

Chapter:

Biochemistry



Aminoacids, Proteins & Nucleic Acids

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Chemistry with MJS

Chemistry Preparation by MJS

Amino Acids

The proteins in all living species from bacteria to humans, are made up of 20 amino acids.

Amino acids contain two group

Amino group = $-NH_2$

Carboxylic group = $-COOH$

Classification of Amino acids:

Amino acids can be classified in various ways.

(i) Either Acidic, Basic or Neutral

(ii) Either Non-polar or Polar R group.
Polar R group is further divided into:
-ve charged and +ve charged

(iii) Essential or Non Essential -

(iv) α - β - γ , δ Amino acids.

Classification - A (at pH=7)

• Acidic Amino acids: (2 out of 20 A. acid)

having polar side chain $-COOH$ group \Rightarrow $-NH_2$ group

e.g. glutamic acid, Aspartic acid etc.

- Basic Amino acids: (3 → out of 20)

↓
having
polar side
chain

Amino groups > carboxylic group

e.g Arginine, lysine, ~~etc~~, Histidine

- Neutral Amino acids: (15 out of 20)

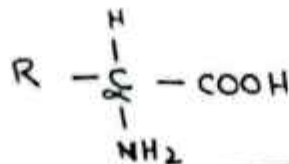
↓
(may have polar or non polar side chain)

Amino groups = carboxylic group

✓ classification- B :

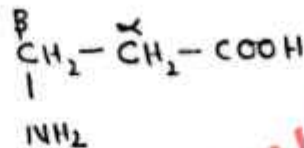
General
formula

• α-Amino Acid

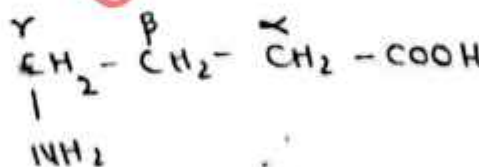


∴ Almost all naturally occurring amino acids are α-A. Acids

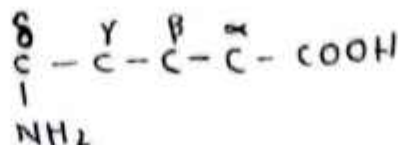
• β-Amino acid



• γ-Amino acid



• δ-Amino acid



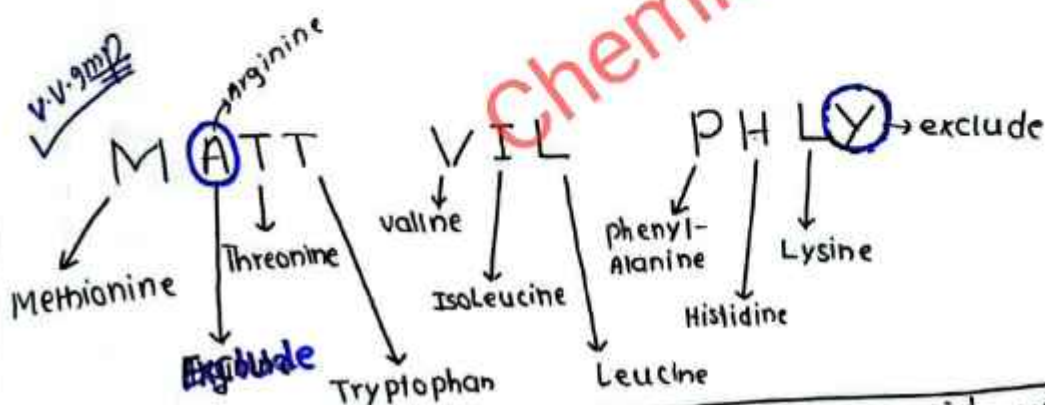
Chemistry with MJS

Classification - C :

(i) Essential Aminoacids : (out of 20 \rightarrow 9 are essential) amino acids

\downarrow
which can not be prepared by our body.
 \rightarrow deficiency of essential amino acids may cause diseases

Trick to learn essential-Aminoacids:



NOTE: Arginine is non-essential amino acid which is called a (conditionally-essential amino acid)

(ii) NON-ESSENTIAL Amino acids :

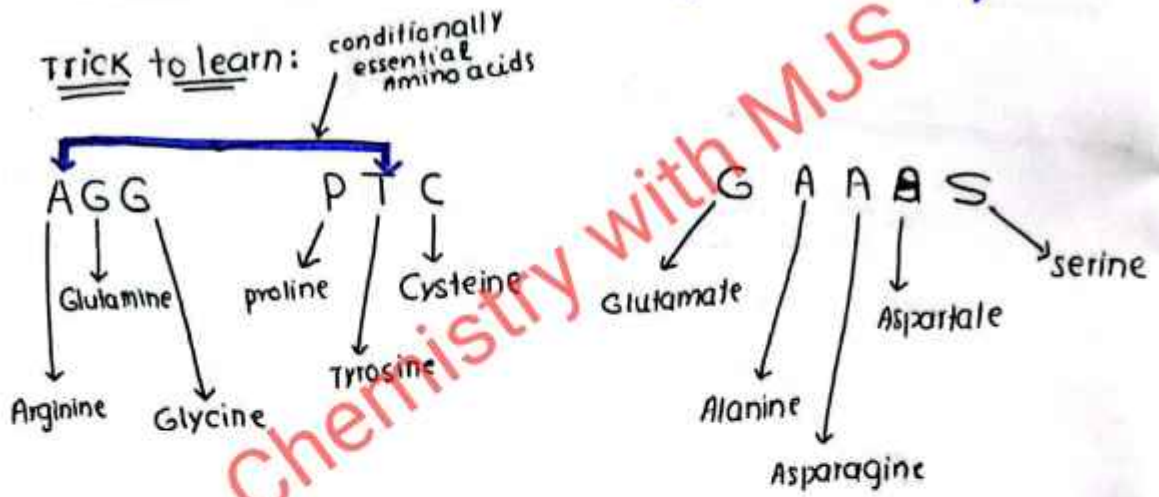
\downarrow
which can be synthesized through our body called non-essential amino acids

\rightarrow Left 11 amino acids are non-essential. but out of 11 \rightarrow 6 amino acids are conditionally essential amino acids.

Conditionally Essential Aminoacids : \rightarrow (Also in non-essential category)
 (are six)

\downarrow
Their synthesis can be limited under special pathophysiological conditions such as prematurity in the infant or individuals in severe catabolic distress or sometimes during illness.

✓: NON essential Amino Acids: → (11 out of 20)



NOTE

Arginine is a non-essential A-Acid which is called a conditionally-essential amino acid. Arginine is abundant in protamines and histones - both proteins associated with nucleic acids and was 1st isolated in 1895 from animal horn

Classification-D: → (classification based on polarity & non polarity of side chain (R))

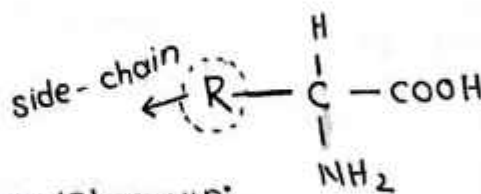
4-categories

∩ Polar ∴ non polar = 9 ✓
 ∴ polar = 11 ✓

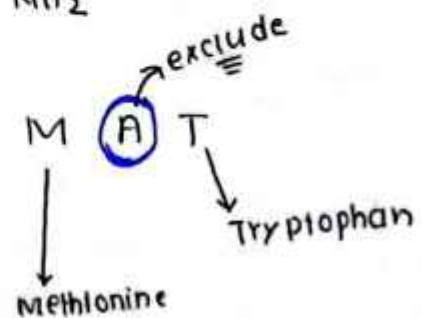
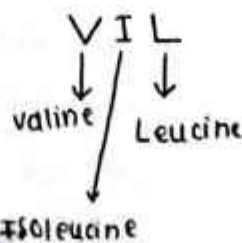
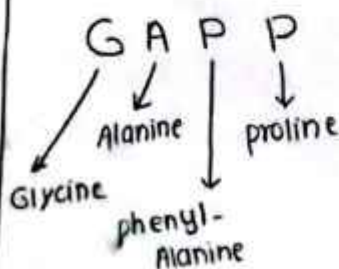
(i) Neutral NON-polar Amino acids: → (9)

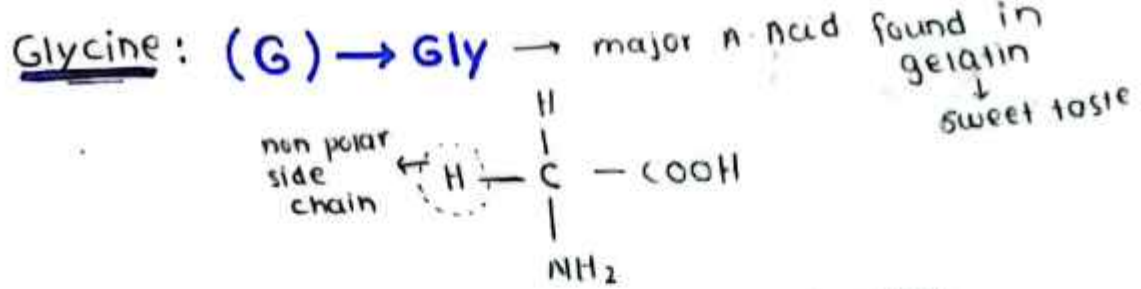
(Hydrophobic)

General Formula

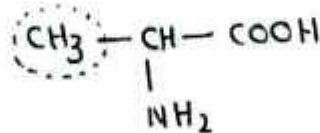
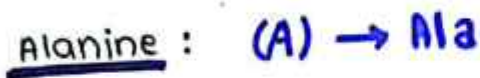


Trick for non-polar (R) → group:

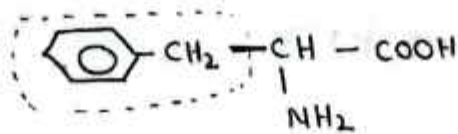




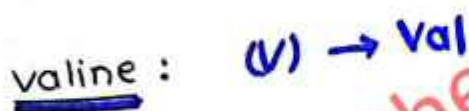
- ∴ only Amino acid which has no-chiral carbon
- ∴ simplest Amino acid



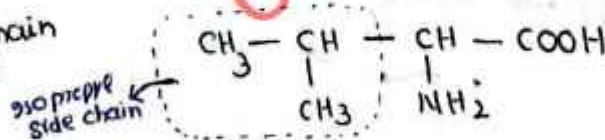
- Simplest A. Acid after glycine.
- It convert glucose into Energy
- Eliminate excess toxins from your liver.



- → classified as aromatic amino acid
- It is found in mother milk & no. of foods.
- It is studied for its effects on depression, pain & skin disorder.



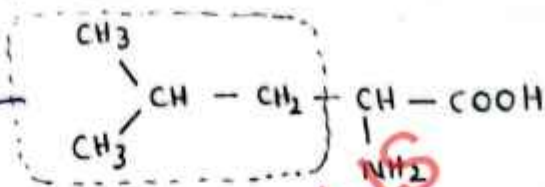
→ Branched chain Amino acid



- First isolated from casein in 1901 by Emil Fischer
- The name comes from valeric acid
 ↓
 (5-C)

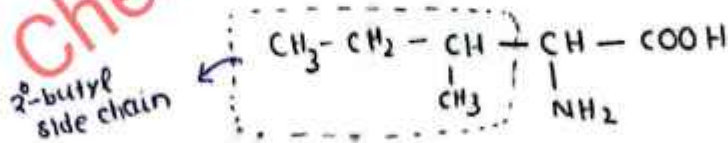
Leucine: (L) → Leu

→ Branched chain
Amino acid
↳ bulky side chain



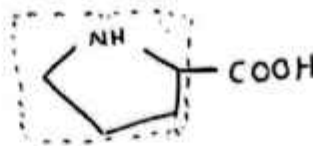
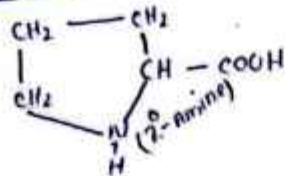
- It is encoded by the codons UUN, UUG, CUU etc.

ISO-Leucine: (I) → Ile



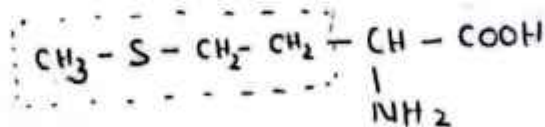
- An essential amino acid b/c most animals can not synthesize branched-chain amino acids.
- particularly recommended to professional athletes & body builders
- primary function is to boost up energy level.

proline: (P) → Pro



- → contains a 2°-amine group, referred to as an α -imino acid
- plays important role in molecular recognition particularly intracellular signalling.

Methionine: (M) → Met

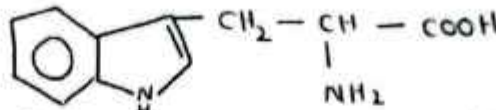


- side chain functions as a methyl group donor
- produce cystein & other sulphur-containing amino acids

Tryptophan: (W) → Trp

- its purpose is to balance the N in adults & growth in infants

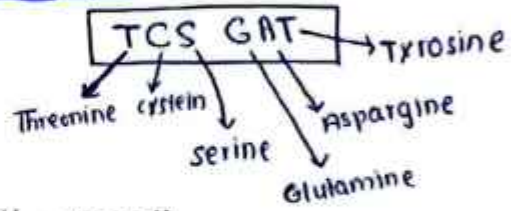
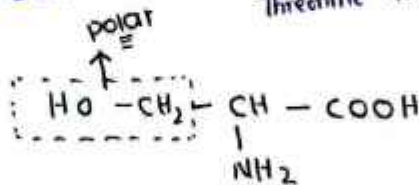
→ Aromatic Amino Acid.



- it creates niacin → which create neurotransmitter serotonin

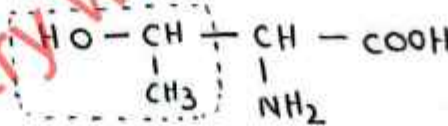
(ii) Neutral polar Amino-acids: → (6)

Serine: (S) → Ser



- It is found at the active site of many enzymes.
- Important in metabolism & participates in the biosynthesis of purines & pyrimidines
- It is precursor to several A-Acids e.g. glycine, Cysteine & Tryptophan in bacteria.

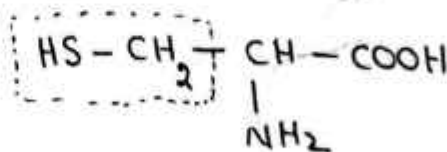
Threonine: (T) → Thr



- Named for its similarity to the sugar Threose
- It supports CNS, cardiovascular, liver & Immune Functioning

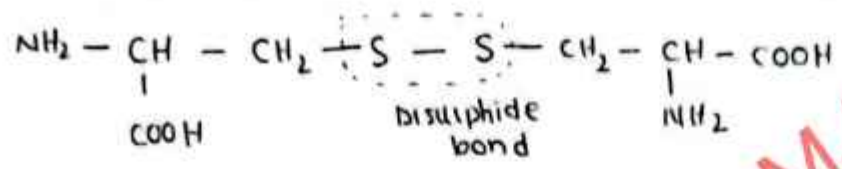
Cysteine: (C) → Cys

- play imp. role in communication b/w immune systems cells

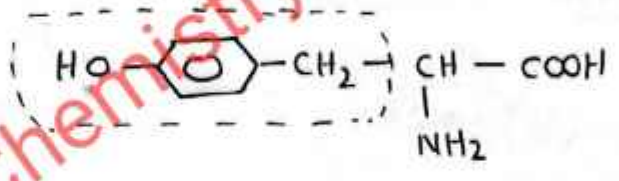


- oxidation of two cysteine molecules linked by di-sulfur bond → yields cystine

Cystine



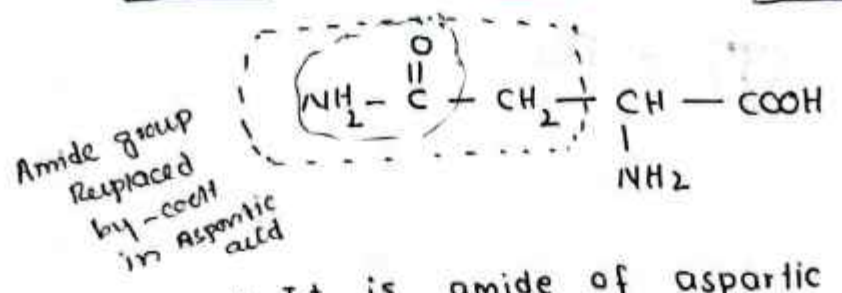
Tyrosine: (Y) → Tyr



- Aromatic amino acid.
- play imp. role in photosynthesis.
- produce brain chemical that help nerve cells communicate.

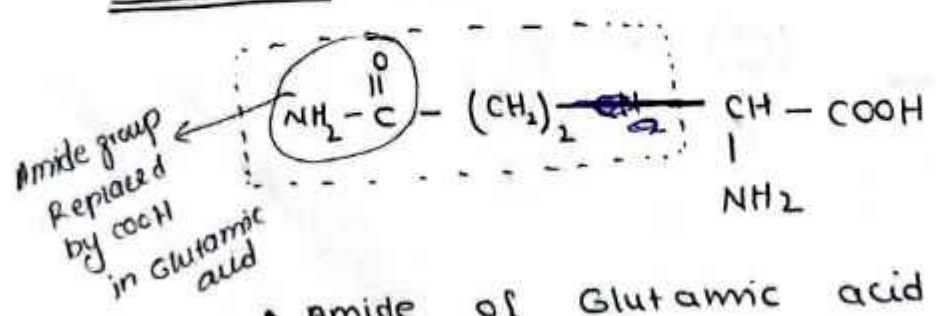
* Asparagine: (N) → Asn

↳ 1st A. acid that was isolated in 1806 → was found in asparagus juice.



- It is amide of aspartic acid
- play imp. role in the synthesis of NH
- Required for the development & function of the brain

Glutamine: (Q) → Gln

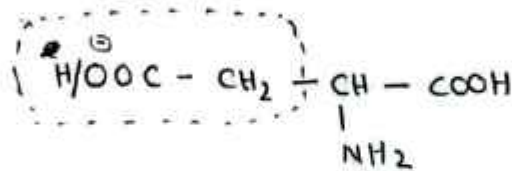


- Amide of Glutamic acid
- critical part of immune system
- play imp. role in intestinal health.

(iii) Acidic polar Amino acids: → (2)

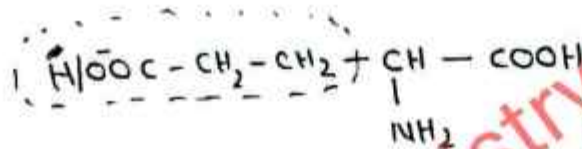
↓
Amino acids with -vely charged
R- Group

Aspartic acid: (D) → Asp



- carboxylic groups are ionized at physiological pH; Also known as aspartate
- play imp. Role in Hormone production & release

Glutamic acid: (E) → Glu



- carboxyl groups are ionized at physiological pH; Also known as Glutamate
- important Neurotransmitter.

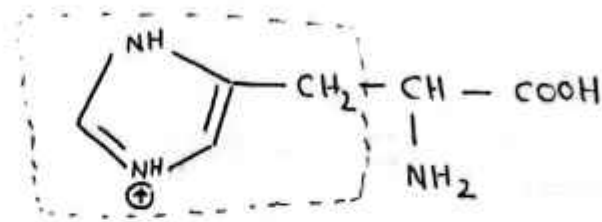
Classification of 20 Amino acids:

- | | |
|---------------------|------------------------------------|
| → Acidic A.A = 2 | → non-essential A.A = 11 |
| → Basic A.A = 3 | → α-A.A (all A.A) = 20 |
| → Neutral A.A = 15 | → optically inactive = 1 (Glycine) |
| → polar A.A = 11 | → Aromatic A.A = 3 |
| → non-polar A.A = 9 | → Aliphatic A.A = 17 |
| → Essential A.A = 9 | → Ring containing = 5 |
| | → S-containing A.A = 2 |

(iv) Basic polar Amino-acids: → (3)

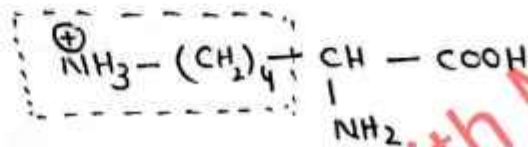
↓ Amino acids with +vely charged R-group

Histidine:



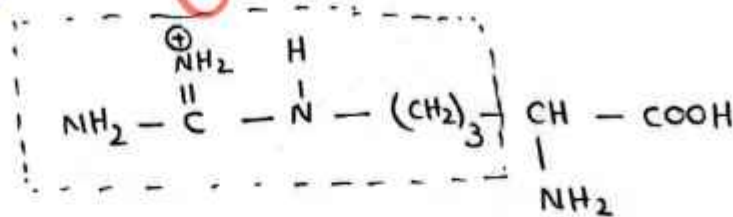
- has imidazole F. group.
- imidazole → common participant in enzyme catalysed Reaction
- it is vital in hemoglobin.

Lysine:



- Elevated in blood level cause disease called Hyperlysinemia

Arginine:



- Almost as strong base as NaOH
- ~~Hyperlysinemia is caused by the deficiency of enzyme~~
- It is precursor for the synthesis of NO play imp. Role in Regulation of blood pressure
- imp. Role in the Removal of NH₃ from the body.

✓ Configuration:

All amino acids except glycine has chiral carbon has configuration and found in two forms (i) L-form (ii) D-form

NOTE

✓✓

The Amino acids found in proteins are (L-Amino acids)

✓ Physical properties:

(i) Amino acids are colourless, crystalline substances having sweet taste.

(ii) They melt with decomposition at higher temperature (more than 200°C)

(iii) They are soluble in H_2O but insoluble in organic solvent

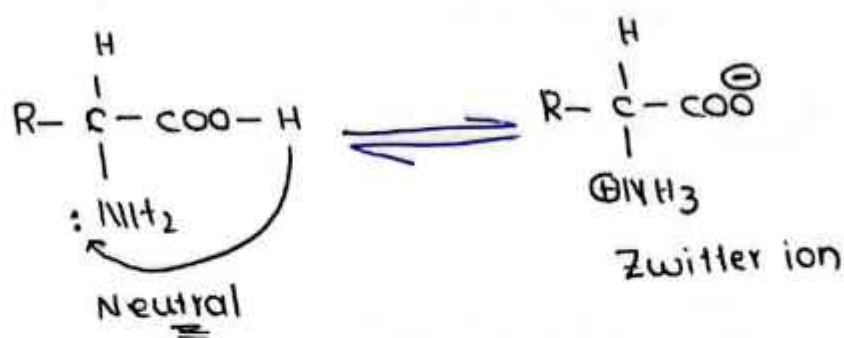
(iv) Optical Activity:

Except Glycine All α -Amino acids are optically active and have a chiral centre. They have two optical isomers called enantiomers (D & L form)

(v) Zwitter ion & Isoelectric point:

$-\text{NH}_2$ group is basic & $-\text{COOH}$ group is acidic. In neutral solution it exists as an internal ionic form called a Zwitter ion where proton of COOH group transfer to

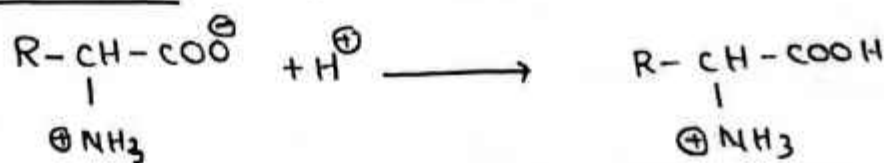
-NH₂ group to form inert salt also called dipolar ion



Zwitter ion is dipolar charged but overall electrically neutral, → contains both +ve & -ve charge

(vi) Acidic & Basic character:

Basic character: → acceptance of H⁺



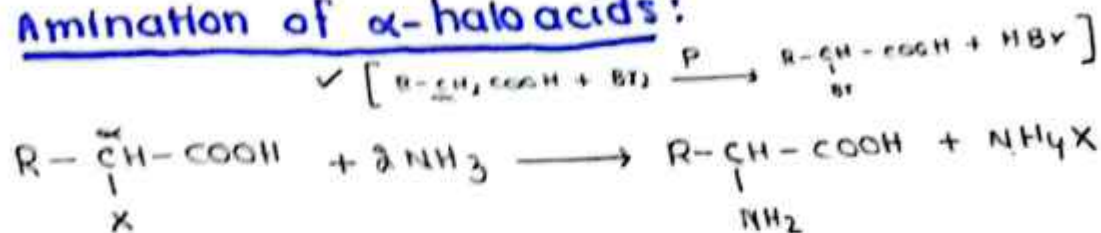
Acidic character: → donation of H⁺



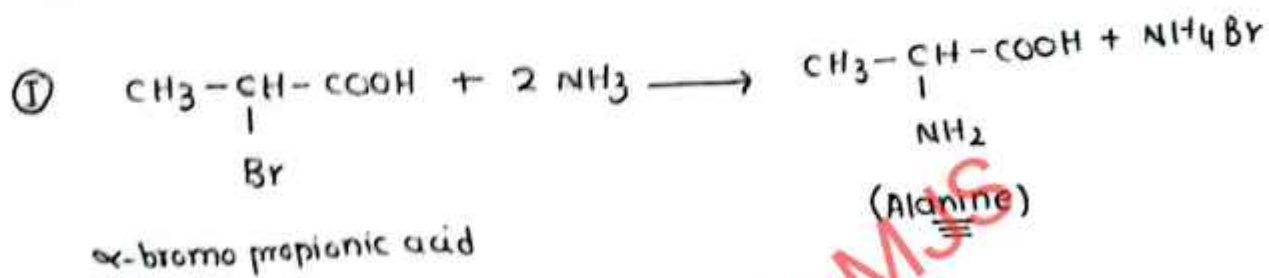
Chemistry with MJS

Preparation of Amino Acids:

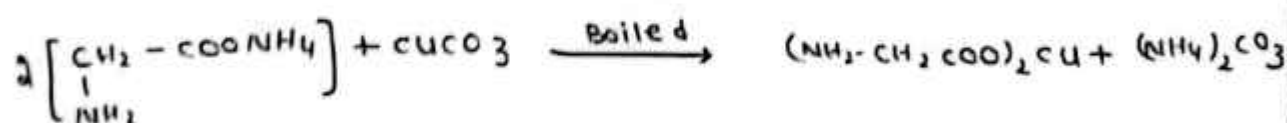
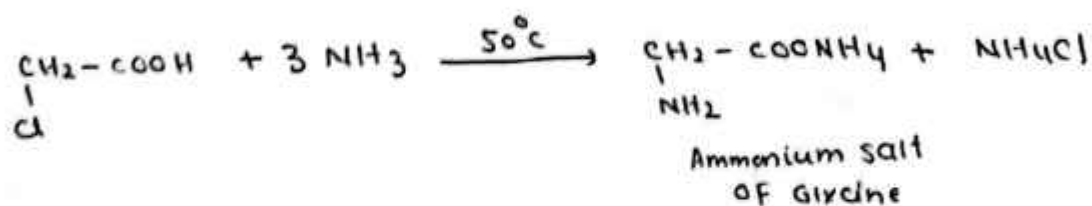
1) Amination of α -haloacids:



Example

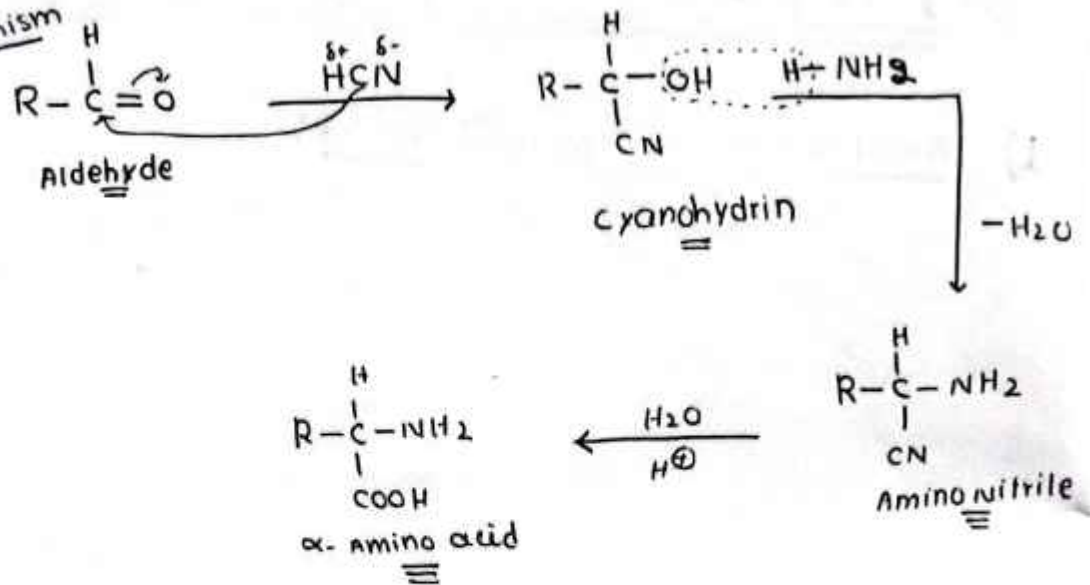


③ Lab preparation of glycine:

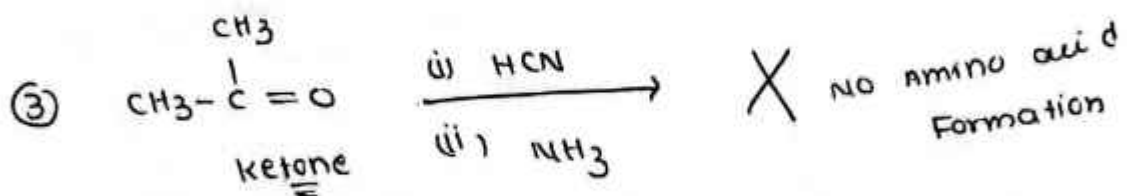
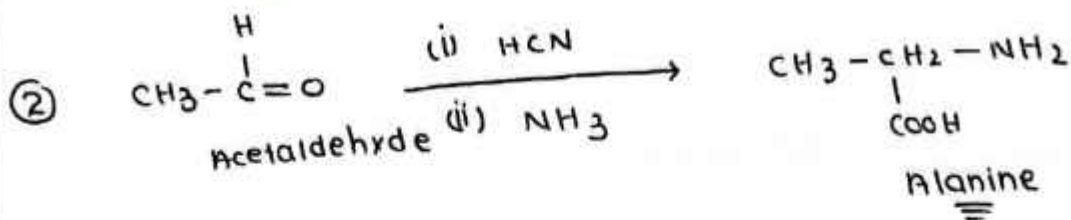
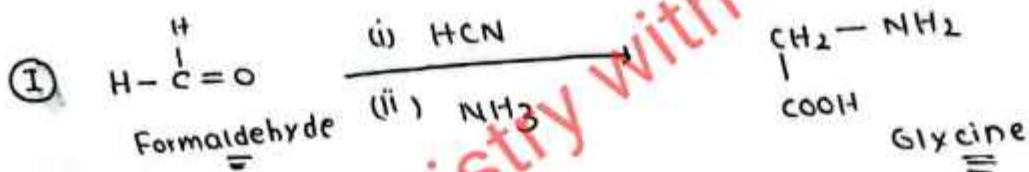


2) Strecker synthesis: (from Aldehydes)

General mechanism

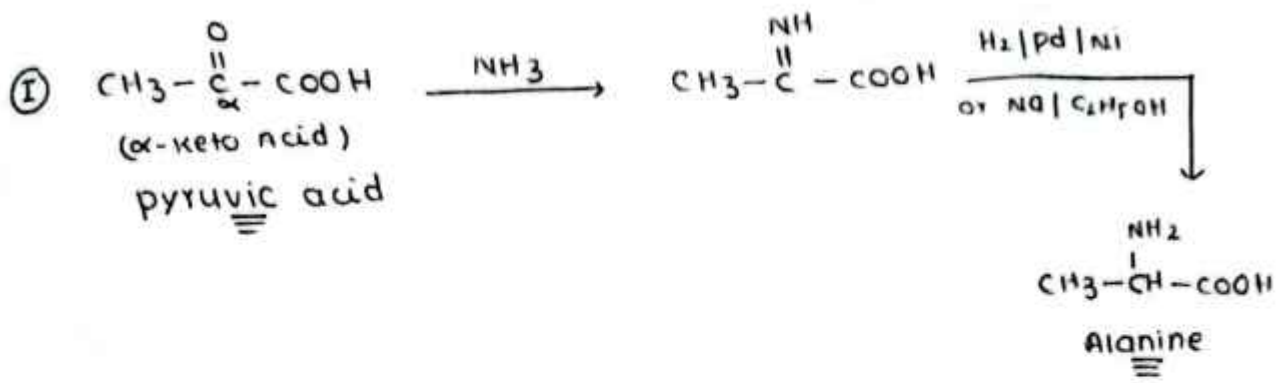


Examples

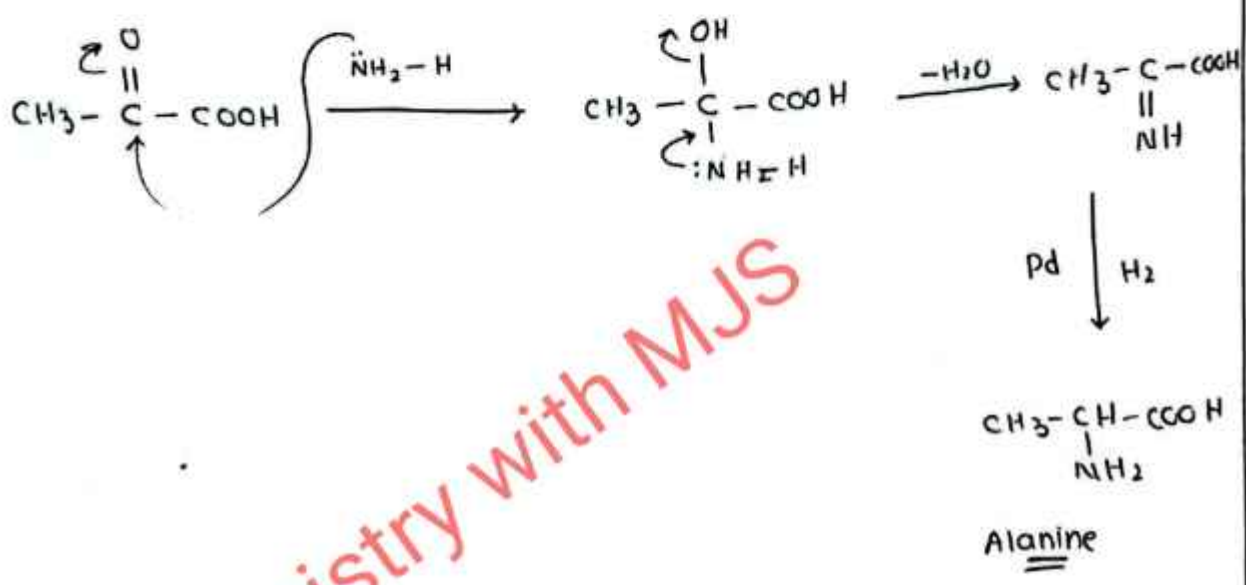


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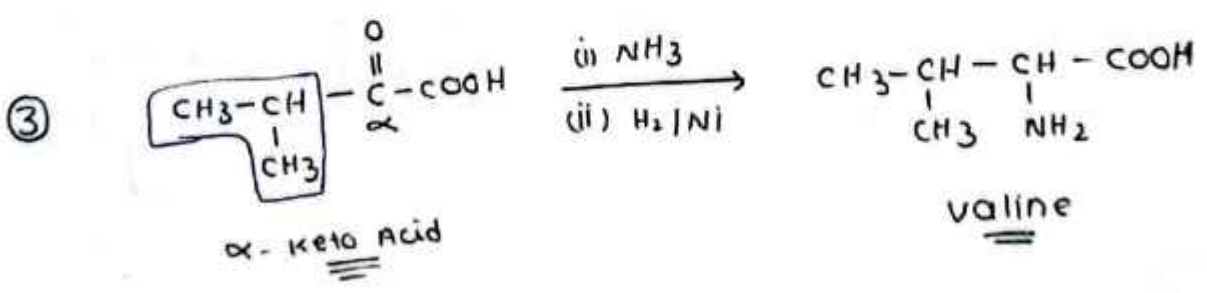
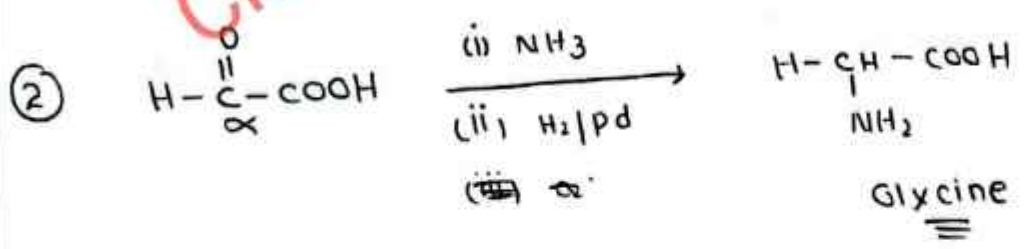
3) Koop synthesis: (from α -keto acids)



Mechanism



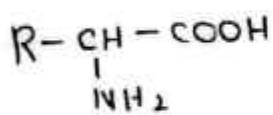
Chemistry with MJS



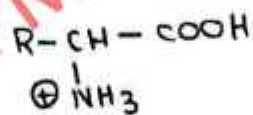
CHEMICAL PROPERTIES OF AMINO ACIDS

Amino acids are amphoteric in nature.

Depending on pH of the solution A. Acids can donate or accept proton

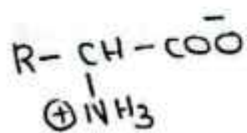


(Neutral)



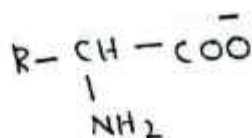
pH=0

(cation is fairly acidic medium)



(pH=7)

(Zwitter ion in neutral medium)



pH=11

(Anion is fairly basic solution)

• When an ionized form of Amino acid is placed in an electric field, it will migrate towards the opposite electrode depending upon pH of the medium

→ AT low pH:

+ve ions move towards cathode.

→ AT High pH:

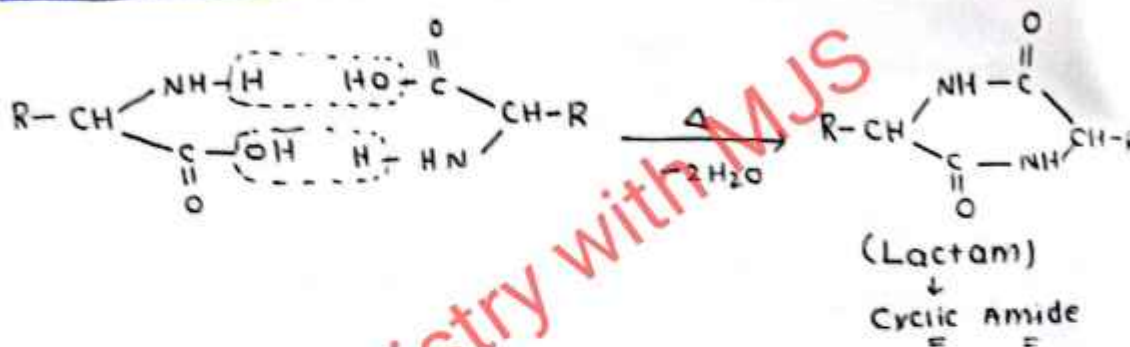
-ve ions moves toward anode

→ AT pH=7: → called isoelectric point.

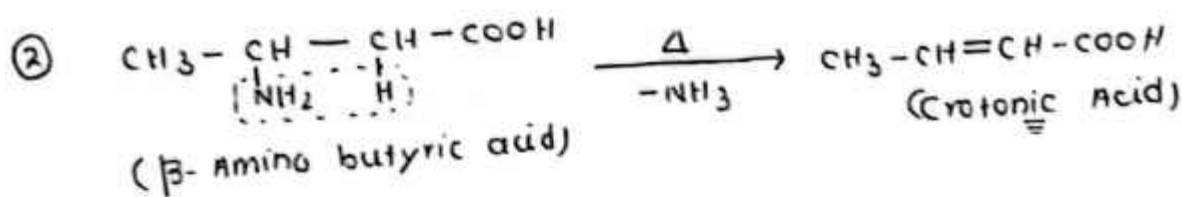
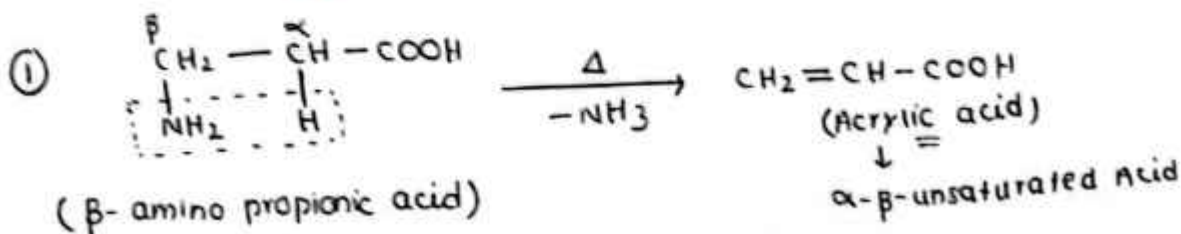
Zwitter does not move toward any electrode.

1) Effect of Heat:

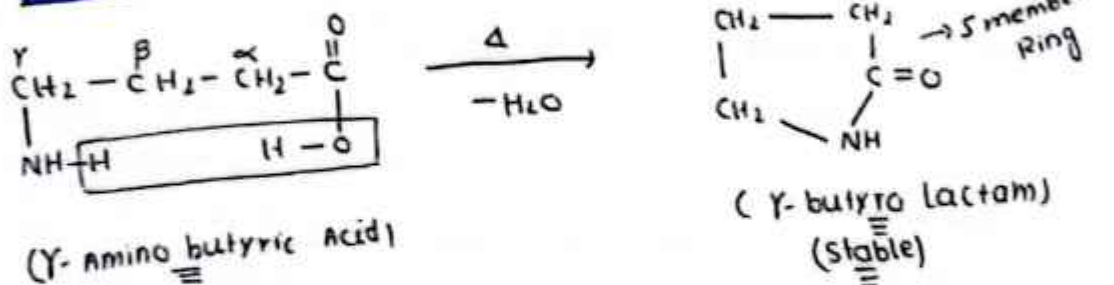
* α-Amino acids:



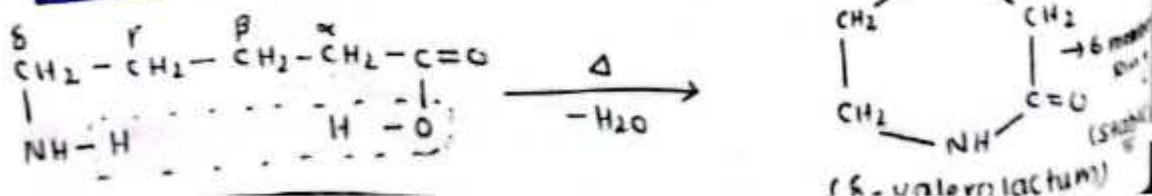
* β-Amino acids:



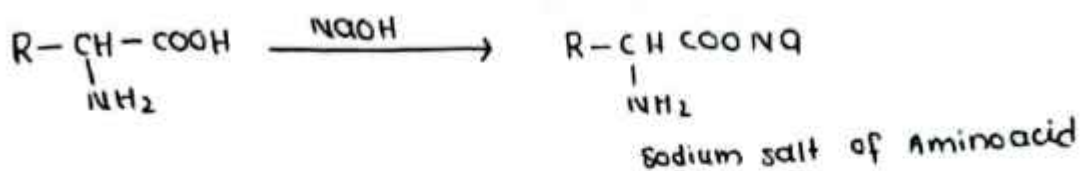
* γ-Amino acids:



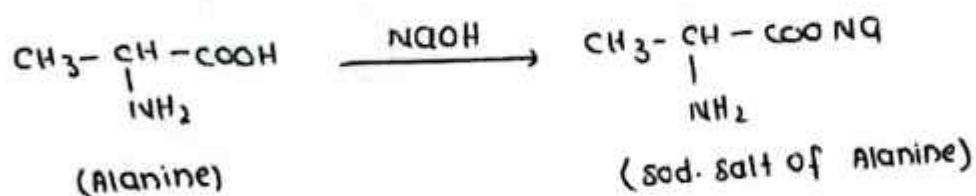
* δ-Amino acids:



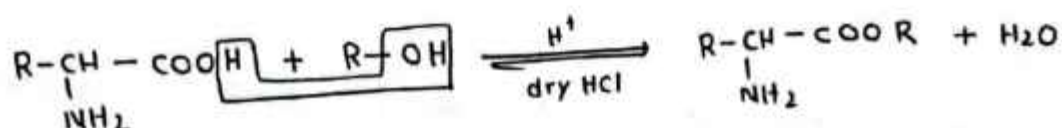
2) with Bases:



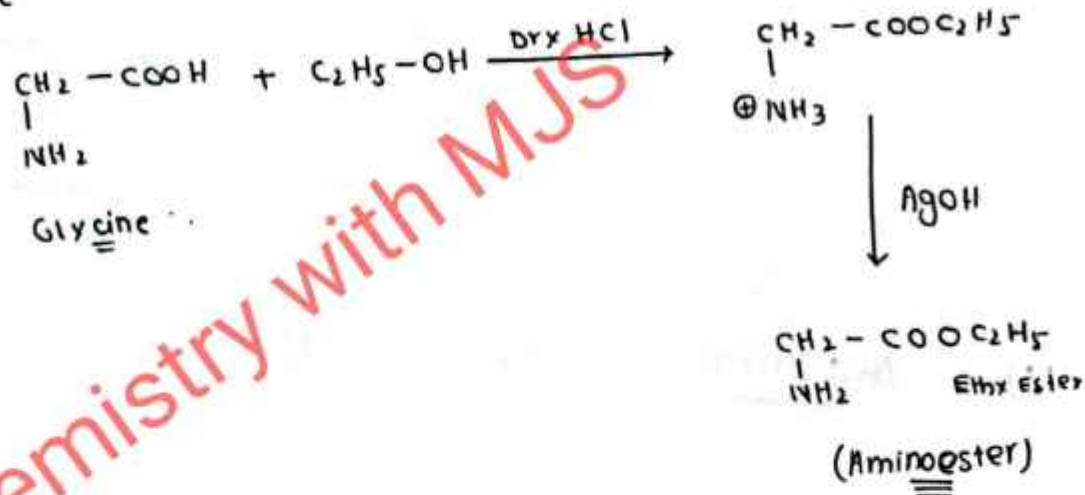
Example



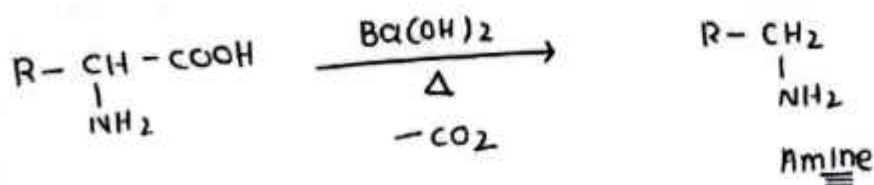
3) Esterification:



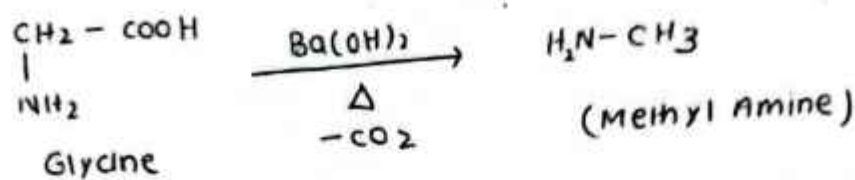
Example



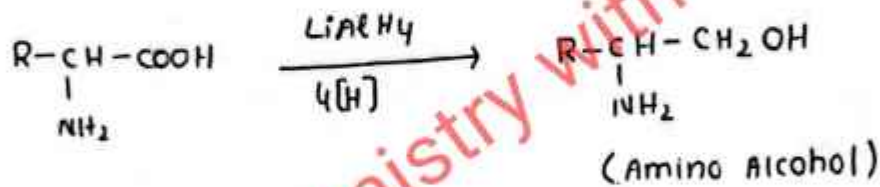
4) Decarboxylation:



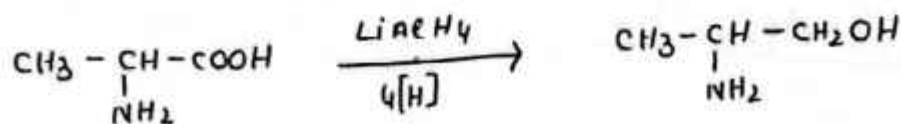
Example



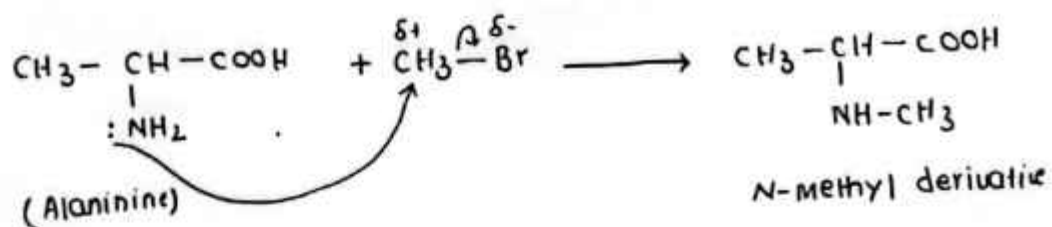
5) Reduction of Amino acids:



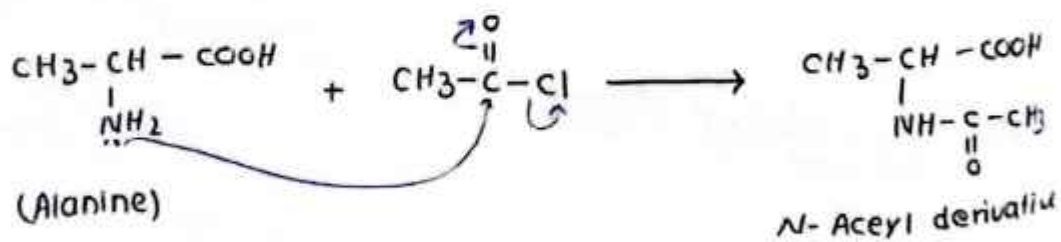
Example



6) Alkylation of Amino Acids: (E⁺-substitution)



7) Acylation of Amino Acids:



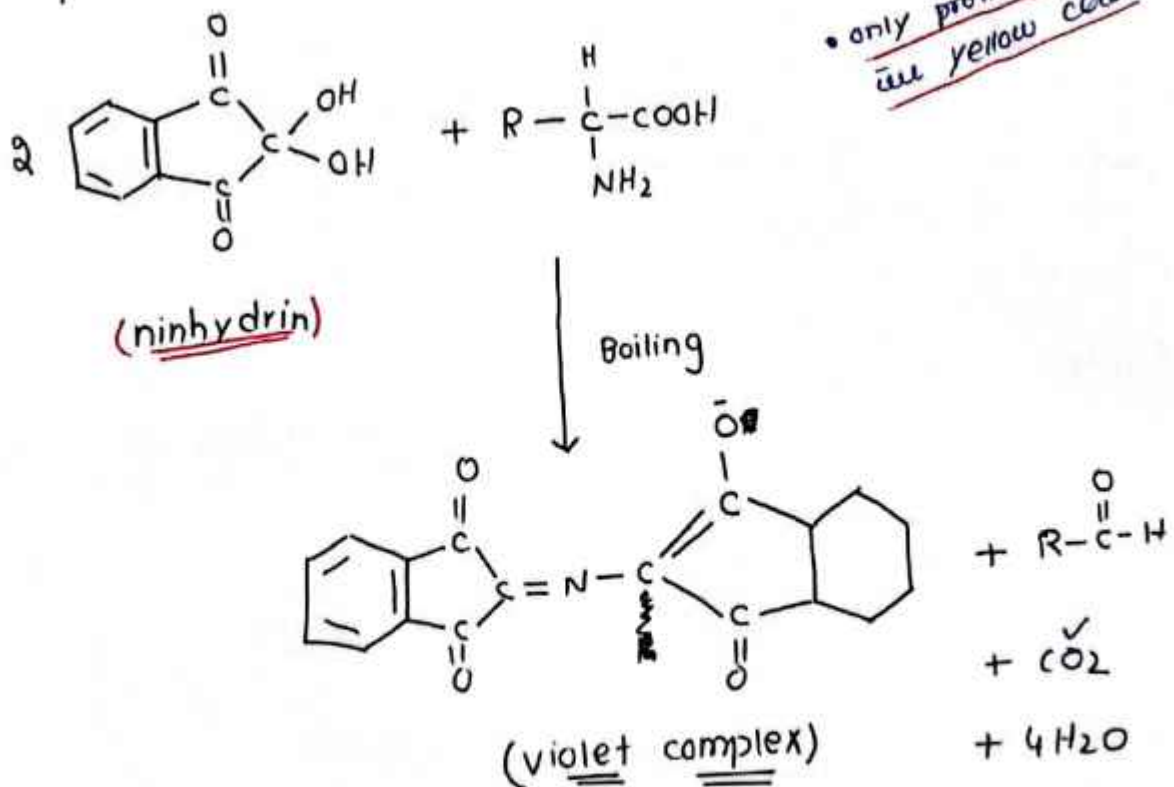
✓: Tests for Amino acids & proteins:

There are 6 tests for the detection of functional groups in amino acids & proteins

- (i) ✓ Ninhydrin test.
- (ii) Biuret test -
- (iii) Xantho proteic test.
- (iv) Millon's test.
- (v) Hopkins-cole Test.
- (vi) Nitroprusside test.

I) Ninhydrin test:

This test is given by All proteins & Amino acids. When they are boiled with a dil. solution of ninhydrin a violet colour is produced.



2) Biuret test:

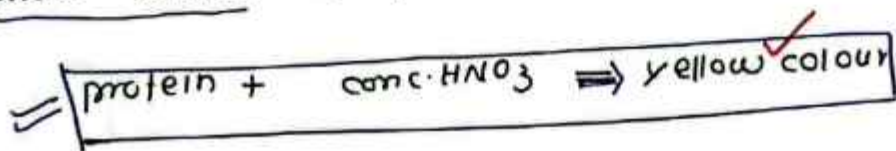
On adding a dil. solution of CuSO₄ to Alkaline soln. of protein, a violet colour is developed.

- This test is due to the presence of peptide $-\overset{\text{O}}{\parallel}{\text{C}}-\text{NH}$ linkage.

3) Xanthoproteic test:

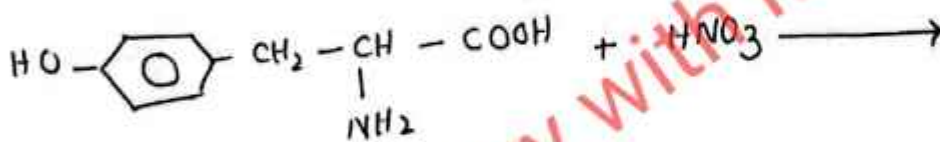
(colour test)

when protein is warmed with conc. HNO₃, a yellow colour is produced

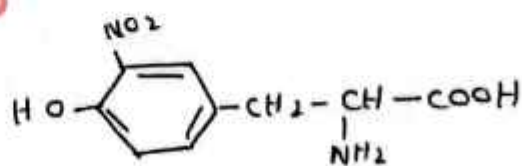


NOTE

This test is given by a protein/aminoacids that containing Aromatic/benzene ring e.g. Tyrosine & Tryptophan. Nitration of Aromatic Ring occur and gives yellow product.



Tyrosine



(yellow coloured)

4) Millon's test: → only for tyrosine. i.e. containing a phenol group

protein + Millon's Reagent = white ppt

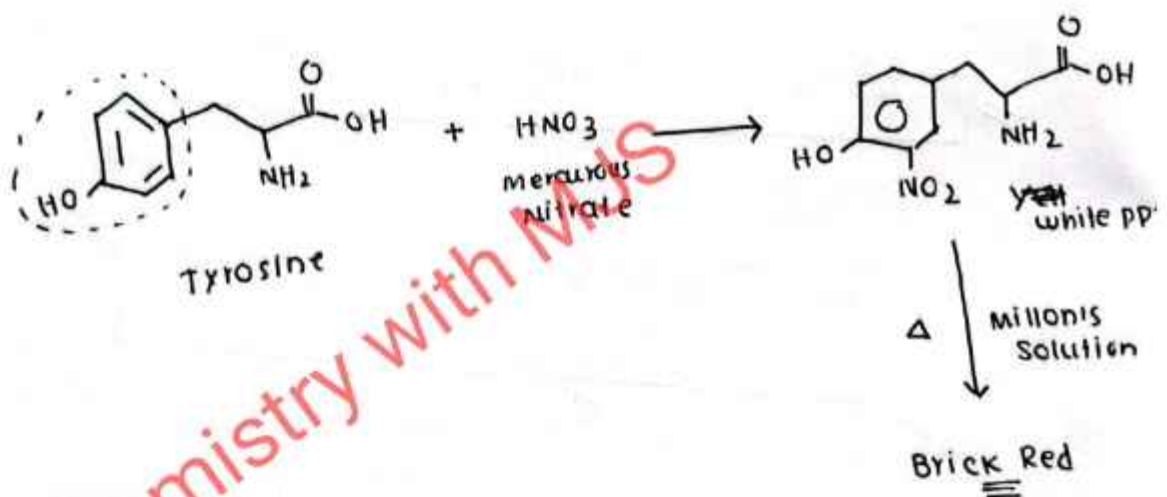
NOTE:

This test is given by protein/A.A. which yield tyrosine on Hydrolysis

Millon's test = mercury dissolved in nitric acid

NOTE

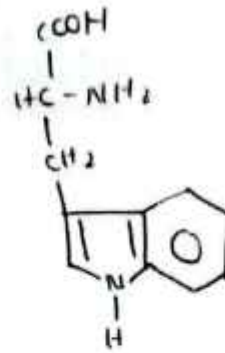
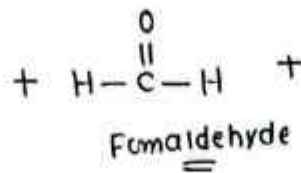
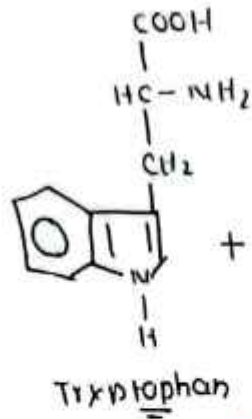
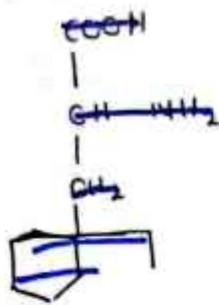
All phenols give +ve Millon's Test



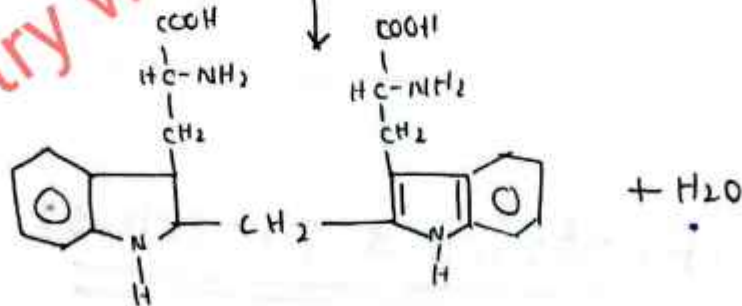
5) Hopkin's - cole test:

↓
used for the detection of specific Amino acid like Tryptophan.

⇒ on adding conc. H₂SO₄ down side containing a solution of protein/A.A., a violet colour is observed



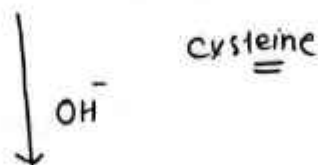
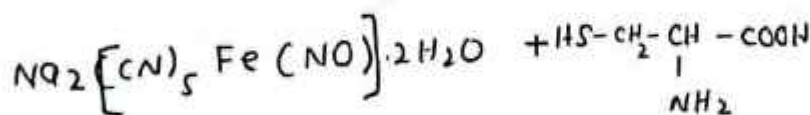
Chemistry with MJS



violet colour

6) Nitroprusside test:

↓
 It is specific for proteins/ amino acids having sulphur (-SH) group e.g. cysteine & cystine gives a red colour called "Morner test"



Red complex

7) Molisch's test: → Generally this test used for carbohydrates

This test is given by those proteins which contain carbohydrate residue.

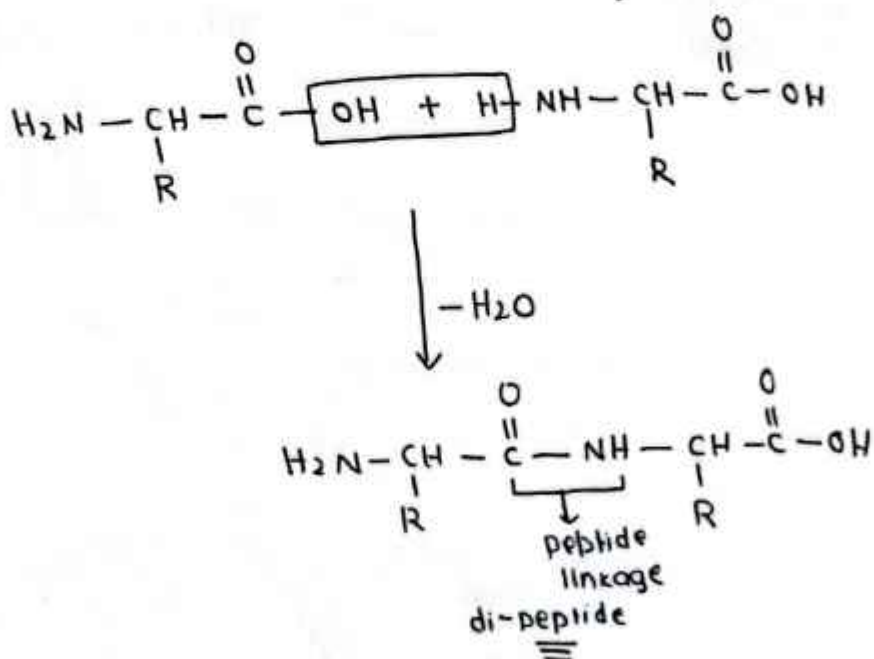
∴ Alc. soln. of α -naphthol + conc. H_2SO_4 + Glyco-protein

↓
Violet Ring

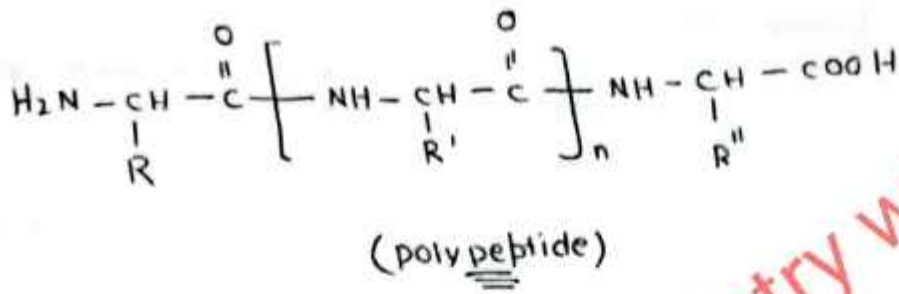
✓✓ Peptides

The bond b/w two amino acids called peptide bond and thus elimination of a water molecule occur and peptide formation occur.

peptides are classified; di, tri, tetra or penta peptides.

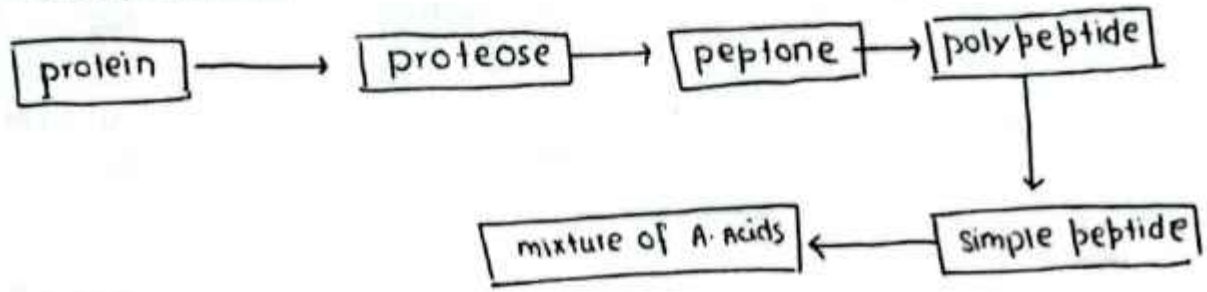


poly peptide: → when a large no. of A acids combine.



Hydrolysis of proteins:

simple proteins are hydrolysed by acids, alkalis or enzymes to produce amino acids
Hydrolysis order:



نوٹ: peptide bonds = Peptides - 1

poly peptide:

- ① it is single linear chain of upto 50 amino-acids.
 < 50 A. acids
 • mol. wt upto 10000 Dalton.
- ② unbranched chain
- ③ formed from A. acids
- ④ composed of peptide bonds

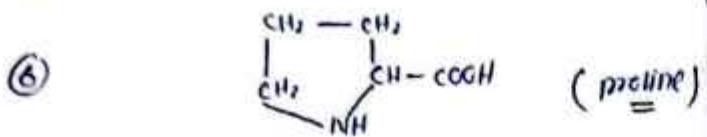
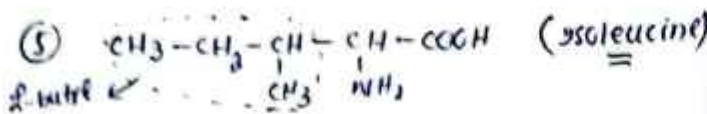
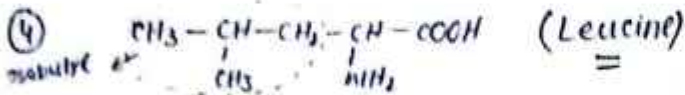
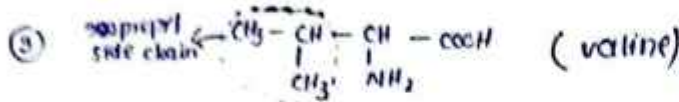
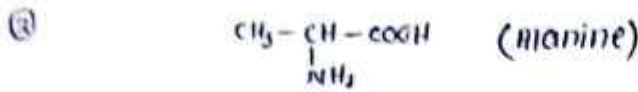
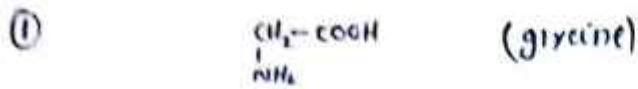
proteins

- ① it consist of one or more polypeptides (more than about 50 A. acids
 > 50 A. acids
 • mol. wt > 10000 Da
- ② complex structure
- ③ formed from polypeptides
- ④ composed of several types of bond e.g. peptide bonds, disulphide bonds, ionic bond & vanderwaals

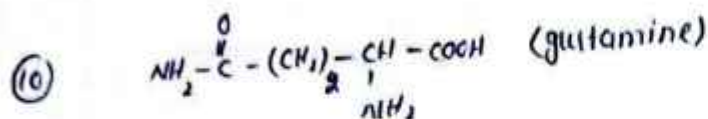
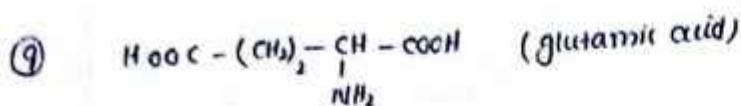
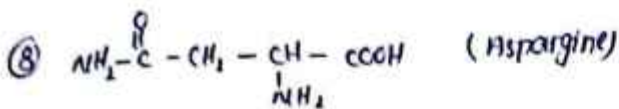
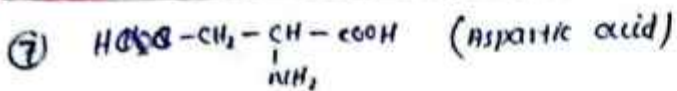
Chemistry with MJS

How to draw all amino acids structures easily

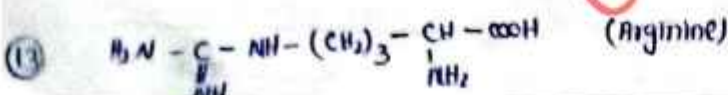
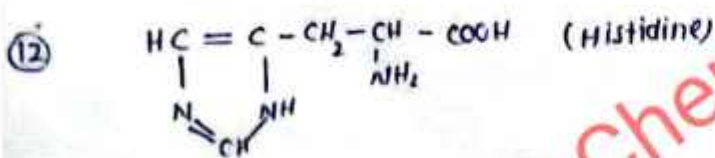
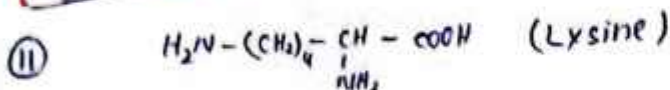
(learn series)



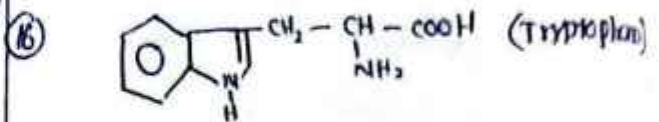
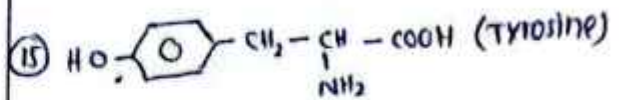
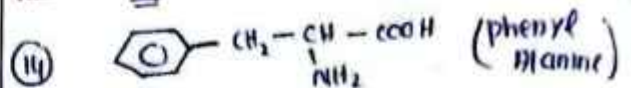
* Acidic Amino Acids & Their Derivatives



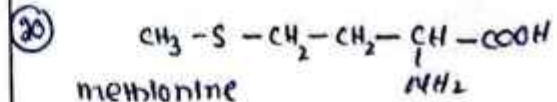
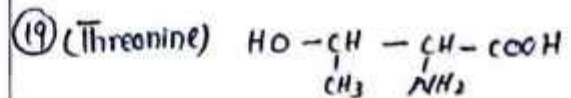
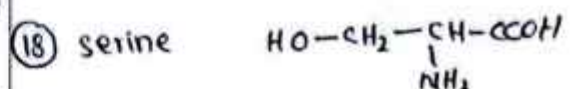
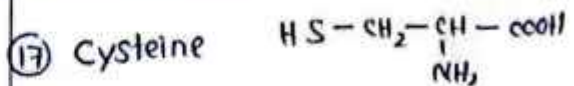
Basic Amino Acids



Aromatic Amino Acids



Sulphur containing & Their Resembling



BEST OF LUCK

Chemistry with MJS

Proteins

- proteins are macromolecules of Amino acids together by peptide Bond.
- 20 amino acids commonly found in proteins.
- proteins are molecules having a very high molecular weight, ranging from 5000 to several millions.

Functions:

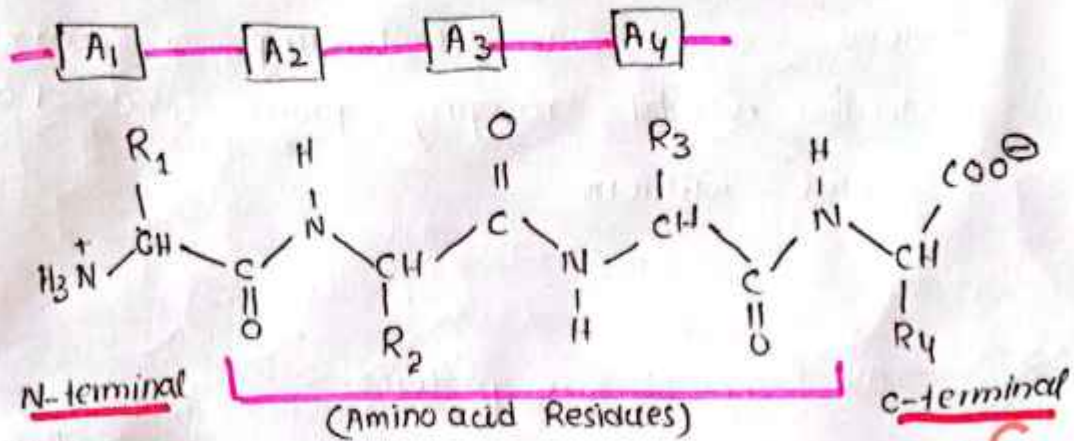
- All Enzymes are proteins.
- Transport of molecules e.g Hemoglobin (carrier of O₂) & Lipoproteins (carrier of Lipids)
- Structural Significance e.g cell membrane made up of Lipoproteins & skin, bones (collagen)
- Hormones are proteins e.g insulin
- Antibodies (immunoglobulins) protein in nature
- Keratin (skin) → Hair, nails etc
- prothrombin (Blood clotting)
- Ferritin (storage form of Iron)
- Myosin in muscles & collagen in Tendons.

Conformation of proteins:

protein in its native state has a 3-D structure known as conformation.

1. Primary structure:

- Order or Arrangement of A.A. in poly-peptide chains.
- covalent bond is responsible for primary structure.
 - ↳ (peptide bond)
- No. & Sequence of Amino acids is determined



Sequence of arrangement of Amino acids is determined by terminal Residue Analysis.

2. Secondary structure:

- Spatial Relationship of adjacent Amino acids Residues.
- Hydrogen bonds are responsible for this structure.
 - ↳ (B/w H of -NH group of one amino acid residue & carbonyl oxygen of fourth one)

proline can not form α -helix b/c lack of hydrogen on proline's nitrogen prevents it to form H-bonding - also containing irregular Geometry

Two major conformations

α -Helix

β -pleated sheet

- When side chains in a polypeptides are bulky
- peptide chains twists into a right handed or spiral
- complete turn distance = 0.54 nm
- Each turn contains = 3-6 A.A.

↳ structure formed b/w two or more separate polypeptide chains.

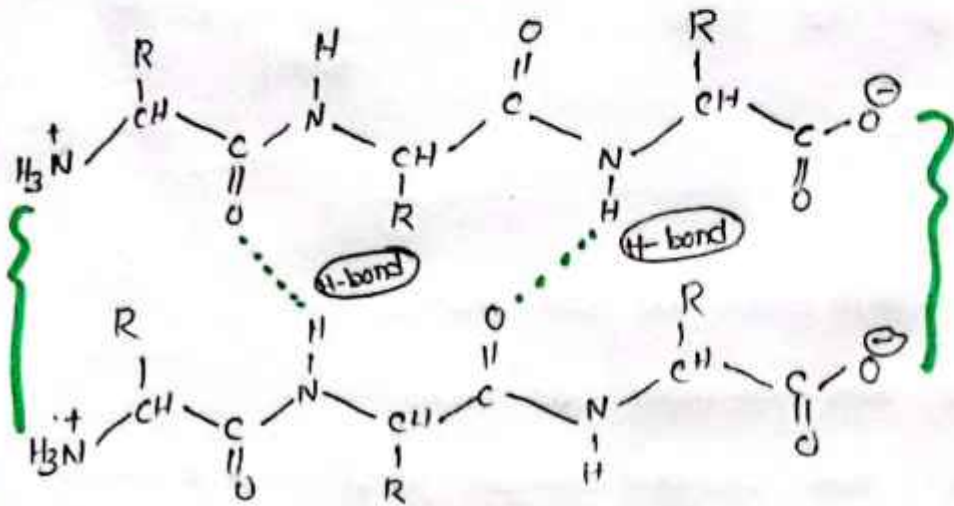
β-pleated sheet

parallel β-sheets

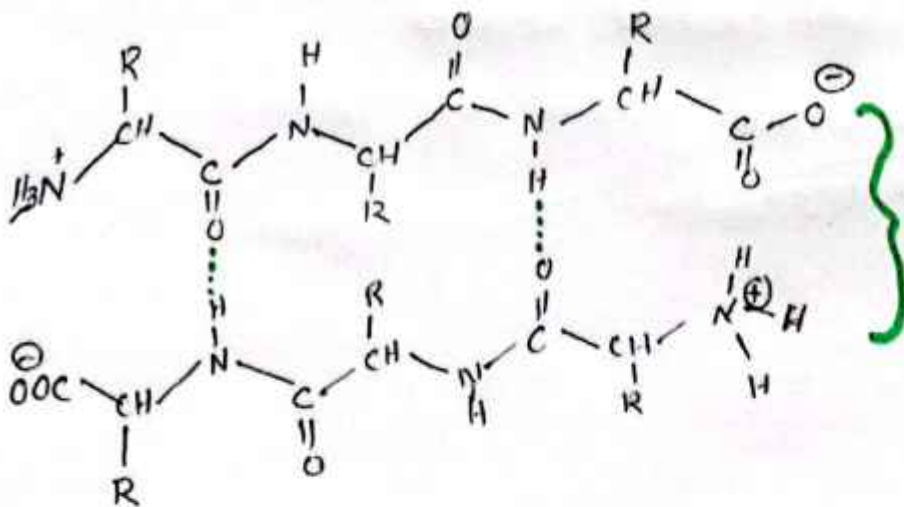
Two polypeptide chains in the same direction

Anti-parallel β-sheets

Two polypeptide chains in opposite directions



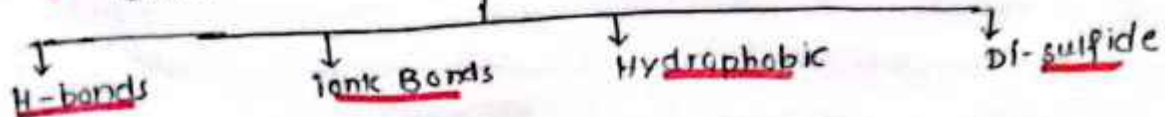
parallel
β-pleated
sheet

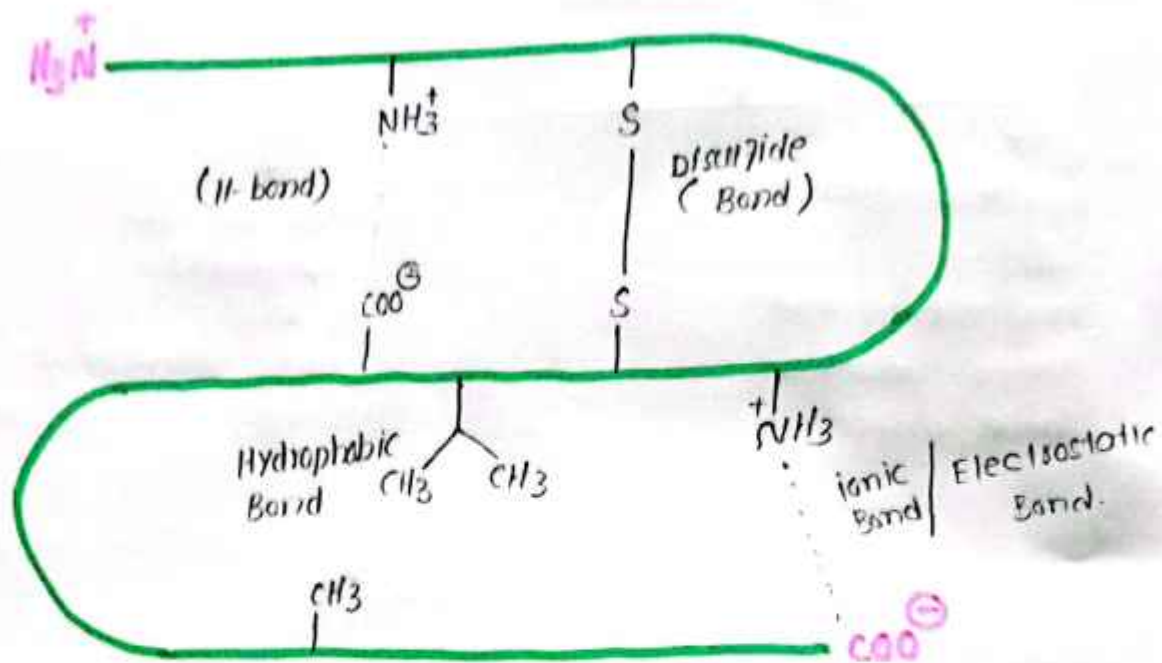


Anti-parallel
β-pleated
sheet

3. Tertiary Structure:

- High Folding & Bending of long peptide chains
- Final Rearrangement of polypeptide chain
- Bonds Responsible



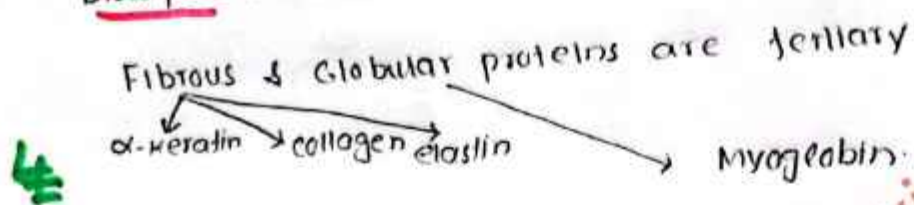


H-bond = within chain or B/W chains

Hydrophobic = B/W non-polar side chains

Ionic bond = B/W oppositely charged group in side chains

Disulfide = B/W cysteine residues



4. Quaternary Structure:

- Many proteins are composed of several polypeptide chains called subunits. Each subunit contains its own independent 3-D conformation.
- Aggregates of polypeptide sub-units.

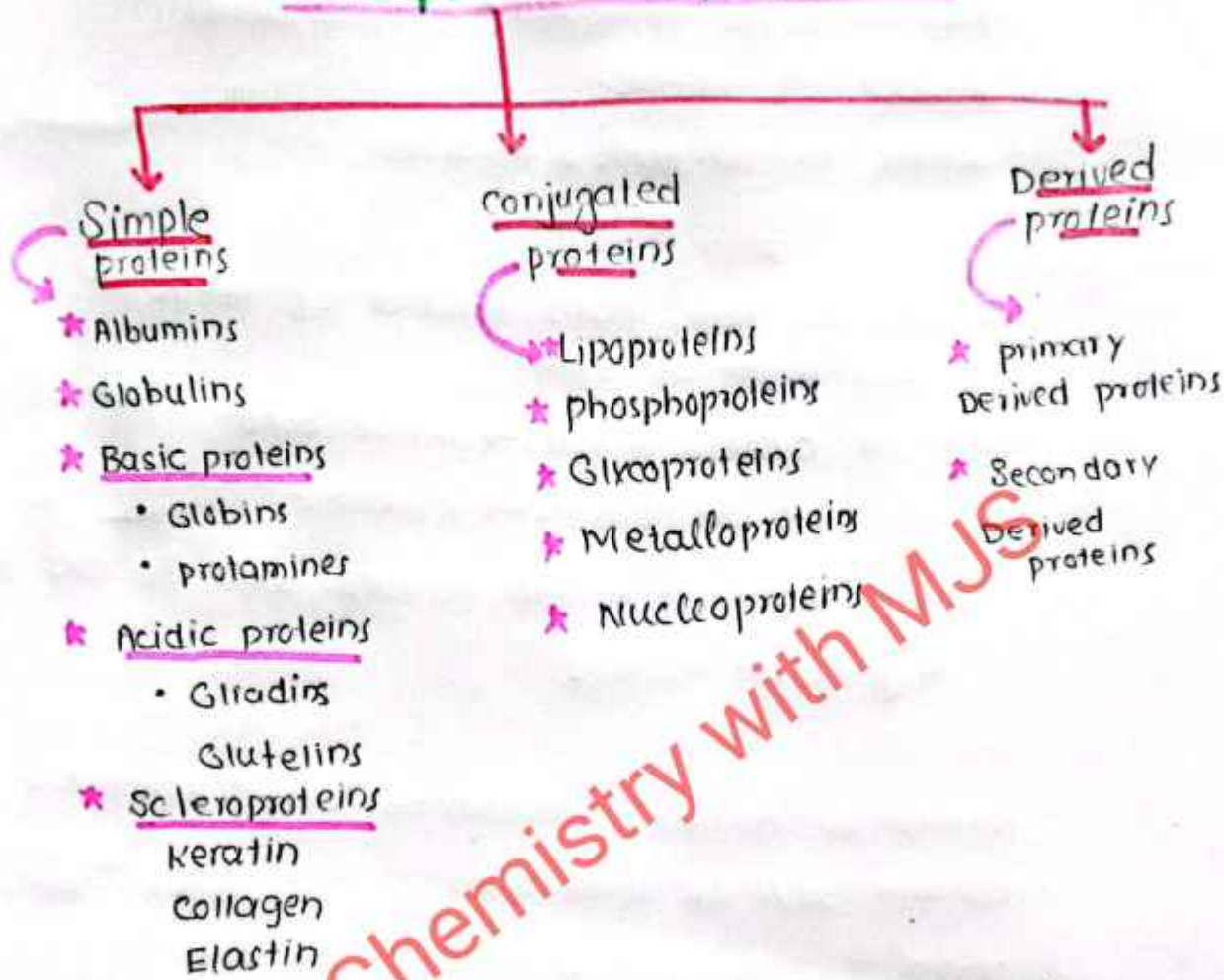
Bonds Responsible:

1. H-bonds
2. Electrostatic bond.
3. Hydrophobic bond

Examples:

- * insuline → 2 subunits
- * globin of hemoglobin → 4 subunits.

Classification of proteins



1) Simple proteins: → on Hydrolysis yield only Amino acids.

Albumins & Globulins:

- Both coagulated by heat
- Both present in blood, milk & egg.
- Albumin soluble in water while globulin soluble in salt solution.
- Albumin m.wt = 68,000 Da while globulins 150,000 Da

Globins: (Basic protein) → H1stone protein

- constituent is H1stidine (Basic A. Acid)
- Forms Haemoglobin when combine to heme.

Glutelins: (Acidic protein)

- Rich in acidic amino acid (glutamic acid)
- present in cereals.
- Soluble in dil. acids & alkalis.

Keratins:

- Found in hair, nails, enamel of teeth & out-layer of skin
- Rich in cysteine (A. Acid) → sulphide bond
- Hydrophobic amino acids → insoluble
- Genetic defect in keratin causes skin diseases
e.g. Hyperkeratosis

Collagens:

- Formed by connective tissues cell called Fibroblasts.
- contain 30% of total body proteins (most abundant protein of body)
- consist of 3 polypeptide chains, each chain contains 1050 A. Acids → held together by Hydrogen bonds
- composition:
33% glycine + 10% proline + 10% Hydroxyproline + 7% Hydroxylysine
- collagen diseases:
Scurvy → deficiency of vitamin C
Osteogenesis imperfecta: → mutation in gene that codes for α -chain of collagen

Elastin: (Connective tissue protein)

- Rubber like
- present in lungs, blood vessels, elastic ligament.
- formed by 4 polypeptide chains

2) Conjugated proteins:

phosphoproteins:

- protein + phosphate group

Examples:

- Casein: milk protein
- vitellin: egg yolk protein

Lipoproteins:

- protein + Lipid part
- plasma lipoproteins
- transport the lipids in blood
- passes through the substances in cell membrane

Glycoproteins:

- Carbohydrate part + protein

Examples:

- Blood Group, Enzymes, Mucin
- proteoglycans (cell membrane)

Nucleo proteins:

proteins + nucleic acid (DNA or RNA)

Examples:

Chromosomes, Ribosomes

Metalloproteins:

Iron:

Haeme → iron containing
e.g. Hemoglobin
myoglobin
cytochromes
ferritin
Transferrin

copper:

ceruloplasmin → Cu containing
↳ in plasma protein
Erythrocyteins → Cu
↳ in Red cells

Zinc:

insuline hormone
Carbonic Anhydrase

Chemistry with MJS

3) Derived proteins:

Primary Derived proteins

Denaturing proteins

Examples:

- ① Coagulated Albumins & Globulins
- ② Gelatin derived by boiling of collagen

Secondary Derived proteins

Hydrolytic proteins

proteins → proteases

↓
polypeptides ← peptides

↓
amino acids.

TECHNIQUES FOR SEPARATION OF PROTEINS & Amino acids:

Chromatography

Dialysis

Electrophoresis

precipitation

ultracentrifugation.

Chemistry with MJS



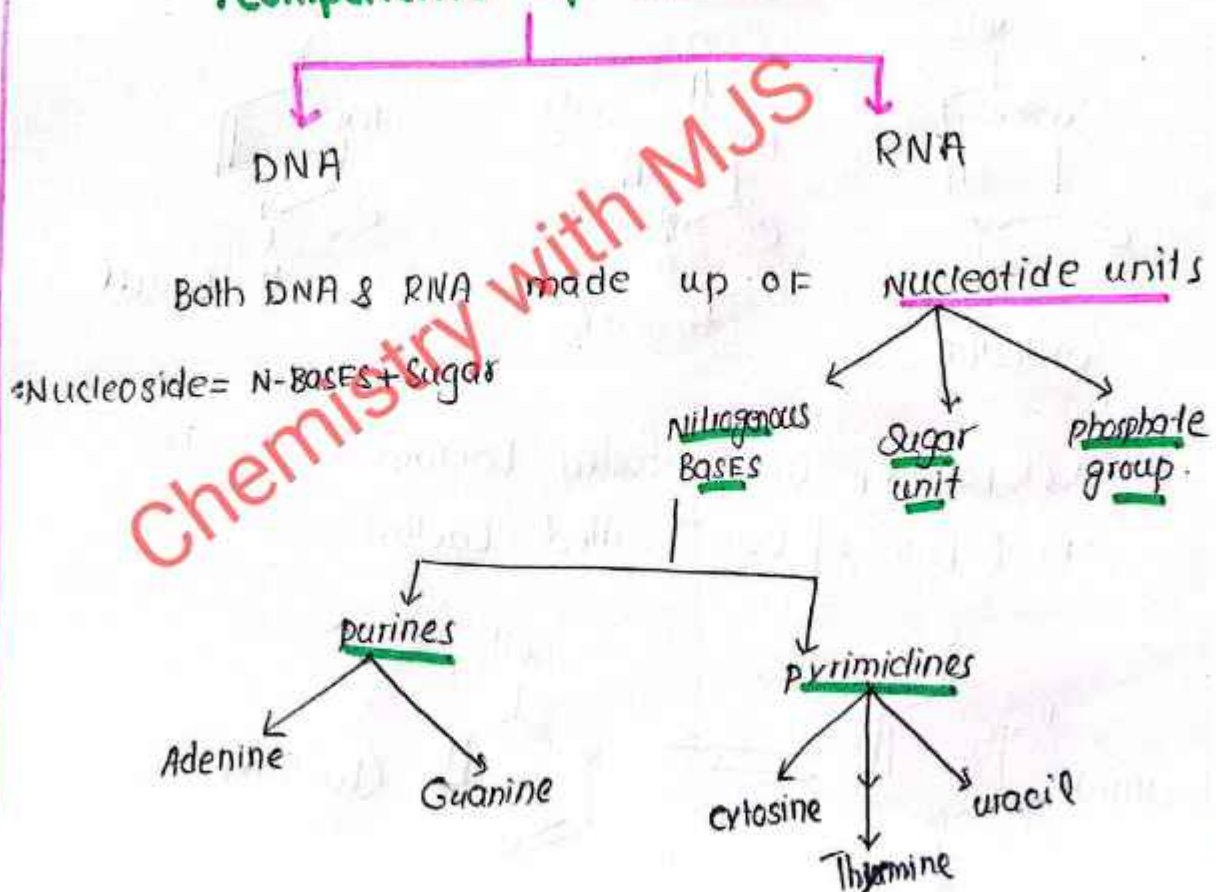
Good Luck
MJS



: Nucleic Acids:

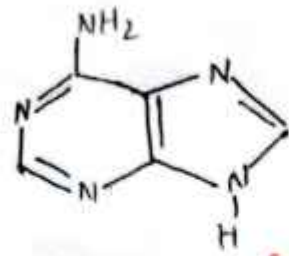
- Nucleic Acids are the poly-nucleotides
- 1st were demonstrated in the nuclei of pus cells in 1868 & in sperm cells in 1872 by Friedrich Miescher, called nuclein.
- present in every living cell as well as in viruses
- Essential substance of genes (Role in the transfer of genetic information.)
- Typical strand of Human DNA contains approx. 10^8 nucleotides.
- Direct the synthesis of proteins including Enzymes
- cancer research involves the extensive study of nucleic acids.

: Components of Nucleic Acids:

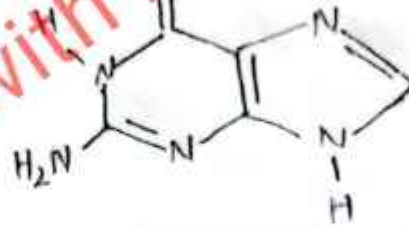


Nitrogenous Bases:

purines:



Adenine
(6-Aminopurine)



2-amino-6-oxypurine
(Guanine)

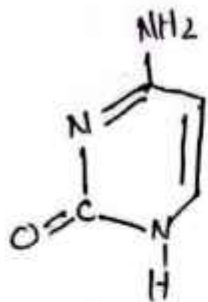
Both Adenine & Guanine Found in Both DNA & RNA.

Both pyrimidine & purines undergo keto-enol tautomeric shifts.

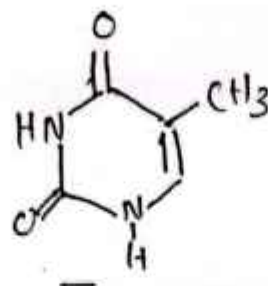
pyrimidines:

↳ in DNA (Cytosine & Thymine)

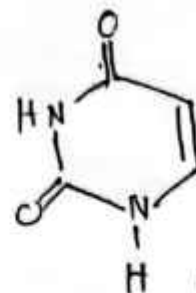
↳ in RNA (Cytosine & Uracil)



Cytosine



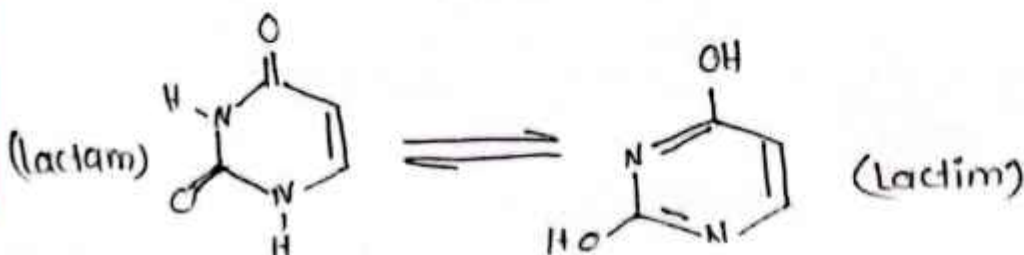
Thymine



Uracil

Keto form of uracil called Lactam

Enol form of uracil called Lactim



Nucleosides:

- Nitrogenous Base + Sugar unit = nucleoside
↓
pentose sugar
- Ribonucleosides = Nit. Base + pentose-D-Ribose
- Base linked to sugar through glycosidic Bond.
- Nucleosides are more water soluble than the free bases b/c of hydrophilicity of sugar moiety.
- pyrimidine nucleosides are also resistant to acid hydrolysis.
- purines nucleosides are easily hydrolyzed in acid to yield the free base & pentose.

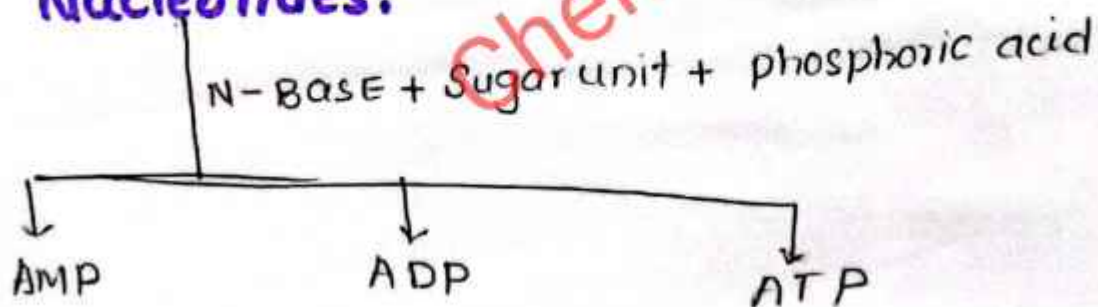
* cytidine = cytosine + pentose

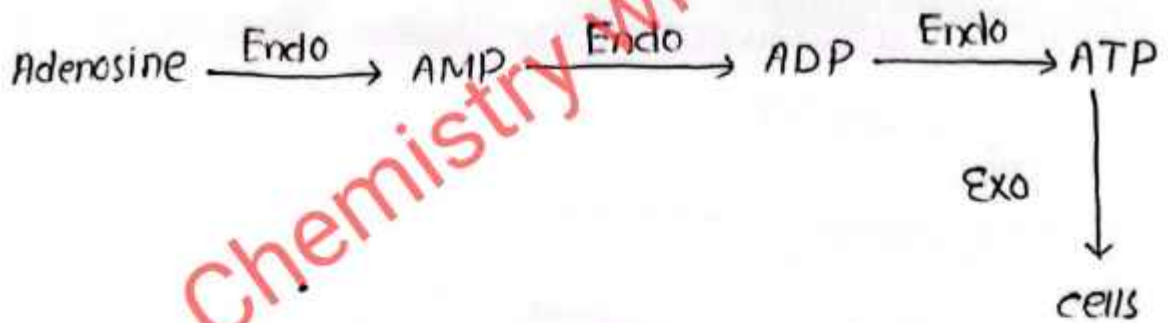
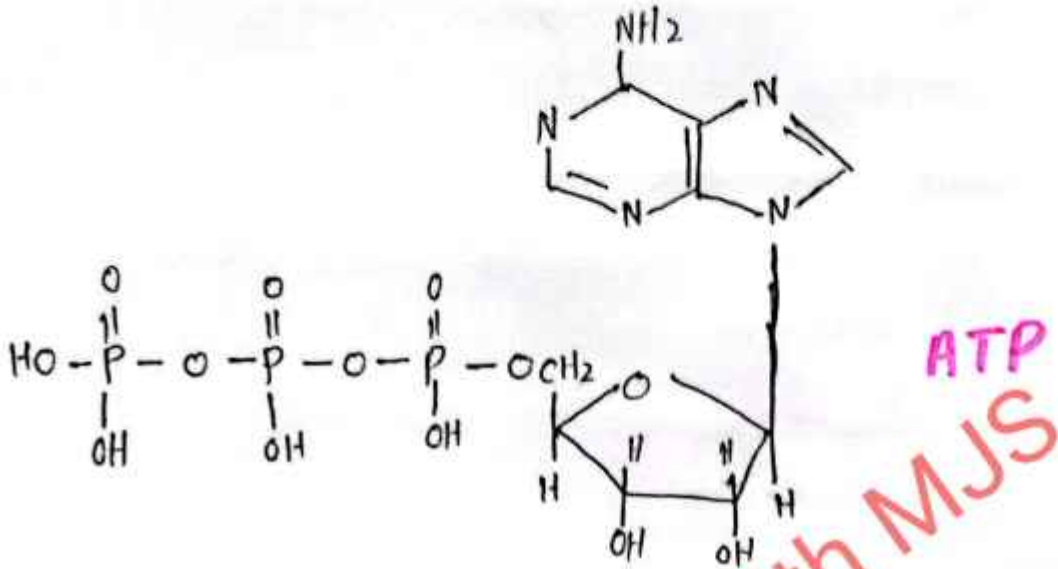
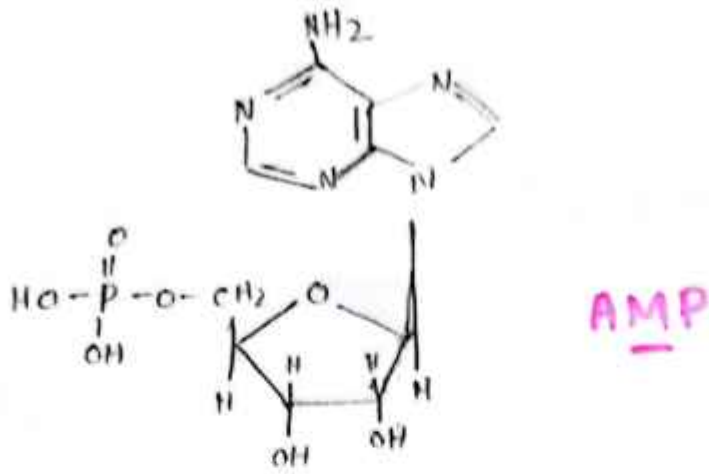
* Uridine = uracil + pentose

* Adenosine = Adenine + pentose

* Guanosine = Guanine + pentose

Nucleotides:





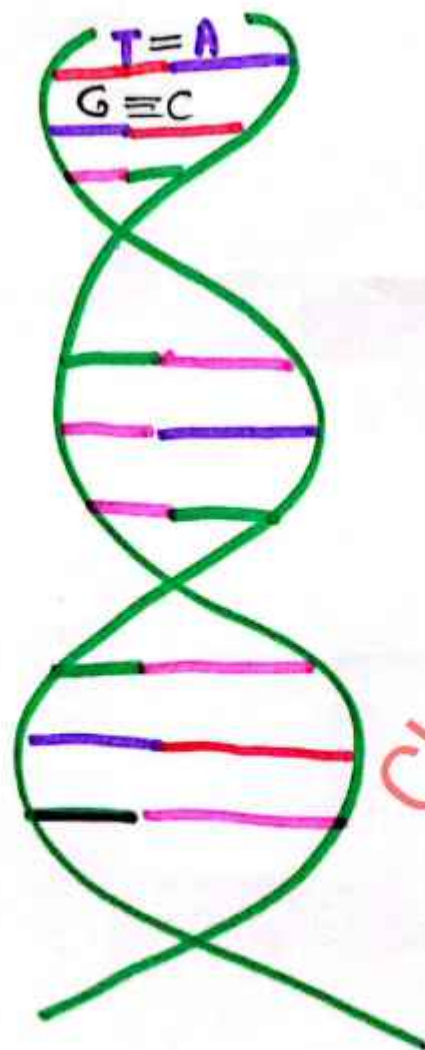
1 mole of ATP \longrightarrow ADP + phosphate
 -30 KJ/mole
 \hookrightarrow E release

Nucleic Acids are poly-nucleotides

- polymers of Ribonucleotides \rightarrow RNA
- polymers of Deoxyribonucleotide \rightarrow DNA

Fundamental structure OF DNA:

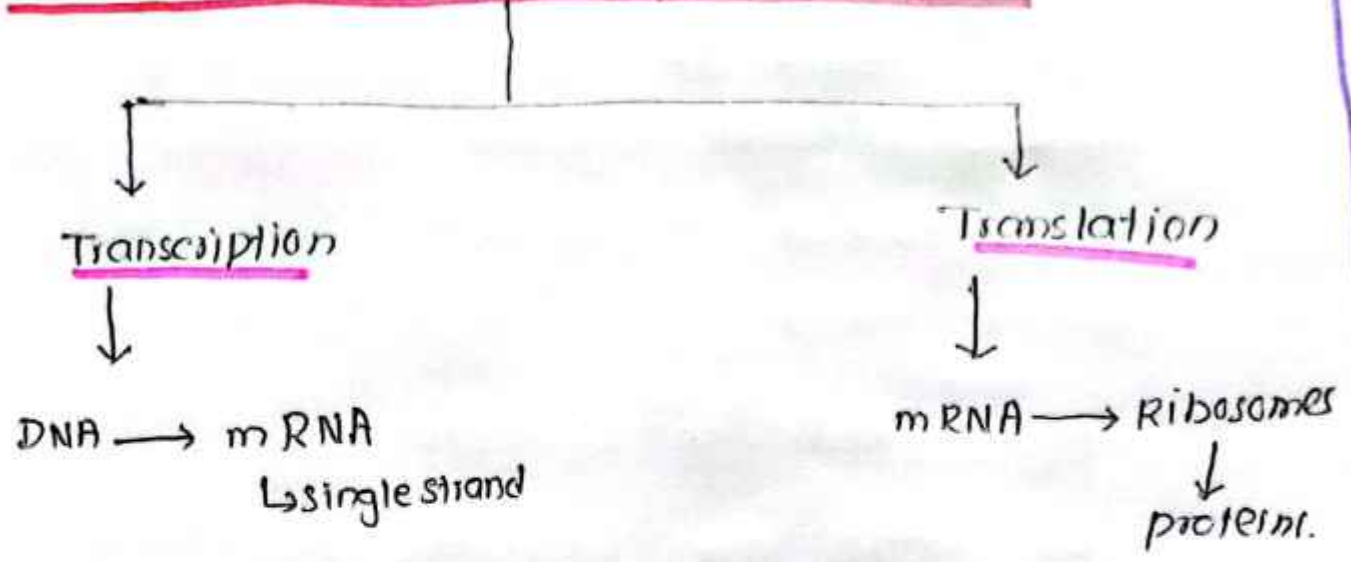
- DNA consists of two polynucleotides strands wound together to form a long, helical structure called DNA Double Helix
- Strands run in opposite direction.
- Two strands are held together by hydrogen bonds.
- DNA copied itself through DNA polymerase Enzyme.



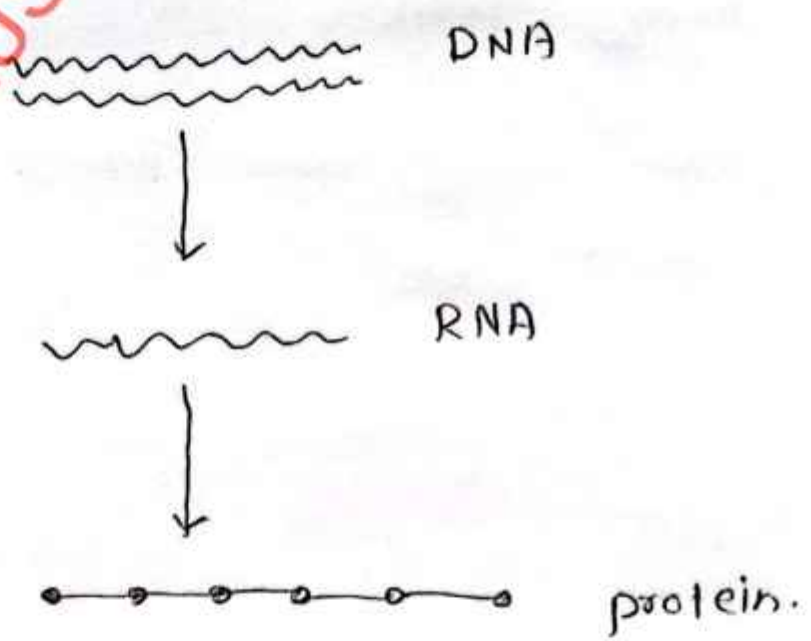
T = A
G ≡ C

Chemistry with MJS

DNA - Directed protein Biosynthesis:



Chemistry with MJS



Good Luck
MJS

