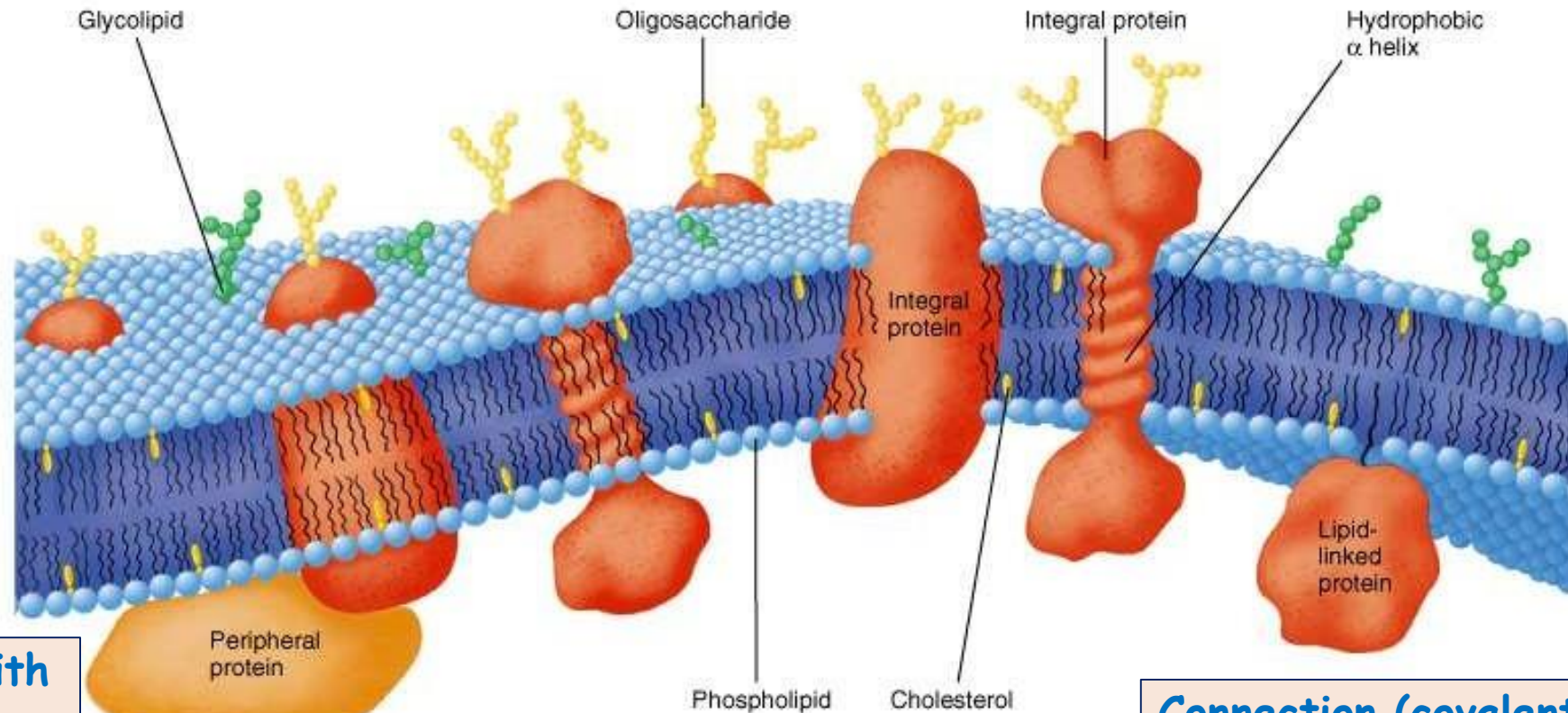
- Peripheral proteins:
  - are associated with **Embedded with the membrane** via noncovalent interactions
- Integral membrane proteins:
  - anchored into membrane via **Integrate with the membrane**
- Lipid-anchored:
  - associated via a lipid group

**Electrostatic or  
hydrogen bond**

**NOTE:** Lipid anchors are attached to the lipid anchor to proteins via a covalent bond whereas the peripheral proteins are attached to the cytosolic domains of integral membrane proteins through non-covalent interactions.



# Membrane proteins



Non covalent with others protein or lipid and still attach or near the membrane

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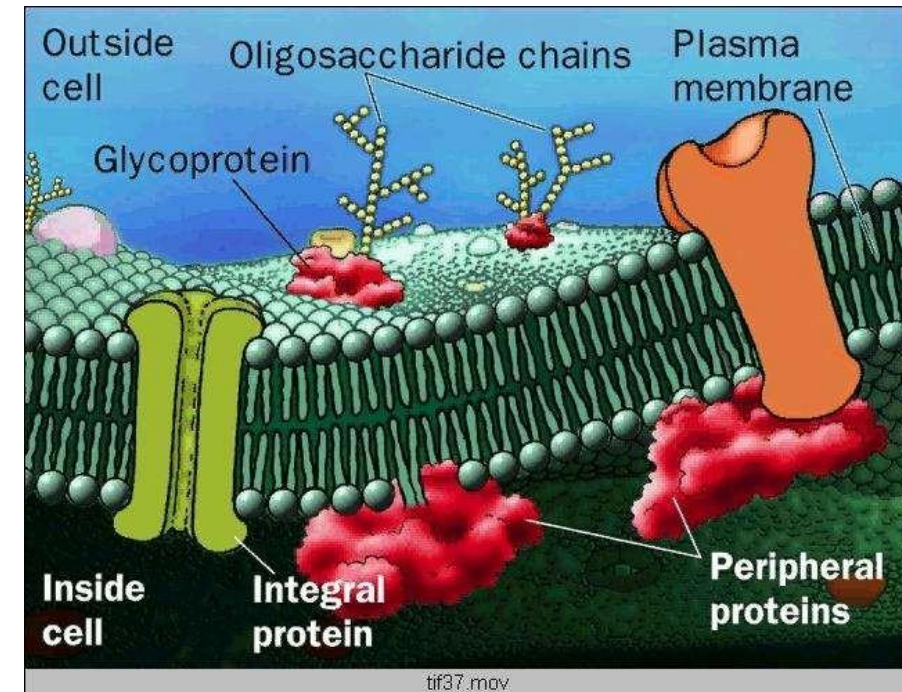
Connection (covalent) between fatty acid and protein



# Peripheral membrane proteins

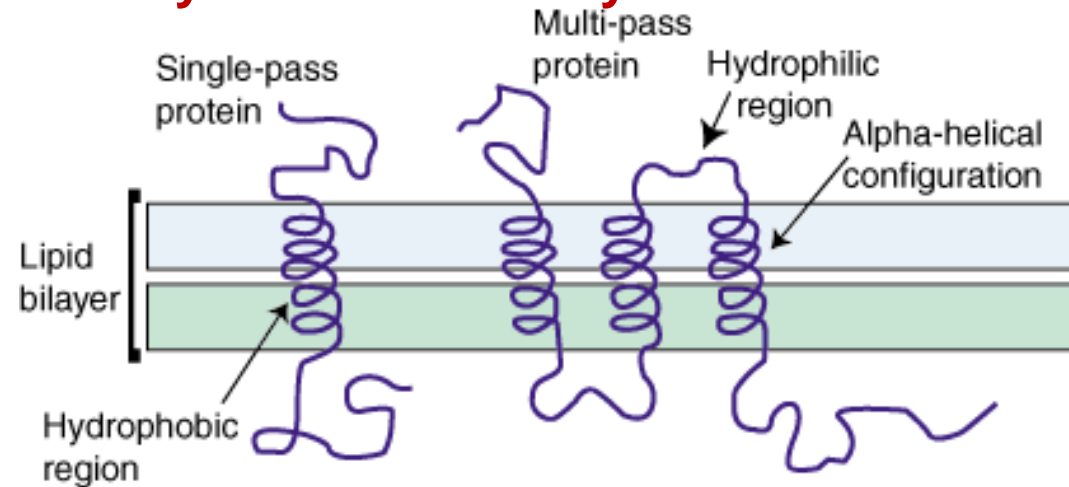


- They are associated with membranes but do not penetrate the hydrophobic core of the membrane.
  - They can be associated with integral membrane proteins.
- They are not strongly bound to the membrane and can be removed without disrupting the membrane structure.
  - Treatment with mild detergent



# Integral membrane proteins

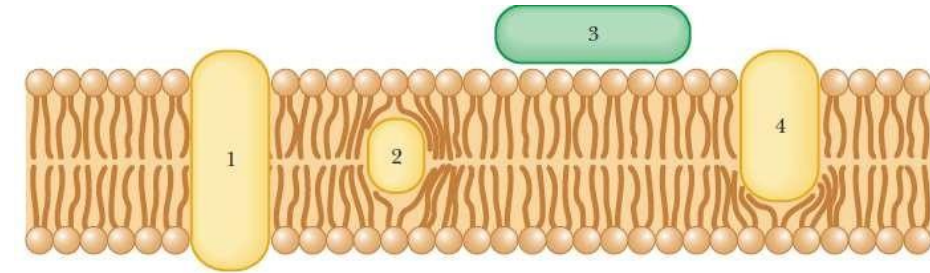
The integral membrane proteins can be associated with the lipid bilayer in several ways.



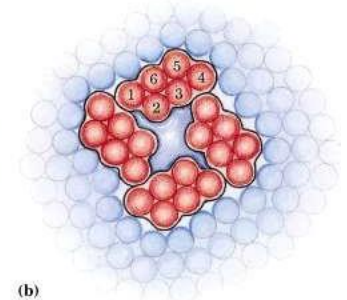
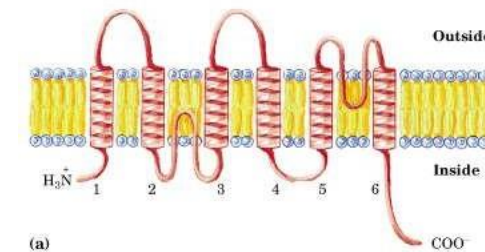
Some can form channels.

Very hard to study this structure so we can build a hybrid form to simulate it .

Inserted fully in plasma membrane they might be exposed on one or both sides of the cell membrane



The membrane integral domains are:  
1. Single or multiple  
2.  $\alpha$ -helix (human) or  $\beta$ -sheet (bacteria)





# Structure-Function of Membranes (some of functions)

- Transport: like channels
  - Membranes are impermeable barrier
  - Proteins can be carriers or channels
- Signaling
  - Protein receptors and small molecules (some can be lipids themselves)
- Catalysis
  - Enzyme-linked receptors

In both membrane protein and soluble protein .

Accelerate the speed of reaction



# Amino acids

Summer 2023

Amino acid name comes from :  
Amino from the amine, acid comes from the carboxylic acid



# General structure

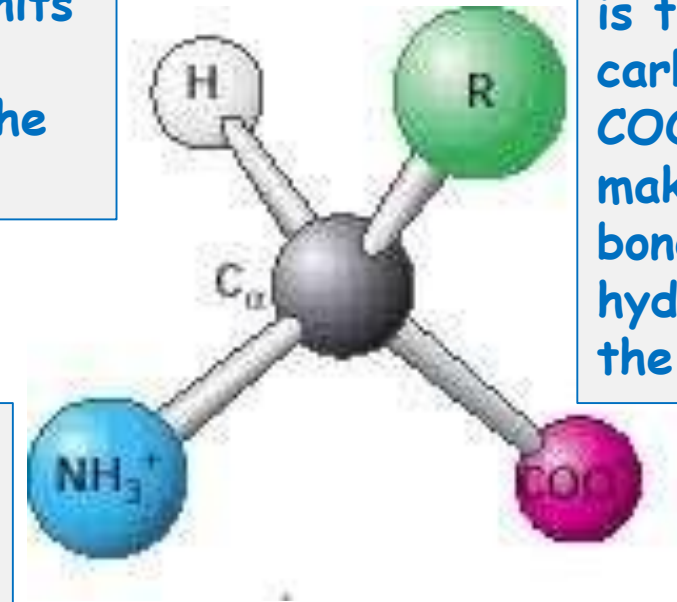


- Proteins are polymers of  $\alpha$ -amino acids (or amino acids).
- An amino acid consists of:
  - a central carbon atom, called the  $\alpha$  carbon, linked to four groups
  - an amino group ( $-\text{NH}_2$ ),
  - a carboxylic acid group ( $-\text{COOH}$ ),
  - a hydrogen atom, and
  - a specific R group (the side chain)

We can distinguish the amino acids by R group (side chain)

The first 4 units are the main monomer of the protein

Alpha carbon is considered chiral if the R group differs from the other groups



Alpha carbon is the first carbon after  $\text{COOH}$  and it makes bonds with hydrogen and the R group

# L and D isomers



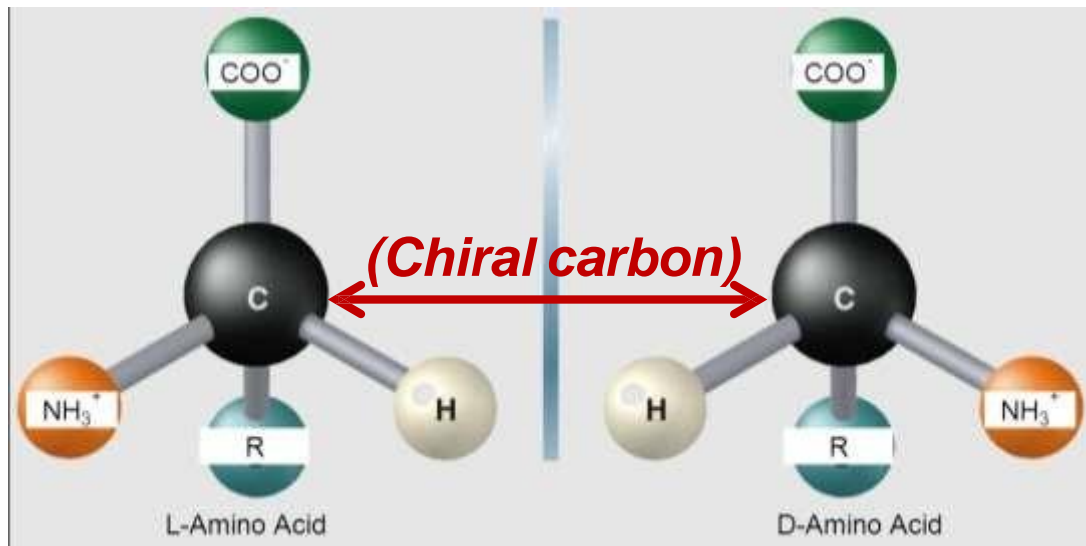
Depends on the amino group

AMINE GROUP إذا دوجوم

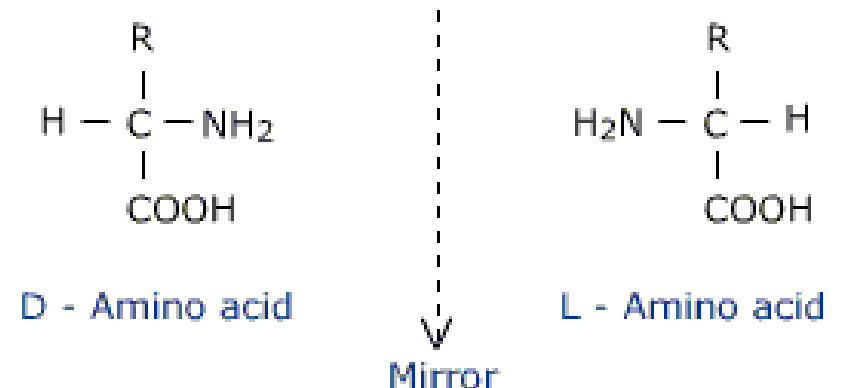
D isomers نوكت نيميلاع

L isomer نوكت لامشلاع دوجوم اذا

- The  $\alpha$ -carbon atom is chiral and, thus, amino acids can be present in two forms that are mirror-images of each other (they are enantiomers).
- The amino acids are chiral (when a central carbon is bonded to four different groups).
- They are called L isomer and D isomer.
- Only L amino acids naturally make up proteins.



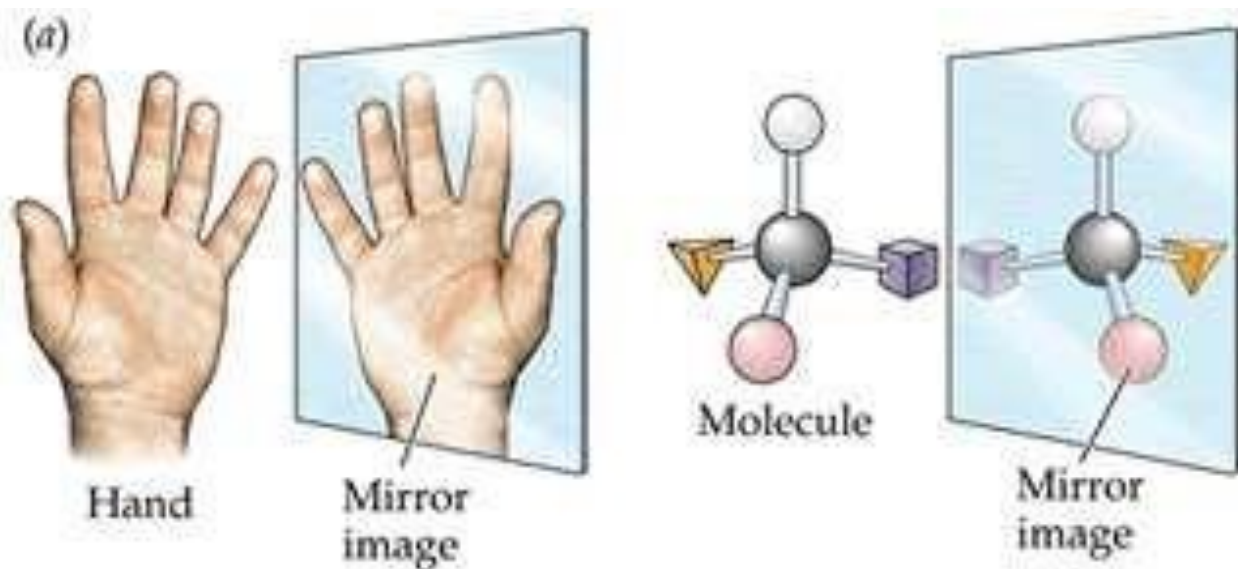
The amine group of L-amino acids occurs in the left-hand side when drawn in the Fischer projection, keeping the carboxylic acid group on bottom (or top) and the carbon chain in the top, whereas the amine group of the L-amino acids occurs in the left.





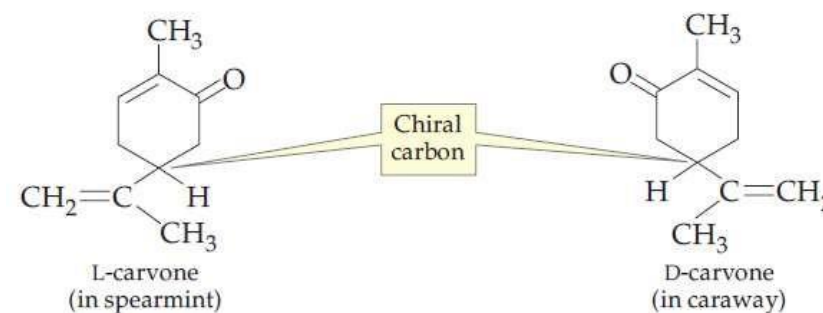
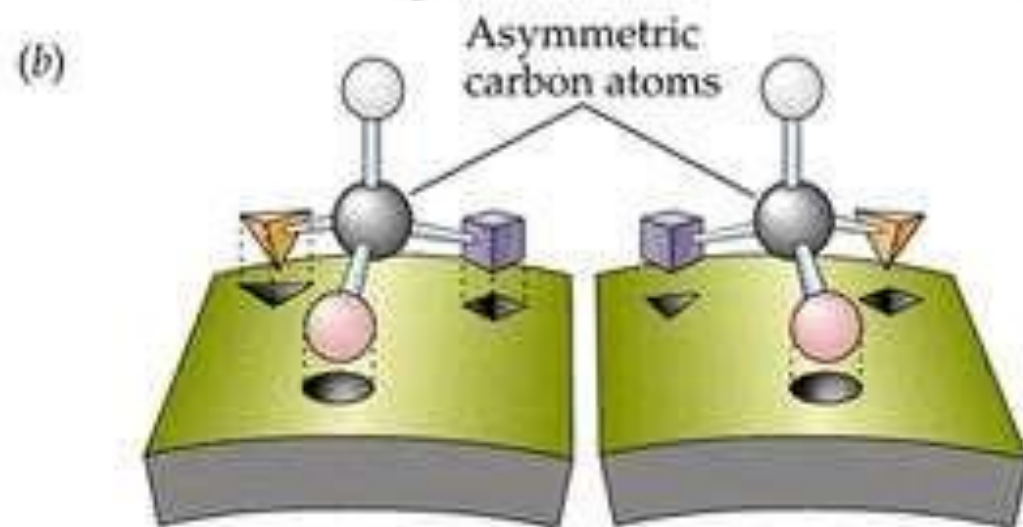
- The amino acids that exist in the human body are L isomers, but that doesn't mean that our bodies can't deal with D isomers, no; our body can deal with it because we may take it in the drugs so our body doesn't excrete it
- There are so many amino acids in our body, but there are only 20 in our protein structure, these 20 amino acids do other functions, not just the functions they do in the protein such as their function as neurotransmitters, stains, hormones and allergic mediators
- The presence of the amine group in the structure of the amino acid helps it to do these functions

The amine group is the backbone



There are 19 amino acids that has D and L isomers except glycine which is the simplest amino acid that doesn't has isomers

D and L isomers is very important because the differentiation in isomerism makes a huge changes such as carvone in the above picture



Used in the toothpaste



# Types of amino acids



- There are twenty kinds of amino acids depending on the side chains varying in
  - Size
  - Shape
  - Charge
  - Hydrogen-bonding capacity
  - Hydrophobic character
  - Chemical reactivity

There are so many classifications of amino acids but the only one we will talk about in the biochemistry is **THE POLARITY**

# Classification (according to R group)



They are both polar and charged, the positive one basic and the negative one acidic

Non-polar	Polar	Charged (positive)	Charged (negative)
Alanine	Serine	Lysine	Glutamate
Valine	Threonine	Arginine	Aspartate
Leucine	Glutamine	Histidine	
Isoleucine	Asparagine		
Methionine	Cysteine		
Tryptophan	Tyrosine		
Phenylalanine			
Proline			
Glycine			

Glycine is the simplest one



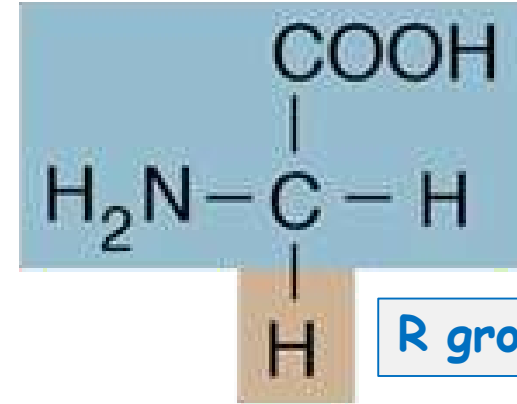
- The polar amino acids could be charged or uncharged
- charged could be positive and that means that the side chain in the amino acid has accepted protons and acts as a BASE ,if it is negative that means the amino acid has lost protons and acts as a ACID
- Uncharged don't transfer the full charge and they make dipole bond

# Glycine



- Glycine is a derivative of acetic acid.
- It could be considered a derivative of aminoethane.

This is the back bone



R group

Glycine (gly)

We abbreviate  
the amino acids  
by 3 letters

**Is it chiral?** no





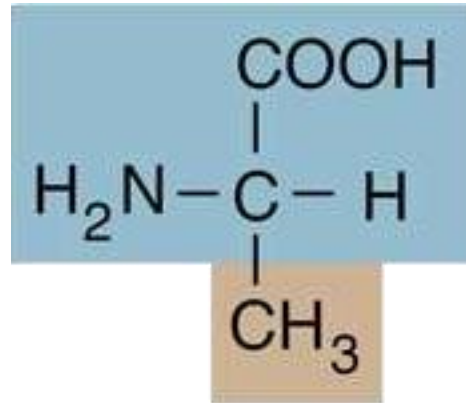
# Non-polar, aliphatic amino acids

# Alanine



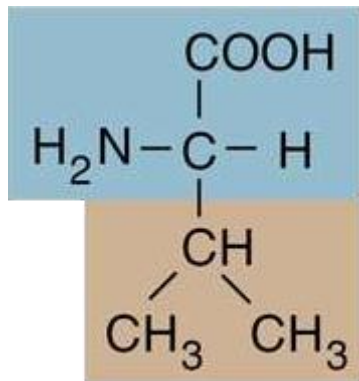
- Alanine, the next simplest amino acid, has a methyl group (-CH<sub>3</sub>) as its side chain.

The methyl group is hydrocarbon(non polar)

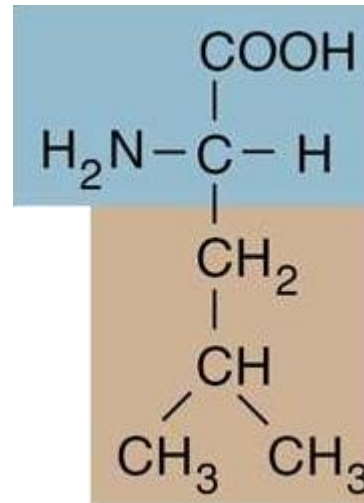


Alanine (ala)

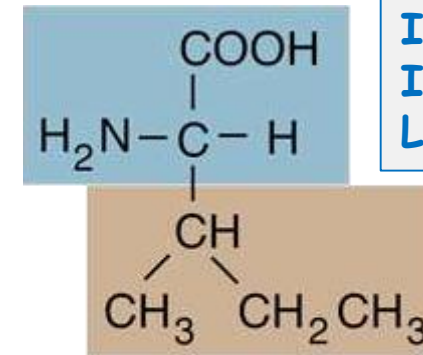
# Valine, leucine, and isoleucine



Valine (val)\*



Leucine (leu)\*



Isoleucine (ile)\*

Ile abbreviation:  
I comes from iso  
Le from leucine

Isoleucine and  
leucine are  
isomers

They differ  
in the  
attachment  
carbon

They are branched amino acids.

These are *essential amino acids in the sense that the body cannot synthesize them.*

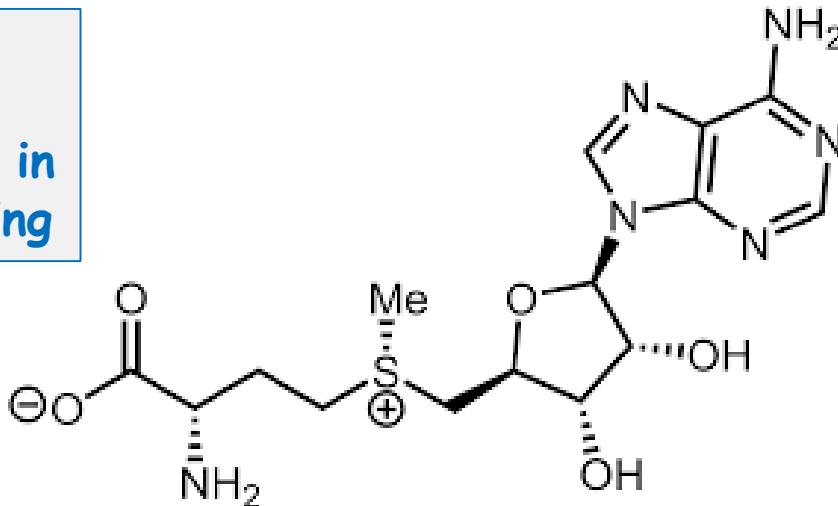
# Methionine

Non polar amino acids

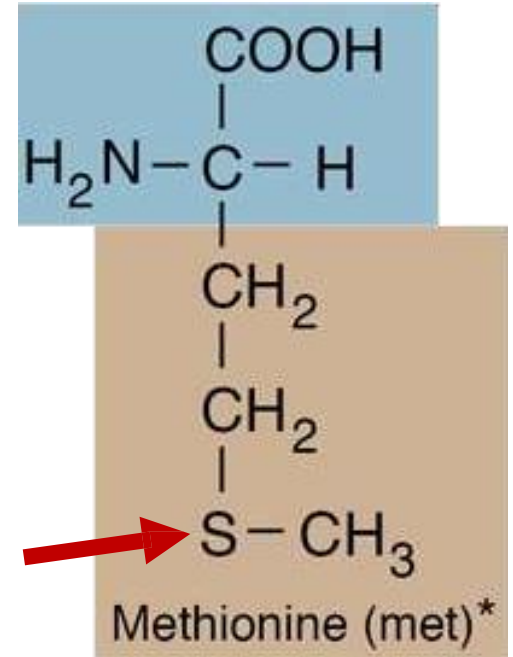
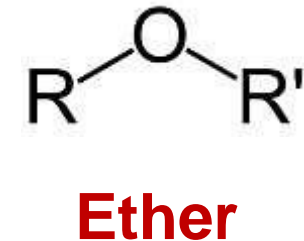
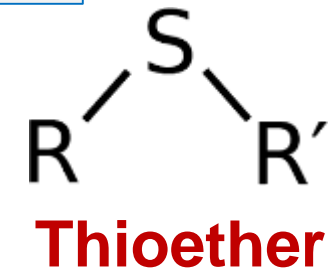
It can react to form S-Adenosyl-L-Methionine (SAM) which serves as a methyl donor in reactions.

During the degradation of methionine SAM is produced, which is used in methylation reactions that's why it serves as a methyl donor

Its so important and always in the beginning



It's has a polar atom but overall its nonpolar cuz of the net charge is zero



R chain is non polar

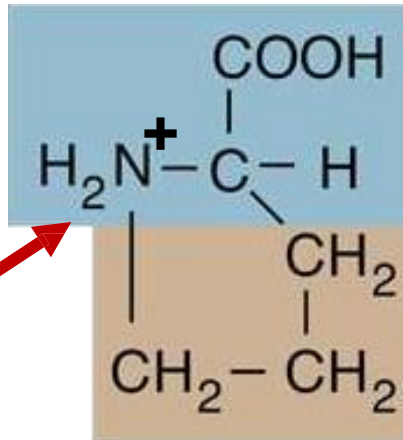




# Proline (the only cyclic amino acid)

Proline is the only secondary amino acid

**Secondary  
nitrogen**



Proline (pro)

Proline is the only amino acid that makes a ring structure with the back bone, which gives the amino acid new characteristics that we will talk about later

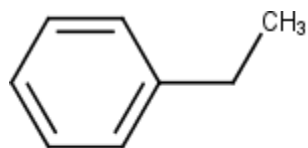


# Phenylalanine and Tryptophan

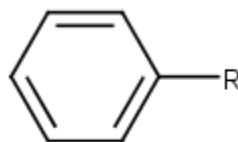


- Phenylalanine contains a phenyl ring.
- Tryptophan has an indole ring; the indole group consists of two fused rings and an NH group.

Aromatic could be either polar uncharged or nonpolar and it is polar when it has a side group

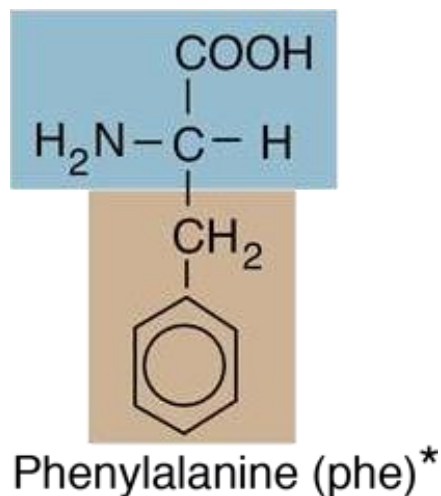


benzyl

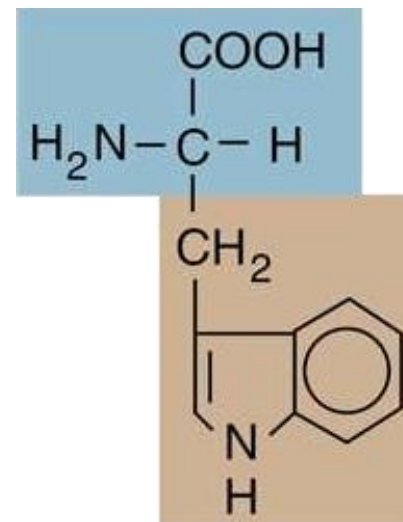


phenyl

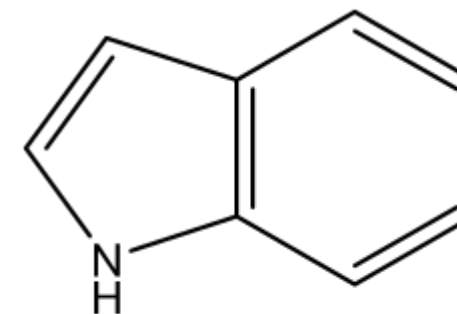
TRP abbreviation we use it because there are another amino acid with TYR abbreviation so we change it to distinguish them



Phenylalanine (phe)\*



Tryptophan (trp)\*



Indole

Benzene fused with 5 carbons ring



# Positively-charged amino acids



- Because the side chains on the lysine and arginine have a basic characteristics,  $pK_a$  values for these bases are low
- When you compare the conjugate acid for these basic amino acids with acidic chain in amino acid you will find that the  $pK_a$  of the real acidic chains is higher than the conjugate acid though it is low but the conjugate acid is lower
- $pK_a$  for lysine and arginine is high, higher than the physiological  $pK_a$  in the body
- So you will find that these side chains are mostly protonated, because their  $pK_a$  is higher than 7.4, so the desired  $OH$  is acidic compared to it, so usually they will be having a charge

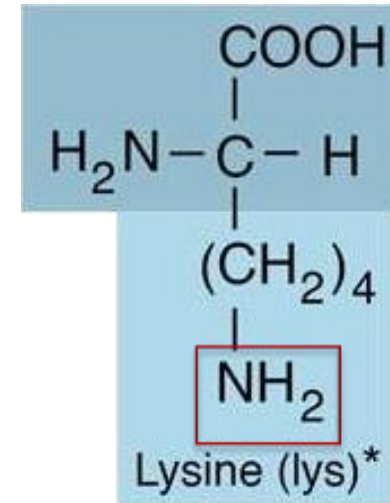
# Lysine and arginine



- Lysine and arginine have relatively long side chains that terminate with groups that are positively charged at neutral pH.
- Lysine ends with a primary amino group and arginine by a guanidinium group.

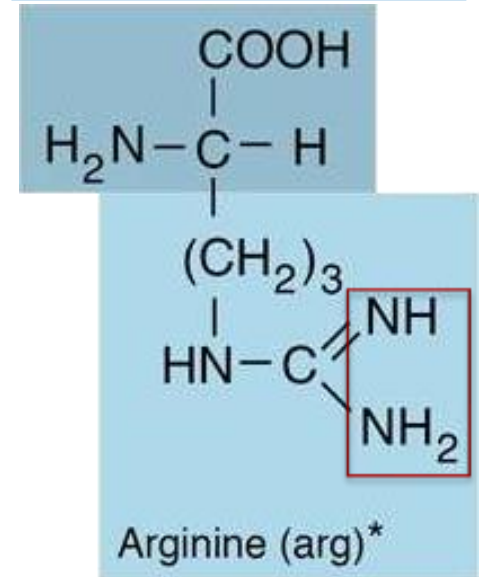
Just identify the structures

The presence of  $\text{NH}_2$  which is a base group, that makes a basic side chain



This fork shaped structure is called guanidinium

You have to identify the structure, just identify

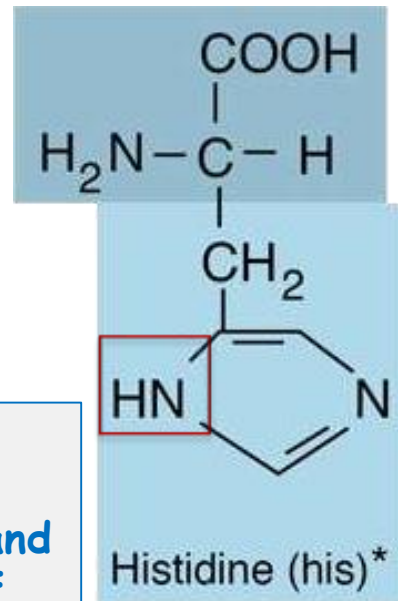


Though it has 3 nitrogen, only one c an accept proton

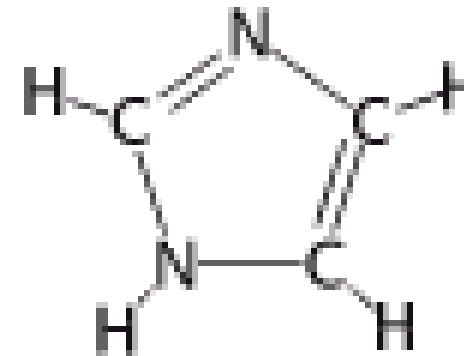
# Histidine



- Histidine contains an imidazole group, an aromatic ring that also can be positively charged.



It has two different nitrogens, and only one of them can accept proton



Imidazole

This is the nitrogen that accepts proton

Pka of the histidine is very close to the physiological ph





- Histidine is a basic amino, but not as lysine and arginine because they are very far from the physiological pH, but histidine pH is very close to the physiological pH
- So we are close to 1 ratio which means pH and pKa are very close, which means a very small change in pH will lead to take or remove protons from histidine
- Histidine can take or lose protons more than any amino acid
- The presence of histidine in the structure of proteins, leads them to work as buffers
- Hemoglobin has 38 histidines more than albumin that has 18 histidines, hemoglobin is a better buffer because it has more histidine which means more buffering capacity



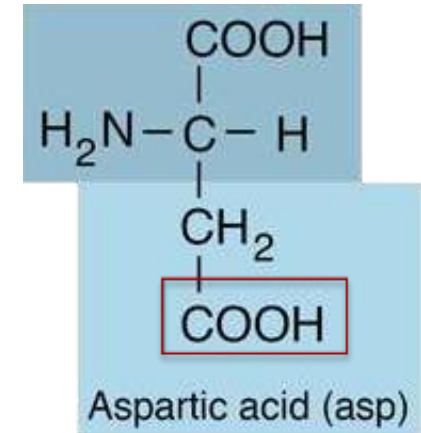
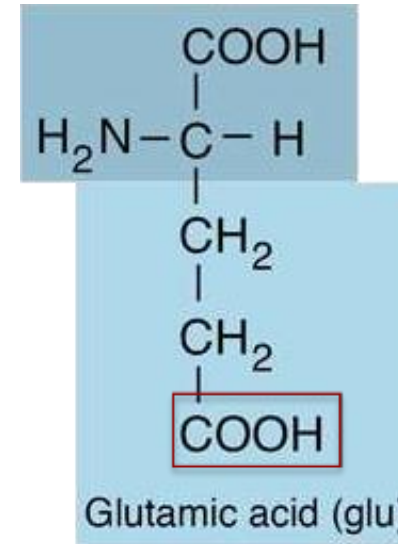
# Negatively-charged amino acids

# Aspartic acid and glutamic acid



Mostly they are deprotonated

- Two amino acids contain acidic side chains: aspartic acid and glutamic acid.
- These amino acids are often called aspartate and glutamate when they are charged.



We can call them aspartate and glutamate because usually they donate the proton and pka values for their side chains are very low, lower than the physiological pH, so they are charged under the physiological conditions.

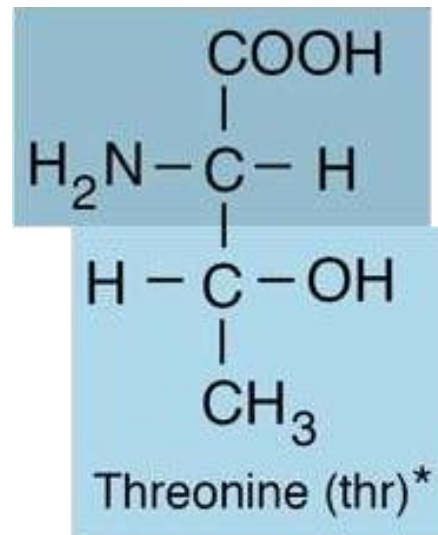
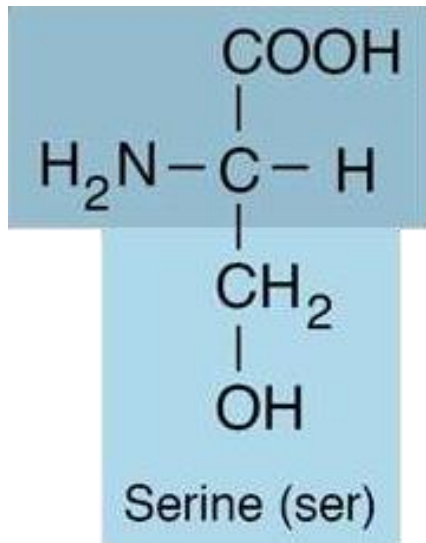


# Polar, hydrophilic, neutral amino acids

# Serine and threonine



- Serine and threonine, contain aliphatic hydroxyl groups.
- The hydroxyl groups on serine and threonine make them hydrophilic and reactive.



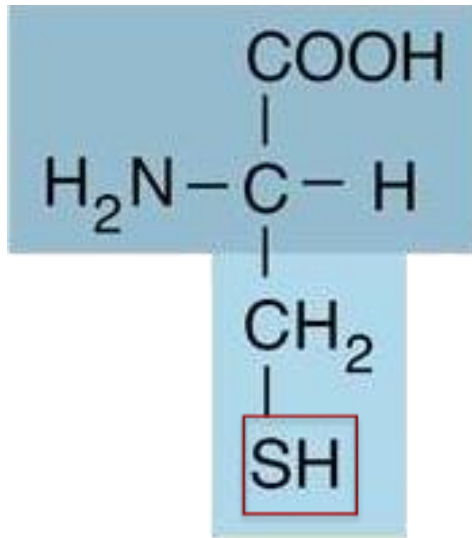
They are polar, aliphatic amino acids not charged but they contain an amino group (polar group) because of that they are considered reactive (they enter in many reactions)  
**both** contain hydroxyl group



# Cysteine (Cys)



- Cysteine contains a sulfhydryl or thiol (-SH), group.
- The sulfhydryl group is reactive.

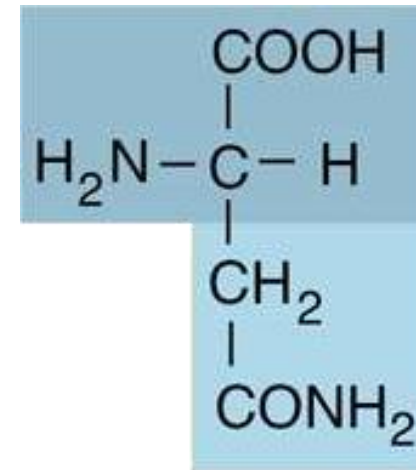
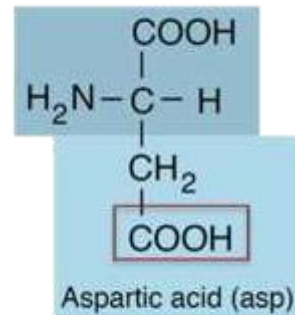
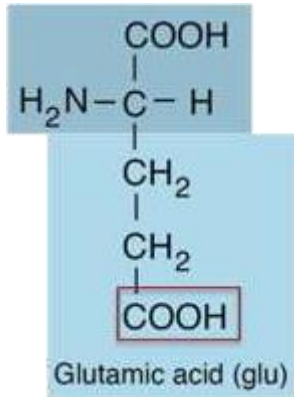


Cysteine (cys)

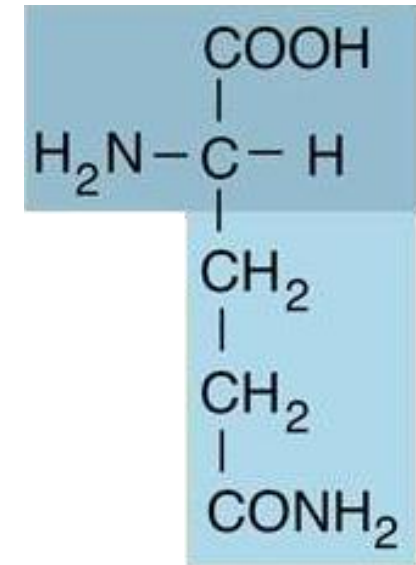
Sulfur is a thiol terminal group( at the end of the chain )-->it is polar because of that it is very active

# Asparagine and glutamine

- Asparagine and glutamine are uncharged derivatives of aspartate and glutamate.
- Each contains a terminal carboxamide in place of a carboxylic acid.



Asparagine (asn)



Glutamine (gln)

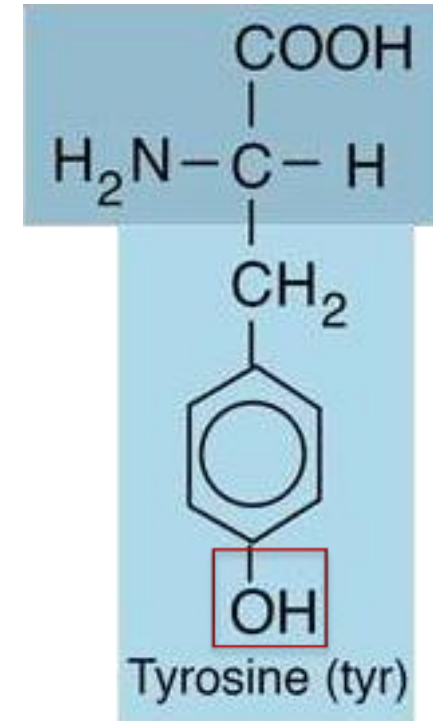
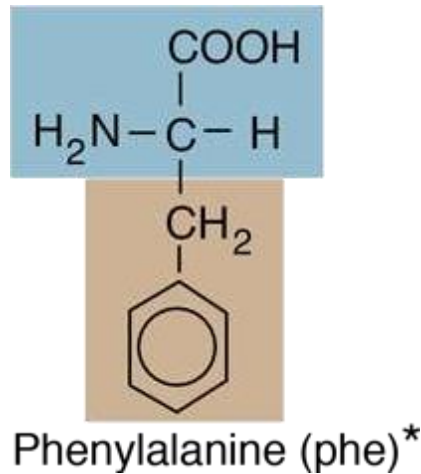
Glutamate and aspartate  
Instead of carboxyl group in them ,  
we remove oh and put amino group  
which isn't charged but they are  
polar , reactive groups

# Tyrosine



Is the 3rd amino acid that contain hydroxyl group with serine and threoline  
It looks like any amino acid ? Phenylalanine but we take phenyl group and put oh group ( phenol)

- The aromatic ring of tyrosine contains a hydroxyl group.
- It is derived from phenylalanine.
- This hydroxyl group is reactive.

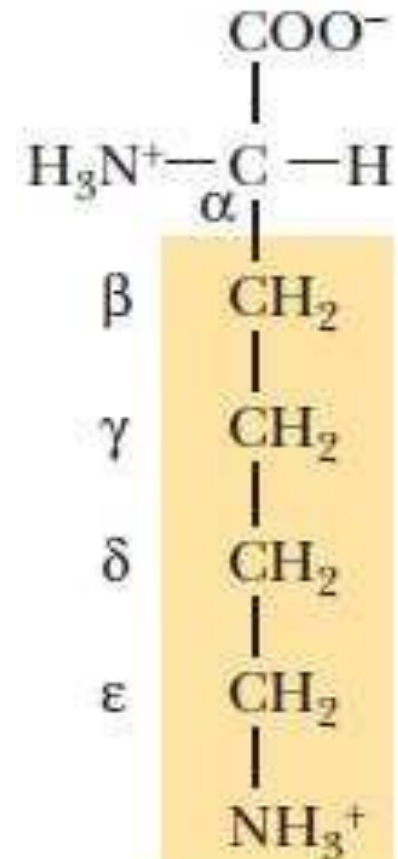


# Designation of carbons



- Side-chain carbon atoms are designated with letters of the Greek alphabet, counting from the  $\alpha$ -carbon. These carbon atoms are, in turn, the  $\beta$ -,  $\gamma$ -,  $\delta$ -, and  $\epsilon$ -carbons.
- If a carbon atom is terminal, it is referred to as the  $\omega$ -carbon.

The last carbon in the chain is called ( omega carbon)



# Questions



- Two amino acids are negatively-charged:
- The following amino acid is achiral:
- ...etc.





# Specialized and uncommon amino acids

## ► Important :-

Amino acids not only produce proteins , they may do another functions like neurotransmitter, signaling molecule but some of them are converted to new structure with a specific function .

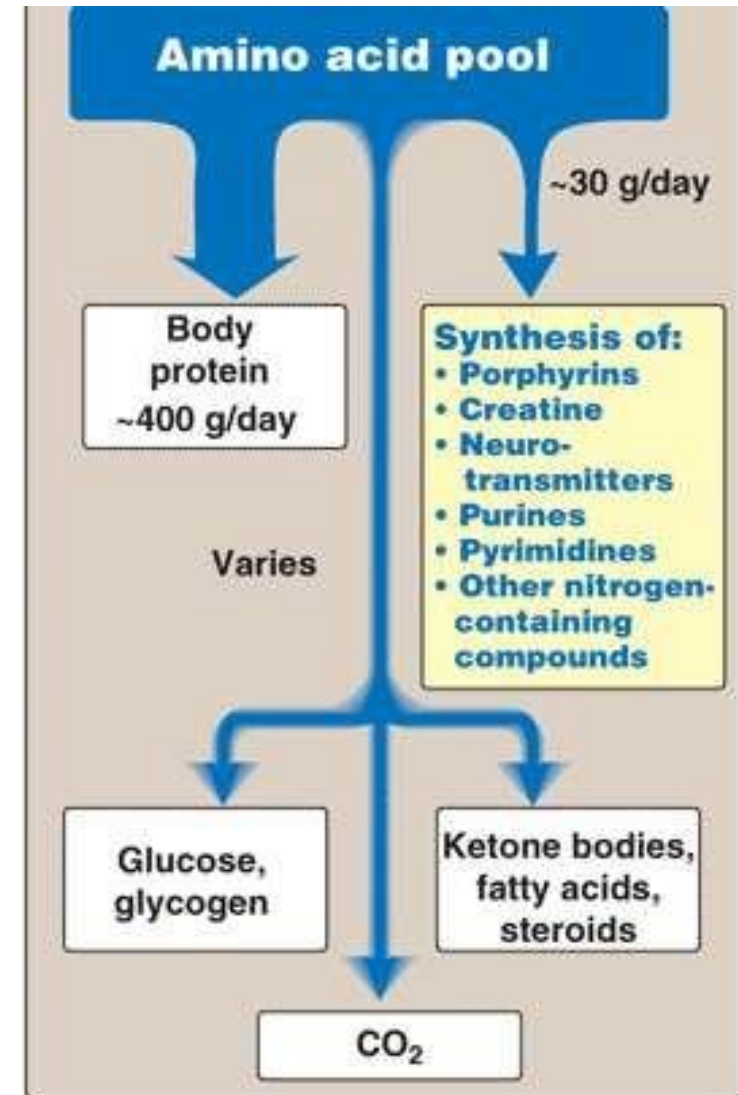
We can use them in metabolism

Amino acid --> sugar (like glucose) **we will study them later ;)**

# Biological significance of amino acids



- $\alpha$ -nitrogen atom of amino acids is a primary source for many nitrogenous compounds:
  - Hormones
  - Neurotransmitters
  - Biologically active peptides



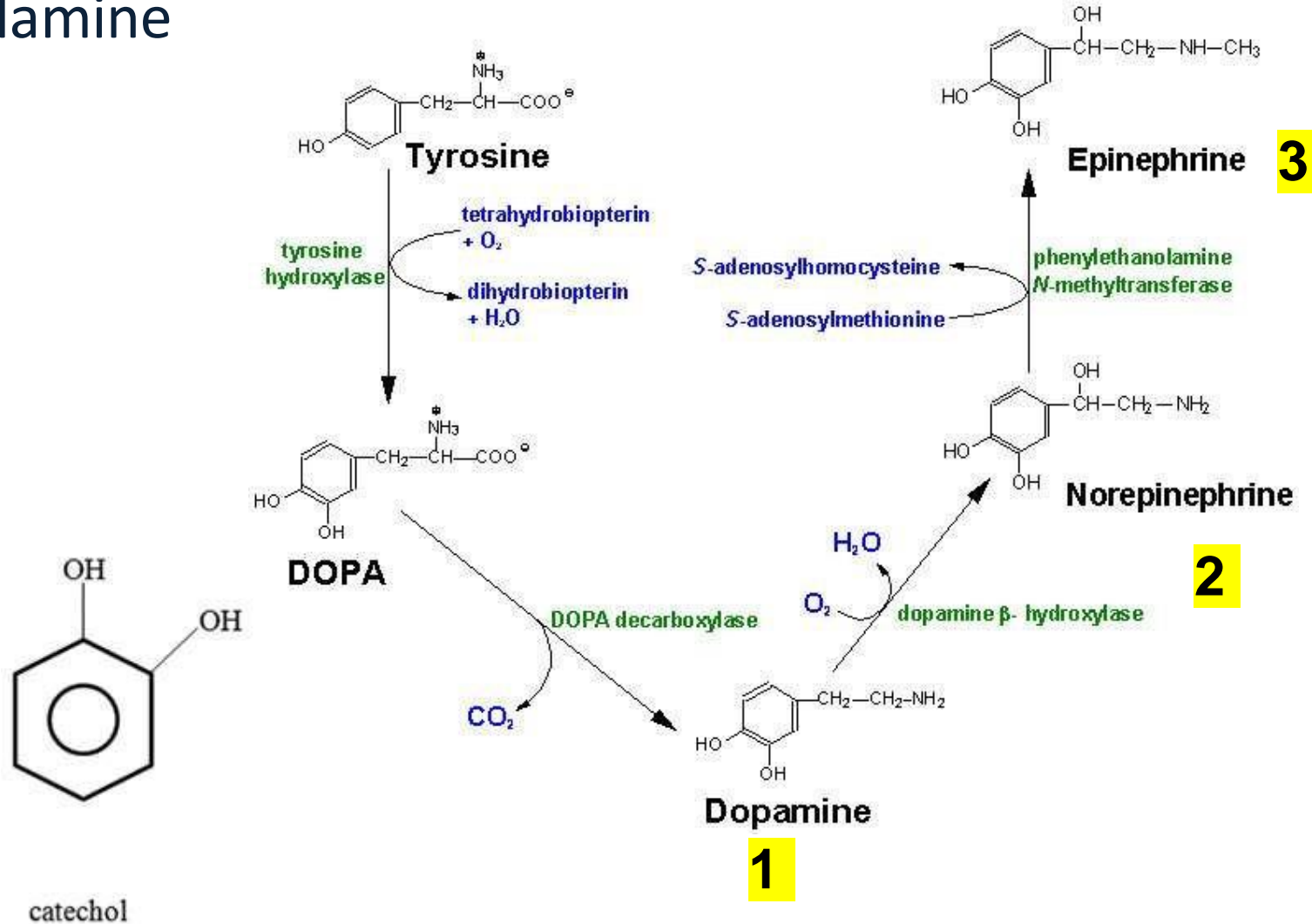
# Tyrosine (1)



- It is converted into catecholamine neurotransmitters

- Dopamine
- Norepinephrine
- Epinephrine
  - flight or fight

**1+2+3 are neurotransmitters**  
Catecholamines because they contain catechol group (a benzene with 2 hydroxyl groups and an amino group from tyrosine )





Further explanation regarding the previous slide:

You aren't required to memorize the structure or the enzyme(not important now)

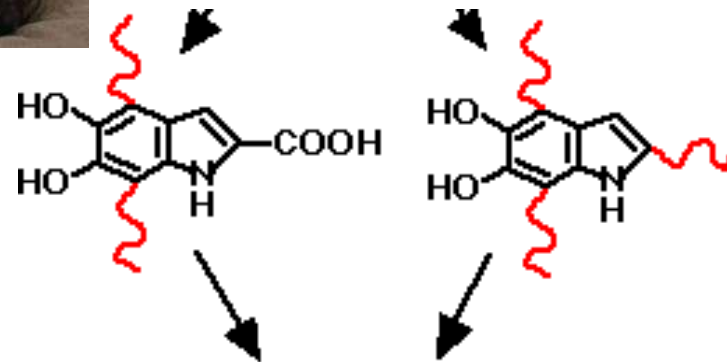
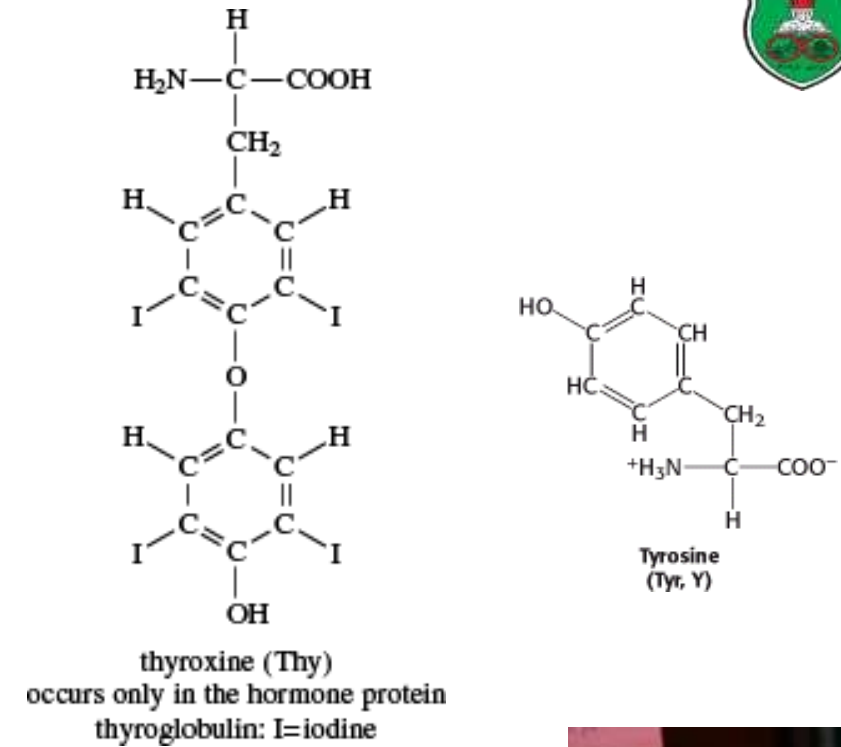
#But you must now that this molecule is derived from this amino acid and know its general function

**Epinephrine = adrenaline** its function is production of energy as dopamine . like excitement

# Tyrosine (2)



- Tyrosine is converted into
  - Melanin (skin colour)
  - Thyroxine (hormone) a thyroid gland hormone



Brownish color of the skin

EUMELANINS

PHEOMELANINS

Red color of ( skin , hair)

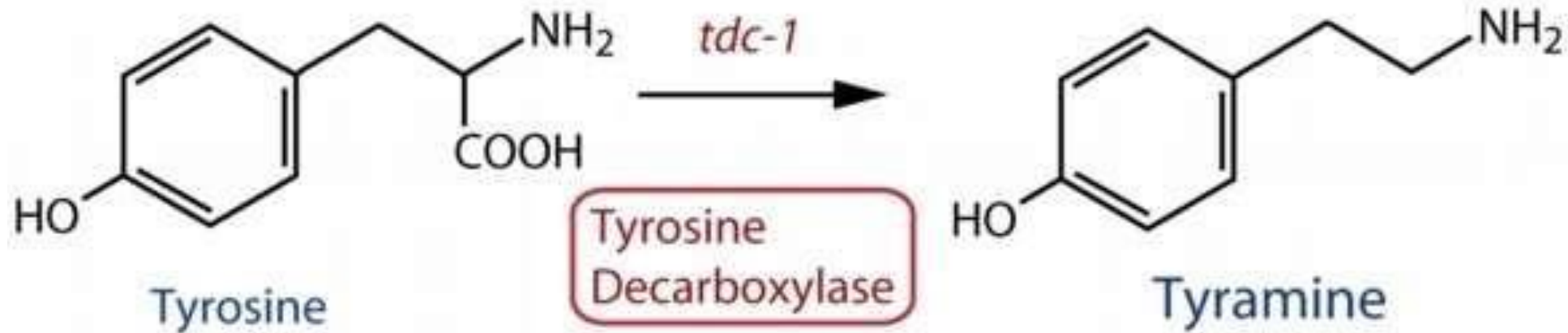


# Tyrosine and life



Tyramine is mainly founded in our daily food like cheese  
# its function like epinephrine which gives energy .

- Cheese contain high amounts of tyramine, which mimics epinephrine; for many people a cheese omelet in the morning is a favorite way to start the day.





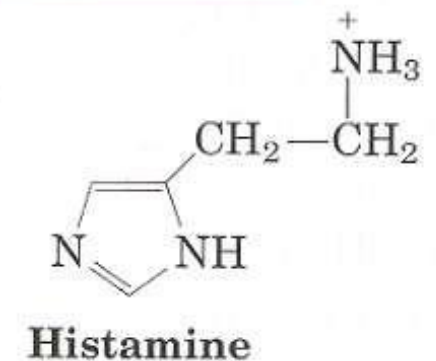
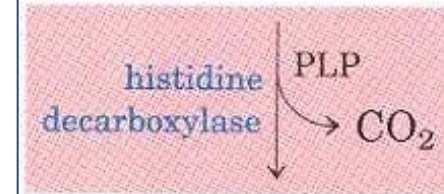
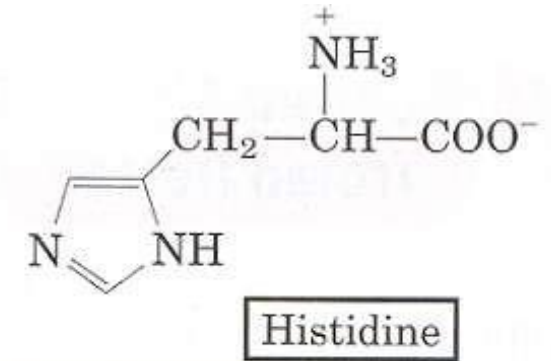
# Histamine

a chemical your immune system releases. It's mainly known for its role in causing allergy symptoms



- Regulates physiological function in the gut
- Acts as a neurotransmitter
- Causes allergic symptoms (a major cause for asthma)
- Contributes to inflammatory response
- Causes constriction of smooth muscle

It is From histidine amino acid (His) +ve charge amino acid with an imidazole group, an aromatic ring that also can be positively charged.

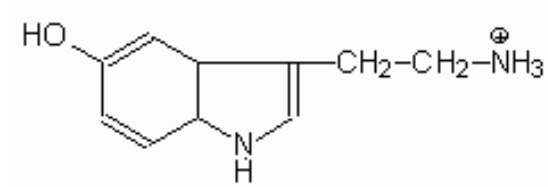


# Tryptophan

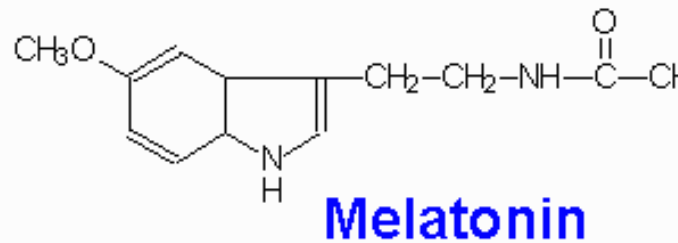
Trp ; bulk , hydrophobic despite of nitrogen that is in one ring  
it has 2 rings



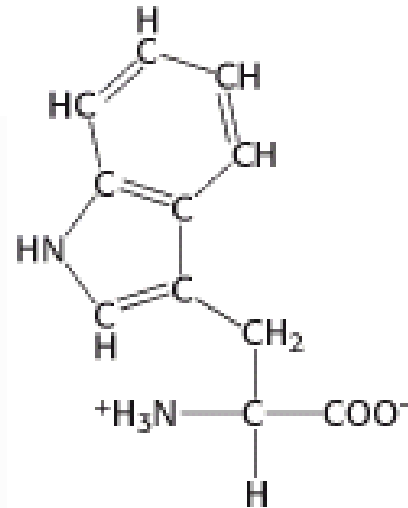
- Tryptophan serves as the precursor for the synthesis of Neurotransmitters
  - Serotonin (neurotransmitter-sedative)
  - Melatonin (day-night cycle)



**Serotonin**  
(5-hydroxytryptamine)



**Melatonin**



**Tryptophan**  
(Trp, W)

## From a gland in brain

قَسْلًا لُوصَفَ وَابْنًا مَوْلَا رَبِّي رَوْعَ شَلَا نَع لَوْ سَمَلَا وَه  
Infants have high levels of this hormone because of that they sleep for long hours not like old ages (low rate of this hormone)

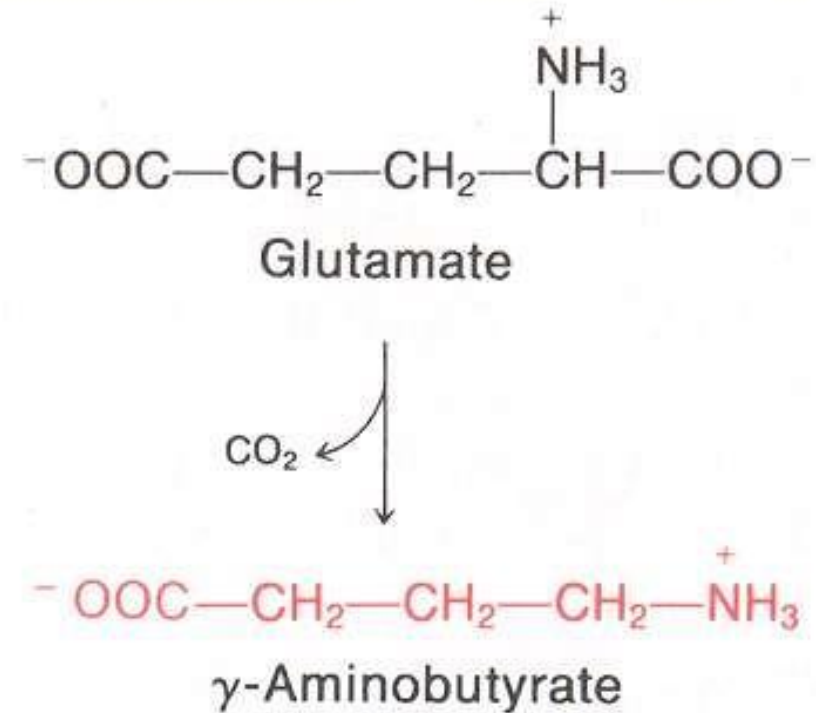
نَع مَوْلَا لِيَعْلَى هَمَلَخْتَسَا مِي لُضِيَا مَقْلَلَا مِي رَهْلَا  
عَم لُودَلَا بَدَل نِيَقْلَلَا فَلَاحَا

It is typically safe for adults to take melatonin every night for short periods of time. Several factors affect whether an individual should take melatonin every day.

# Glutamate



- It is a precursor of  $\gamma$ -aminobutyric acid (GABA) In anxiety
- It is an inhibitory neurotransmitter (CNS) that reduces neuronal excitability.
- It is synthesized in brain because it does not cross the BBB.
- It has relaxing, anti-anxiety, and anti-convulsive effects.



# $\gamma$ -carboxyglutamate (Gla)

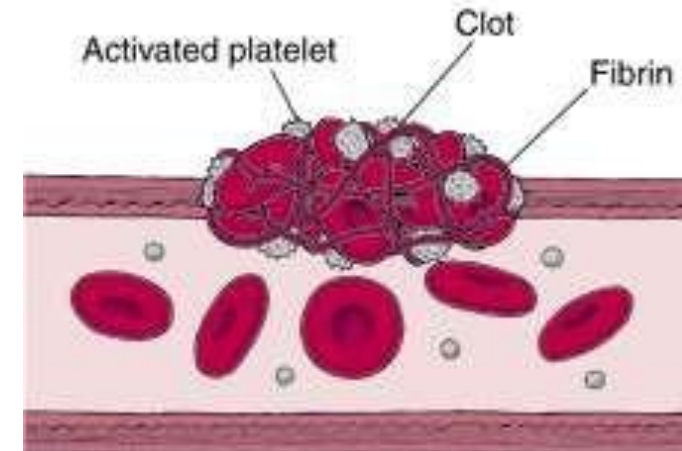
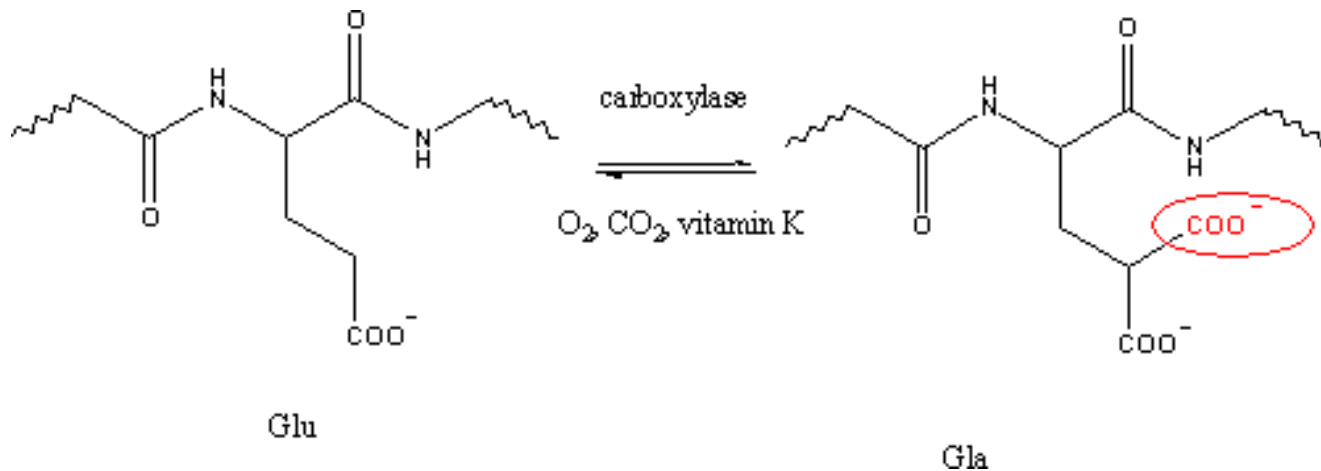


It has a carboxyl group

- The glutamate residues of some clotting factors are carboxylated to form  $\gamma$ -carboxyglutamate (Gla) residues.
  - Vitamin K is essential for the process
- This carboxylation is essential for the function of the clotting factors.

مدلا

والتخثر

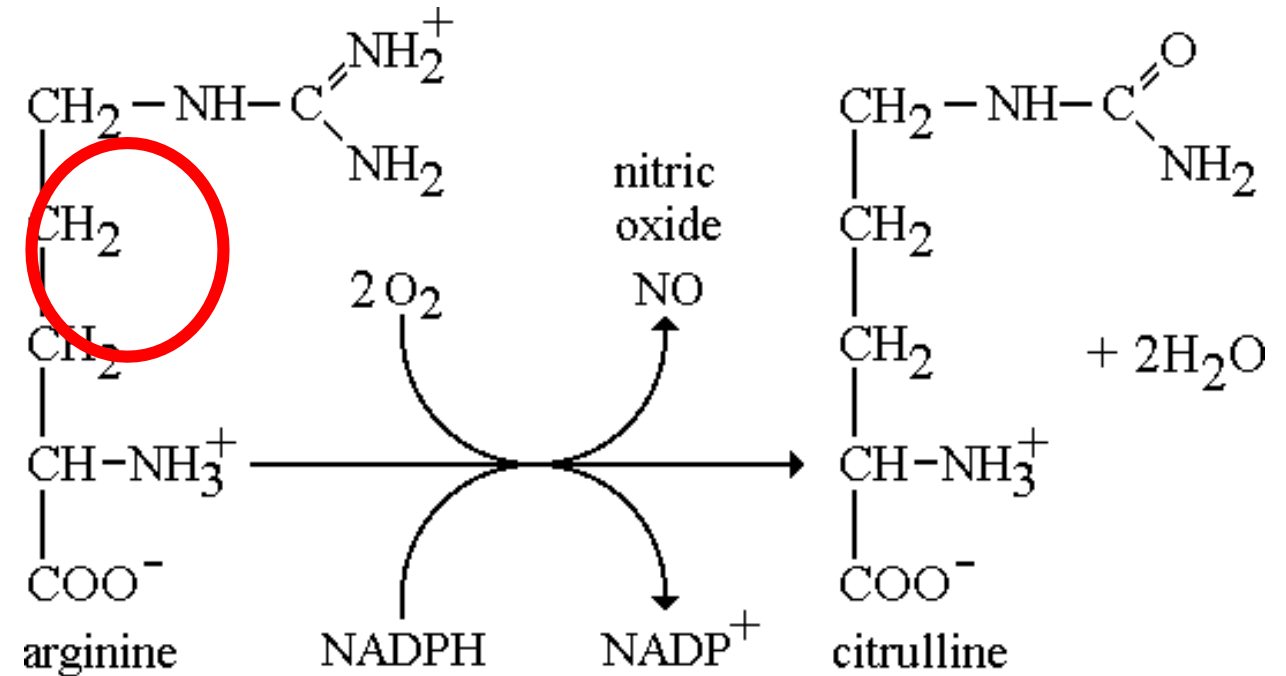


# Arginine



- L-arginine is the precursor of nitric oxide (NO)
- NO functions:
  - Vasodilation, inhibition of platelet adhesion, inhibition of leukocyte adhesion, antiproliferative action, scavenging superoxide anion (anti-inflammatory)

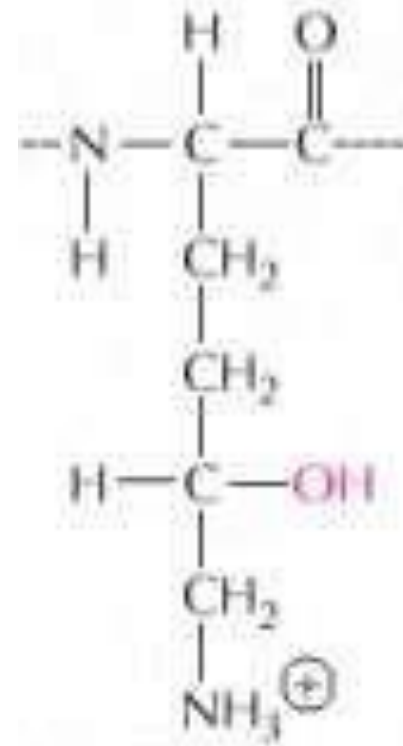
It has a +charge amino acid that contain guanidium group.  
We can extract a gas from R group we take one of these 2 N and take nitrile oxide it is a gas from endothelial cells  
It is a vasodilator



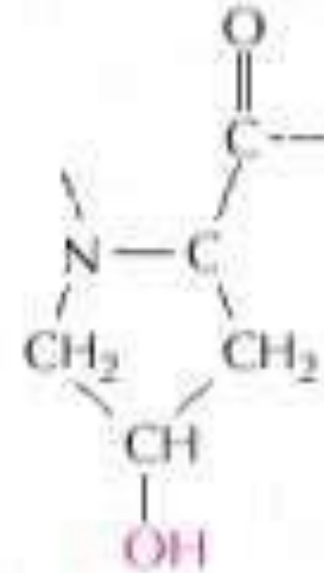


# Hydroxylysine and hydroxyproline

- Both are hydroxylated and are part of collagen structure.
- Both are derived from the common amino acids.
- Both are produced by modification of the parent amino acid after protein synthesis, posttranslational modification.



hydroxylysine  
in protein



hydroxyproline  
in protein

They are modified lysine and proline respectively with an oh group  
Lysine positive charge amino acid + oh --> Hydroxylysine  
Proline non-polar cyclic and rigid amino acid R+ N group  
They are presented in collagen ( we will take it later)

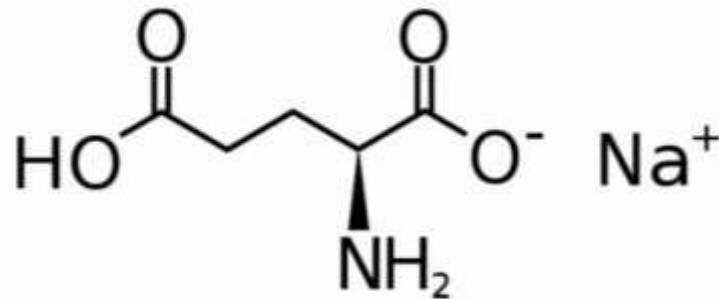




# Biochemical applications: Monosodium glutamate (MSG)



MONOSODIUM GLUTAMATE



SODIUM SALT OF GLUTAMIC ACID

Glutamic acid derivative

Flavor enhancer, Asian food.

MSG causes a physiological reaction in some people (chills, headaches, and dizziness)

Chinese restaurant syndrome.



**Monosodium glutamate (MSG):-**

**Is a meat tenderizer**

لهسي لكلاو غضملا

The goal of a meat tenderizer is to make meat softer and easier to chew (غضملا). it is used for Tough meat is difficult to chew and digest



- Amino acid and protein molecular weight. **Amino acids make the structure of protein**

- The average molecular weight of an amino acid residue is about 110.
  - The molecular weights of most proteins are between 5500 and 220,000.

**Important in the process of separating proteins in order to identify and discover them**

- We refer to the mass of a polypeptide in units of Daltons.
  - A 10,000-MW protein has a mass of 10,000 Daltons (Da) or 10 kilodaltons (kDa).

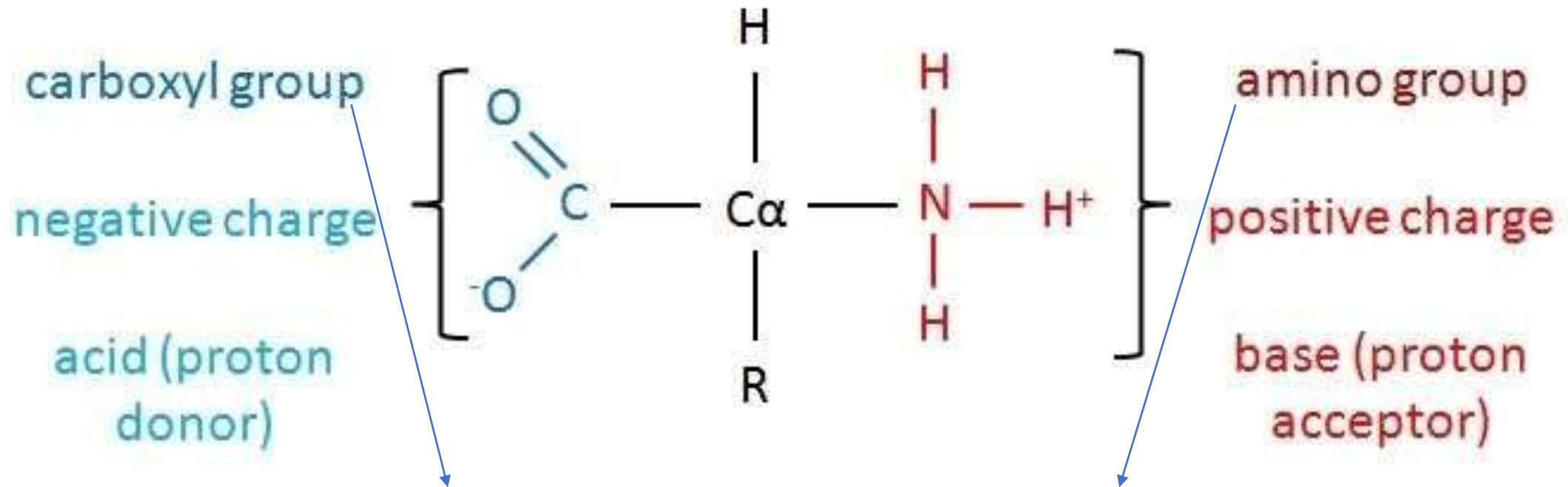


# Ionization of amino acids

Because we are talking about compounds that have acidic and basic functional groups, whether in side chains or backbones, we expect them to have acidic, basic and ionizing properties, so we will link them with the first topic we took, which is acids and bases



# Why do amino acids get ionized?



under physical conditions it loses protons (**Donner proton**) because the pka is low

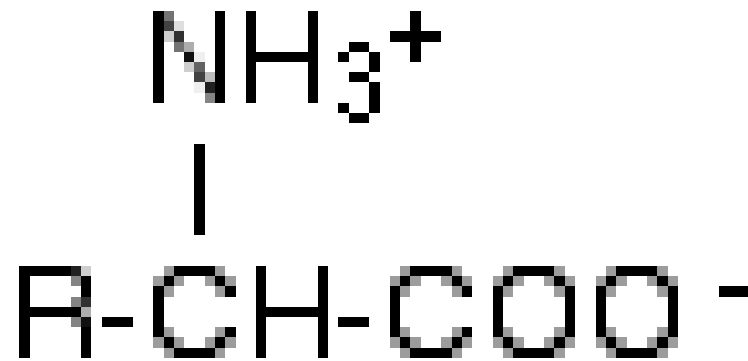
under physical conditions, it doesn't lose protons (**proton acceptor**) because the pka is high



# Zwitterion and isoelectric point

- At physiological pH, amino acids (without ionizable groups) are electrically neutral.
- Zwitterion: a molecule with two opposite charges and a net charge of zero.

**Zwitterion:** Amino acid has charges in different places but the over total charge equal zero



Zwitterion is formed at a point called the **isoelectric point**.  
**Isoelectric point:** the pH when the molecule is in the zwitterion form

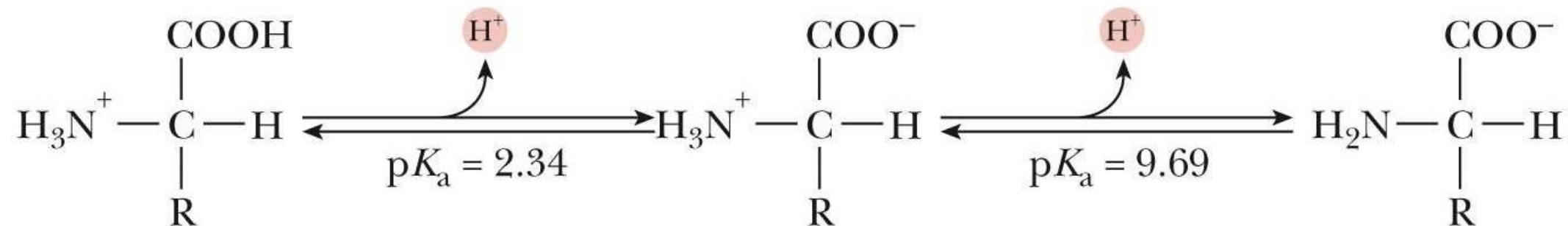
**a zwitterion**



# Effect of pH

The group or the charges each unit has its own pka While when we talk about the **isoelectric point** here, we talk about the compound

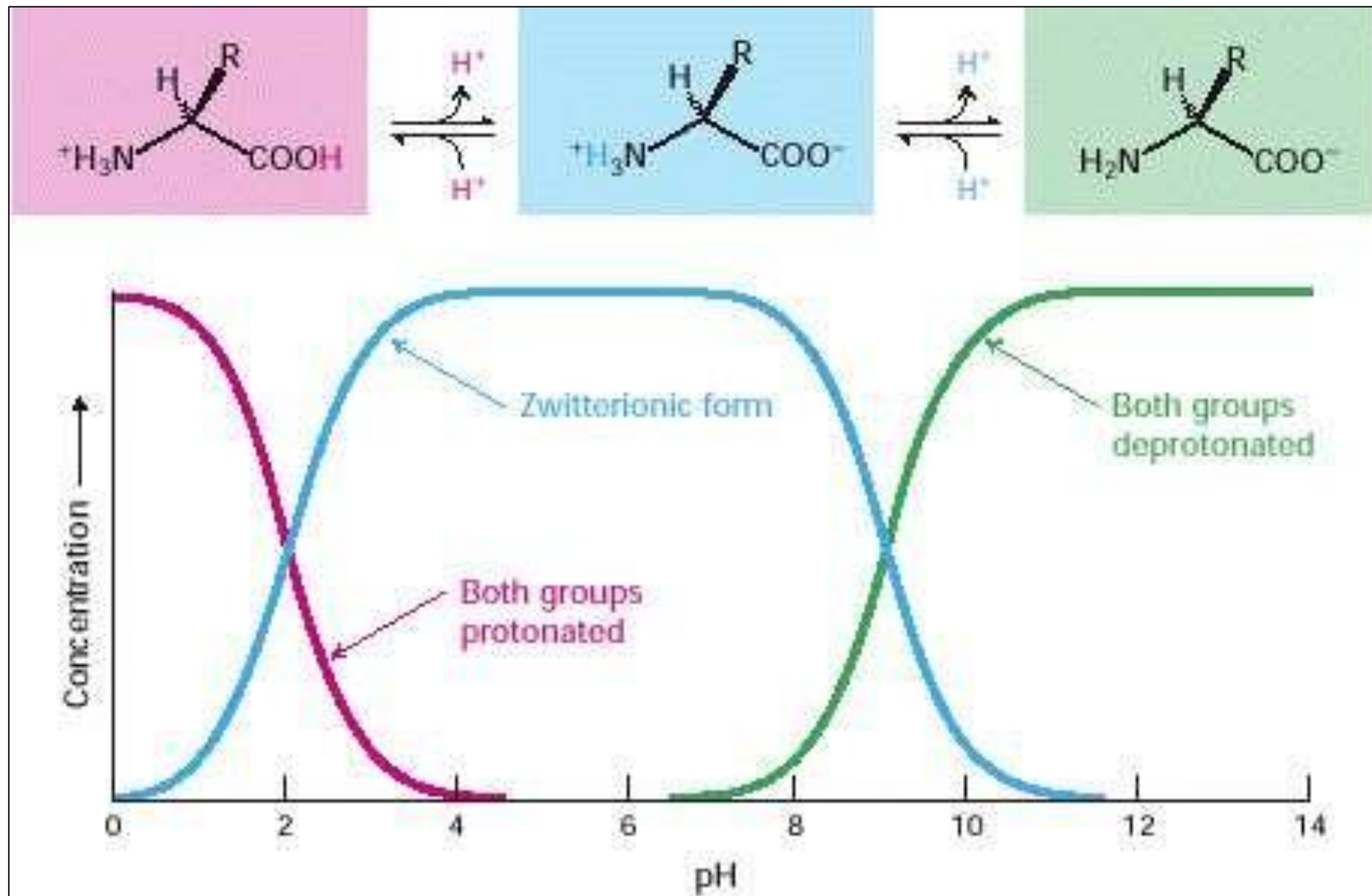
## Isoelectric zwitterion



Over total  
charges  
equal +1

Over total  
charges  
equal zero

Over total  
charges  
equal -1



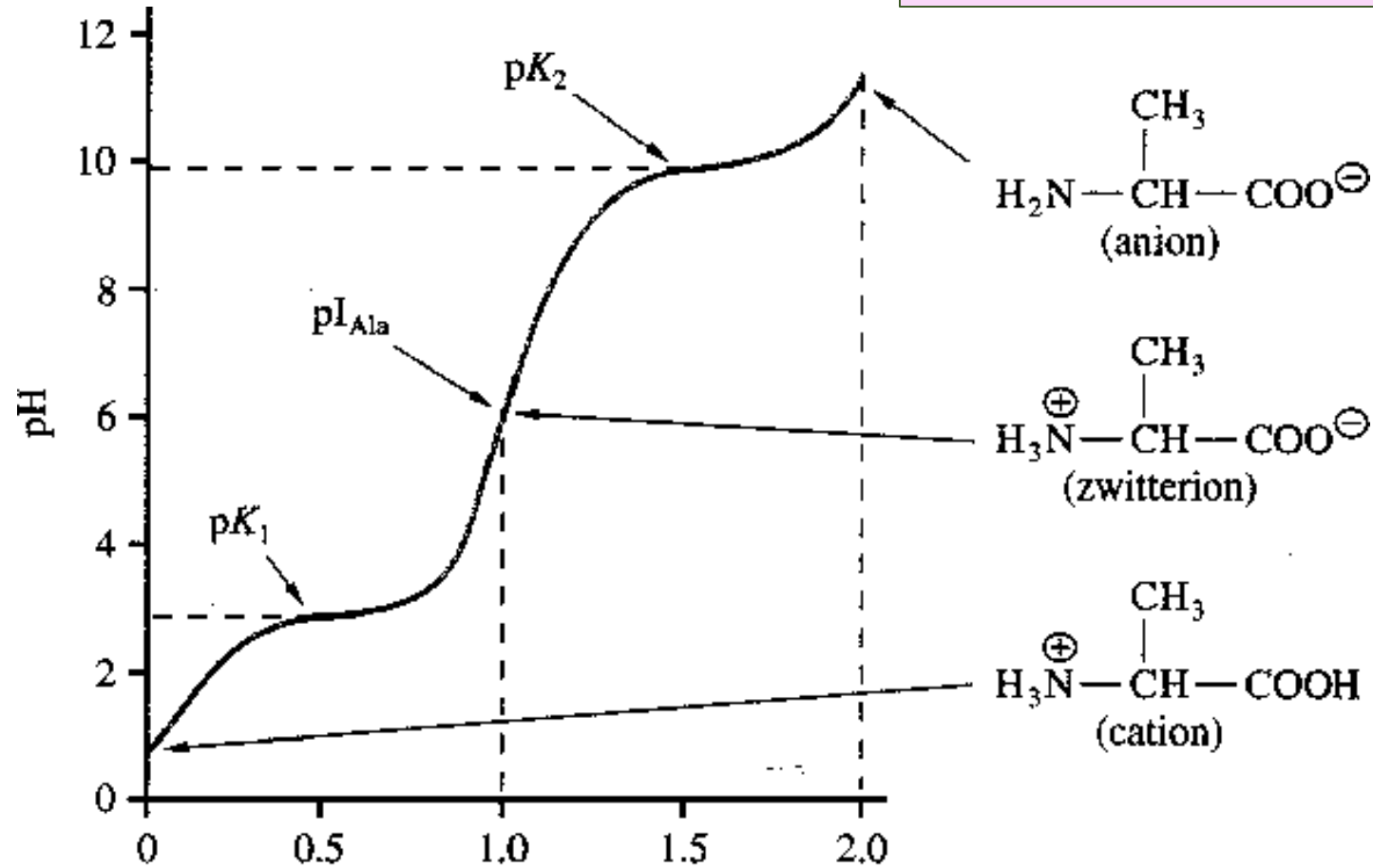
When the pH is below the pka (constant) the group will be protonated

امّ امت سڪلاو



# Example 1 (alanine)

قروتكد حرش بئي عجرن حار ياه ةديلا سلا  
ياجلا يفيوملاب اهلا نومأم روتكو لايد



$$\text{pH} = \text{pK}_a + \log \frac{[\text{conjugate base}]}{[\text{weak acid}]}$$



# Isoelectric Point

The isoelectric point of an amino acid is the point at which the amino acid has no net electrical charge

- The pH where the net charge of a molecules such as an amino acid or protein is zero is known as isoelectric point or pI.
- For the nonpolar and polar amino acids with two pKa's, the isoelectric point is calculated by taking the numerical average of the carboxyl group pKa and the α-amino group pKa.

$$pI = \frac{pK_{a1} + pK_{a2}}{2}$$



# Ionization of side chains

- Nine of the 20 amino acids have ionizable side chains.
- These amino acids are tyrosine, cysteine, arginine, lysine, histidine, serine, threonine, and aspartic and glutamic acids.
- Each side chain has its own pKa values for ionization of the side chains.

## Extra information (it wasn't mentioned) :-

Seven amino acid side chains contain groups that ionize between pH 1 and 14. For Asp, Glu, Tyr, and Cys, the ionizable groups are uncharged below their pK and negatively charged above their pK. For His, Lys, and Arg, the ionizable groups are positively charged below their pK and uncharged above their pK.



# pH of amino acids

## Isoelectric Point

Amino Acid	Side Chain $pK_a^3$	pI
Arginine	12.5	10.8
Aspartic Acid	4.0	3.0
Cysteine	8.0	5.0
Glutamic Acid	4.1	3.2
Histidine	6.0	7.5
Lysine	11.0	10

Aspartic & Glutamic Acids are much lower than physiological PH (more acidic) so they must be deprotonated

Histidine is close to physiological PH

*Let's consider  $pK_a$  of  $-NH_2 = 9$  and  $pK_a$  of  $-COOH = 2$  for all amino acids*



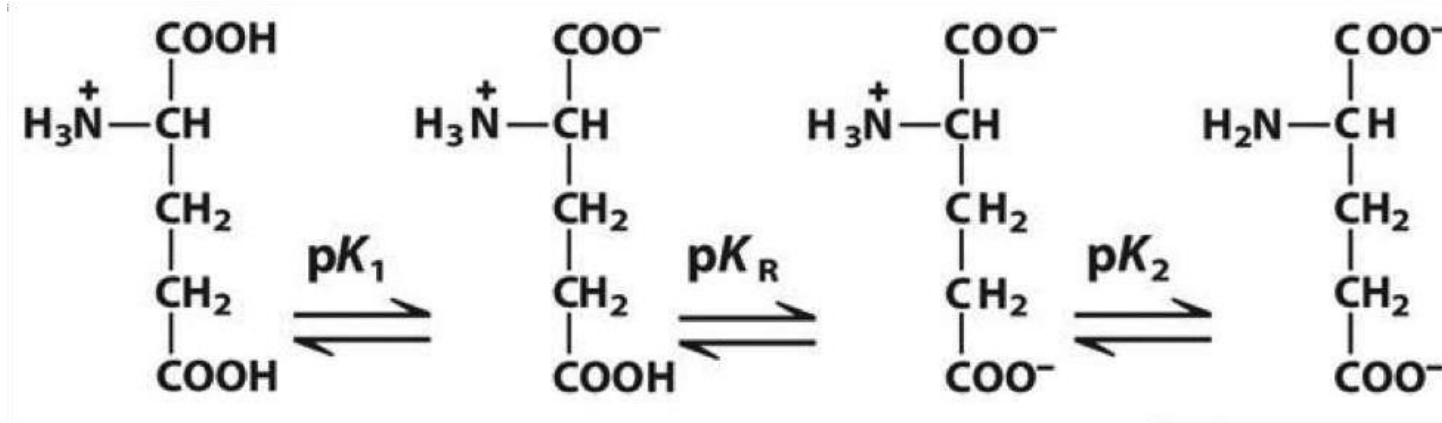
# Calculation of pI of amino acids with ionizable R groups

- The isoelectric point for these amino acids is calculated by taking the average of the pKa's of the groups with same charge when ionized
- In this case, the total charge on the groups with like charge must equal one (1) so that it can be balanced by the one (1) opposite charge present on the molecule

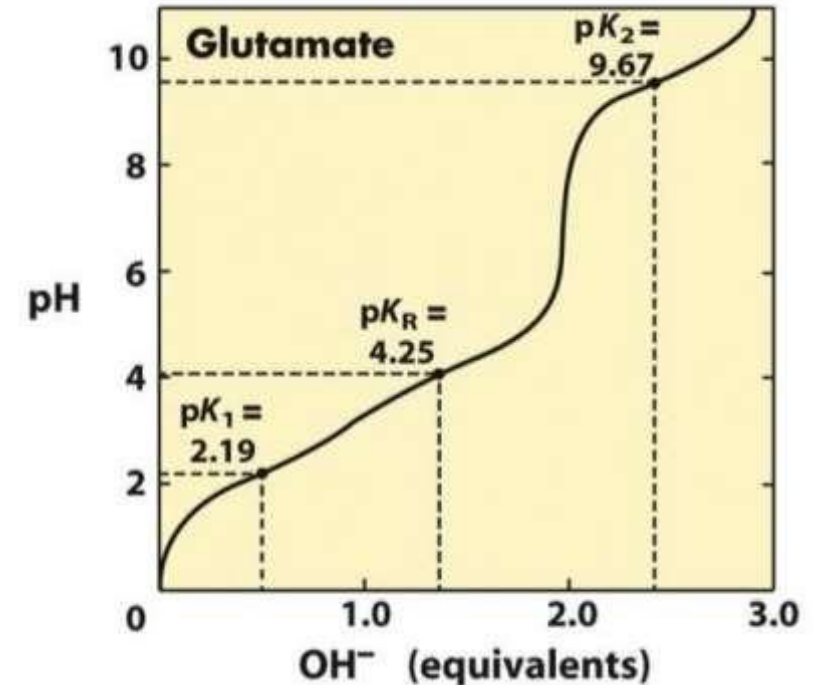
# Example: Glutamate



It is like example 1 , slide 46 :-  
It will be explained in more details  
in the next modified



- To calculate the isoelectric point of Glu, the pKa's of the two carboxyl groups are averaged.

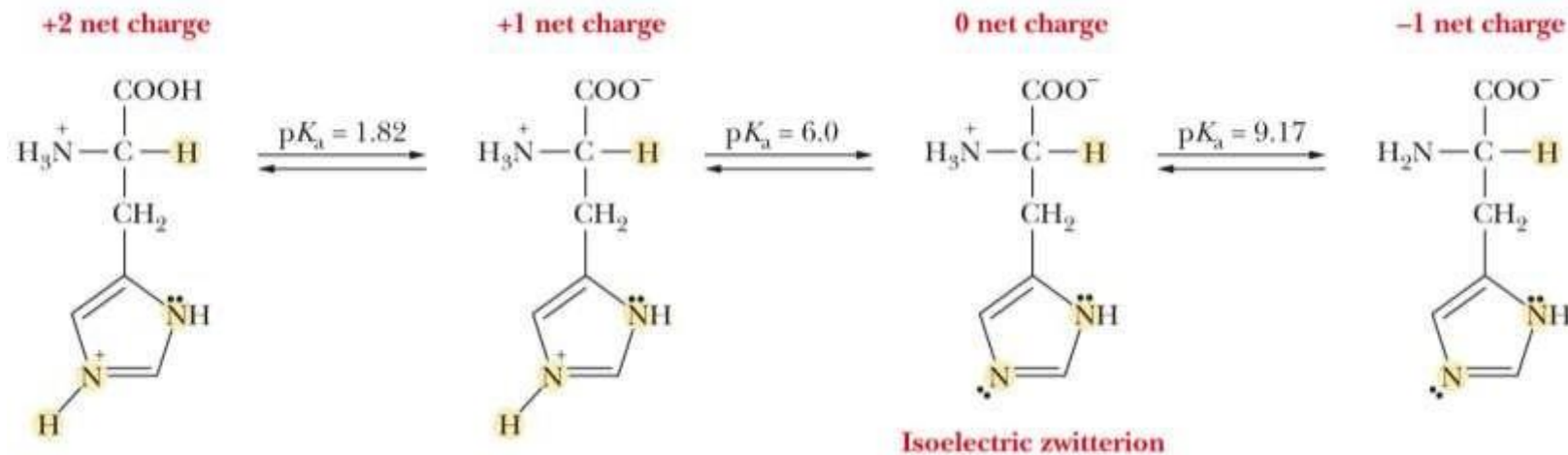
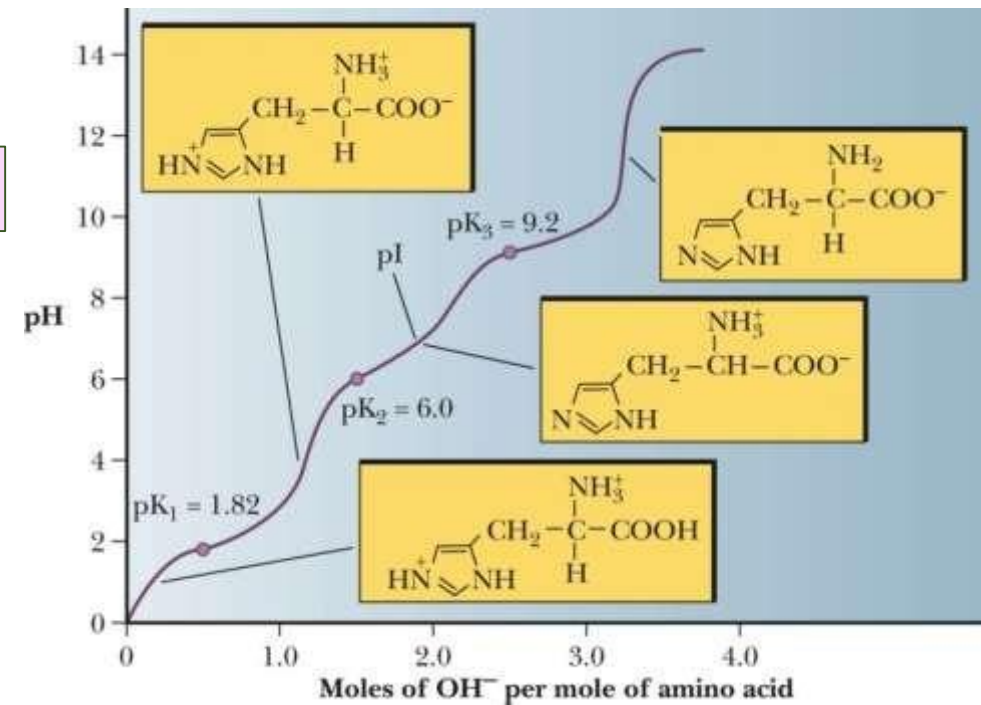




# Histidine

A basic amino acid

- $\text{pH} \approx 7.5$  (The imidazole group can be uncharged or positively charged near neutral pH).



Further explanation in next slide:-





## Explanation to the previous slide:-

R groups for histidine play an important role (buffering) because it is around physiological PH value.

Its  $P_k=6$  in an individual histamine.

# If histamine was a part of a large molecule (like large protein) histamine's R group will be affected by the environment (like hydrogen , electrostatic bond , van der Waals by taking or giving electrons to histidine R group)

# So, when histamine is apart from protein its pka will not stay 6, it will change to 7.3.

# If this protein is in cell ,blood ,cell surface, it is going to accept or donate protons and it can regulate a physiological ph.

# Albumin is a plasma protein ( which is the most abundant protein in blood 60%) contains 16 histamine residues or (16 R group) which plays an important role in buffering.



# Questions

- Draw the titration curve of histidine.
- What is the ratio of conjugate base/acid of glutamate at pH 4.5?
- What is the total charge of lysine at pH 7?