

Volume of Distribution

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Volume of Distribution (Vd)

- **Defined as the total volume of body fluids in which a drug appears to be distributed according to its concentration in the blood or plasma.**
- **It is an apparent volume
e.g chloroquine 13000L/70kg**

The major body fluid compartments are

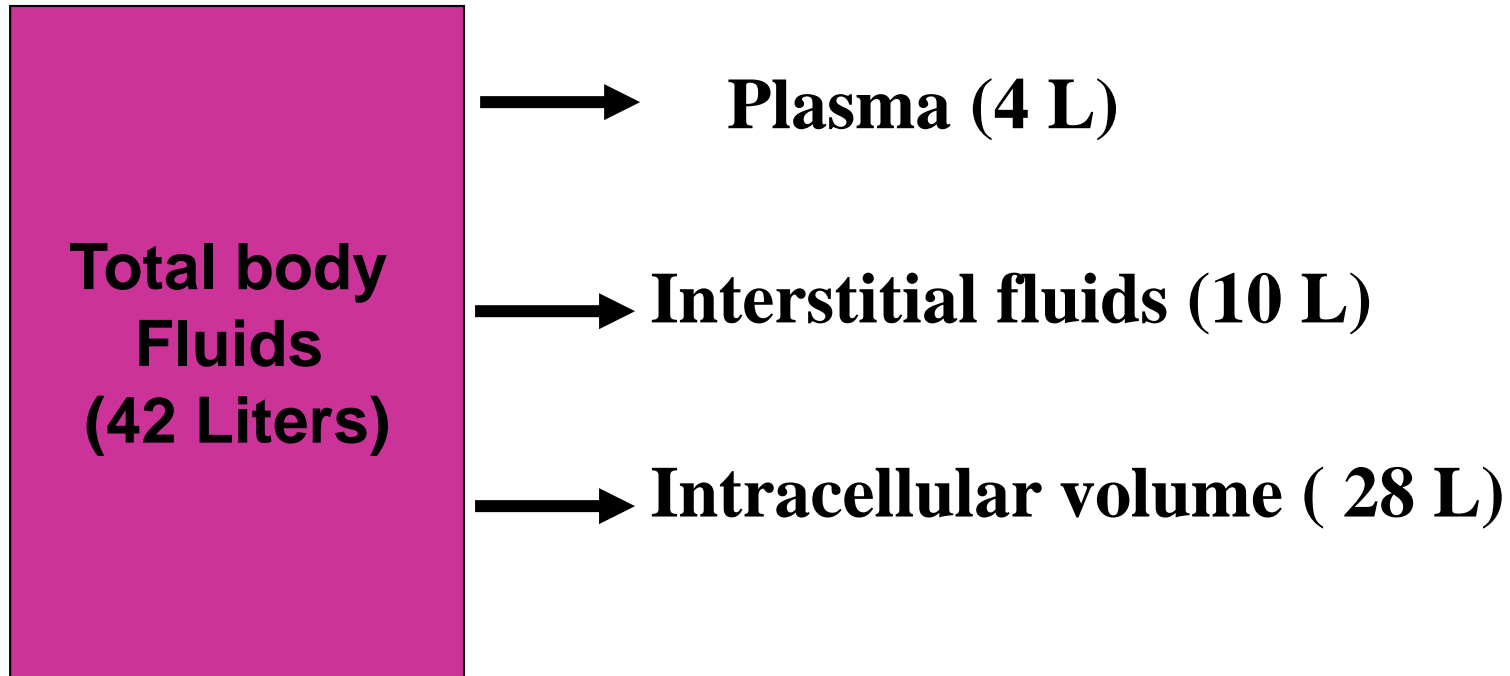
➤ **Extracellular fluid (22%)**

- Plasma (5 % of body weight = 4 liters).
- Interstitial fluid (16 % = 10 liters).
- Lymph (1 %).

➤ **Intracellular fluid (35 %)**

- fluid present inside cells (28 L).

Total body fluids (60% of body weight in 70-kg individual)



Volume of Distribution (Vd)

- It relates the amount of drug in the body to the concentration of drug in blood and is calculated by the following formula:

$$Vd \text{ (L)} = \frac{\text{Total amount of drug in body (mg)}}{\text{Concentration of drug in blood (mg/L)}}$$

Volume of Distribution (V_d)

- **V_d can vastly exceed any physical volume in the body because it is the volume apparently necessary to contain the amount of drug homogeneously at the concentration found in blood**
- **It represents balance between drug which is distributed in body tissue and the plasma.**

Volume of Distribution (V_d)

- V_d is directly correlated with the amount of drug distributed into tissue
- a higher V_d indicates a greater amount of tissue distribution.
- V_d greater than the total volume of body fluids (approx 42 L in humans) indicate that the drug is highly distributed into tissue.

Factors affecting the V_d of a Drug:

- ▶ **lipid solubility**

↑ lipid sol ↑ V_d

- ▶ **Plasma protein binding**

↑ PPB ↓ V_d

- ▶ **Tissue Binding**

↑ BT ↑ V_d

- ▶ **Body fat**

In obese ↑ V_d e.g. theophylline and ↓ Digoxin

- ▶ **Diseases like CCF, Renal failure**

↑ V_d e.g. Gentamicin a hydrophilic drug

Volume of Distribution (Vd)

Drugs with high Vd

- ▶ Have higher concentrations in tissues than in plasma.
- ▶ Relatively lipid soluble.
- ▶ Distributed intracellularly
- ▶ Not efficiently removed by haemodialysis.
e.g. phenytoin, morphine, digoxin

Volume of Distribution (Vd)

Drugs with low Vd

- ▶ confined to plasma & interstitial fluid.
- ▶ distributed in extracellular compartments.
- ▶ Polar drugs or lipid insoluble drugs. e.g. Carbenicillin, gentamycin.
- ▶ High MW e.g. heparin – insulin.
- ▶ High plasma protein binding e.g. warfarin.
- ▶ Do not cross BBB or placental barriers.
- ▶ Easily Removed by Haemodialysis

THANKS

The End