

# Isotonicity

Colligative Properties and E values



# Isotonic solutions

## Objectives

- Understand and use the dissociation factor( $i$ ).
- Use the NaCl E value in adjusting a solution isotonicity, and performing other isotonicity related calculations (dilution and adjustment).
- Understand colligative properties and its use in adjusting solutions isotonicity.

# Isotonic solutions

Why prepare isotonic solution?

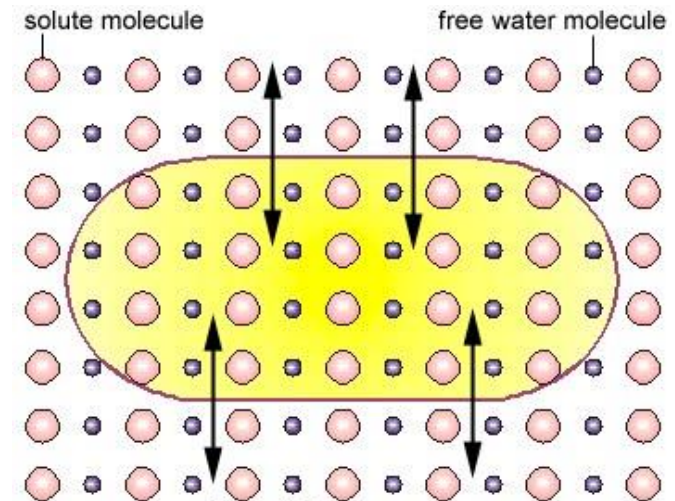
Which pharmaceutical solutions are isotonic?

Parenteral solutions?

Nasal solutions

Ophthalmic solutions

Enemas



Isotonic Environment:  
The solute concentration and the free water  
concentration are the same inside and  
outside the cell.

Water flows in and out of the cell at an equal rate.



# Isotonicity

Name  
another IV  
isotonic  
solution ?

- Solutions having the same osmotic pressure as that of **0.9% NaCl** are considered isotonic.
- **NaCl equivalent E** of a substance: The amount (in grams) of NaCl with equivalent osmotic pressure as that of 1 g of the substance.



# Isotonicity

You should be able to calculate the % concentration of an isotonic solution of any **substance** if you know its E

- Isotonic sodium chloride solution contains 0.9% NaCl, if a sodium chloride equivalent of boric acid is 0.52, what is the percentage strength of an isotonic solution of boric acid?



*E value of Boric acid is 0.53*

*0.53g NaCl  $\approx$  1g Boric acid*

*0.9g NaCl  $\approx$  Xg Boric acid*

$$x = \frac{1 \times 0.9}{0.53} = 1.7$$

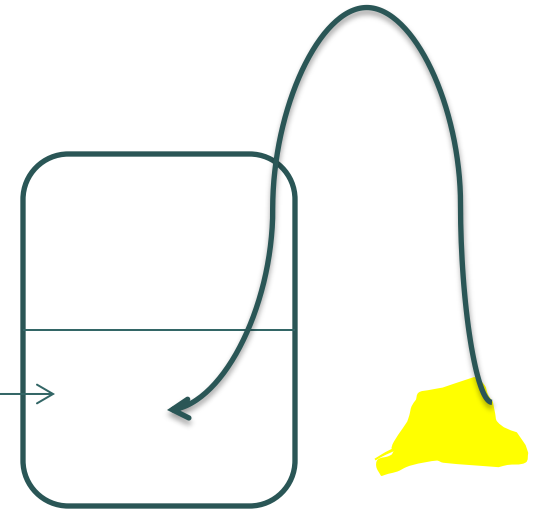
A solution of 1.7 % of Boric acid is isotonic

# Isotonic Solutions

- Useful equation (White Vincent)

$$V = w \times E \times 100 / 0.9$$

isotonic



$V$  = volume of the isotonic solution prepared by  $w$  (grams of the substance)

How many mL can be rendered isotonic with 1g Atropine sulfate, ( $E=0.13$ )?

What is the concentration of the isotonic solution of Atropine sulfate? ( what is the % strength of an isotonic solution of Atropine?)

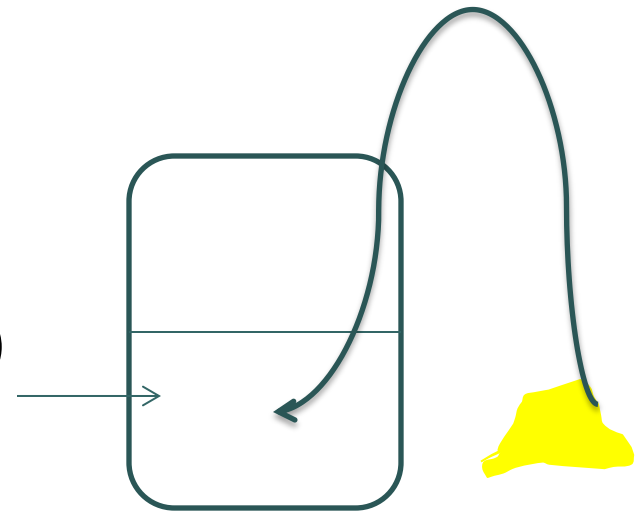
How can you prepare 1% isotonic Atropine sulfate solution?

# Isotonic Solutions

- Useful equation (White Vincent)

$$V = w \times E \times 100 / 0.9$$

isotonic



How many mL can be rendered isotonic with  
1g Atropine sulfate, ( $E=0.13$ )?

$$V = 1g \times 0.13 \times 100 / 0.9 = 14.4 \text{ mL}$$

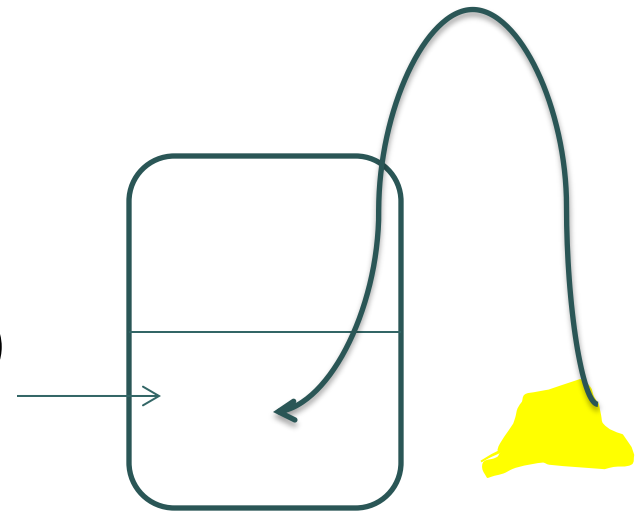


# Isotonic Solutions

- Useful equation (White Vincent)

$$V = w \times E \times 100 / 0.9$$

isotonic



What is the concentration of the isotonic solution of Atropine sulfate? ( what is the % strength of an isotonic solution of Atropine?)

● ● ● | *E value of Atropine acid is 0.13*

*0.13g NaCl  $\approx$  1g Atropine*

*0.9g NaCl  $\approx$  Xg Atropine*

$$x = \frac{1 \times 0.9}{0.13} = 6.9$$

A solution of 6.9% of atropine is isotonic

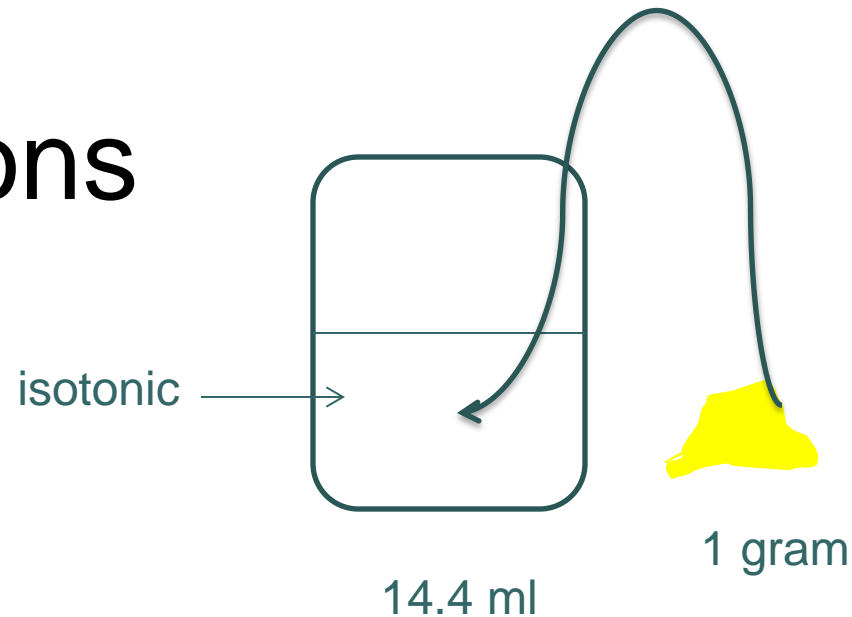
$$V = w \times E \times 100/0.9$$

$$100 = w \times 0.13 \times 100/0.9 = 6.7 \text{ g}$$

# Isotonic Solutions

- Useful equation (White Vincent)

$$V = w \times E \times 100 / 0.9$$



How can you prepare 1% isotonic Atropine sulfate solution?

$$V = 1g \times 0.13 \times 100 / 0.9 = 14.4 \text{ ml}$$

What about the rest of the 100 ml?

We can add NaCl or another substance to adjust the isotonicity.

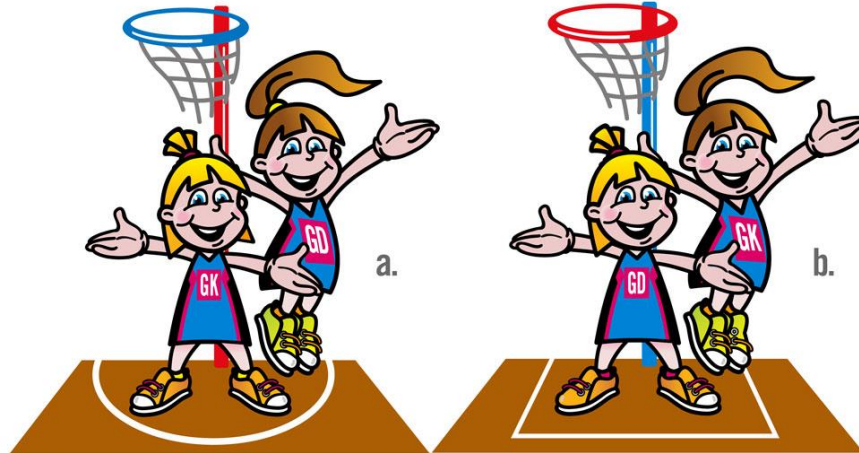
We have  $100 - 14.4 = 85.6$  ml to adjust its isotonicity

If we use NaCl the  $0.9\%$  of the  $85.4$  ml =  $0.77$  g

# Isotonicity

Naphazoline HCl (E=0.27)	1%
NaCl	qs
Water	qsad 30 ml
Dispense isotonic sol	

Naphazoline HCl (E=0.27)	1%
NS [normal saline]	qs
Water	qsad 30 ml
Dispense isotonic sol	





# Isotonicity

Sometimes we need to adjust the isotonicity using another agent other than NaCl. Why?

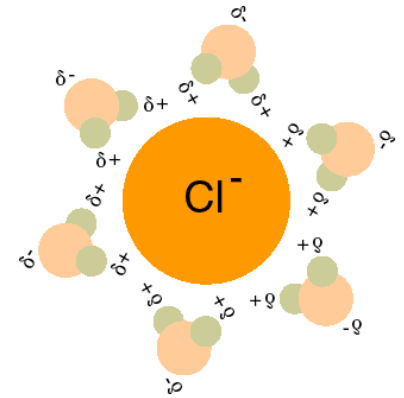
		E
Phenacaine HCl	1%	0.2
Chlorobutanol	0.5%	0.24
Boric acid	qs	0.52
Purified water	qs 60 mL	
Prepare isotonic solution		



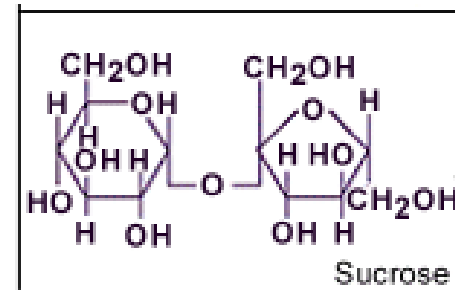
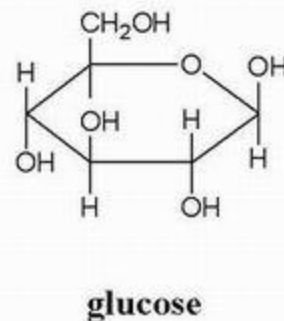
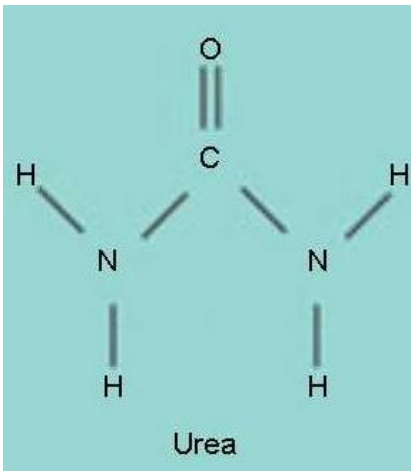
## Extra problem

- How many milligrams of NaCl are required to prepare the following prescription . **Note: Epinephrine solution is isotonic.**
- Atropine Sulfate ( $E = 0.14$ ) 1%
- Epinephrine solution 1:2000 8 mL
- Purified water Qs 30 mL

# Physical Properties of solutions: Colligative Properties



- Definition: Properties that depend on the concentration of the solute molecules or ions, not the identity of the solute.





# Colligative Properties

- Elevation of boiling point
- Depression of the vapor pressure
- **Depression of freezing point**
- **Osmotic pressure**





# Freezing Point Depression

- Definition: It is the point at which the liquid and the solid phases coexist in equilibrium at one atmosphere.
- Freezing points of a pure solvent is higher than the freezing points of a solution.
- The depression of the freezing point depends on the molal concentration of the solute.

# Freezing Point Depression

○  $\Delta T_f = 1.86 \times m$

*For water  $K_f = 1.86 \frac{\text{deg}}{\text{kg.mole}}$*

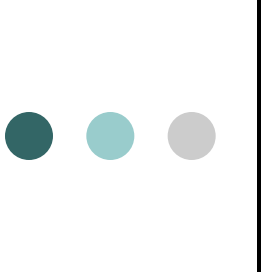
*( in diluted solutions  $m=M$ ).*

*What does this equation mean?*

*It means, but adding **one mole** of particles the freezing point of the 1 kilogram solvent (water) drops by  $1.86^\circ\text{C}$ .*

# Freezing Point Depression

- If you have 0.1 M solution of dextrose will result in  $\Delta T_f = 0.1 \times 1.86 = 0.186^\circ$ .
- 0.1 M solution of NaCl will result in  $\Delta T_f = i \times 0.1 \times 1.86$ . If  $i = 1.9$  then  $\Delta T_f = 0.353$ .



Calculate the  $i$  of solution of  $\text{Na}_2\text{SO}_4$   
(the dissociation percentage is 85%).  
Can you find  $\Delta T_f$  for 0.1  $\text{Na}_2\text{SO}_4$ ?



# Isotonic Solutions

- Freezing point depression,
  - An isotonic solution has a freezing point of  $-0.52^{\circ}\text{C}$  .

The isotonicity of parenteral fluids can be adjusted with NaCl or glucose till the solution acquires freezing point of  $-0.52^{\circ}\text{C}$  .

(Zatz Pharmaceutical Calculations book pg 244)



## Preparation of isotonic solution

$$w = \frac{0.52 - a}{b}$$

*0.52 is the  $\Delta T_f$  of isotonic solutions*

*w = weight% of the adjusting solution*

*a = freezing point (depression) of the solution to be adjusted.*

*b =  $\Delta T_f$  of 1% of the adjusting solution*



# Problem

- Calculate the NaCl needed to adjust the isotonicity of a 50 mL solution of 0.5% lidocaine HCl
- 1% lidocaine HCl causes a  $\Delta T_f$  of  $0.13^\circ\text{C}$ .
- 1% NaCl causes a  $\Delta T_f$  of  $0.576^\circ\text{C}$ .



# References

- Amiji Applied Physical Pharmacy book
- Physical Pharmacy by Martin
- [www.chemguide.co.uk](http://www.chemguide.co.uk)
- Physicochemical Principles of Pharmacy Alexander Florence Chapter 2 and 3.