

Its like blood juice 😊

هذه المذكرة هي عبارة عن كويز البيو الثاني
سلايدات الدكتور + MCQ + اضافات من الكتاب وأعوام سابقة

Done by

أحمد العقيل

Special thanks for abo yosra
I appreciate ur great help

&

Thanx for Hegaz group
I add some notes from them

BLOOD CELLS :

- **☐ Erythrocytes** (Red blood cells)
- **Leukocytes** (White blood cells)
 - **Granulocytes**
 - Neutrophils
 - Basophils
 - Eosinophils
 - **Monocytes**
 - **Lymphocytes**
 - T
 - B
- **Megacaryocyte** (Platelets)

RBC

- Men 4.6 – 6.2 x 10¹²/l.
- Women 4.2 – 5.4 x 10¹²/l
- Total number of red cells in circulation = 2.5x10¹³

WBC

- Men and women 5-7 x 10⁹/l.

Platelets

- Men and women 250 x 10⁹/l

Hb

- Men 14 – 16 g/l
- Women 12 – 16 g/l

PCV (Haematocrit)

- Men 0.42 – 0.52 l/l
- Women 0.37 – 0.47 l/l

*** RBC :**

- **Biconcave disks:** **Highly specialized**
 - **Diameter** 6 - 9 μ m
 - **Thickness:** 1 - 2 μ m
 - **Volume;** ~ 88 fl.
- **Deformable -** i.e. can change shape to transverse smallest blood vessels.
- **Contain haemoglobin (~ 33%). (MCQ)**
- **No nucleus or mitochondria.**
- **Function: Transport of O₂ and CO₂.**
- **Normal Range:**
 - 5.5 \pm 1.0 x 10¹²/L
 - 4.8 \pm 1.0 x 10¹²/L

- Deliver oxygen to tissues and CO₂ from tissues to lungs.
- Red cell life span is 120 days.
- Senescent red cells are destroyed by spleen and replaced by juvenile cells released by bone marrow.
- An average 70 Kg adult male produces 2.3x10⁶ red cell/sec.

ERYTHROCYTE STRUCTURE

- **Biconcave shape. Spherical.**
- **Simple structure:**
 - Membrane surrounding cytoplasm.
 - Almost 95% of solutes in cytosol is haemoglobin. (MCQ)
- **No intracellular organnels. No nucleus no synthetic activities**
- **Has a cytoskeleton, which plays an important role in determining shape give the cell its deformability.**

- Red cells contain 35 % solids. (MCQ)
- Hemoglobin, the chief protein of the red cells. (MCQ)
- **Potassium, magnesium, and zinc concentrations in red cells are much higher than in the plasma.** (MCQ)

Erythrocytes Composition

- Major cation: -K⁺ (MCQ)
- Other cation: -Na⁺, Ca⁺⁺, Mg⁺⁺
- Major anion:
 - Cl⁻
 - HCO₃⁻
 - Hb
 - Inorganic phosphate
 - 2,3 bisphosphoglycerate

HAEMOGLOBIN

Haemproteins :-

Haem proteins include :

- **Cytochrome**(electron carrier)
- **Enzymes** as catalase (active site)
- **Haemoglobin**
& **myoglobin**

(the 2 most abundant haem proteins in humans)

Haemoglobin

- Major solute in red cells.(MCQ)
- Globular protein
- Conjugated protein: globin + haem.
- **Made of 4 subunits (Quarternary structure)**
4 globins + 4 haems → haemoglobin.
- **Binds O₂ to haem group to form oxyhaemoglobin**
Hb + 4 O₂ → Hb (O₂)₄.

In Adults

Hb A :	~97%	$\alpha_2 \beta_2$
Hb F :	<1%	$\alpha_2 \gamma_2$
Hb A ₂ :	2.5 – 3.5%	$\alpha_2 \delta_2$

At Birth

HbF :	70 %	$\alpha_2 \gamma_2$
Hb A :		$\alpha_2 \beta_2$

During Embryonic life

Hb Gower 1
Hb Gower 2
Hb Portland

GLOBIN CHAINS OF HAEMOGLOBIN

Amino acids in globin chains

α Globin

141 a.a.

β -like globin chains:

146 a.a.

Structure of globin chains

- Globular, compact structure
- ~75% α -helices
- Have a hydrophobic cavity for binding haem.

HAEM GROUP

- Protoporphyrin IX + Fe
- Has tetra pyrrole rings linked together by methylene bridges.
- Fe⁺⁺ coordinates with 4 Nitrogen of the 4 pyrrole rings:
 - **Bind with coordinate covalent bond to Histidine F8.(MCQ)**
 - **Binds to O₂ between Fe⁺⁺ and His E7. (MCQ)**
- If Fe is oxidized to ferric (Fe⁺⁺⁺) the Hb is known as met Hb, which cannot binds O₂ (MCQ).

- The active site in the haemoglobin and myoglobin molecules is a non-protein group called **haem**.
- The haem consists of a flat organic ring surrounding an iron atom.
- The organic part is a porphyrin ring based on a tetrapyrrole ring, and is the basis of a number of other important biological molecules, such as **chlorophyll** and **cytochrome**.
- The ring contains a large number of conjugated double bonds, which allows the molecule to absorb light in the visible part of the spectrum.

Structure and Function of Myoglobin

- Myoglobin present in heart & skeletal muscle (MCQ) ; acts as a reservoir for oxygen and as an oxygen carrier. (increase O₂ transport rate) (MCQ)
- Consist of a single polypeptide chain .
- Myoglobin is a compact molecule where **80% of its polypeptide chain is folded into eight stretches of alpha- helix (A-H)** .The a helical regions are terminated either by the presence of proline whose 5 –membered ring cannot be accommodated in an alpha- helix or by b- bends & loops stabilized by hydrogen bonds & ionic bonds.
- The interior of the molecule is composed entirely of **nonpolar amino acids** which are packed closely forming a structure stabilized by hydrophobic interactions
- On the surface there are **polar amino acids** where they form hydrogen bonds with each other & with water
- The crevice which contains the **heme group** is lined by **nonpolar amino acids except for 2 histidines** .

- Iron forms **six bonds**; four with porphyrin nitrogens, plus two additional bonds one above & one below, the planar porphyrin
- The proximal histidine (F helix) binds directly to the iron while the distal histidine (E helix) doesn't bind directly but helps stabilize the binding of oxygen to the ferrous oxygen
- The protein portion permits reversible oxygenation (binding of 1 molecule oxygen) (oxygenation)
- The simultaneous loss of electrons by the ferrous iron (oxidation) occurs only rarely .

- Iron forms **six bonds**; four with porphyrin nitrogens, plus two additional bonds one above & one below, the planar porphyrin
- One of these is coordinated to the side **ring chain of a histidine residue** of the globin molecule whereas the other position is available to **bind oxygen** (MCQ)

Haemoglobin

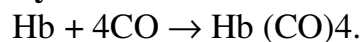
- Found exclusively in the red blood cells
- Heterotetrameric formed of 2 identical dimers
- HbA is **the major hemoglobin in adults** formed of 2 α - chains & 2 β - chains held together by noncovalent bonds (MCQ)
- It can transport CO₂ from tissues to the lungs & O₂ from the lungs to the tissues
- The oxygen binding properties are regulated by interaction with allosteric effectors
- The hemoglobin molecule is an assembly of **four globular protein** subunits.
- Each subunit is composed of a **protein chain** tightly associated with a **non-protein heme**
- group present in a pocket (crevice).

Interactions Holding the Haemoglobin Tetramer

- **Within the dimer** : interchain hydrophobic interactions (primarily) + ionic & hydrogen bonds
- **Between dimers**: mainly polar bonds
- **2 forms: T-form & R form**
- T-form: taut (tense) form is the deoxy form of hemoglobin. It is the low **oxygen-affinity form (MCQ)**
- R form: relaxed form ; the **high-affinity form (MCQ)**

- Affinity for O₂ depends on partial pressure of O₂, CO₂, and H⁺, 2,3 DPG levels. (MCQ)
- Binds CO₂ to N-terminal of β-globin chain → to form carbamino Hb.

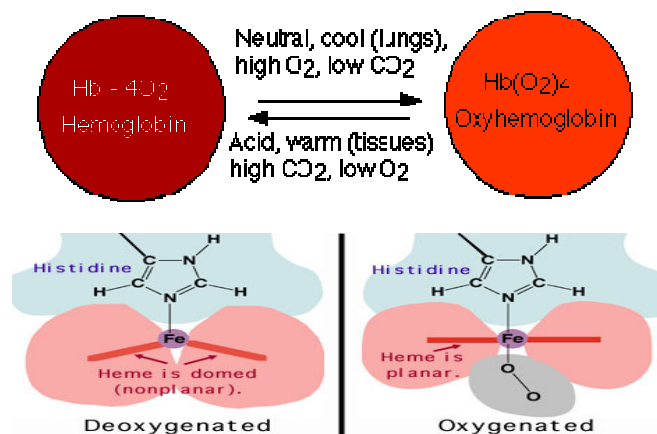
- **Carboxy Hb.**



(Has high affinity for CO₂)

- Allosteric protein: has 4 O₂ binding sites
- O₂ binding curve of Hb is **sigmoidal**. (MCQ)
- Coopertive effect: i.e. binding of some O₂ molecules makes it easy for other O₂ molecules to bind.
- O₂ affinity of Hb is affected by pO₂, pCO₂, H⁺, 2,3 DPG.(MCQ)

Note : P₅₀ for myoglobin = 1mm Hg
 P₅₀ for HB = 26mmHg
 (the higher the O₂ affinity , the lower the P₅₀)



Allosteric Effects : (MCQ)

- The ability of Hb to bind to oxygen is affected by allosteric effectors:
 - pH
 - pO₂ (through heme-heme interaction)
 - pCO₂
 - 2,3-bisphosphoglycerate

Note : The binding of O₂ to myoglobin is NOT influenced by the allosteric effectors of HB

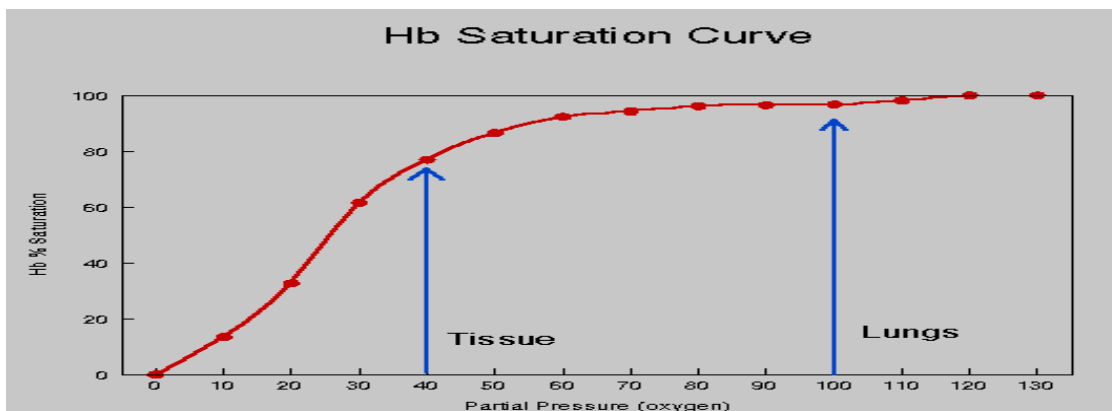
Binding of oxygen & Oxygen Dissociation Curve

- Myoglobin: reversible bind one molecule of oxygen (hyperbolic shape) (MCQ)
- Note : it bind O that released by Hb at low PO₂ then released in muscle in response to O₂ demand .
- Hemoglobin : cooperative binding of oxygen (sigmoidal in shape) due to heme-heme interaction (MCQ)

Notes :

Heme-heme interaction :

- cooperative binding . (the binding of last O₂ is 300 greater than its affinity for the 1st O₂)
- sigmoid curve



Notes :

- Hb in lung is saturated (loaded)
- HB in tissue is less saturated (unloads) so here release O₂ .

But myoglobin that has hyperbolic curve have maximum O₂ affinity in lung & tissue
So, No O₂ release but when we decrease PO₂ a lot(i.e : to a much lower level than the PO₂ of the lung) then it release O₂ .

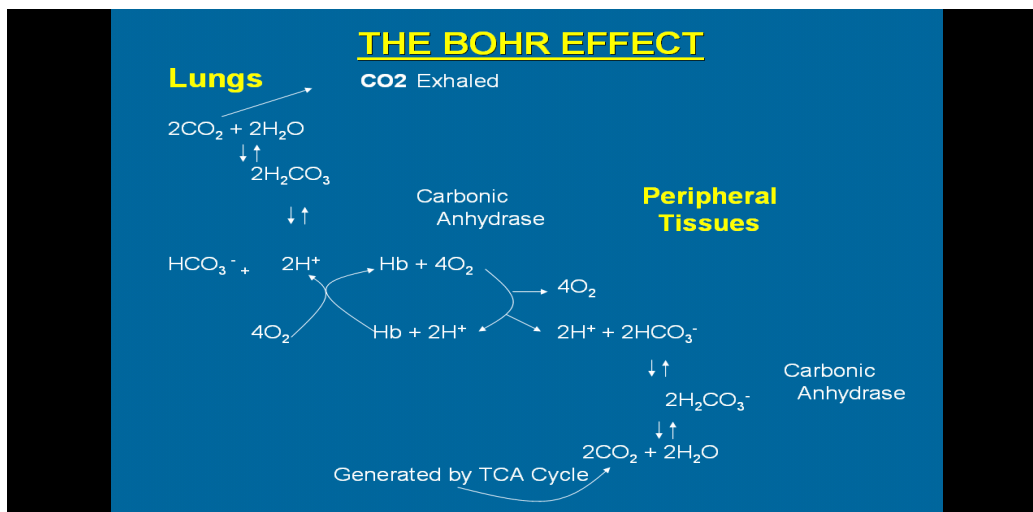
THE BOHR EFFECT (MCQ)(IMP)

In Lungs:

- High PO_2 , $\downarrow H^+$, $\downarrow CO_2$ → high affinity of Hb for O_2 (O_2 dissociation curve shifts to left).

In Tissues

- Low $P O_2$, $\uparrow H^+$, $\uparrow CO_2$, $\uparrow 2,3 DPG$ → Low affinity of Hb for O_2 (O_2 dissociation curve shifts to right)



Notes : Source of protons that lower the PH :

- concentration of CO_2 & Hb in tissue capillaries is higher than alveolar capillaries .

- $H_2CO_3 \rightleftharpoons H^+ + HCO_3^-$

These protons lower the PH (lung have higher PH than tissue) favors the unloading of O_2

In the peripheral tissue , and loading in the lung .

(this property make Hb more efficient transporter of O_2)

Remember : deoxy Hb has higher affinity for protons than oxyHb So , deoxyHb can form ionic (salt) bond that stabilized & decrease O_2 affinity .

• **BINDING OF 2,3 BISPHTHOSGLYCERATE**

- one molecule of 2,3 DPG /Hb molecule. (approximately same concentration)
- 2,3 DPG negative charged P binds between (center) of 2 β -chains (positive) of Hb A to form ionic bond of deoxyHb. (mutation produce Hb with abnormal high O_2 affinity .
- It binds to Hb and converted to deoxyHb So, decreases affinity for O_2 .
- NOT bind to oxyhemoglobin .
- Hb F cannot bind 2,3 DPG and ,therefore,has higher affinity for O_2 .(MCQ)
 - can be transport O_2 from mother to fetal blood

Notes :

- **increase 2,3BPG in response to chronic hypoxia (high altitude- emphysema)**
 - **increase 2,3BPG in response to chronic anemia .**
- So, low affinity for O₂ ---→ unloading of O₂ in the tissue capillaries .**
(normal 2,3BPG level = 5mmol/ l)

Notes :

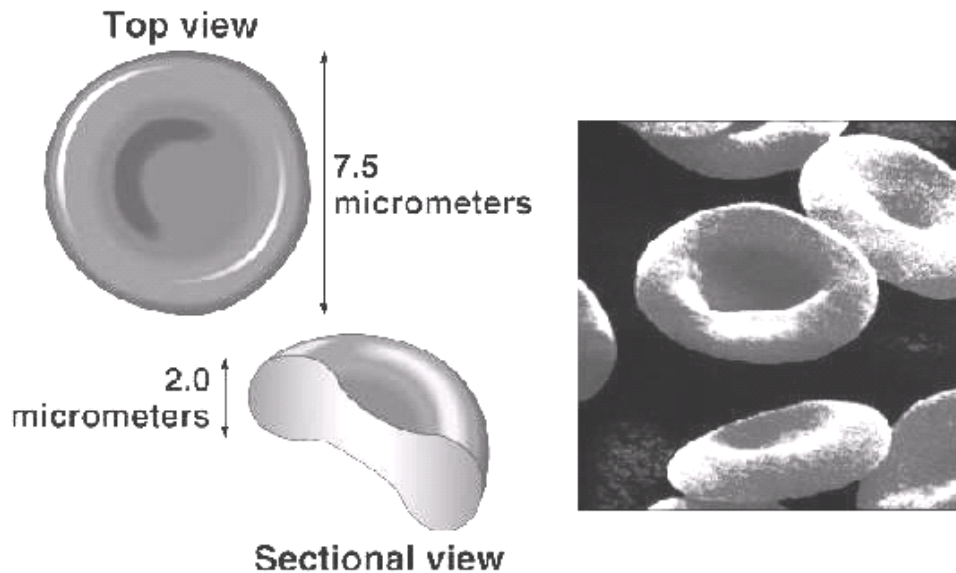
- **Storing blood in acid-citrate dextrose → decrease 2,3BPG → low affinity .**
- Hb deficient in 2,3BPG act as an O₂ trap rather than transport system .
- when 2,3 BPG is depleted ,it restore within 24 – 48 h .
- ill Patient is seriously compromised if transfused with large amount of 2,3 BPG "stripped blood "

Notes :

- **Storing blood in acid-citrate dextrose → decrease 2,3BPG → low affinity .**
- Hb deficient in 2,3BPG act as an O₂ trap rather than transport system .
- when 2,3 BPG is depleted ,it restore within 24 – 48 h .
- ill Patient is seriously compromised if transfused with large amount of 2,3 BPG "stripped blood "

The Red Blood Cells

- **RBC** are not true cells in the strict sense .
- Contain no Nucleic Acid & cannot reproduce .
- They contain no cell organelles & possess no synthetic activities .



Composition of the RBC :

- Diameter 6.9 μm
- Thickness 1-2 μm
- Range 5.5 +/- 1.0 $\times 10^{12}/\text{L}$ ♂
4.5 +/- 1.0 $\times 10^{12}/\text{L}$ ♀
- in Children 4.0 +/- 0.8 $\times 10^{12}/\text{L}$
(10-12 yrs) 4.7 +/- 0.7 $\times 10^{12}/\text{L}$

- Red Cells contain 35% Solids (33% Heamoglobin)
- **Heamoglobin** is the chief protein of Red Cells
- Other protein are present in combination with Lipids & Oligosaccharides chain forming Stroma & Cell membrane

" Spend less time worrying about what you need and more time enjoying what you have"

Erythrocyte composition :

Cation : K^+ (main) - Na^+ - Ca^{++} - Mg^{++}

Anion : Cr - HCO_3^- - Hb - InOrganic Phosphorous - 2,3 DPG

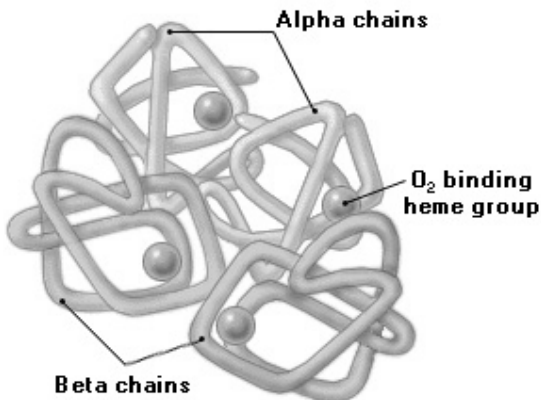
- Potassium - Magnesium - Zinc Conc. In Red Cell are much **Higher** than Plasma.

Gross Composition of Plasma & Red Cells :

Component	Plasma	RBC
Water	91-95%	65%
Solids	8-10%	35%
Protein	6-8%	31-33%
Specific Gravity	~ 1.026	-

NB.

-Heamoglobin is a tetramer : 2α & 2β
in a Diameres $(\alpha - \beta)_1$ & $(\alpha - \beta)_2$



-Bonds within the same Diameres are:
(mainly)
1.Hydrophobic Bonds
2.Hydrogen Bonds
3.Ionic Bonds

-Bonds between two Diameres are weaker than Hydrophobic Bonds :
(Polar Bonds)
1.Hydrogen bonds
2.Ionic Bonds

-In Heam of the Hb , Iron should be in Ferrus state Fe^{++}

(in Error) **Metheamoglobin** :Acquired or Congenital (ferric cannot bind to O2)

Wilson Disease : Serioloplasmin (Copper contained)

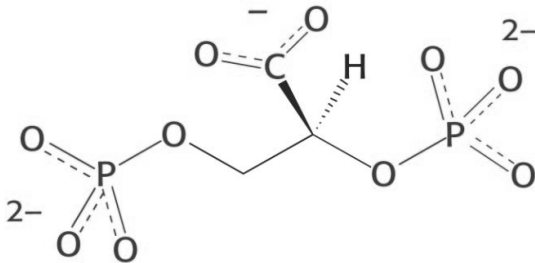
- Iron in Heam have 6 bonds : 4 Pyrol Ring
1 Oxygen
1 Histadine

Heamoglobin :

- Hb the main component of RBC (the chief protein)
- Forms about 33% of RBC
- not undergo degradation & resynth. during life of the cell

Responsible for :

1. carrying Oxygen from Lungs → Tissue
2. carrying Carbon Dioxide in the opposite direction
3. Buffering of Carbon Dioxide

2,3 Diphosphoglycerate (Anion) :

- one molecule of 2,3BPG binds to each Hb in tetramer
- it's conc. In the erythrocyte is nearly **identical** to that of Hb .

**2,3-Bisphosphoglycerate
(2,3-BPG)**

- An **increase** in 2,3BPG : (bind Hb & decrease its affinity to O₂)
 1. promotes the release of Oxygen
 2. stabilize the T-state (deoxygenated Form) of Hb by "**cross linking**" the two β-globulin subunit through multiple salt bridge .
 3. occur in response to tissue (**Hypoxia**)
e.g Anemia – Pulmonary dysfunction – Cigarette smoking
– High altitude

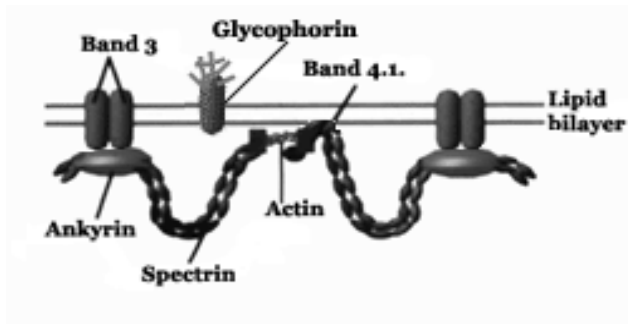
NB.

- **Hb F** keep Oxygen not in favor for tissue **more than Hb A** .
- 2,3BPG cause **shift to the right**

RBC Membrane Structure :

- **RBC** ,must be able to squeeze through some tight spot in Microcirculation, for that RBC must Easily & Reversibly Deformable , it's membrane must be both Flexible & Fluid (**Unlike the Sickle cell Anemia**)
- **About :**
 - 50% of membrane is Protein
 - 40% Fat – up to 10% Carbohydrate
- RBC membrane comprise :
 - >**lipid bilayer** (which determine the membrane fluid)
 - >**proteins** (responsible for flexibility) , either Peripheral or Integral penetrating the lipid bilayer
 - >**carbohydrate** occur only on the external surface

- Major lipid classes are : Phospholipids & Cholesterol
Glycosphingolipids as Gasngliosides & Complex series including ABO blood group substances constitute 5-10% of total lipids
- Glycophorins A,B,C are **Transmembrane Glycoproteins**
- **Glycophorin A (more -ve)** contain binding site for Influenza Virus & Plasmodium Falciparum
- Defect of protein may explain some of the abnormalities of shape of RBC membrane as **Hereditary Spherocytosis & Elleptocytosis**
- Alteration in lipid composition because of Congenital or Acquired abnormalities in plasma cholesterol or phospholipids (PL) , may be associated with other membrane abnormalities as **Target Cell** .

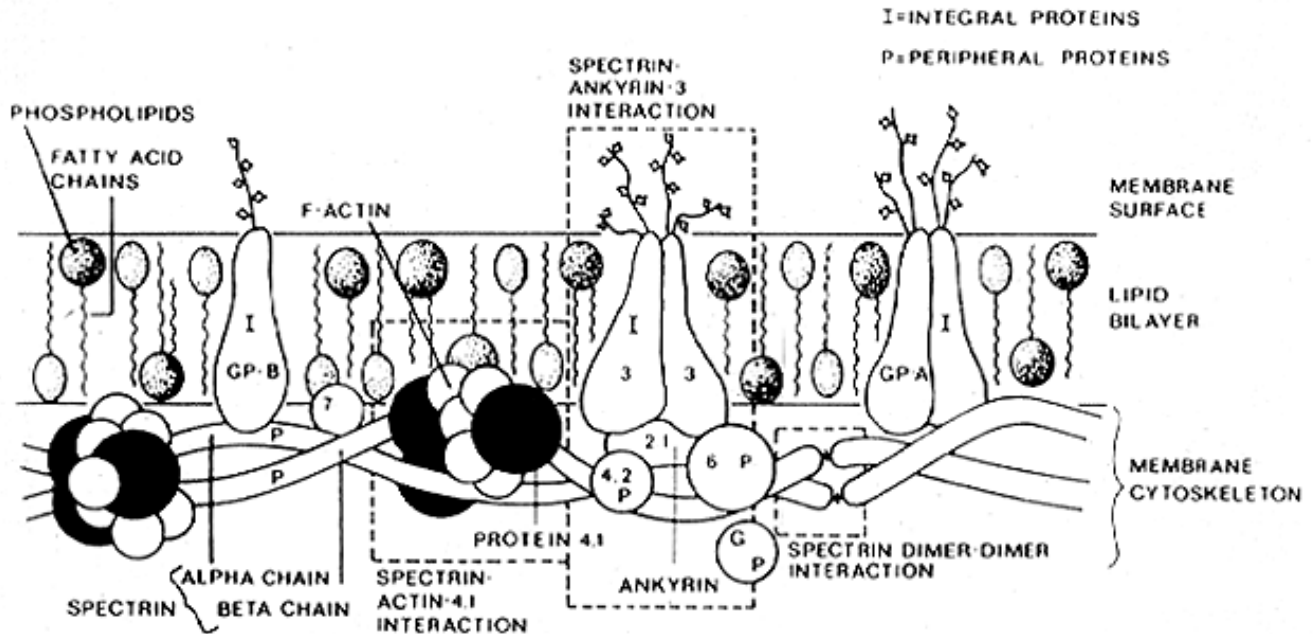


- The membrane Skeleton is **4 structural** proteins that include :
 1. A & B Spectrin
 2. Ankyrin
 3. Protein 4.1
 4. Actin

- 4 links :
 1. Head-Head tetramer called spectrin head association
 2. Ankyrin : pear-shape that attached to Band 3
 3. Protein 4.1 & 4.Actin both at the same time attached to bilayer through Glycophorin
- **Spectrin** is major protein of the cytoskeleton & it's two chains (α & β) are lined in antiparallel manner .
- (α & β) chains are loosely interconnected forming a Dimer , One Dimer interact with another forming a Head-to-Head tetramer
- Ankyrin binds spectrin & in turn binds highly to Band 3 secure attach of spectrin to membrane (**Spectrin Ankyrin-3 Interaction**)
- Band 3 : is Anoin exchange protein, permits exchange of Cl^- for HCO_3^-

"You have to have confidence in your ability, and then be tough enough to follow through"

- Actin binds to the tail of spectrin & to protein 4.1 while in turn binds to integral protein Glycophorins A&C (**Spectrin Actin 4.1 Interaction**)



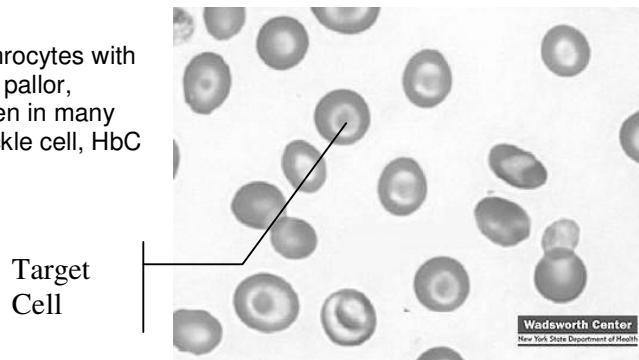
NB.

Hereditary Spherocytosis :

- Autosomal Dominant , present as Hemolytic Anemia
- Decrease surface area (osmotic fragility test)
- Pigments stone , Splenomegaly

Target cell :

- Congenital or Acquired abnormalities in cholesterol & phospholipids
- An abnormal red blood cell with a ringed appearance; associated with anemia
- seen in Liver Disease
- Target cells (codocytes) are erythrocytes with a central color spot in the area of pallor, resembling a target. They are seen in many hemolytic anemias, especially sickle cell, HbC disease, and thalassemia.



"The secret of success is constancy of purpose"

In Laboratory :

- when you take your blood sample keep it in an **Isotonic Solution** .
- Chloroform – Salt – Hypotonic solution
→ Hemolysis (Ghost Appearance for RBC)

Red Cell Aging : (MCQ)(IMP) (IMP)(IMP)

Example of changes occur in Aging Red Cell :

☺	↑ in Old Cells	↓ in Old Cells
Hb	Glycosylated Hb * use to follow up DM treatment	Bisphoglycerate
Membrane	Osmotic fragility Na ⁺ Binding to IgG	Siatic Acid K ⁺ Lipids Proteins
Enzymes	-	G6PD Pyruvate Kinase(dehydrogenase) Hexokinase Others
General	Cell density Spheriatty	Deformability Disc-like shape

General points about Hb :

- 1) allosteric protein (bind to 4 O2)&(show sigmoid curve)
- 2) cooperative effect
- 3) Affinity for O2 depend on :
- PO₂ , PCO₂ & H₂ & 2,3 DPG level .
- 4) Carboxy Hb (has high affinity to CO)
- 5) HB → oxyhemoglobin
Need : neutral PH , cool (lung) , high O₂ , low CO₂
- 6) oxyhemoglobin → Hb
Need : Acid PH , warm (tissue) , high CO₂ , low O₂
- 7) The Bohr effect :
In lung : High pO₂ , low H&CO₂ → high affinity for O₂
(O₂ dissociation curve shift to left)
In tissue : low pO₂ , high H&CO₂&2,3BPG → low affinity of Hb for O₂ .
(O₂ dissociation curve shifts to right)
- 8) Hb F cannot bind 2,3 DPG & has higher affinity for O₂ .

RBC Metabolism

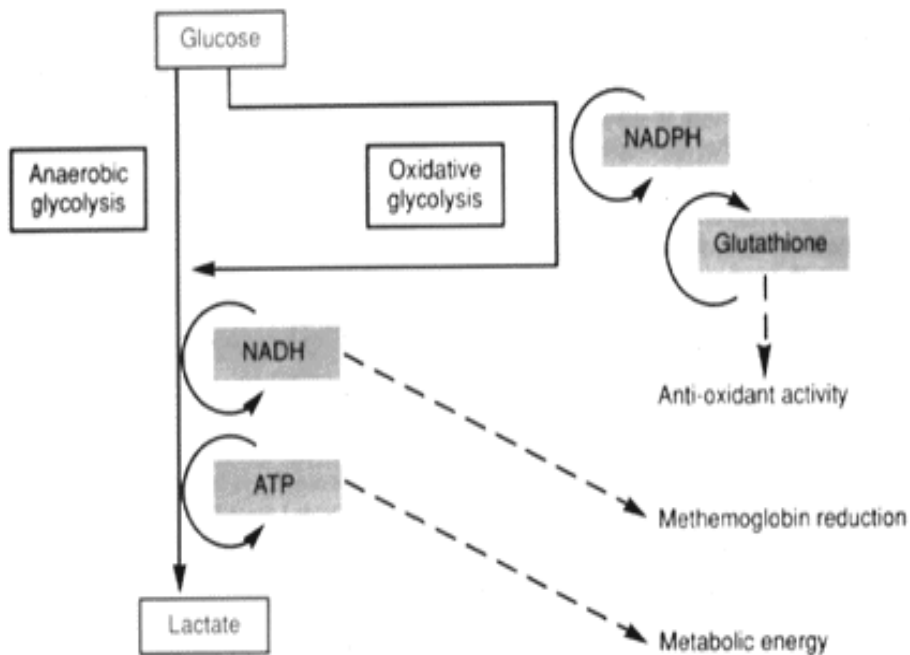
❖ **Erythrocyte contain :**

1. no mitochondria , so there is no respiratory chain .
2. no citric acid cycle .
3. no oxidation of fatty acid or ketone body .

❖ **Energy** is obtained from the Glycolytic Breakdown of glucose with production of lactate (anerobic) .

N.B: RBC needs ATP via Glycolysis and there are 3 ways :

1. Embden-Meyerhof Pathway
end with: Lactate + 2ATP
2. Rapapord- Luebering Pathway
3. HMP Pathway
mainly In producing : reducing equivalents



❖ **ATP** producing being used for :

- Na⁺ pump
- RBC membrane structure & flexibility

❖ **Rapoport & Luebering** described a special enzyme in glycolysis
bisphosphoglycerate mutase

▪ Bisphosphoglycerate Mutase **converts** :

1,3 bisphosphoglycerate (1,3BPG) → 2,3 bisphosphoglycerate (2,3BPG)

- this reaction waste the high energy bond in 1,3BPG without generate of ATP
- this explain the fact that the RBC utilize more glucose than is required to maintain their vitality

▪ **RBC** contain an active Pentose Phosphate Pathway that supplies
NADPH

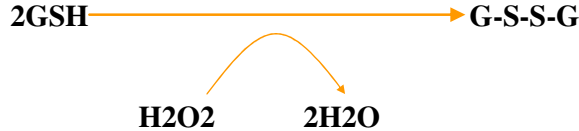
- NADPH is important in keeping glutathione in Reduced Glutathione
- Reduced Glutathione plays a very important role in survival of RBC
(membrane structure – Hb sulfhydryl oxidation ; Hinz body)

SUMMARY OF RED CELL METABOLISM

- No synthesis of glycogen, fatty acids, proteins or nucleic acids in red cells
- Highly dependant on glucose as energy source.
- Glucose is metabolized by:
 - Glycolysis (~ 95%)
 - Pentose phosphate pathway (~ 5%)
- Glycolysis produces lactate + ATP
 - 2,3 DPG regulates O₂ affinity of Hb.
- PPP produces NADPH, necessary for keeping red cells in reduced state.

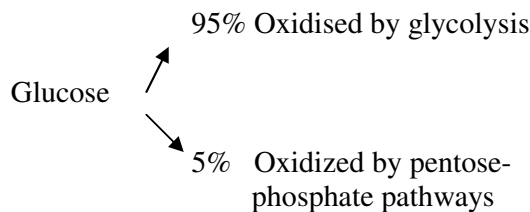
.....Contd

- Reduced glutathione is important as it keeps the:
 - red cells and other proteins in reduced state.
 - reduces oxidizing radicals (peroxides) generated in red cells



- Iron of Hb is kept in reduced state (Ferrous, Fe⁺⁺) by NADH-dependant methaemoglobin reductase.
- Glucose uptake by red cells is by facilitated diffusion.
- Proteins involved in **facilitated diffusion** of glucose are glucose transporters (~ 2% to membrane protein of RBC).
- Almost 7 different glucose transporters have been identified in different tissue.
- Glucose transporters in red cells membrane are **insulin-independent**.

Glucose Metabolism in Erythrocytes :



The role of glycolysis in the functional requirements of mature red cells:

<u>Function</u>	<u>EMP</u>	<u>PPP</u>
- Maintenance of shape	ATP	
- Membrane structure and Function		GSH
- Regulation of O ₂ transport	2,3-DPG ATP	
- Reducing potential	NADP	GSH NADPH

PRODUCTION OF POWERFUL OXIDANT IN RED CELLS DURING METABOLISM :

- **During metabolism, there is production of:**
 - **Superoxides (O₂):** $O_2 + e \rightarrow O_2^-$
 - **Hydrogen peroxide (H₂O₂)**
 $O_2 + O_2 + 2H \rightarrow H_2O_2 + O_2$
 - **Peroxyl radicals (ROO)**
 - **Hydroxyl radicals (OH^{*})**
- **These oxidizing radicals are highly reactive molecules and can react with proteins, nucleic acids, lipids and other mol. to alter their structure and produce tissue damage.**
- **Red cell need several reducing reactions to keep it in reduced state and protect it from damage by oxidizing radicals.**

PROTECTION OF RED CELLS FROM HAEMOLYSIS

By:

- **Super oxide dismutase**
 $O_2^- + O_2^- + 2H^+ \rightarrow H_2O_2 + O_2$
- **Catalase:**
 $H_2O_2 + 2H^+ \rightarrow 2H_2O$
- **Glutathione**
 $2GSH + RO-OH \rightarrow GSSG + H_2O + ROH$
Glutathione Oxidised Glutathione

Glucose-6-Phosphate Dehydrogenase (G-6-PD)

- **G-6-PD is the first enzyme of the Pentose Phosphate Pathway.**
- **Catalyses the following reaction:**

$$\text{G-6-P} + \text{NADP} + \xrightarrow{\hspace{2cm}} \text{6-Phosphogluconolactone} + \text{NADPH} + \text{H}^+$$
- **NADPH is necessary for the red cell integrity and stability.**
- **Co-enzyme for glutathione reductase which converts oxidised glutathione to reduced glutathione. This reduces oxidising radicles and protects red cells from damage.**
 - **Deficiency of G-6-PD leads to hemolytic anaemia under oxidative stress (e.g. antimalarial drugs, fava beans, infections, diabetic acidosis)**

❖ **Defecting in G-6-P dehydrogenase leads to reduced RBC survival.**

- **The erythrocyte** contain Carbonic Anhydrase enzyme
 - CO_2 combine with H_2O only after it enters the RBC
 - Hb is the most important **buffer** for the resulting Carbonic Acid is present

N.B : Carbon dioxide reacts with water to give bicarbonate, carbonic acid freed protons via the reaction, which is catalyzed by carbonic anhydrase:



❖ **RBC** also contain " Rhodanese Enzyme " responsible for the detoxification of Cyanide

* An average 70Kg adult male produce 2.3×10^6 red cell / sec .

Other Blood Cells

PLATELETS (Thrombocytes)

- Discoid, anucleated cells with agranular cytoplasm.
 - Diameter = 3 μm
 - Thickness = 1 μm
 - Volume = 7 fl.
- 250x10⁹ platelets/litre.
- Synthesis increased by thrombopoietin. (MCQ)
- Synthesised from megakaryocytes.(MCQ)
- Survival in circulation 10-12 days.
- Primary role:
- in haemostasis: stick to the edges of wounds and form a plug to arrest blood loss.
- Platelets also involved in development of atherosclerosis and hence can lead to thrombosis.

White Blood Cells (Leucocytes)

Two Main Groups

a. The Phagocytes :Play a role in protecting the body against infection by phagocytosis.

i- Granulocytes:

- Neutrophils
- Eosinophils
- Basophils

ii- Monocytes

b. The Lymphocytes (immunocytes)-

- i- B-Lymphocytes-----Provide humoral immunity.**
- ii-T- Lymphocytes-----Provide cellular immunity.**

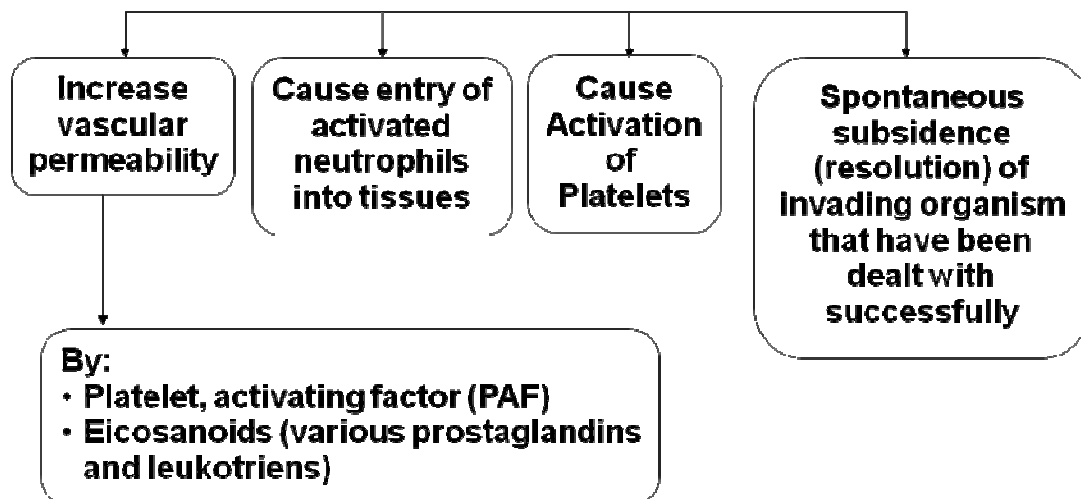
Total leucocytes: 4.00-11.0x 10⁶/l

GRANULOCYTES

- Have numerous lysosomes and granules (secretory vesicles).
 - Also known as polymorphonuclear leukocytes (PMN) as they have multilobular nuclei
 - Types of granulocytes:
 - Neutrophils,
 - basophils and
 - eosinophils
- are distinguished by their morphology and staining properties of their granules.

NEUTROPHILS

Responsible for acute inflammatory response



FUNCTIONS OF MONOCYTES:

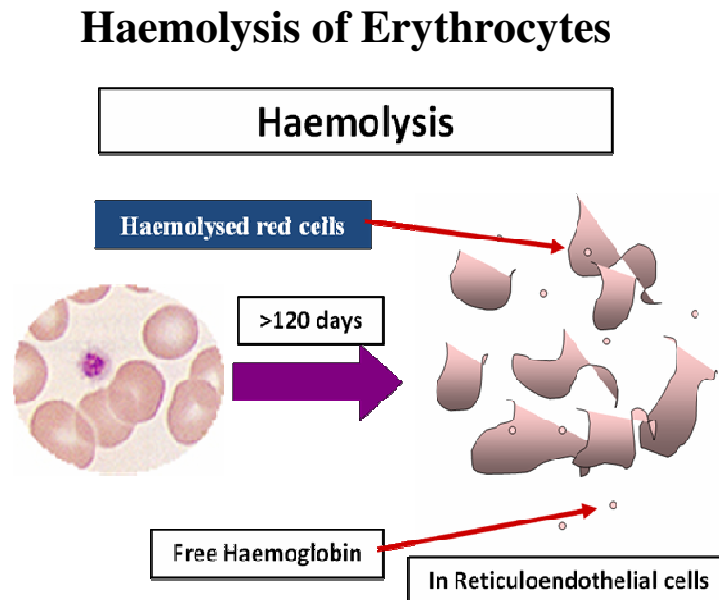
- Monocytes are precursors of macrophages, which are actively involved in phagocytosis.

FUNCTIONS OF LYMPHOCYTES:

- **B-Lymphocytes:**
 - Synthesize and secrete antibodies (humoral immunity)
- **T-Lymphocytes:**
 - Involved in cellular immune mechanism e.g.
 - ✓ killing virally infected cells and some cancer cells.
 - ✓ activate B cells to make antibodies.

PLATELETS

- Involved in coagulation of blood



Haemolysis of Erythrocytes :

- ◆ After a life span of 120 days, erythrocytes are haemolysed

- ◆ **In:**

- Spleen
- Bone marrow
- Other REC

- ◆ **Signal for haemolysis:**

- Loss or alteration of:
 - ✓ Cytoskeleton structure
 - ✓ Active ion pump
 - ✓ Membrane lipids
 - ✓ Membrane glycoproteins
- Most intracellular components are reutilized.