26.5: Disorders of Acid-Base Balance

Skills to Develop

- Identify the three blood variables considered when making a diagnosis of acidosis or alkalosis
- Identify the source of compensation for blood pH problems of a respiratory origin
- Identify the source of compensation for blood pH problems of a metabolic/renal origin

Normal arterial blood pH is restricted to a very narrow range of 7.35 to 7.45. A person who has a blood pH below 7.35 is considered to be in acidosis (actually, "physiological acidosis," because blood is not truly acidic until its pH drops below 7), and a continuous blood pH below 7.0 can be fatal. Acidosis has several symptoms, including headache and confusion, and the individual can become lethargic and easily fatigued (Figure 26.5.1). A person who has a blood pH above 7.45 is considered to be in alkalosis, and a pH above 7.8 is fatal. Some symptoms of alkalosis include cognitive impairment (which can progress to unconsciousness), tingling or numbness in the extremities, muscle twitching and spasm, and nausea and vomiting. Both acidosis and alkalosis can be caused by either metabolic or respiratory disorders.

As discussed earlier in this chapter, the concentration of carbonic acid in the blood is dependent on the level of CO\textsubscript{2} in the body and the amount of CO\textsubscript{2} gas exhaled through the lungs. Thus, the respiratory contribution to acid-base balance is usually discussed in terms of CO\textsubscript{2} (rather than of carbonic acid). Remember that a molecule of carbonic acid is lost for every molecule of CO\textsubscript{2} exhaled, and a molecule of carbonic acid is formed for every molecule of CO\textsubscript{2} retained.
Figure 26.5.1: Symptoms of Acidosis and Alkalosis. Symptoms of acidosis affect several organ systems. Both acidosis and alkalosis can be diagnosed using a blood test.

**Metabolic Acidosis: Primary Bicarbonate Deficiency**

**Metabolic acidosis** occurs when the blood is too acidic (pH below 7.35) due to too little bicarbonate, a condition called primary bicarbonate deficiency. At the normal pH of 7.40, the ratio of bicarbonate to carbonic acid buffer is 20:1. If a person’s blood pH drops below 7.35, then he or she is in metabolic acidosis. The most common cause of metabolic acidosis is the presence of organic acids or excessive ketones in the blood. Table 26.5.1 lists some other causes of metabolic acidosis.

*Acid metabolites from ingested chemical.

**Table 26.5.1**

**Common Causes of Metabolic Acidosis and Blood Metabolites**

<table>
<thead>
<tr>
<th>Cause</th>
<th>Metabolite</th>
</tr>
</thead>
<tbody>
<tr>
<td>Diarrhea</td>
<td>Bicarbonate</td>
</tr>
<tr>
<td>Uremia</td>
<td>Phosphoric, sulfuric, and lactic acids</td>
</tr>
<tr>
<td>Diabetic ketoacidosis</td>
<td>Increased ketones</td>
</tr>
<tr>
<td>Strenuous exercise</td>
<td>Lactic acid</td>
</tr>
<tr>
<td>Methanol</td>
<td>Formic acid*</td>
</tr>
<tr>
<td>Paraldehyde</td>
<td>β-Hydroxybutyric acid*</td>
</tr>
<tr>
<td>Isopropanol</td>
<td>Propionic acid*</td>
</tr>
<tr>
<td>Ethylene glycol</td>
<td>Glycolic acid, and some oxalic and formic acids*</td>
</tr>
<tr>
<td>Salicylate/aspirin</td>
<td>Sulfasalicylic acid (SSA)*</td>
</tr>
</tbody>
</table>
The first three of the eight causes of metabolic acidosis listed are medical (or unusual physiological) conditions. Strenuous exercise can cause temporary metabolic acidosis due to the production of lactic acid. The last five causes result from the ingestion of specific substances. The active form of aspirin is its metabolite, sulfasalicylic acid. An overdose of aspirin causes acidosis due to the acidity of this metabolite. Metabolic acidosis can also result from uremia, which is the retention of urea and uric acid. Metabolic acidosis can also arise from diabetic ketoacidosis, wherein an excess of ketones is present in the blood. Other causes of metabolic acidosis are a decrease in the excretion of hydrogen ions, which inhibits the conservation of bicarbonate ions, and excessive loss of bicarbonate ions through the gastrointestinal tract due to diarrhea.

**Metabolic Alkalosis: Primary Bicarbonate Excess**

*Metabolic alkalosis* is the opposite of metabolic acidosis. It occurs when the blood is too alkaline (pH above 7.45) due to too much bicarbonate (called primary bicarbonate excess).

A transient excess of bicarbonate in the blood can follow ingestion of excessive amounts of bicarbonate, citrate, or antacids for conditions such as stomach acid reflux—known as heartburn. Cushing’s disease, which is the chronic hypersecretion of adrenocorticotrophic hormone (ACTH) by the anterior pituitary gland, can cause chronic metabolic alkalosis. The oversecretion of ACTH results in elevated aldosterone levels and an increased loss of potassium by urinary excretion. Other causes of metabolic alkalosis include the loss of hydrochloric acid from the stomach through vomiting, potassium depletion due to the use of diuretics for hypertension, and the excessive use of laxatives.

**Respiratory Acidosis: Primary Carbonic Acid/CO₂ Excess**

*Respiratory acidosis* occurs when the blood is overly acidic due to an excess of carbonic acid, resulting from too much CO₂ in the blood. Respiratory acidosis can result from anything that interferes with respiration, such as pneumonia, emphysema, or congestive heart failure.

**Respiratory Alkalosis: Primary Carbonic Acid/CO₂ Deficiency**

Respiratory alkalosis occurs when the blood is overly alkaline due to a deficiency in carbonic acid and CO₂ levels in the blood. This condition usually occurs when too much CO₂ is exhaled from the lungs, as occurs in hyperventilation, which is breathing that is deeper or more frequent than normal. An elevated respiratory rate leading to hyperventilation can be due to extreme emotional upset or fear, fever, infections, hypoxia, or abnormally high levels of catecholamines, such as epinephrine and norepinephrine. Surprisingly, aspirin overdose—salicylate toxicity—can result in respiratory alkalosis as the body tries to compensate for initial acidosis.
Watch this video to see a demonstration of the effect altitude has on blood pH. What effect does high altitude have on blood pH, and why?

Compensation Mechanisms

Various compensatory mechanisms exist to maintain blood pH within a narrow range, including buffers, respiration, and renal mechanisms. Although compensatory mechanisms usually work very well, when one of these mechanisms is not working properly (like kidney failure or respiratory disease), they have their limits. If the pH and bicarbonate to carbonic acid ratio are changed too drastically, the body may not be able to compensate. Moreover, extreme changes in pH can denature proteins. Extensive damage to proteins in this way can result in disruption of normal metabolic processes, serious tissue damage, and ultimately death.

Respiratory Compensation

Respiratory compensation for metabolic acidosis increases the respiratory rate to drive off CO₂ and readjust the bicarbonate to carbonic acid ratio to the 20:1 level. This adjustment can occur within minutes. Respiratory compensation for metabolic alkalosis is not as adept as its compensation for acidosis. The normal response of the respiratory system to elevated pH is to increase the amount of CO₂ in the blood by decreasing the respiratory rate to conserve CO₂. There is a limit to the decrease in respiration, however, that the body can tolerate. Hence, the respiratory route is less efficient at compensating for metabolic alkalosis than for acidosis.

Metabolic Compensation

Metabolic and renal compensation for respiratory diseases that can create acidosis revolves around the conservation of bicarbonate ions. In cases of respiratory acidosis, the kidney increases the conservation of bicarbonate and secretion of
H⁺ through the exchange mechanism discussed earlier. These processes increase the concentration of bicarbonate in the blood, reestablishing the proper relative concentrations of bicarbonate and carbonic acid. In cases of respiratory alkalosis, the kidneys decrease the production of bicarbonate and reabsorb H⁺ from the tubular fluid. These processes can be limited by the exchange of potassium by the renal cells, which use a K⁺-H⁺ exchange mechanism (antiporter).

### Diagnosing Acidosis and Alkalosis

Lab tests for pH, CO₂ partial pressure (pCO₂), and HCO₃⁻ can identify acidosis and alkalosis, indicating whether the imbalance is respiratory or metabolic, and the extent to which compensatory mechanisms are working. The blood pH value, as shown in Table 26.5.2, indicates whether the blood is in acidosis, the normal range, or alkalosis. The pCO₂ and total HCO₃⁻ values aid in determining whether the condition is metabolic or respiratory, and whether the patient has been able to compensate for the problem. Table 26.5.1 lists the conditions and laboratory results that can be used to classify these conditions. Metabolic acid-base imbalances typically result from kidney disease, and the respiratory system usually responds to compensate.

#### Table 26.5.2: Types of Acidosis and Alkalosis

<table>
<thead>
<tr>
<th></th>
<th>pH</th>
<th>pCO₂</th>
<th>Total HCO₃⁻</th>
</tr>
</thead>
<tbody>
<tr>
<td>Metabolic acidosis</td>
<td>↓</td>
<td>N, then ↓</td>
<td>↓</td>
</tr>
<tr>
<td>Respiratory acidosis</td>
<td>↓</td>
<td>↑</td>
<td>N, then ↑</td>
</tr>
<tr>
<td>Metabolic alkalosis</td>
<td>↑</td>
<td>N, then ↑</td>
<td>↑</td>
</tr>
<tr>
<td>Respiratory alkalosis</td>
<td>↑</td>
<td>↓</td>
<td>N, then ↓</td>
</tr>
</tbody>
</table>

Reference values (arterial): pH: 7.35–7.45; pCO₂: male: 35–48 mm Hg, female: 32–45 mm Hg; total venous bicarbonate: 22–29 mM. N denotes normal; ↑ denotes a rising or increased value; and ↓ denotes a falling or decreased value.

Metabolic acidosis is problematic, as lower-than-normal amounts of bicarbonate are present in the blood. The pCO₂ would be normal at first, but if compensation has occurred, it would decrease as the body reestablishes the proper ratio of bicarbonate and carbonic acid/CO₂.

Respiratory acidosis is problematic, as excess CO₂ is present in the blood. Bicarbonate levels would be normal at first, but if compensation has occurred, they would increase in an attempt to reestablish the proper ratio of bicarbonate and carbonic acid/CO₂.

Alkalosis is characterized by a higher-than-normal pH. Metabolic alkalosis is problematic, as elevated pH and excess bicarbonate are present. The pCO₂ would again be normal at first, but if compensation has occurred, it would increase as the body attempts to reestablish the proper ratios of bicarbonate and carbonic acid/CO₂.
Respiratory alkalosis is problematic, as CO$_2$ deficiency is present in the bloodstream. The bicarbonate concentration would be normal at first. When renal compensation occurs, however, the bicarbonate concentration in blood decreases as the kidneys attempt to reestablish the proper ratios of bicarbonate and carbonic acid/CO$_2$ by eliminating more bicarbonate to bring the pH into the physiological range.

**Chapter Review**

Acidosis and alkalosis describe conditions in which a person's blood is, respectively, too acidic (pH below 7.35) and too alkaline (pH above 7.45). Each of these conditions can be caused either by metabolic problems related to bicarbonate levels or by respiratory problems related to carbonic acid and CO$_2$ levels. Several compensatory mechanisms allow the body to maintain a normal pH.

**Interactive Link Questions**

Q. Watch this video to see a demonstration of the effect altitude has on blood pH. What effect does high altitude have on blood pH, and why?

Answer: Because oxygen is reduced, the respiratory rate increases to accommodate, and hyperventilation removes CO$_2$ faster than normal, resulting in alkalosis.

**Review Questions**

Q. Which of the following is a cause of metabolic acidosis?

A. excessive HCl loss
B. increased aldosterone
C. diarrhea
D. prolonged use of diuretics
Answer: C

Q. Which of the following is a cause of respiratory acidosis?
A. emphysema
B. low blood K+
C. increased aldosterone
D. increased blood ketones
Answer: A

Q. At a pH of 7.40, the carbonic acid ratio is ________.
A 35:1
B. 4:1
C. 20:1
D. 3:1
Answer: C

Q. Which of the following is characterized as metabolic alkalosis?
A. increased pH, decreased pCO₂, decreased HCO₃⁻
B. increased pH, increased pCO₂, increased HCO₃⁻
C. decreased pH, decreased pCO₂, decreased HCO₃⁻
D. decreased pH, increased pCO₂, increased HCO₃⁻
Answer: B

Critical Thinking Questions

Q. Case Study: Bob is a 64-year-old male admitted to the emergency room for asthma. His laboratory results are as
follows: pH 7.31, pCO₂ higher than normal, and total HCO₃⁻ also higher than normal. Classify his acid-base balance as acidosis or alkalosis, and as metabolic or respiratory. Is there evidence of compensation? Propose the mechanism by which asthma contributed to the lab results seen.

A. Respiratory acidosis is present as evidenced by the decreased pH and increased pCO₂, with some compensation as shown by the increased total HCO₃⁻. His asthma has compromised his respiratory functions, and excess CO₂ is being retained in his blood.

Q. Case Study: Kim is a 38-year-old women admitted to the hospital for bulimia. Her laboratory results are as follows: pH 7.48, pCO₂ in the normal range, and total HCO₃⁻ higher than normal. Classify her acid-base balance as acidosis or alkalosis, and as metabolic or respiratory. Is there evidence of compensation? Propose the mechanism by which bulimia contributed to the lab results seen.

A. Metabolic alkalosis is present as evidenced by the increased pH and increased HCO₃⁻, without compensation as seen in the normal pCO₂. The bulimia has caused excessive loss of hydrochloric acid from the stomach and a loss of hydrogen ions from the body, resulting in an excess of bicarbonate ions in the blood.

Glossary

- **metabolic acidosis**: condition wherein a deficiency of bicarbonate causes the blood to be overly acidic
- **metabolic alkalosis**: condition wherein an excess of bicarbonate causes the blood to be overly alkaline
- **respiratory acidosis**: condition wherein an excess of carbonic acid or CO₂ causes the blood to be overly acidic
- **respiratory alkalosis**: condition wherein a deficiency of carbonic acid/CO₂ levels causes the blood to be overly alkaline

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Updated: Mon, 04 May 2020 13:58:42 GMT

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