Chemical Reactions in Dialysis

NANT 33rd Annual Symposium March 22nd, 2016 John Sweeny Hemodialysis Technical Consultant St. Petersburg, FL

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• Mole

- International measure of quantity
- 6.023 x 10²³ (about 600 septillion)
- Known as Avogadro's Number
- Used to measure a quantity of atoms or molecules
- Generally expressed in moles per liter for aqueous solutions

McGraw Hill Dictionary of Scientific and Technical Terms, 2nd edition, Editor - Daniel N. Lapedes, © 1978, McGraw Hill Book Company, ISBN 0-07-045258-X, p 1041, 132.

Gram Atomic Weight

The weight of one mole of a particular atom expressed in grams.

• Gram Atomic Weights of dialysate atoms:

Sodium (Na) = 23.00Chlorine (Cl) = 35.45Potassium (K) = 39.10Carbon (C) = 12.01Hydrogen (H) = 1.008Oxygen (O) = 16.00Magnesium (Mg) = 24.31Calcium (Ca) = 40.08

Handbook of Chemistry and Physics, 58th Edition, Editor - Robert C. West, Ph D, © 1977-78, CRC Press Inc., ISBN 0-8493-0458-X, p F-106

Gram Molecular Weight

- The weight of one mole of a particular atom expressed in grams.
- Molecular weights are the sum of the atomic weights of the atoms in the molecule.
- Sodium Chloride (NaCl) molecular weight =

Na + Cl = 23.00 + 35.45 = 58.45 grams

Sodium Bicarbonate (NaHCO₃) =
23.00 + 1.008 + 12.01 + 3(16.00) = 84.018 grams

Handbook of Chemistry and Physics, 58th Edition, Editor - Robert C. West, Ph D, © 1977-78, CRC Press Inc., ISBN 0-8493-0458-X, p F-106

Equivalent (Eq):

- One mole of charge positive or negative.
- Charges of the dialysate ions:

Na⁺, K⁺, Mg⁺⁺, Ca⁺⁺, Cl⁻, HCO₃⁻, CH₃COO⁻

- One mole of Na⁺ ions = 1 Equivalent
- One half mole of Ca⁺⁺ ions = 1 Equivalent
- 1/1000 Equivalent = 1 milliEquivalent = mEq
- 1 milliEquivalent/liter = mEq/L

College Chemistry, Donald C. Gregg, © 1961, Allyn and Bacon Inc., p 220.

Chemistry Topics

- CO₂ in Dialysate The mystery ingredient
- Hardness What's "as CaCO₃" mean?
- Urea vs. BUN They aren't the same
- Amino Acids The Lego set for proteins
- The 1st Membrane An explosive story

The Acid/Bicarbonate Reaction

• When the Acetic Acid in the Acid concentrate mixes with the Bicarbonate ion in the Bicarb concentrate the following reaction occurs:

$\begin{array}{cccc} CH_{3}COOH + HCO_{3}^{-} \longrightarrow CO_{2} + H_{2}O + CH_{3}COO^{-} \\ \hline Acetic & Bicarb & Carbon & Water & Acetate \\ \hline Acetate & Diamondation & Carbon & Ca$

Aciu	ION	Dioxide		Ion
4 mM/L	4 mEq/L	4 mM/L	4 mM/L	4 mEq/L

For 45X concentrates the 37 mEq/L bicarbonate concentrate becomes 33 mEq/L in the final dialysate solution.

How much CO₂ pressure will be in the dialysate?

- The reaction with the acetic acid produced 4 mmol of carbon dioxide.
- That's 1/250th (4/1000) of a mole/liter. (0.1.76 gm of CO_2 /liter)
- From Avogadro's law, one mole of a gas occupies 22.4 liters at one atmosphere of pressure (760 mmHg)
- Boyle's Law: $P_1 \times V_1 = P_2 \times V_2$ hence

760 mmHg x 22.4 liters = 17,024 mmHg x 1 liter

17,024 mmHg / 250 = 68 mmHg

- Normal blood CO₂ pressure is 40 mmHg hence:
- CO₂ will go from the dialysate to the blood
- This is a good thing because CO₂ levels promote breathing and as a result, proper Oxygen levels.

Blood pH and the Bicarbonate Ion

- Blood pH is directly related to the concentrations of bicarbonate and carbon dioxide in the blood.
- The relationship is defined by the Henderson-Hasselbalch Equation:

$$pH = pK + log \frac{\left[HCO_{3}^{-}\right]}{0.03 \times pCO_{2}}$$

 Where: pK = the blood pH from acids in the blood HCO₃⁻ = bicarb concentration (mEq/L) pCO₂ = carbon dioxide pressure (mmHg)

Solving the equation

- Blood normal values for pK, HCO_3^- , and pCO_2 are: pK = 6.1, HCO_3^- = 24 mEq/L, and pCO_2^- = 40 mmHg
- Placing these values in the equation yields:

Blood pH = 6.1 + log
$$\left(\frac{24 \text{ mEq/L}}{0.03 \times 40 \text{ mmHg}}\right)$$

$$\mathsf{Blood}\,\mathsf{pH} = 6.1 + \log\left(\frac{24}{1.2}\right)$$

Blood pH = $6.1 + \log(20) = 6.1 + 1.3 = 7.4$

What is hardness?

- Hardness is defined by the various minerals of Calcium and Magnesium dissolved in water.
- There are 8 specific compounds:
 - Calcium Carbonate CaCO₃ chalk, limestone, marble
 - Calcium Bicarbonate Ca(HCO₃)₂
 - Calcium Sulfate CaSO₄ gypsum
 - Calcium Chloride CaCl₂
 - Magnesium Carbonate MgCO₃ chalk, limestone, marble
 - Magnesium Bicarbonate Mg(HCO₃)₂
 - Magnesium Sulfate MgSO₄ Epsom salts
 - Magnesium Chloride MgCl₂
- Other elements such as Iron and Manganese also add to total hardness when in high concentrations which is rare.

The hardness unit: grain per gallon (gpg)

- A grain is 1/7000 of a pound = 64.8 milligrams (453.6 gm = 1 lb.)
- There's about 25,000 grains of rice to a pound.
- A gallon of water weights 8.34 pounds.
- 1 grain/gallon = $0.000143/8.34 = 1.71 \times 10^{-5}$
- To convert this ratio to parts per million (ppm) multiply by 1 million:

 $(1.71 \times 10^{-5}) \times 10^{6} = 1.71 \times 10^{1} = 17.1 \text{ ppm}.$

1 ppm = 1 mg/L

How hard is hard?

Hardness Level*	Grains per Gallon	Parts per Million
Soft	0-3.5	0 - 60
Medium	3.5 - 7.0	61 - 120
Hard	7.0 – 10.5	121 - 180
Very Hard	Over 10.5	181 +

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* Defined by the United States Geological Survey

Grains per gallon as CaCO₃

- Rather than list each mineral in water for its individual contribution to hardness, the minerals are converted to the equivalent amount of Calcium Carbonate and then added together.
- This conversion is done using the gram equivalent weights of the minerals.
- For Calcium and Magnesium compounds the equivalent weight of each compound is half it's gram molecular weight because both Calcium (Ca⁺⁺) and Magnesium (Mg⁺⁺) have double charges.

Molecular and Equivalent Weights

	Molecular Weight	Equivalent Weight
Calcium carbonate	100.090	50.045
Calcium bicarbonate	162.114	81.057
Calcium sulfate	136.142	68.071
Calcium chloride	110.986	55.493
Magnesium carbonate	84.322	42.161
Magnesium bicarbonate	146.346	73.173
Magnesium sulfate	120.374	60.187
Magnesium chloride	95.218	47.609

Equivalent Conversion Example

The formula:

Amt. of Mineral ×
$$\frac{\text{Eqv. Wt.}(\text{CaCO}_3)}{\text{Eqv. Wt.}(\text{Mineral})}$$
 = Amt. Of Mineral as CaCO₃

 Convert 10.0 gpg of Magnesium sulfate to gpg as Calcium carbonate.

10 gpg MgSO₄ ×
$$\left(\frac{50.045}{60.187}\right)$$
 = 8.31 gpg as CaCO₃

Urea vs. BUN

 Urea - A compound formed in the liver by the process known as the urea cycle. It's formula and gram molecular weight is:

$CO(NH_2)_2$

 $12.01 + 16.00 + [14.008 + (1.008 \times 2)] \times 2 = 60.058$ Daltons

 BUN = Blood Urea Nitrogen = Amount of nitrogen from the urea in the blood.

Ratio: Urea/Urea Nitrogen =

60.058/2 x 14.008 = 60.058/28.016 = 2.14

Dorland's Illustrated Medical Dictionary, Editor: Elizabeth J. Taylor, 27th Edition,1988, W. B. Saunders Company, pp 59, 1789.

Other Nitrogen Compounds

- Ammonia NH₃
- Subtract a Hydrogen and add a Chlorine and you get:
- Mono-Chloramine NH₂Cl
- Subtract a Hydrogen and add a Chlorine and you get:
- Di-Chloramine NHCl₂
- Subtract a Hydrogen and add a Chlorine and you get:
- Nitrogen Trichloride NCl₃

Organic Chemistry - α–Amino Acids

 All amino acids must have an amino group and an carboxyl group

AMINO CARBOXYL
(NH₂-) (COOH+)

There are 20 amino acids used to build all human proteins.

- 9 are essential (You must eat them.)
- 11 are nonessential (You can make them.)

Dorland's Illustrated Medical Dictionary, Editor: Elizabeth J. Taylor, 27th Edition,1988, W. B. Saunders Company, pp 59, 1370.

The α -Amino Acids

Essential

- Histidine
- Isoleucine
- Leucine
- Lysine
- Methionine
- Phenylalanine
- Threonine
- Tryptophan
- Valine

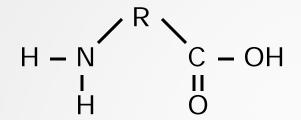
Non-essential

- Alanine
- Arginine
- Asparagine
- Aspartic Acid
- Cysteine
- Glutamic Acid
- Glutamine
- Glycine
- Proline
- Serine
- Tyrosine

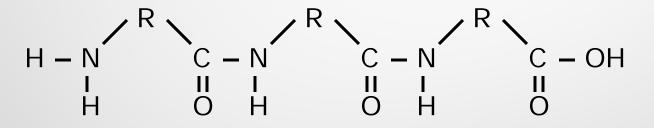
Proteins are...

- Complex organic compounds containing carbon, hydrogen, oxygen, nitrogen and sulfur.
- The main ingredient in the protoplasm of all living cells.
- High molecular weight molecules consisting of α amino acids in peptide linkages.
- Each has a genetically defined sequence of amino acids which determine its shape and function.
- Water/salt solution soluble types:
 - Albumins, globulins, histones, protamines
- Water insoluble types:
 - Collagens, elastins, keratins, actins, myosin

Amino Acid Structure/Bonding



For the smallest amino acid, Glycine, R = CH - H



Formation of a peptide (miniature protein) is called peptide linking. Note that each middle amino acid in the chain has lost a water molecule (H_2O)

Organic Chemistry, M. A. Fox and J. K. Whitesell, Copyright 1994, Jones and Bartlett, pp 619-621

Amino Acid Possibilities

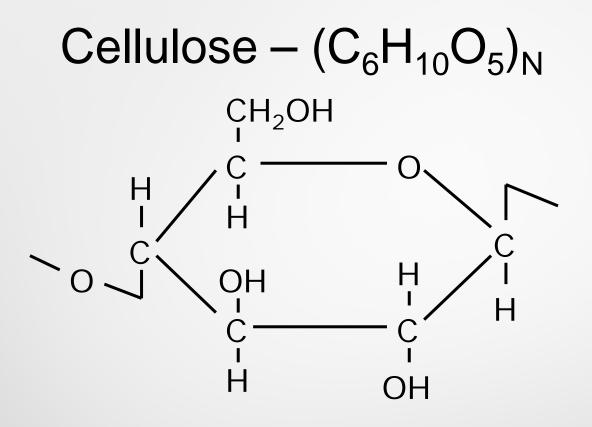
Amino Acid Chain	Combinations	
2	400	
3	8,000	
4	160,000	
10	1.02 x 10 ¹³	
100	1.27 x 10 ¹³⁰	

The first practical membrane

- 1845 Friedrich Schoenbeim spills a combination of nitric and hydrochloric acids and wipes it up with his wife's cotton apron.
- The reaction synthesizes cellulose trinitrate (gun cotton).
- He patents the process and forms his own company making the product which proved to be a fairly safe explosive for construction and mining.
- Adapted by Alfred Nobel of Sweden into a controlled explosive called dynamite.
- Nobel uses his wealth to create the Nobel Prizes.
- Schoenbeim also discovered cellulose dinitrate which could be dissolved in ether and then dried to make thin films of the material.

The development of Collodion

- Sheets of cellulose dinitrate were used as surgical dressings and the basis for photographic film until 1935.
- The material was called "collodion" from the Greek word "Kolla" for glue like substance.
- In the laboratory it became known as a good semipermeable membrane and was used by Fick to study diffusion.
- 1913 It was used by Able, Roundtree, and Turner to perform the first in vivo studies of blood. They made the collodion into tubes by dipping glass rods into the collodion solution.
- 1924 Haas dialyzed patients using collodion for two years but none of the patients lived.
- 1929 Collodion was replaces by cellulose acetate when the Visking company in Chicago started making tubes of cellulose acetate for sausage casing.



The main polysaccharide of plant cell walls. It's a polymer of β -D-glucose with a water molecule removed to form links between units resulting in chains of between 2000 and 4000 units long.

McGraw Hill page 263