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Introduction into Biochemistry



Course information

- Recommended textbooks
 - Marks' Basic Medical Biochemistry: A Clinical Approach 5th Edition, by Michael Lieberman (Author), Alisa Peet MD (Author), 2018
 - Biochemistry 8th edition by Mary Campbell (Author) and Shawn Farrell (Author)
- Online:

https://themedicalbiochemistrypage.org/

- Instructors
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Outline

- Introduction
- Acids and bases, pH, and buffers
- Macromolecules
 - Carbohydrates, lipids, and amino acids, peptides, and proteins
- Protein structure-function relationship
 - part I: fibrous proteins: collagen, elastin, and keratins
 - part II: globular proteins (plasma proteins, myoglobin, hemoglobin, and immunoglobulins)
 - part III: Regulation of hemoglobin
- Enzymes
 - structural features and classification, kinetics, mechanisms of regulation, cofactors
- Protein purification and analysis



Biochemistry & chemical composition of living organisms

Biochemistry = understanding life

- Know the chemical structures of biological molecules
- Understand the biological function of these molecules
- Understand the interaction and organization of different molecules within individual cells and whole biological systems
- Understand bioenergetics (the study of energy flow in cells)

Biochemistry in medicine:

- explains all disciplines
- diagnose and monitor diseases
- design drugs (new antibiotics, chemotherapy agents)
- understand the molecular bases of diseases



Chemical elements in living creatures

- The human body is composed mainly of ~30 elements.
- Four primary elements: carbon, hydrogen, oxygen, and nitrogen (96.5% of an organism's weight)
- Then, calcium and phosphorus (that's 98.5%).
- Others exist in trace amounts but are essential, elements Just Kin r Burgaret Na. (mostly metals).

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Na	Mg Mg Mapasium											ta Al ximi Alminim	"Si	Paprove	"S		MAR Sister Argon
* K Nate	an Ca catom Catom	SC 	22 Ti Tianium		"Cr	Mn Lan	Fe	"Co	, Ni]]	· Cu	* Zn	Ga salar	Ge	As States	Se	Br nas brane	M Kr Nypten
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Bulk biological elements

Trace elements believed to be essential for bacteria. plants or animals



Possibly essential trace elements for some species bo men

	TABLE 2.1	Elements of the Human Body								
	Name		Symbol	Percentage of Body Weight						
	Major Elements (Total 98.5%)									
_	Oxygen -		0	65.0 18.0 10.0						
S	Carbon -		С							
Know e	Hydrogen -	Large	н							
	Nitrogen -		N	3.0						
unovic	Calcium		Ca	1.5						
	Phosphorus .	-	Р	1.0						
• •	Lesser Elements (Total 0.8%)									
tart to	Sulfur -		S	0.25						
e less	Potassium -		к	0.20						
	Sodium 🥣		Na	0.15						
	Chlorine		CI	0.15						
	Magnesium		Mg	0.05						
	Iron		Fe	0.006						
		Trace E	lements (Total 0.7%)	but essential						
	Chromium	Cr	Molybdenum	Mo						
	Cobalt	Co	Selenium	Se						
	Copper	Cu	Silicon	Si						
	Fluorine	F	Tin	Sn						
	lodine	1	Vanadium	v						
	Manganese	Mn	Zinc	Zn						



Important properties of bonds

- Bond strength (amount of energy that must be supplied to break a bond)
- Bond length: the distance between two nuclei
- Bond orientation: bond angles determining the overall geometry of atoms

Important G have we one talking about the angle of the atom? its position in space

The three-dimensional structures of molecules are specified by the bond angles and bond lengths for each covalent linkage. three things
 The strength of the bond



Polarity of covalent bonds

- Covalent bonds in which the electrons are shared unequally in this way are known as polar covalent bonds. The bonds are known as "dipoles".
 - Oxygen and nitrogen atoms are electronegative
 - Oxygen and hydrogen
 - Nitrogen and hydrogen
 - Not carbon and hydrogen

Both water and CO_2 contain polar bonds, but only water is a polar molecule.





What are non-covalent interactions?

• They are reversible and relatively weak.

<u>Electrostatic interactions</u> (charge-charge interactions):

- They are formed between two charged particles.
- These forces are quite strong in the absence of water .







van der Waals interactions

Unequal distribution of electronic charge around an atom changes with time.

The strength of the attraction is affected by distance.

Hydrophobic interactions "it's important because it determine the state molecule"

- it's important because it determine the structure of the molecule

- Self-association of nonpolar compounds in an aqueous environment
- Minimize unfavorable interactions between nonpolar groups and water





Properties of noncovalent interactions

- Reversible -> because at any point of time it can change it interaction
- Relatively weak
- Molecules interact and bind specifically.
- Noncovalent forces significantly contribute to the structure, stability, and functional competence of macromolecules in living cells.
- Can be either attractive or repulsive
- Involve interactions both within the biomolecule and between it and the water of the surrounding environment



Carbon

The road to diversity and stability



Properties of carbon (1)

life is all about diversity

diversity

- It can form four bonds, which can be single, double, or triple bonds.
- Each bond is very stable. Int of energy
 - strength of bonds: triple > double > Single)
- They link C atoms together in chains and rings. *shudures*
 - These serve as a backbones. of the molecule





Properties of carbon (2)

- Carbon bonds have angles giving molecules three-dimensional structures.
- In a carbon backbone, some carbon atoms rotate around a single covalent bond producing molecules of different shapes.
- The electronegativity of carbon is between hydrogen other atoms. Git's intermediate it makes non polar covalent bond with H
- have weak
- E-vity Compaired bo

Carbon

It can form polar and non-polar molecules.
 Pure carbon is not water soluble, but when carbon forms covalent bonds with other it elements like O or N, the molecule that makes carbon compounds is soluble.

when by forming certain bonds it can become soluble at

Polar covalent bond

Nonpolar covalent bond

each bransh it structured by it own

Surrounding enviorment

@ they Can votate & bind







Water

Properties of water (1)

- Water is a polar molecule as a whole because of:
 - the different electronegativities between Hydrogen and oxygen
 - dt is angular.
- Water is highly cohesive.
- Water molecules produce a network.
- Water is an excellent solvent because it is small, and it weakens it makes electrostatic forces and hydrogen with bonding between polar molecules.



Solution

Note



Dipole-charge interaction





Properties of water (3)

Water molecule is very reactive molecule 8 it's involved in a lot of reactions. Because O have the pair electrons that's ready to give it up. Oxygen is ready to give up easily on Hydrogen



• It is reactive because it is a nucleophile.

 A nucleophile is an electron-rich molecule that is attracted to positivelycharged or electron-deficient species (electrophiles).



when we talk about encymes

enzymes in general

be involved So they take the substrates away from the water in the active site because water can ruin the reaction & after that the product win be released out from the active site

Water is reactive and it Can vuin the reaction

Properties of water (4)

o it can d

 Water molecules are ionized to become a positively-charged hydronium ion (or proton), and a hydroxide ion:

$$H_2O + H_2O \longleftrightarrow H_3O^{\oplus} + OH^{\odot}$$

• it can dissociate to hydronum ion ' H_3O°
which is like proton § hydroxide ion " OH^{\circ}
Note: $H_3O^{\dagger} = H^{\dagger}$





Types of acids and bases

Arrhenius acids and bases

OH

- Acid: a substance that produces H+ when dissolved in water
- H+ Reacts with water-producing hydronium ion (H_3O^+).

$$\begin{array}{cccc} H \longrightarrow Cl &+ & H \longrightarrow \ddot{O}: & \longrightarrow & \left[H \longrightarrow \ddot{O} \longrightarrow H \\ & & H \\ & & H \end{array} \right]^{+} + & Cl^{-} \\ H & & H \end{array}$$

Base: a substance that produces OH⁻ when dissolved in water.

Types of acids and bases



give H⁺

- The Brønsted-Lowry acid: any substance (proton donor) able to give a hydrogen ion (H⁺-a proton) to another molecule.
 - Monoprotic acid: HCI, HNO3, CH3COOH Imole -> Imole
- $\begin{array}{c} \underline{\text{Diprotic acid: } H_2SO_4} & \text{I mole} \rightarrow 2 \text{ moles} \\ \hline \underline{\text{Triprotic acid: } H_3PO_3} & \text{I mole} \rightarrow 3 \text{ moles} \\ \end{array}$
- Brønsted-Lowry base any substance that accepts a proton (H⁺) from an acid.
 - NaOH, NH₃, KOH Ammonia

Water = amphoteric - it means both



- Substances that can act as an acid in one reaction and as a base in another are called amphoteric substances.
 - Example: water
- With ammonia (NH₃), water acts as an acid because it donates a proton (hydrogen ion) to ammonia.

 $NH_3 + H_2O \leftrightarrow NH_4^+ + OH^-$

• With hydrochloric acid, water acts as a base. HCl+ $H_2O \rightarrow H_3O^+ + Cl^-$ if water reacts with strong acid then water reacts as a base
if water reacts with Strong base then it reacts as an acid



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Rule



the product we have

- The stronger the acid, the weaker the conjugate base.
- Strong vs. weak acids
 - Strong acids and bases are one-way reactions

 $HCI \rightarrow H^+ + \underline{CI^-} \rightarrow the ability to associate to proton is weak it likes to be in form of chloride ion NaOH <math>\rightarrow Na^+ + OH^-$

• Weak acids and bases do not ionize completely

 $HC_{2}H_{3}O_{2} \leftrightarrow H^{+} + C_{2}H_{3}O_{2}^{-}$ $NH_{3} + H_{2}O \leftrightarrow NH_{4}^{+} + OH^{-}$

Equilibrium constant and Acid dissociation constant



- Acid/base solutions are at constant equilibrium.
- We can write equilibrium constant (K_{eq}) for such reactions



- The value of the K_a indicates the direction of the reaction.
 - When K_a is greater than 1 the product side is favored.
 - When K_a is less than 1 the reactants are favored.

Ka>1 the products higher -> Strong acid than the reactanc -> Strong acid Ka<1 the reactanc higher -> Weak acid than the products -> Weak acid it's mainly likes to be in the associated form "HA form"

Ocomparing Formic acid & Acetic acid which one of them is stronger as an acid Formic acid - Stronger because we have more protons. It dissociate to the Conjugate base. unlike Acetic acid TABLE 2.4 Dissociation constants and physolutions at 25°C	a? p K a walues of weak acids in a	aqueous	- log K • HCL doesn't dissociate 10 we still have small amount of TABLE 9.4 K	a D% HCL really small AND PKA VALU	Strong Acids the Stronge Stronge	e lower the rthe acid ACIDS
Acid	<i>K</i> _a (M)	p <i>K</i> a	Name it's in the product form than	Formula	Ka	рKa
HCOOH (Formic acid) -> We have move formic acid	than 1.77 × 10 ⁻⁴ Stronger	3.8	Hydrochloric acid	HCl	1.0×10^7	-7.00
CH3COOH (Acetic acid) > 1 " weak acid"	$k_{\alpha} < 1$ 1.76×10^{-5}	4.8	Phosphoric acid	H_3PO_4	7.5×10^{-3}	2.12
CH ₃ CHOHCOOH (Lactic acid)	1.37×10^{-4} than	3.9	Hydrofluoric acid	HF	$6.6 imes 10^{-4}$	3.18
H ₃ PO ₄ (Phosphoric acid)	7.52×10^{-3} stronger than	2.2	Lactic acid	CH ₃ CH(OH)CO ₂ H	$I = 1.4 \times 10^{-4}$	3.85
$H_2PO_4^{\ominus}$ (Dihydrogen phosphate ion)	6.23 × 10 ⁻⁸) stronger thour	7.2	Acetic acid	CH ₃ CO ₂ H	$1.8 imes 10^{-5}$	4.74
(HPO ₄ (Monohydrogen phosphate ion)	2.20×10^{-7}	12.7	Carbonic acid	H ₂ CO ₃	$4.4 imes 10^{-7}$	6.36
H_2CO_3 (Carbonic acid)	4.30×10^{-11}	0.4 10.2	Dihydrogenphosphate ion	$H_2PO_4^-$	$6.2 imes 10^{-8}$	7.21
NH₄⊕ (Ammonium ion)	5.62×10^{-10}	9.2	Ammonium ion	NH4 ⁺	5.6×10^{-10}	9.25
$CH_3NH_3^{\oplus}$ (Methylammonium ion)	2.70×10^{-11}	10.7	Hvdrocvanic acid	HCN	4.9×10^{-10}	9.31
which one is stronger H3PO4 or H2PO4 PO4 A3PO4 → stronger H3PO4 → stronger Tecause the ability of H2PO4 to release a proton is h H3PO4 H3PO4	avder than		 Scientist used the (log) to reasiev to read & understand instaid of using Ka -> pl The lower the pKa the strong the lower the pKa the strong the stron	make numbers $K_{\alpha} = -log(K_{\alpha})$ ngev the acid	it's hard for a negativly d to release a proton	navged molecule

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Molarity of solutions

- Solutions can be expressed in terms of its concentration or molarity.
- Moles of a solution are the amount in grams in relation to its molecular weight (MW or a.m.u.).

moles = grams / MW

• A molar solution is where the number of grams equal to its molecular weight (moles) in 1 liter of solution.

M = moles / volume (L)

 Since (mol = grams / MW), you can calculate the grams of a chemical you need to dissolve in a known volume (L) of water to obtain a certain concentration (M) using the following formula:

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grams = M x volume (L) x MW
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• Acids and bases can also be expressed in terms of their normality (N) or equivalence (Eq).







- When it comes to acids, bases and ions, it is useful to think of them as equivalents.
- An equivalent is the amount of moles of hydrogen ions that an acid can donate .
 - or a base can accept.
- A 1 g-Eq of any ion is defined as the molar mass of the ion divided by the ionic charge.

Inde HCL = 12 equivalent of H2SO4

Examples

- For acids:
- 1 mole HCl = 1 mole [H⁺] = 1 equivalent
- 1 mole $H_2SO_4 = 2$ moles $[H^+] = 2$ equivalents

• 1 eq of $H_2SO_4 = \frac{1}{2}$ mol (because 1 mole gives two moles of H^+ ions) 1 mole $\rightarrow 2$ eq. while $\frac{1}{2}$ mole $\rightarrow 1$ eq.

- For ions:
- One equivalent of Na⁺ = 23.1 g
- One equivalent of Cl⁻ = 35.5 g
- One equivalent of $Mg^{2+} = (24.3)/2 = 12.15 g$

Remember: One equivalent of any acid neutralizes one equivalent of any base.

it's the refeliction of how many protons - Can accept

one equivalent of a certain ion = MW / charge when I would to equalize acids and bases I have to take in consideration equivalents "not molarity or volume" Because one equivalent of (A) should be equal of one equivalent of (B) regardless the charges





One equivalent of any acid neutralizes one equivalent of base.

Based on the equation above, since x eq of an acid is neutralized by the same x eq of a base, then (n x M x vol) of an acid is neutralized by (n x M x vol) of a base.





• Note that each one produces 1 mole of the ions (H⁺ or OH⁻), so 1M of HCl is equal to 1M of NaOH. $E_{q} = n \times M \times V(L)$

Eq of base = Eq of acid n x M1 x Vol1 = n x M2 x Vol2 $1 \times M1 \times 12 = 1 \times 0.12 \times 22.4$

 $M1 = (0.12 \times 22.4) / 12$



$$E_{2} = n \times M \times V(L) \qquad 0.0224 = \frac{22.4}{1000}$$

$$E_{3} = M \times 0.12 \times 22.4 \text{ ml} \qquad 0.012 = \frac{12}{1000}$$

$$E_{3} = M \times M \times 12.0 \text{ ml} \qquad 0.012 = \frac{12}{1000}$$

$$M = \frac{0.12 \times 0.0224}{0.012} = 0.224 \text{ M}$$



Problem 2 10.93 What volume of 0.085 M (HNO₃ is required to titrate v(r) 15.0 mL of 0.12 M Ba(OH)₂ solution?

- Note that 1 mole of HNO₃ produces 1 mole of H⁺, but 1 mole of Ba(OH)₂ produces 2 moles of OH⁻. In other words, the n is different.
- Also, remember that Equivalents = n x M x volume (L), where n is the number of charges or the number of H + (or OH-) the acid or base can produce or accept.

• Titration means that we an acid to a base slowly. At one point during titration, the acid and the base neutralize or cancel each other. In other words, "to titrate" means "to neutralize". At the point of neutralization, the concentration of H+ is equal to the concentration of OH-. The best way to calculate how much acid is needed to neutralize a base (or the opposite) is to calculate the equivalents.

$$E_2 = n \times M \times V(L)$$

$$HNO_3 = 1 \times 0.085 \times V(L)?$$

$$Ba(cH)_2 = 2 \times 0.12 \times 15 ml$$

Eq of acid = Eq of base N x M1 x Vol1 = n x M2 x Vol2 1 x 0.085 x Vol = 2 x 0.12 x 15 Vol = (2 x 0.12 x 15) / 1 x 0.085 Vol = 42.35 mL

$$V(L) = \frac{2 \times 0.12 \times 0.015}{0.085}$$

42.35mL = 0.04235L



Ionization of water

• Water dissociates into hydronium (H_3O^+) and hydroxyl (OH^-) ions.

 \odot

 For simplicity, we refer to the hydronium ion as a hydrogen ion (H⁺) and write the reaction equilibrium as:

((it exist in an equilibrium form))

$$H_2O \Longrightarrow$$

@ wate must reach equilibrium

0 Thoes should increase

everytime
$$H_2O$$
 dissociate the products make joint forces to form H_2O at a certian equilibrium



Equilibrium constant

• The equilibrium constant Keq of the dissociation of water is:



The equilibrium constant for water ionization under standard conditions is
 1.8 x 10⁻¹⁶ M.
 if we multipled H₂O & Keg we will have a constant

if I have 1 mole of HzO then I have 1.8 × 10⁻¹⁶ of [H^{*}][OH^{*}] which is really little much much much less than HzO
That's why water mainly exist in HzO Form not as it ions
Which means if I have 1 mole of HzO I should have 1 of [H^{*}][OH^{*}] to give us 1.8×10⁻¹⁶ M
Again if I have pure water I have concentration like you put pure water in water what's it molarity? it's constant if we multipled H2O 8 Keg we will have a constant this also will lead that the molarity / moles will be constant the concentration of water isn't effected by anything "when it's a water-water reaction" Because the concentration will be the same and if you add water or remore water only the volum will change but not the concentration

pure



Kw

 Since there are 55.6 moles of water in 1 liter, the product of the hydrogen and hydroxide ion concentrations results in a value of <u>1 x 10⁻¹⁴</u> for:

water concentration
$$\rightarrow K_{eq} (55.5 \text{ M}) = [H^{\oplus}] [OH^{\Theta}]$$

• This constant, Kw, is called the ion product for water

$$K_{w} = [H^{\oplus}][OH^{\ominus}] = 1.0 \times 10^{-14} M^{2}$$

Kws-equilibrium constant/ionic product of water
 which is s- [H^Φ]·[OH^Φ] = 1.0 X10¹⁴M²



- For pure water, there are equal concentrations of [H⁺] and [OH⁻], each with a value of 1 x 10⁻⁷ M.
- Since Kw is a fixed value, the concentrations of [H⁺] and [OH⁻] are inversely changing.
- If the concentration of H⁺ is high, then the concentration of OH⁻ must be low, and vice versa. For example, if [H⁺] = 10⁻² M, then [OH⁻] = 10⁻¹² M

