

Theoretical Physiology

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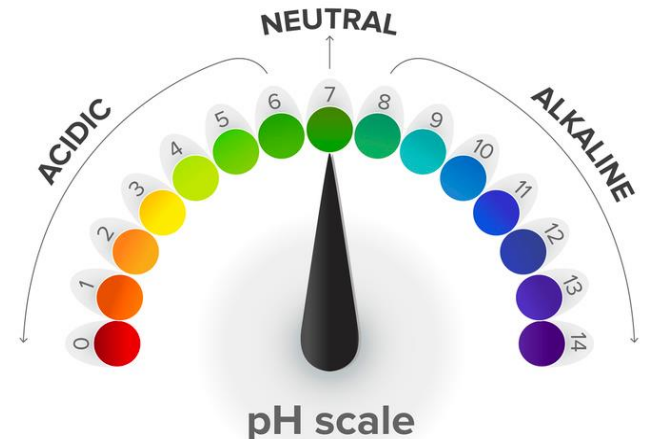
Lecture 9 , 10

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First Stage

*Acidic Base Balance

*Buffer System



Lec 9

Introduction

The concept of **Acid-base balance** is concerned with maintaining a normal hydrogen ion concentration in the body fluids. Acid-base homeostasis is essential to **allow normal tissue and organ system function**. Specifically, intracellular enzyme systems require an appropriate PH for maintenance of activity. Regulation of hydrogen ion (H^+) balance is similar in some ways to the regulation of other ions in the body. For instance, to achieve homeostasis, there must be a balance between the intake or production of H^+ and the net removal of H^+ from the body. **Extracellular fluid pH is normally maintained around a level of 7.35-7.45**. This is achieved through **chemical buffer systems** and through homeostatic responses mediated by **the lungs and the kidney**.

The limit of extracellular fluid pH compatible with life are about **6.8 to 8**

The hydrogen ion (H^{*}) concentration of the body

And pH of Body Fluid

fluids The hydrogen ion (H^{*}) concentration of the body fluids is **extremely low**. In arterial blood, the H^{*} concentration is **40 x 10⁻⁹ Eq/L (0.00000004 Eq/L or 40 nEq/L)**.

(H^{*}) concentration is routinely expressed as pH The pH is the negative logarithm to base 10 of H ion activity

$$\text{pH} = -\log_{10}[\text{H}^*] \quad \text{pH} = -\log_{10}[40 \times 10^{-9} \text{ Eq/L}] \quad \text{pH } 7.4$$

* The pH scale is an exponential scale, where a 0.3 unit fall in pH reflects a doubling of hydrogen ion concentration

***As H⁺ concentration increases, pH decreases, and conversely.**

ACIDOSIS - ALKALOSIS

ALKALOSIS

KICKIN' THE PH UP

PH
↑ 7.4

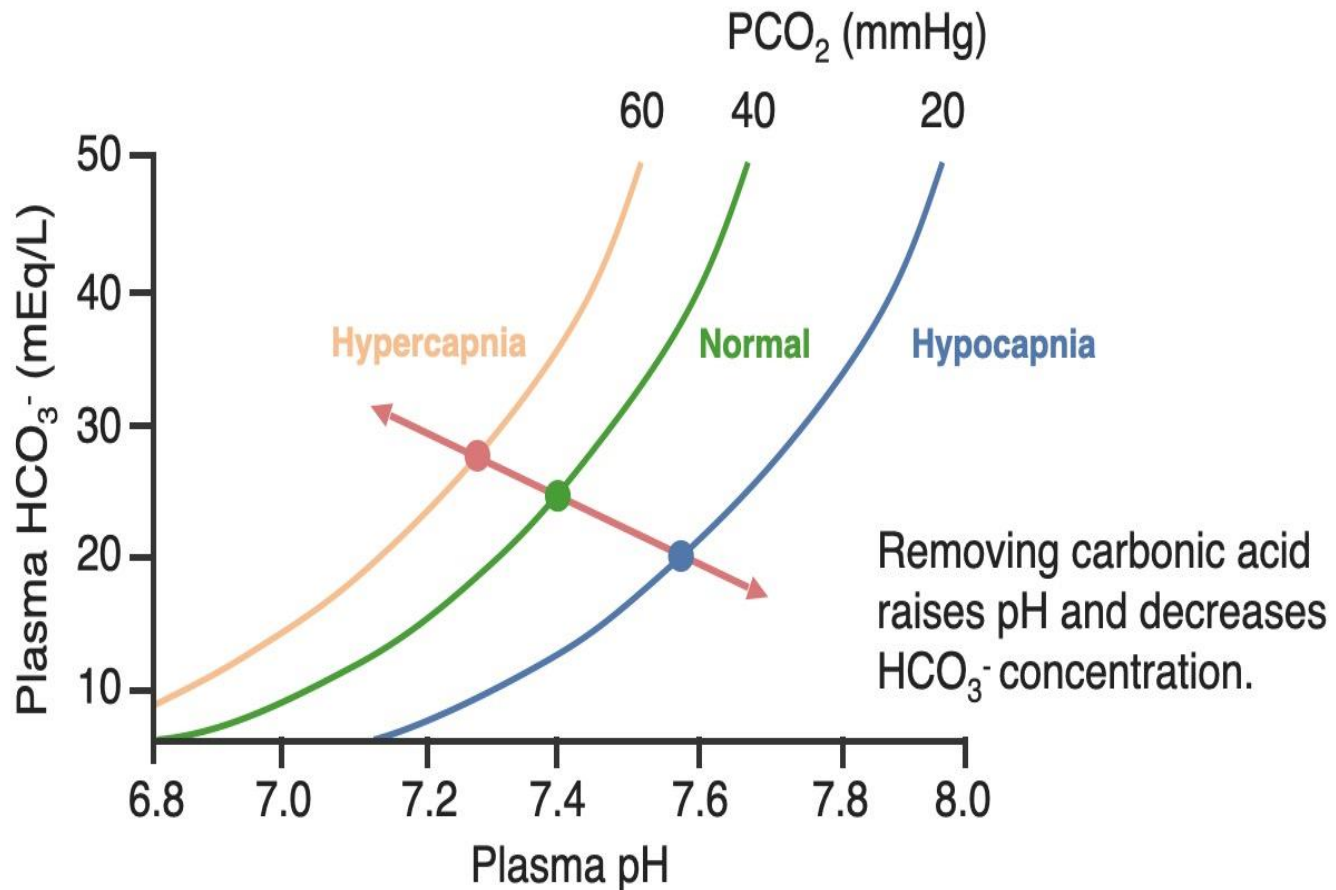
ACIDOSIS

PH
↓ 7.4

SLIDIN' THE PH DOWN

SJ MILLER

Adding carbonic acid to plasma lowers pH and increases HCO_3^- concentration:



Acidosis and Alkalosis

Normal human physiological pH is **7.35 to 7.45**. A decrease in pH below this range is **Acidosis**, an increase over this range is **Alkalosis**.

Alkalosis is defined as a disease state where the body's pH is elevated to greater than 7.45 secondary to some metabolic process.

HCO₃ functions as an alkalotic substance. CO₂ functions as an acidic substance. Therefore, increases in HCO₃ or decreases in CO₂ will make blood more alkalotic. The opposite is also true where decreases in HCO₃ or an increase in CO₂ will make blood more acidic. CO₂ levels are physiologically regulated by the pulmonary system through respiration, whereas the HCO₃ levels are regulated through the renal system with reabsorption rates. Therefore, metabolic alkalosis is an increase in serum HCO₃

The primary pH buffer system in the human body is the bicarbonate (HCO_3)/carbon dioxide (CO_2) chemical equilibrium system. Where:

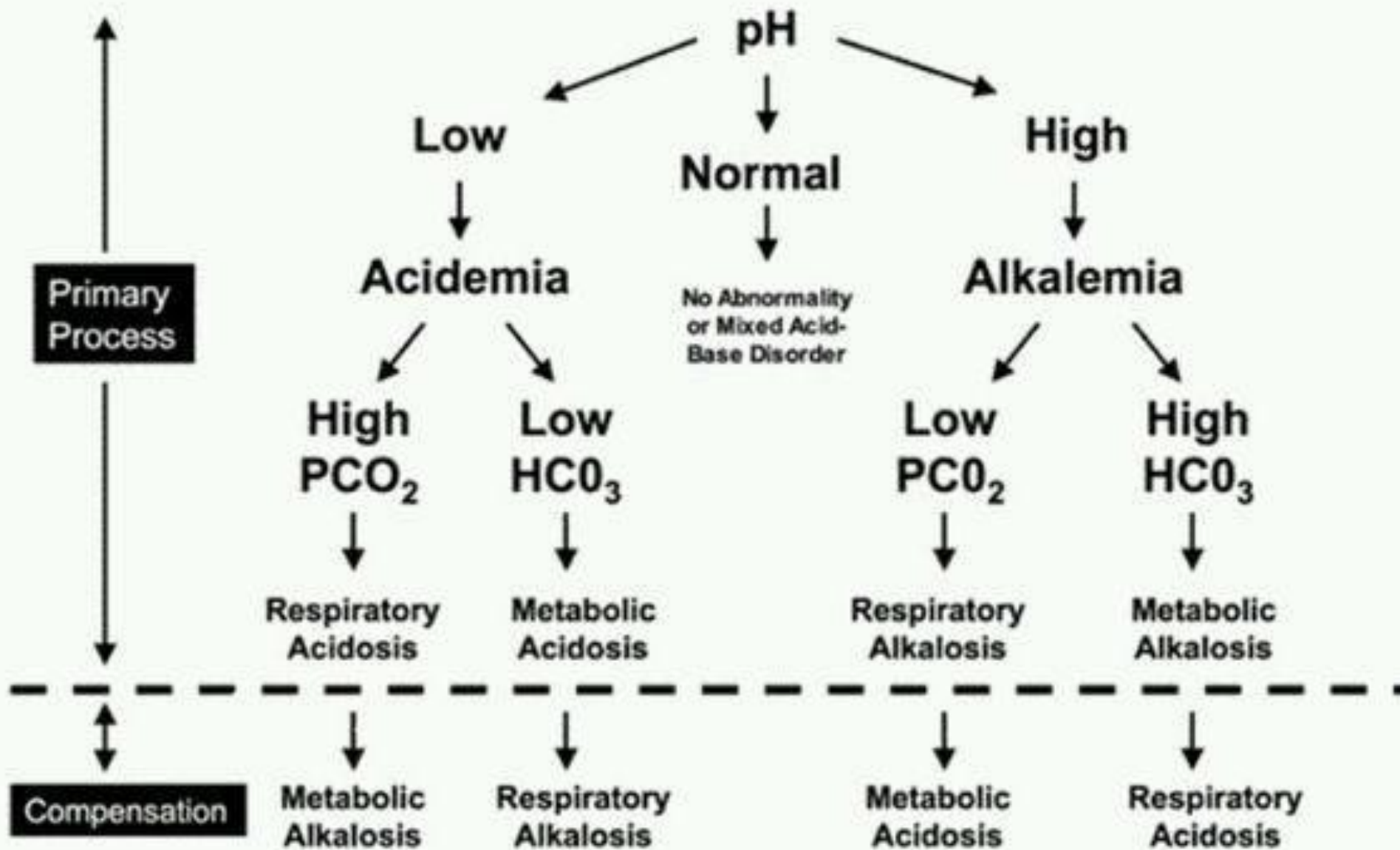


Measurement of Arterial pH (ABGO)

An Arterial Blood Gas is a laboratory test used for the measurement of arterial pH, the arterial partial pressure of O_2 (PaO_2), the arterial partial pressure of CO_2 (PaCO_2), bicarbonate (HCO_3), base excess, total CO_2 , and O_2 saturation

Venous blood can be used for the test, but the result will be more acidic (more than normal)

ARTERIAL BLOOD GAS INTERPRETATION



ARTERIAL BLOOD GAS INTERPRETATION

1° DISORDER	pH	P_aCO_2	$[HCO_3^-]$	COMPENSATION
AG/non-AG Metabolic Acidosis	↓	↓ (2°)	↓ (1°)	$P_aCO_{2, \text{ expect }} = 1.5 [HCO_3^-] + 8 \pm 2$ If $P_aCO_{2, \text{ actual }} < P_aCO_{2, \text{ expect }}$ also 1° respiratory alkalosis If $P_aCO_{2, \text{ actual }} > P_aCO_{2, \text{ expect }}$ also 1° respiratory acidosis
AG Acidosis "Delta/Delta"	For AG metabolic acidosis, calculate $\Delta AG / \Delta [HCO_3^-] = (AG - 12) / (24 - [HCO_3^-])$ if < 0.8, non-AG acidosis; if > 2, metabolic alkalosis			
Metabolic Alkalosis	↑	↑ (2°)	↑ (1°)	$P_aCO_2 = 0.7 \times [HCO_3^-] + 20 \pm 5$ If $P_aCO_{2, \text{ actual }} < P_aCO_{2, \text{ expect }}$ also 1° respiratory alkalosis If $P_aCO_{2, \text{ actual }} > P_aCO_{2, \text{ expect }}$ also 1° respiratory acidosis
Respiratory Acidosis	↓	↑ (1°)	↑ (2°)	For each ↑ 10 mmHg in P_aCO_2 Acute: ↑ $[HCO_3^-]$ 1 mmol/L and ↓ pH 0.08 Chronic: ↑ $[HCO_3^-]$ 4 mmol/L and ↓ pH 0.03
Respiratory Alkalosis	↑	↓ (1°)	↓ (2°)	For each ↓ 10 mmHg in P_aCO_2 Acute: ↓ $[HCO_3^-]$ 2 mmol/L and ↑ pH 0.08 Chronic: ↓ $[HCO_3^-]$ 5 mmol/L and ↑ pH 0.03

Primary disorder (1°), compensation (2°); arrows relative to "normal" baseline values:
 pH 7.35 - 7.45, P_aCO_2 35 - 45 mmHg and $[HCO_3^-]$ 22 - 26 mEq/L

***Causes Of Alkalosis**

1. Gastrointestinal Loss of Hydrogen

Stomach fluids are highly acidic at a pH of approximately 1.5 to 3.5. Hydrogen secretion is accomplished via parietal cells in the gastric mucosa. Therefore, the large volume loss of gastric secretions will correlate as a loss of hydrogen chloride, an acidic substance, leading to a relative increase in bicarbonate in the blood, thus driving alkalosis. Losses can occur pathologically via vomitus or nasogastric suctioning.

2. Intracellular Shift of Hydrogen

Anytime that hydrogen ions are shifted intracellularly, this imbalance in the buffer system has a relative increase in bicarbonate. Processes that drive hydrogen intracellularly include hypokalemia.

3. Renal loss of hydrogen

Hydrogen is used within the kidneys as an antiporter energy gradient to retain a multitude of other elements. Of interest here, sodium is reabsorbed through an exchange for hydrogen in the renal collecting ducts under the influence of aldosterone. Therefore, pathologies that increase the levels of mineralocorticoids or increase the effect of aldosterone, such as Conn syndrome will lead to hypernatremia, hypokalemia, and hydrogen loss in the urine. In a similar vein of thought, loop and thiazide diuretics are capable of inducing secondary hyperaldosteronism by increasing sodium and fluid load to the distal nephron, which encourages the renin-angiotensin-aldosterone system .

4.Retention/Addition of Bicarbonate

Several etiologies lead to increases in bicarbonate within the blood. The simplest of which is an overdose of exogenous sodium bicarbonate in a medical setting. Milk-alkali syndrome is a pathology where the patient consumes excessive quantities of oral calcium antacids, which leads to hypercalcemia and varying degrees of renal failure. Additionally, since antacids are neutralizing agents, they add alkaline substances to the body while reducing acid levels thus increasing pH. A pathology that is in line with normal physiology is the body's natural compensation mechanism for hypercarbia .

When a patient hypoventilates, CO₂ retention occurs in the lungs and subsequently reduces pH. Over time, the renal system compensates by retaining bicarbonate to balance pH. This is a slower process. Once the hypoventilation is corrected, such as with a ventilator-assisted respiratory failure patient CO₂ levels will quickly decrease, but bicarbonate levels will lag in reducing. This causes post-hypercapnia metabolic alkalosis, which is self-correcting

Alkalosis

is a relatively common diagnosis in medicine. The biological effects of metabolic alkalosis are directly resultant to associated problems such

as hypovolemia and potassium and chloride depletion.

These changes lead to decreased myocardial contractility, arrhythmias,

decreased cerebral blood flow,

confusion,

increased neuromuscular excitability,

hypoxia

Acidosis

Any process that increases the serum hydrogen ion concentration is an acidotic process. The term acidemia is used to describe serum that is abnormally acidic, and this can be due to a respiratory acidosis, which involves changes in carbon dioxide, or a metabolic acidosis which is influenced by decreased bicarbonate. Metabolic acidosis is characterized by an increase in the hydrogen ion concentration in the systemic circulation that results in an abnormally low serum bicarbonate levels

Strong Acidic and Strong Alkaline

Acids are molecules that release H^+ in solution

Alkaline (Bases) are molecules that can accept H^+ .

Strong Acid is one that rapidly dissociates and releases large amounts of H^+ in solution. Ex .. **HCl**- Strong Bases

Strong Base is one that reacts rapidly and strongly with H^+ in solution. And quickly remove H^+ from solution EX **HCO₃⁻**

Buffer System

Any substance that can reversibly bind H^+ . buffer system Is a system one or more compounds that act to resist change in PH.

There are three systems that regulation of H concentration.

1.The chemical acid base buffer system of the body fluids.

A. Bicarbonate buffer 24 to 26(CO_2 , H_2O)

pka of the system the nearer the value of pka of buffer system to blood PH 7.4 the greater the buffering effect of the system ,the pka of bicarbonate buffer system is 6.1 so its far from PH ,so its weak buffer system.(Bicarbonate is ECF Buffer)

B . Phosphate buffer

ICF buffer (found in the cells) , it has pka 6.8. play role in renal tubular fluid

C. Protein buffer

Its ICF buffer system, protein can function as both acids and bases (amphoteric), pka of protein fairly close to 7.4

It's the best chemical buffer system.

2.The respiratory system(the lungs) (change ventilation)

The second line of defense against acid-base disturbances is control of extracellular fluid CO₂ concentration by the lungs. Reflex changes in ventilation help to defend blood pH. By changing the PCO₂ and, hence, [H₂CO₃] of the blood, the respiratory system can rapidly and profoundly affect blood pH. An elevated arterial blood PCO₂ is a powerful stimulus to increase ventilation; it acts on both peripheral and central chemoreceptors, but primarily on the latter.

CO₂ diffuses into brain interstitial and cerebrospinal fluids, where it causes a fall in pH that stimulates chemoreceptors in the medulla oblongata.

When ventilation is stimulated, the lungs blow off more CO₂, making the blood less acidic. Conversely, a rise in blood pH inhibits ventilation; the consequent rise in blood (H₂CO₃) reduces the alkaline shift in blood pH.

Buffering power of the respiratory system.

Respiratory regulation of acid-base balance is a physiologic type of buffer system because it acts rapidly and keeps the H⁺ concentration . In general, the overall buffering power of the respiratory system is one to two times as great as the buffering power of all other chemical buffers in the extracellular fluid combined.

3. The renal system (the kidney)

Renal Regulation of Acid-Base Balance The kidneys play major role in the maintenance of normal acid-base balance.

The kidneys perform this task by:

- (1) secretion of H^+ .**
- (2) reabsorption of filtered HCO_3^- .**
- (3) production of new HCO_3^- .**

The addition of a bicarbonate to the plasma lowers the plasma hydrogen-ion concentration just as if a hydrogen ion had been removed from the plasma. Sources of hydrogen-ion gain or loss Gain of H^+

Gain of H⁺

Generation of hydrogen ions from CO₂.

Production of nonvolatile acids from the metabolism of protein and other organic molecules.

Gain of hydrogen ions due to loss of bicarbonate in diarrhea.

Gain of hydrogen ions due to loss of bicarbonate in the urine

Loss of H⁺

Utilization of hydrogen ions in the metabolism of various organic anions. Loss of hydrogen ions in the urine , vomitus and Hyperventilation .

Thank you