



Hematology

PHYSIOLOGY



]Handout

Number: 10

Subject: Body Fluids

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***** Before we start,

Please note that Dr. Saleem discussed two problems related to blood groups in the beginning of this lecture. You can find these two problems in the last two pages in this sheet.

***** Topics of this lecture:

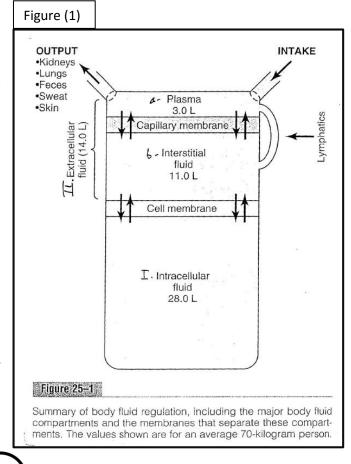
*Body Fluids; compartments and composition.

*Water balance, and Defense of body fluid volume.

*The lymphatic system.

* Body fluids

- I. Major body fluid Compartments Figure (1)
 - In a 70-Kg person, body fluid constitutes about 42 liters.
 - Body fluids occupy two main compartments:
 - The <u>Intracellular</u> compartment → holds most of the total body fluids (28 liters out of 42).
 - 2) The <u>Extracellular</u> compartment, that is further divided into:
 - Interstitial fluid 11 liters.
 - ▶ Plasma -3 liters.



II. Composition of extracellular and intracellular compartments

INTR	OSITION OF	EXTRACELLULAR EXTRACELLULAR LUID COMPARTI less otherwise ind	MENTS	
EXTRACELLULAR FLUID				
AND PROPERTIES	PLASMA	INTERSTITIAL FLUID	FLUID	
Sodium Potassium Calcium Magnesium Chloride Sulfate Bicarbonate Phosphate Organic acids Proteins Glucose (av) Lipids (av) pH	142 4 5 2 101 1 27 2 6 <u>16</u> 90 mg% 0.5 g% 7.4	145 4 5 2 114 1 31 2 7 1 90 mg% -7.4	10 160 2 26 3 20 10 100 <u>65</u> 0-20 mg% 6.7	

Notes concerning the previous table:

• The concentration of each substance in the plasma is similar or near to its concentration in the interstitial fluid. EXCEPT when we talk about proteins. There is a significant difference between the concentration of proteins in plasma (16 meq/L) and the concentration of proteins in the interstitial fluid (1 meq/L).

Additional piece of information:

Meq/L means milliequivalents of solutes per liter of solvent. This unit is especially common for measurement of components in biological fluids.

• The concentration of sodium outside the cell is much higher than its concentration intracellularly. Whereas the concentration of potassium intracellularly is much higher than its concentration outside the cells.

(note: The Na^+/K^+ pump has a key role in maintaining this state, and if this pump does not function properly, the human being will not survive).

• The concentrations of <u>potassium</u>, <u>sulfate</u>, <u>phosphate</u>, and <u>proteins</u> are much **higher inside the cells** compared to their extracellular concentrations. On the other hand, the concentrations of <u>sodium</u> and <u>chloride</u> are much **higher extracellularly** compared to their intracellular concentrations.

III. Distribution of water in various tissues and organs.

	DIS		TABLE 16 OF WAT S AND (ER IN VARIO	US	
Т	TISSUE/ORGAN	PERCENT	·	PERCENT		L. IN. 70
		WATER	ан 8	BODY WEIGH	Т	KG MAN
5	Skin	72.0	· ·	(8.0)		9.07
1	Muscle	75.7		41.7		22.10
S	Skeleton	31.0		(15.9)		3.45
F	Brain	74.8		2.0		1.05
I	Liver	68.3		2.3		1.10
F	Heart	79.2		0.5		0.28
	Lungs	79.0		0.7		0.39
	Kidneys	82.7		0.4		0.23
S	Spleen	75.8		0.2		0.11
E	Blood	83.0		7.7	14	4.47
I	Intestine	74.5		1.8	,	0.94
	Adipose	10.0		9.0		0.63
]	Total body	62.0		100.0		43.40

• Dr. Saleem focused on the encircled values:

A. Water comprises \rightarrow 83 % of blood, 82.7 % of the kidneys (the highest values) And 10 % of adipose tissue (the lowest value)

- B. Skin comprises 18 % of total body weight.
 Muscle → 41.7 % of total body weight.
 Skeleton → 15.9 % of total body weight.
- C. Amount of water In skin \rightarrow 9.07 liters. Amount of water In muscle \rightarrow 22.1 liters. Amount of water In blood \rightarrow 4.47 liters.

IV. Total body water (TBW) in relation to age and sex

Table 1-3. TBW (as percentage of body weight) in relation to age and sex.*)*
Age	Male	Female	
10-18	59%	57%	=02 %
18-40	61%	51%	= 10%
40-60	55%	47%	= 08%
Over 60	52%	46%	= 06%.

• Before the age of 18, there's almost no difference in TBW (as a percentage of body weight) between males and females.

After the age of 18, a significant difference appears between males and females because of the effects of sex hormones.
(*Females have a higher proportion of body fat, as a result, total body water is about 50% of their body weight, whereas about 60% of the body of an adult male is water*).

• After the age of 40, the difference between males and females starts to decline gradually. (note that the 10 % difference in the table declined to 8% then to 6%).

V. Osmolality of plasma

- The total osmolality of a solution is the sum of the osmolality due to each of the constituents.
- The osmolality of plasma is around 290 mOsmol/kg.
- The principle ions (Na⁺, K⁺, Cl⁻, ..., etc.) contribute to 280 mOsmol/kg (about 96% of the osmolality of plasma).
- Glucose, amino acids, and other small non-ionic substances contribute to approximately 10 mOsmol/kg.
- Proteins contribute only around 0.5% to the total osmolality of plasma.

Additional pieces of information

(not mentioned by Dr. Saleem)

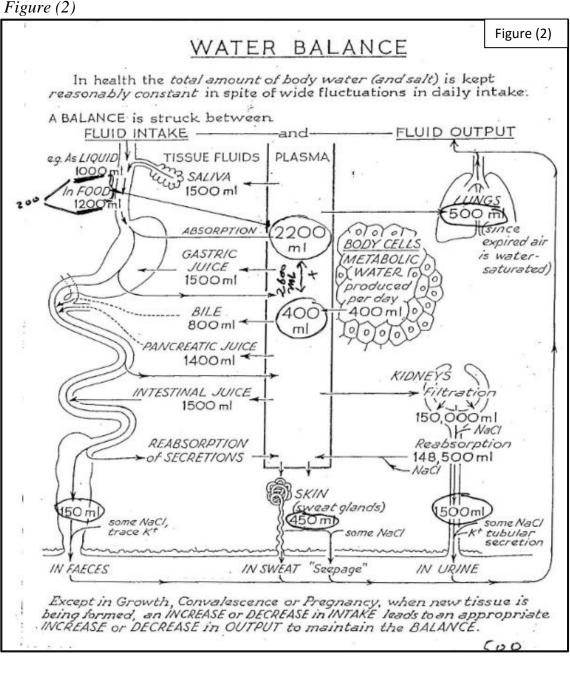
- **Osmol** (unit): the number of moles of solute that contribute to the osmotic pressure of a solution.
- **Osmolality:** a measure of osmols of solute per kg of solvent.
- **mOsmol/kg** means milliOsmols per kg of water. This unit is used when we measure osmolality in the relatively dilute fluids in the body.

* Water Balance

- The body mechanisms must maintain the concentartions of electrolytes and other elements inside and outside the cells in their normal ranges.
- The purpose of keeping body fluids normal is to keep the blood volume normal, **and consequently, maintaining the normal blood pressure.** (the main goal is to normalaize blood pressure).
- One of these body mechanisms include the Na⁺/K⁺ pump. Other methods are through the actions of certain hormones that will be explained in the next section.

Water balance in the body I.





- Fluid intake = Fluid output = 2.6 L
- Fluid intake:
 - Fluid ingested as liquid : 1 L
 - ➢ Fluid in ingested food : 1.2 L
 - ► Fluid from metabolism : 0.4 L

Total fluid intake = 2.6 L

- Fluid output:
 - \succ Through urine : 1.5 L
 - \succ Through expiration : 0.5 L
 - \blacktriangleright Through sweat glands : 0.45 L
 - ➤ Through faeces : 0.15 L
 - Total fluid output = 2.6 L

Note: from the value "fluid in ingested food" (1.2 L), we can tell that the figure refers to populations of the Middle East and Asia. While in European populations, very little amount of the fluid ingested comes from food.

Note 2: the value of fluid output through urine is changeable depending on the amount of the ingested fluid, whereas the values of fluid output through the other three ways are relatively constant.

II. Defense of body fluid volume

Figure (3)

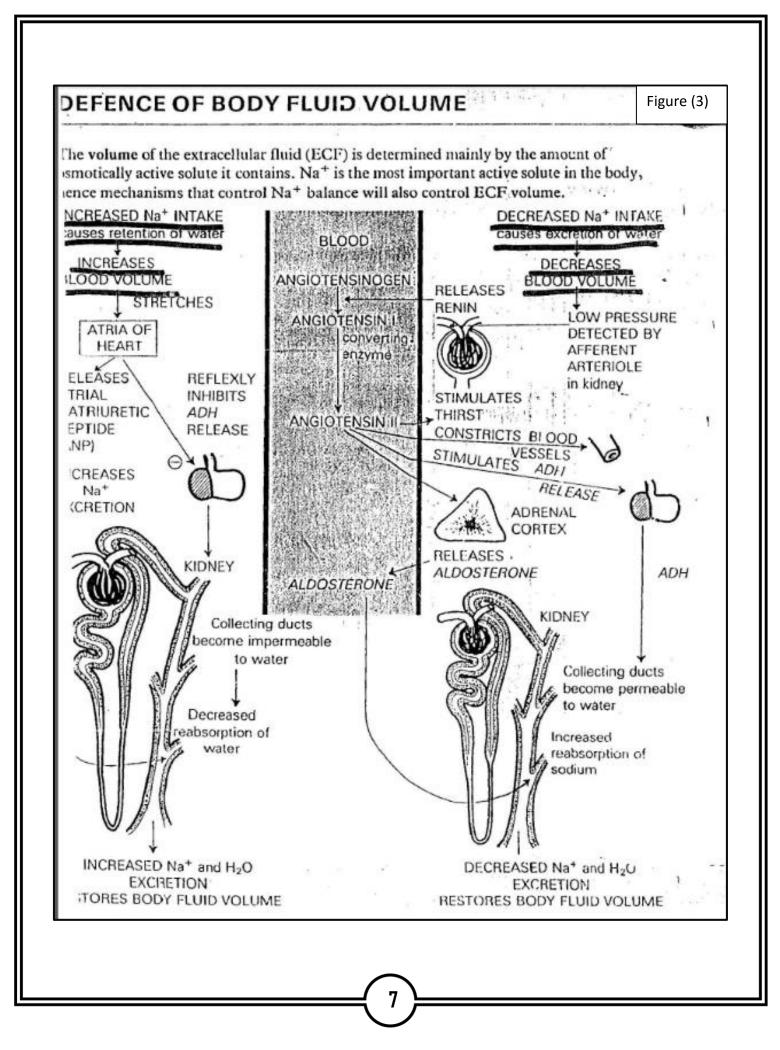
There are defense mechanisms against abnormalities in body fluids (excess or reduced fluid volume).

A. <u>In the case of incresed body fluid volume (or "high blood volume")</u>:

- ➤ A hormone called "Atrial Natriuretic Peptide (ANP)" is released from the atria of the heart. This hormone increases sodium excretion.
- > Reflexly, ADH release from the posterior pituitary is inhibited \rightarrow sodium and water excretion is stimulated.
- \rightarrow The increased Na⁺ and water excretion restores normal body fluid volume.
- B. In the case of reduced body fluid volume
 - Renin will be released from the kidneys. It will convert angiotensinogen into angiotensin I that will be further converted into angiotensin II by a converting enzyme.
 - > Angiotensin II is responsible for:
 - 1. Thirst \rightarrow the individual will ingest more water
 - 2. Constriction of blood vessels
 - 3. Stimulation of the release of ADH
 - 4. Stimulation of the realese of aldosterone from the adrenal cortex.

The previous will lead to increased reabsorption of Na⁺ and water.

→ The decreased excretion of Na⁺ and water restores normal body fluid volume.



III. Dehydration

- If the hemeodynamic mechanisms fail to operate properly, loss of fluid, electrolytes (ions), or both may occur. This is called dehydration.
- Three conditions (*types*) may arise depending on the relative losses of fluid and electrolytes:

1) Isotonic dehydration

(equal loss of fluid and electrolyte)

There is a proportionate loss of fluid and electrolyte so that the total volume of ECF changes but its osmotic pressure remains within normal limits.

2) Hypertonic dehydration

(excessive fluid loss as compared to electrolytes)

More fluid than electrolyte is lost. As a result, the ECF becomes more concentrated. Water thus tends to be drawn from cells.

3) Hypotonic dehydration

(excessive electrolyte loss as compared to fluids)

More solute than fluid is lost, therefore, the ECF becomes diluted. Water thus tends to enter cells.

IV. Hydration (Water Intoxication)

- Hydration is a term referring to the results of;
 - Excessive water intake
 - Decreased loss of water, or
 - Increased reabsorption of water from the kidney because of ADH administration.

(these three are the causes of hydratiom (or water intoxication))

• Excessive water intake may produce the syndrome of water intoxication in which cellular function is disturbed by the dilution of cellular electrolytes.

• Symptoms or consequences of water intoxication include:

Disorientation, convulsions, and coma.

Gastrointestinal dysfunction, muscular weakness, and abnormal cardiac rhythms.

Note: if alcohol is unavailable, alcohol addicts tend to drink too much water (approximately 10 L) because the result (disorientation) would be as if they got drunk due to alcohol intake.

* <u>The Lymphatic System</u>

I. Formation of lymph

Look at *figure (4)* that shows part of a capillary with its two ends (arterial and venous ends).

• <u>At the arterial end:</u>

The blood pressure is 32 mmHg(favoring the movement outwards)The oncotic (plasma protein) pressure is 28 mmHg (favoring the movement inwards)Net pressure = 4 mmHg(outwards)

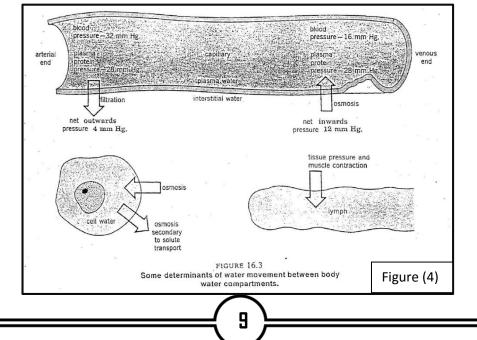
→ The net outwards pressure (4 mm Hg) at the arterial side of the capillary causes filtration (<u>fluid and some proteins</u> exit the capillary to the interstitial fluid).

• <u>At the venous end:</u>

The blood pressure is 16 mmHg	(favoring the movement outwards)	
** decreased compared to the arterial end **		
The plasma protein pressure is 28 mmHg	(favoring the movement inwards)	
** stays the same **		
Net pressure = 12 mmHg (inwards)		

- → The net inwards pressure of 12 mmHg at the venous side causes most of the lost <u>fluid</u> at the arterial side to return back to the capillary (by <u>osmosis</u>).
- the <u>remaining (retained) little amount of fluid</u> and <u>protein</u> that did not go back to the capillary will pass to the lymphatic vessels and this is "the lymph" (fluid and little protein that passed to the lymphatic vessels).

(**note**: we said "little" because it is a small amount *relatively*, *compared to the amount that went back to the capillary*, but in reality, it's not a small amount).



II. The lymphatic system

- The lymphatic system represents an accessory route through which fluid can flow from the interstitial spaces into the blood.
 - 1. Lymph is tissue fluid that enters the lymphatic vessels. It drains into the venous blood via the **thoracic and right lymphatic ducts**.
 - It contains <u>clotting factors</u> and clots on standing in vitro. We know that plasma clots, but does the lymph clot?? Yes (Due to the **presence of clotting factors**), but coagulation of lymph is weak as a consequence of the **absence of platelets**.
 - 3. In most locations and mainly in the liver-, it also contains <u>proteins</u> that traverse capillary walls and return to the blood via the lymph. Its protein content is generally lower than that of plasma, which contains about 7 g/dL, but *lymph protein content varies with the region from which the lymph drains*.
 - 4. <u>Water insoluble fats</u> are absorbed from the intense into the lymphatics, and the lymph in the thoracic duct after a meal in milky because of its high fat content.
 - 5. <u>Lymphocytes</u> enter the circulation principally through the lymphatics, and there are appreciable numbers of lymphocytes in the thoracic duct lymph.

The underlined words above are the answer to the question "what can we find in lymph?"

- Note: in the lymph nodes, there are lymphocytes that capture microbes, and usually, the microbe will not return back to the capillaries, otherwise, it would cause a disease.
- Three organs closely related to the lymphatic system are the **spleen**, **tonsils**, and **thymus**.
- Tissues that lack lymphatic capillaries include:
 - Avascular tissues (such as cartilage, the epidermis, and the cornea of the eye)
 - The central nervous system
 - Portions of the spleen
 - ➢ Bone marrow

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III. Flow of lymph

Lymph, like venous blood, is under relatively low pressure and may not flow readily through the lymphatic vessels without the aid of outside forces. These forces include:

- Contraction of skeletal muscles
- > Pressure changes due to the action of breathing muscles, and
- Contraction of smooth muscles in the wall of larger lymphatic vessels.

IV. The most important functions of the lymphatic system

From the following slide, Dr. Saleem focused on the 4 main headlines, but it's better to read the rest (the details) for better understanding of these 4 main functions.

- 1. Return of excess filtered fluid
- 2. Defense against disease
- **3.** Transport of absorbed fat
- **4.** Return of filtered protein

Following are the most important functions of the lymphatic system:

- Return of Excess Filtered Fluid. Normally, capillary filtration exceeds reabsorption by about 3 liters per day (20 liters filtered, 17 liters reabsorbed) (Fig. 8-17c). Yet the entire blood volume is only 5 liters, and only 2.75 liters of that is plasma. (Blood cells make up the remainder of the blood volume.) With an average cardiac output, 7,200 liters of blood pass through the capillaries daily under resting conditions (more when cardiac output increases). Even though only a small fraction of the filtered fluid is not reabsorbed by the blood capillaries, the cumulative effect of this process being repeated with every heartbeat results in the equivalent of more than the entire plasma volume being left behind in the interstitial fluid each day. Obviously, this fluid must be returned to the circulating plasma, and this task is accomplished by the lymph vessels. The average rate of flow through the lymph vessels is 3 liters per day, compared with 7,200 liters per day through the circulatory system.
- Defense Against Disease. The lymph percolates through lymph nodes located en route within the lymphatic system. Passage of this fluid through the lymph nodes is an important aspect of the body's defense mechanism against disease. For example, bacteria picked up from the interstitial fluid are destroyed by special phagocytic cells located within the lymph nodes.
- Transport of Absorbed Fat. The lymphatic system is important in the absorption of fat from the digestive tract. The end products of the digestion of dietary fats are packaged by cells lining the digestive tract into fatty particles that are too large to gain access to the blood capillaries but that can easily enter the terminal lymphatic vessels.
- Return of Filtered Protein. Most capillaries permit leakage of some plasma proteins during filtration. These proteins cannot readily be reabsorbed back into the blood capillaries but they can easily gain access to the lymphatic capillaries. If the proteins were allowed to accumulate in the interstitial fluid rather than being returned to the circulation via the lymphatics, the interstitial-fluid-colloid osmotic pressure (an outward pressure) would progressively increase while the blood-colloid osmotic pressure (an inward pressure) would progressively fall. As a result, filtration forces would gradually increase and reabsorption forces would gradually decrease, resulting in progressive accumulation of fluid in the interstitial spaces at the expense of loss of plasma volume.

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V. Edema

- Edema is a condition caused by accumulation of fluid (as well as proteins) in the interstitial compartment.
- Probable causes of edema (*figure* (5)):

1. High capillary pressure

 \rightarrow Higher than normal amount of fluid is filtered

 \rightarrow Not the whole filtered blood will return back to the capillary

 \rightarrow Fluid accumulates in the interstitial spaces causing edema

2. Low blood protein (low protein pressure)

→ The low protein pressure causes less than normal amounts of fluid to return back by osmosis from the interstitial spaces

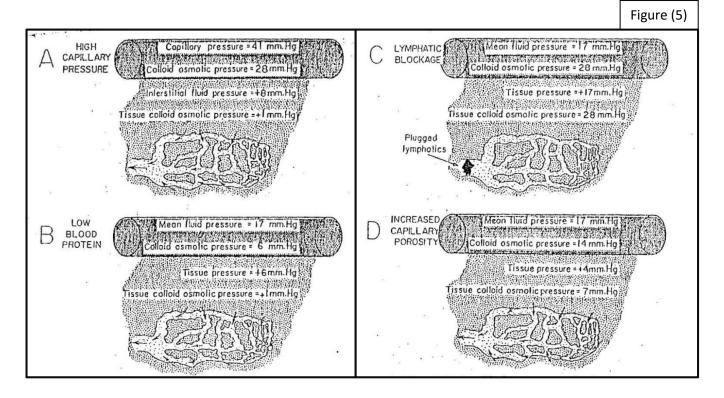
 \rightarrow Fluid accumulates in the interstitial spaces causing edema

3. Lymphatic blockage

Some parasites can block lymphatics and thus prevent fluid from returning back to the circulation.

4. Increased capillary porosity

(increase in the diameter of capillary pores) *Same consequences as in the first cause.*



Final note

For better understanding, it's recommended to study the slides of this lecture (they start from slide 19 in the third handout).

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* <u>Problems related to blood groups and their inheritance</u>

I. A child's blood type is "O NN rh". His mother's blood type is "A MN Rh". a man with a blood type of "A NN Rh" is suspected to be the father. Is it possible for this man to actually be the father of the child or not?

Hints table		
Phenotype	Genotype	
А	AA or AO	
В	BB or BO	
AB	AB	
0	00	
NN	NN	
MM	MM	
MN	MN	
Rh (positive)	RhRh or Rhrh	
rh (negative)	rhrh	

• The first step is to write down the phenotypes (of the child, his mother, and the man) and the probable genotypes.

	The man	the mother	The child
Phenotype	A NN Rh	A MN Rh	O NN rh
Probabilities of the Genotype	AA NN RhRh AO NN RhRh AA NN Rhrh AO NN Rhrh	AA MN RhRh AO MN RhRh AA MN Rhrh AO MN Rhrh	OO NN rhrh

• If we look at the child's Genotype, we can conclude that the two gametes that resulted in this genotype carried the following genes:

ONrh (on the first gamete) & ONrh (on the second gamete)

One of these two is from the mother, the second is from the father. When the paternal gamete fertilizes the maternal one, the child's genotype (OO NN rhrh) will result.

- If we look at the last probable genotype in each of the second and third columns, we'll find out that these two genotypes together could have been what resulted in the child's genotype, because:
 - The possible combinations of gametes resulting from the genotype AO MN Rhrh of the mother include ONrh
 - The possible combinations of gametes resulting from the genotype AO NN Rhrh of the man also include ONrh, meaning that <u>this man</u> <u>could actually be the father of the child.</u>

- II. A man with "B MM rh" blood type got married to a woman with "AB NN Rh" blood type.
 - A) Write down the possible genotypes that could have resulted in the phenotypes of the male and the female.
 - B) Write down the possible combinations of gametes resulting from each genotype you wrote in part A).

The man	The woman
B MM rh	AB NN Rh
1. BB MM rhrh	1. AB NN RhRh
> BMrh	> ANRh
> BMrh	> BNRh
2. BO MM rh rh	2. AB NN Rhrh
> BMrh	> ANRh
➢ OMrh	> ANrh
	> BNRh
	> BNrh
	B MM rh 1. BB MM rhrh $\geq BMrh$ $\geq BMrh$ 2. BO MM rh rh $\geq BMrh$

I apologize for any mistake I may have made.

Wish you all best of luck **:D**