

## **B24 - Intravenous fluids**

**(a) To describe the composition, pH and osmolality of crystalloids and colloids used in clinical practice.**

- role = plasma expanders

### **Colloids**

=

- part way between a suspension and a fluid

(1) albumin

(2) gelatine derivatives

- haemaccel

- gelofusion

(3) polysaccharide derivatives

- dextran 40

- dextran 70

(4) starch solutions

- hetastarch

- pentastarch

### **Crystalloids**

=

(1) hartmans

(2) normal saline

(3) 5% dextrose

(4) dextrose saline (4% dex, 1/5 N/S)

(5) hypertonic saline (7.5% & 3%)

### **Albumin**

#### *Composition*

- protein solution

- clear, straw coloured solution

- 4, 5, 20 & 20% protein
- 96% albumin
- sodium bicarbonate, sodium carbonate and or acetic acid -> adjust pH to 6.4 to 7.4
- no preservatives
- prepared from pooled venous blood
- pasturised at 60 C for 10hrs
- Na<sup>+</sup> 130 to 160mmol/L
- pH = 6.4 to 7.4

*Osmolality* - isoosmotic

### **Haemaccel**

#### *Composition*

- gelatine derivative
- polypeptide produced by the thermal degradation of bovine gelatin -> cross linked with urea
- 500mL plastic bottles containing a sterile, pyrogen free, straw coloured solution.
- pH = 7.2
- 35g of polygeline
- 145 mmol/L of NaCl
- 5.1mmol of K<sup>+</sup>
- 6.25mmol of Ca<sup>2+</sup>

*Osmolality* = 7.4

### **Gelofusion**

#### *Composition*

- gelatine derivative
- Na<sup>+</sup> 154 & Cl 120 mmol/L
- 1.2g of sodium hydroxide
- pH = 7.4

*Osmolality* = 274mosmol/L

### **Dextran 40**

#### *Composition*

- polysaccharide derivative

- polysaccharide derived from sucrose by the action of the bacterium *Leuconostoc mesenteroides* -> further processed by hydrolysis & fractionation.
- clear, colourless 10% solution
- in either 5% dextrose or 0.9% saline
- pH = 3 to 7

*Osmolality* - 300 mosmol/L

## **Dextran 70**

### *Composition*

- polysaccharide derivative
- polysaccharide derived from sucrose by the action of the bacterium *Leuconostoc mesenteroides* -> further processed by hydrolysis & fractionation.
- 6% solution
- with either 5% dextrose or 0.9% saline
- pH = 4 to 7

*Osmolality* = 300mosmol/L

## **Hetastarch**

### *Composition*

- starch solution
- synthetic polymer derived from amylopectin
- hydroxyethyl groups are substituted into the glucose units to retard degradation by serum amylase.
- clear, pale yellow 6% solution of hetastarch in 0.9% saline.
- also contains: NaCl 154 mmol/L

*Osmolality* = 300mosmol/L

## **Normal Saline**

### *Composition*

- 0.9%
- 150mmol of NaCl
- an inorganic salt
- pH = 4.5 to 7
- no preservatives & no microbial agents

## **5% Dextrose**

### *Composition*

- monosaccharide obtained by the hydrolysis of cornstarch
- clear, colourless sterile solution
- contains 5% dextrose in H<sub>2</sub>O
- no buffers or bacteriostatic agents
- pH 3.5 to 6.5

*Osmolality* = 250mosmol/L

### **Dextrose saline (4% dex, 1/5 N/S)**

### *Composition*

- 56mmol/L of dextrose
- pH =

*Osmolality* = 310mosmol/L

### **Hartmans solution**

### *Composition*

- compound sodium lactate
- clear, colourless solution
- Na<sup>+</sup> 131 mmol/L
- Cl<sup>-</sup> 111mmol/L
- Ca<sup>2+</sup> 2mmol/L
- K<sup>+</sup> 5mmol/L
- lactate 29mmol/L -> converted to HCO<sub>3</sub><sup>-</sup> in liver
- pH = 6 to 7.3

*Osmolality* = 274mosmol/L

**(b) To evaluate their effects and fate when used in volume replacement.**

### **Albumin**

- 5% = iso-oncotic
- 20% and above = draws 3 times its volume of tissue fluid into the vascular space in 15min.
- when 1L administered:

- tonicity unaltered
- no change in osmolality
- 20% increase in blood volume

-> detected by volume receptors (above 7-10%) ->

(1) fall in ADH -> excretion of H<sub>2</sub>O

(2) H<sub>2</sub>O loss increases the plasma oncotic pressure -> water moves from ISF to IVF

### **Gelatine derivatives**

- equilibrates throughout the body
- 30% in circulating blood
- 30% in interstitial fluid
- 40% in urine
- > restores haemodynamic status after a period of hypovolaemia

### **Polysaccharide derivatives**

- by its osmotic effect it will hold H<sub>2</sub>O in the vascular space
- 500mL of dextran -> increase the plasma volume by 1000mL
- infusion of 1L of Dextran:
  - increase in blood volume by 20%
  - no change in osmolality
  - sensed by volume receptors (change in >7-10% occurred) -> increased ANP secretion, renal afferent arteriolar vasodilation -
  - > increased GFR, decreased ADH, decreased DCT Na<sup>+</sup> reabsorption -> increased urine output

(1) in the hypovolaemic there will be a return towards normal of haemodynamic parameters.

(2) decrease in ADH secretion

### **Starch derivatives**

See previous for colloids effect.

### **Normal saline**

- effects of 1L of normal saline:

- (1) distributes to the into the plasma where it remains for 30min before being uniformly distributed throughout the ECF (interstitial fluid:plasma - 3:1) -> ISF increases by 750mL and plasma by 250mL = 5% increase in blood volume -> no volume effect as no greater than 7-10% change
- (2) no change in osmolarity -> no osmoreceptor response
- (3) N/S has no protein -> lowered oncotic pressure -> increased GFR & smaller reabsorption of H<sub>2</sub>O in proximal tubule -> increased in urine output.

### **5% dextrose**

= a maintenance fluid

- isotonic

- administration of 1L of Dextrose 5%:

- (1) glucose quickly taken up by cells -> administration of pure H<sub>2</sub>O
- (2) H<sub>2</sub>O distributed throughout TBW in proportions (TBW distribution) -> of 1000mL, 670mL to intracellular fluid & 330mL of ECF
- (3) volume receptors not triggered c/o less than 7-10% change (around 2%)
- (4) 2.5% change in plasma osmolarity -> sensed by osmoreceptors (1-2% change enough) -> ADH decreases -> H<sub>2</sub>O excretion rises.

### **Dextrose saline**

- handle as for dextrose & saline separately

### **Hartmans**

- same as N/S

- except lactate -> HCO<sub>3</sub><sup>-</sup> by liver

### **Hypertonic saline**

- osmolarity is 3 times that of plasma (900mosmol/L)

- (1) Na<sup>+</sup> contents limits distribution to the ECF -> osmolarity increases to 304mosmol/L  
-> increased osmolarity sensed by osmoreceptors -> increased ADH secretion, increase in thirst

(2) increased osmolarity -> draws H<sub>2</sub>O out of cells by decreasing intracellular fluid volume -> increased ECF volume by 2L (only 500mL is in plasma) -> triggers volume receptors

-> ALD secretion will be suppressed

-> ANP will inhibit renin

-> natriuresis & excretion of H<sub>2</sub>O

### **Advantages of Colloids**

- smaller volume to infuse
- higher MW molecules -> longer in plasma -> prolonged increase in plasma volume
- minimal peripheral oedema
- higher systemic O<sub>2</sub> delivery
- may cross BBB less

### **Disadvantages of Colloids**

- greater expense (volume for volume) -> 30x more expensive than crystalloids
- coagulopathy
- pulmonary oedema
- decreased Ca<sup>2+</sup>
- impaired cross-matching (dextran)
- osmotic diuresis
- anaphylactic/oid reactions
- no proven reduction in mortality

### **Advantages of Crystalloids**

- low MW -> freely diffusable
- less expensive
- hypertonic saline good for traumatic brain injury
- greater urinary flow
- replaces sequestered interstitial fluid

### **Disadvantages of Crystalloids**

- short-lived haemodynamic improvement
- peripheral oedema
- pulmonary oedema
- larger infused volume required

**(c) To compare the pharmacology of colloids such as albumin, gelatin derivatives, polysaccharide derivatives and starch solutions with crystalloids such as lactate solutions & normal saline.**

### **Albumin**

#### Composition

- protein solution
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- 4, 5, 20 & 20% protein
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Osmolality - isoosmotic

PK

*Metabolism* - exogenous albumin enters the aa pool & undergoes biotransformation.

*Elimination* -  $t_{1/2}$  = 17 days

PD

*Main action* - plasma expansion

*Mechanism*

- exerts colloid oncotic pressure
- 5% = iso-oncotic
- 20 & 25% = will draw 3 x the administered volume into the circulation in 15 min.

*CVS*

- in context of hypovolaemia -> haemodynamic parameters towards normal
- myocardial depression

*GU*

- increased renal perfusion

*Other adverse effects*

- circulatory overload
- allergic reactions
- aluminium toxicity

## **Gelatin derivatives**

**Haemaccel**



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Osmolality = 7.4

#### *Distribution*

- t<sub>1/2</sub> alpha = 7hrs
- Vd 0.7L/kg

#### *Elimination*

- 80% urinary excretion
- 10% faeces

#### *Other adverse effects*

- provoke histamine release (anaphylactoid 1:1000) -> bronchospasm, tachycardia, rash & severe hypotension.
- don't mix with citrated blood

### **Gelofusion**

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### **Polysaccharide derivatives**

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*Osmolality* - 300 mosmol/L

PK

*Distribution* - Vd = 6L

*Metabolism* - by dextranases present in lung, liver & spleen

*Elimination*

- biphasic
- 70% by renal
- 30% by metabolism to CO<sub>2</sub> & H<sub>2</sub>O
- plasma t<sub>1/2</sub> = 7hrs

PD

*Main action* = plasma expansion

*Metabolic*

- reduces serum lipid concentration
- reduction in serum albumin

*Other side effects*

- severe hypersensitivity reactions in 1:3300
- circulatory overload
- increased capillary oozing
- ARF

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See above for similar details

## **Starch derivatives**

### **Hetastarch**

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- also contains: NaCl 154 mmol/L

Osmolarity = 300mosmol/L

Route - IV

#### Dose

- should not exceed 20mL/kg/day
- increased plasma volume lasts 24-36hrs

PK

#### *Distribution*

- small fraction taken up by spleen

#### *Elimination*

- hydrolysed by serum amylase into fragments
- renally excreted or removed by reticulo-endothelial compartments

PD

*Main action* - plasma expander

*CVS*

- circulatory overload

#### *CNS*

- headache

#### *GI*

- salivary gland enlargement
- vomiting

#### *Haemopoietic*

- interference with platelet function
- inhibits endothelial activation -> preventing neutrophil adhesion
- prolongation of APTT
- decreased FVIII, vWF & fibrinogen

#### *Other adverse effects*

- pyrexia
- itching
- anaphylactoid reaction 0.0005%

#### **Hartmans**

- overtransfusion -> hypernatraemia, pulmonary oedema & metabolic alkalosis
- not recommended in lactic acidosis

#### **Normal saline**

- overtransfusion -> oedema, hypernatraemia & hyperchloraemic acidosis.