

# Analgesia *et al.*

Dr William Dooley



# Plan

- Pain assessment
- Acute vs. Chronic pain
- Overdose / Toxicity

- 
- Other common meds
    - Anti-emetics
    - Laxatives

- Some calculations



# Pain assessment

The 5<sup>th</sup> Vital Sign

History, History, History

**S**ite

**O**nset

**C**haracter

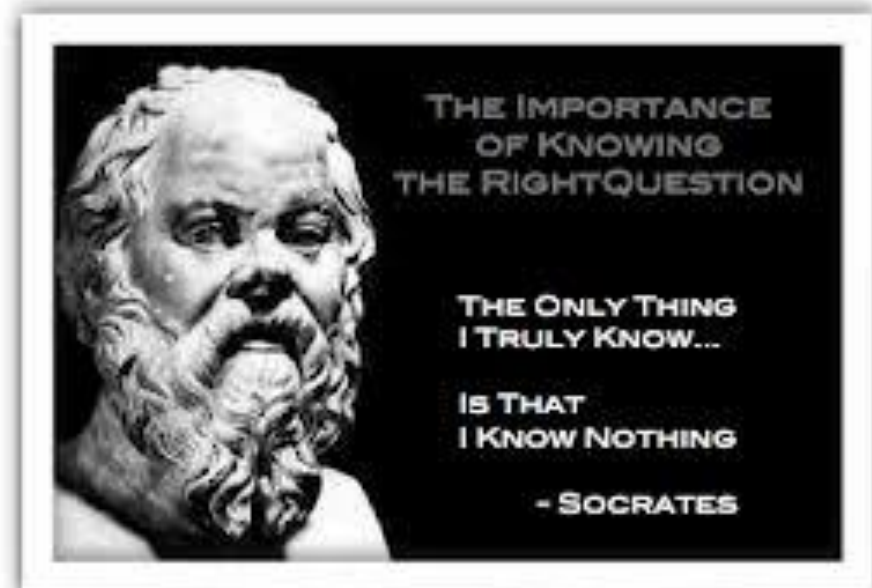
**R**adiation

**A**ssociated symptoms

**T**ime

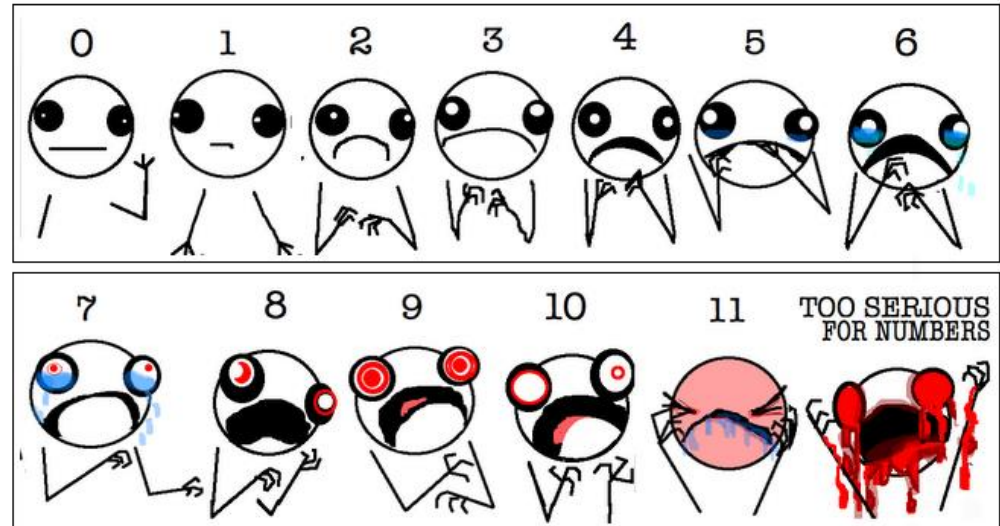
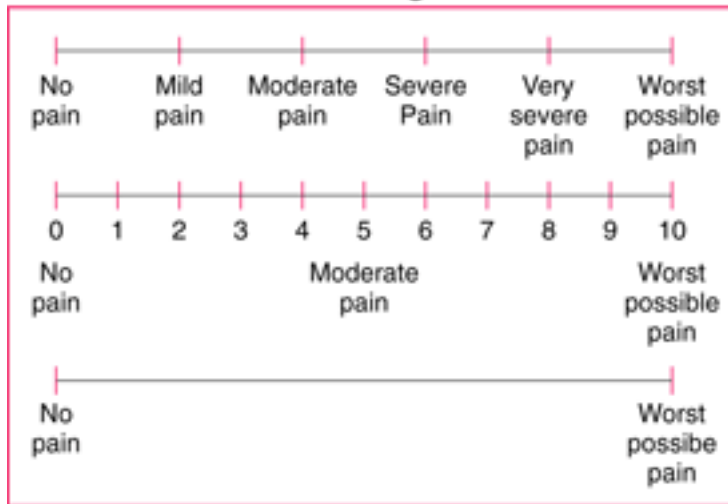
**E**xacerbating/Relieving factors

**S**everity



# Pain assessment

Visual Analog Scale



<http://hyperboleandahalf.blogspot.com>

Graphic Scale



Verbal Scale

"On a scale of 0 to 10, with 0 meaning no pain and 10 meaning the worst pain you can imagine, how much pain are you having now?"

Functional Pain Scale

- 0 = No pain
- 1 = Tolerable and pain does not prevent any activities
- 2 = Tolerable and pain prevents some activities
- 3 = Intolerable and pain does not prevent use of telephone, TV viewing, or reading.
- 4 = Intolerable and pain prevents use of telephone, TV viewing, or reading.
- 5 = Intolerable and pain prevents verbal communication

# Common Analgesia

- Paracetamol
- NSAIDs
- Weak opioids
- Strong opioids



# Paracetamol (*Acetaminophen*)

**500mg-1g QDS (Max. 4g/24hrs)**

**PO/PR/IV**

**Analgesia / Anti-pyrexial**

**Onset of action 10-30 mins**

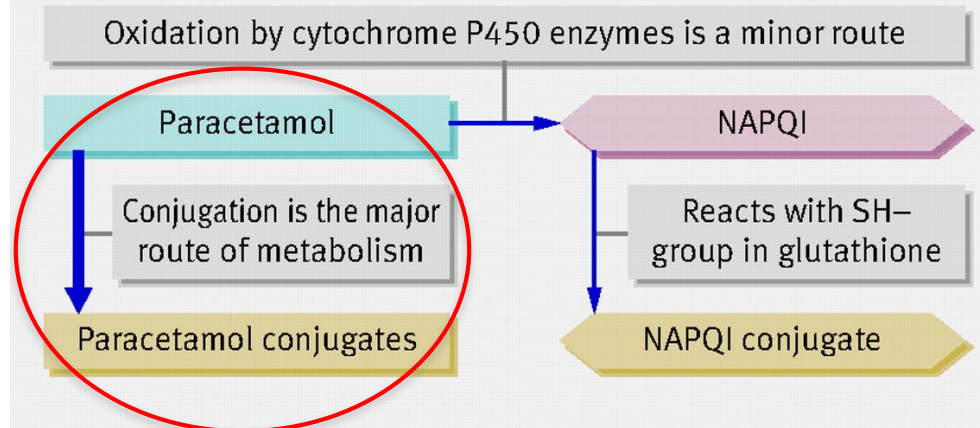
**Half life 1-4 hours**

**Metabolised by the liver and is hepatotoxic**

**Metabolites are then excreted by the kidney**



### At therapeutic doses





# Case of Overdose



What are his risk factors for suicide?

What more would you ask in the history??



# Paracetamol Overdose

History / History / History

## HPC

Dose- ?reliable historian / mixed

Time – once or staggered overdose

Symptoms – nausea and vomiting → RUQ pain

Psych- intentional / support / planned / alone /  
cry for help / warning / letter

## PMH

ETOH Abuse

Previous DSH / Suicide attempts

Chronic Liver Disease

## DH

Enzyme inducing drugs

(e.g. Carbamazepine, phenobarbital, phenytoin,  
rifampicin, St John's wort etc.)



# Paracetamol Overdose

## Examination

Low BMI / malnourished

Liver damage- jaundice / reduced GCS / asterixis / RUQ tenderness

## Investigation

Serum Paracetamol level – measured between 4-16hrs post ingestion

LFTs (particularly transaminases)

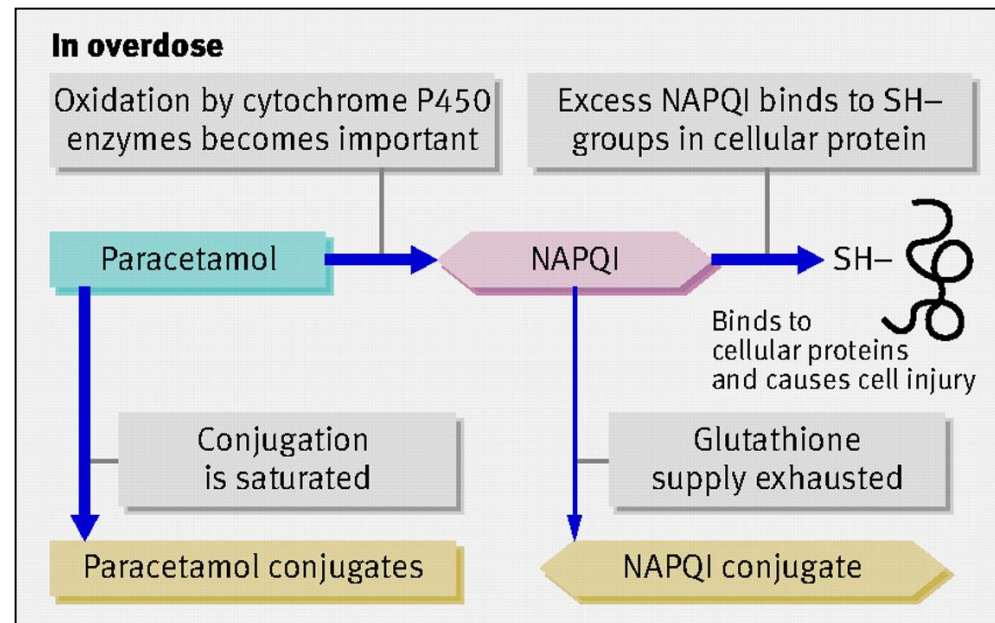
Clotting and INR

U+E

# Factors that increase the risk of liver injury

## High chance of glutathione depletion:

- Malnourished
- Eating disorders (anorexia or bulimia)
- Failure to thrive or cystic fibrosis in children
- Immunocompromised / AIDS
- Cachexia
- Chronic Alcoholism



# Treatment of Paracetamol OD

**Activated charcoal**

**Decreases absorption of paracetamol**

**Needs to be given quickly (within 1 hr)**

**N-acetylcysteine (NAC)**

**Antedote**

**Acts as precursor to glutathione to increase levels and reduce liver damage**

*Adverse effects*

**Nausea and vomiting**

**Anaphylactoid reaction which is histamine mediated**



# NAC use

1. Do they need treatment

2. Dose of treatment



# Case

**54yo female, PMH- depression (previous suicide attempts), chronic ETOH abuse, malnourished. Found unconscious with multiple packets of meds.**

**1.Do they need treatment**

**2.Dose of treatment**

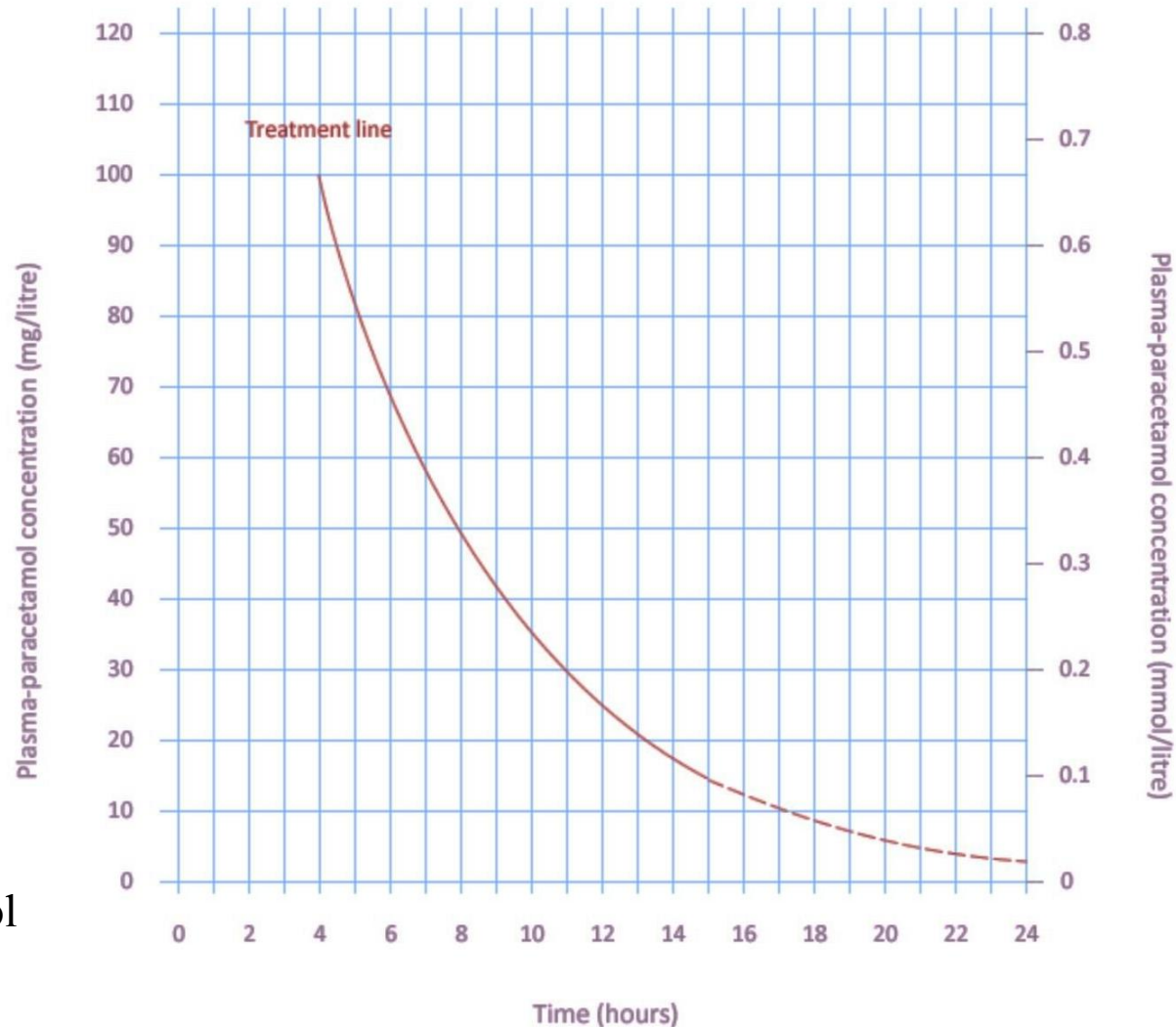


# Would you treat these?

1. OD 15x500mg tablets at 12:00.  
Blood at 18:00.  
PPC= 100mg/litre
2. OD 24x 500mg tablets at  
09:00. Blood at 17:00  
PPC= 30mg/litre
3. OD at 20:00 ?amount. Blood  
at 07:00. PPC= 30mg/litre

# What would you do ?

4. Staggered OD between 0900 and  
1600. Admits taking 50 Paracetamol  
tablets. Presents at 1630.
5. OD at 0800. Admits taking 4  
tablets with ETOH XS. Presents at  
1100.





# Would you treat these?

1. OD 15x500mg tablets at 12:00.  
Blood at 18:00.  
PPC= 100mg/litre

**TREAT**

2. OD 24x 500mg tablets at  
09:00. Blood at 17:00.  
PPC= 30mg/litre

**DON'T TREAT**

3. OD at 20:00 ?amount. Blood  
at 07:00. PPC= 30mg/litre

**TREAT**

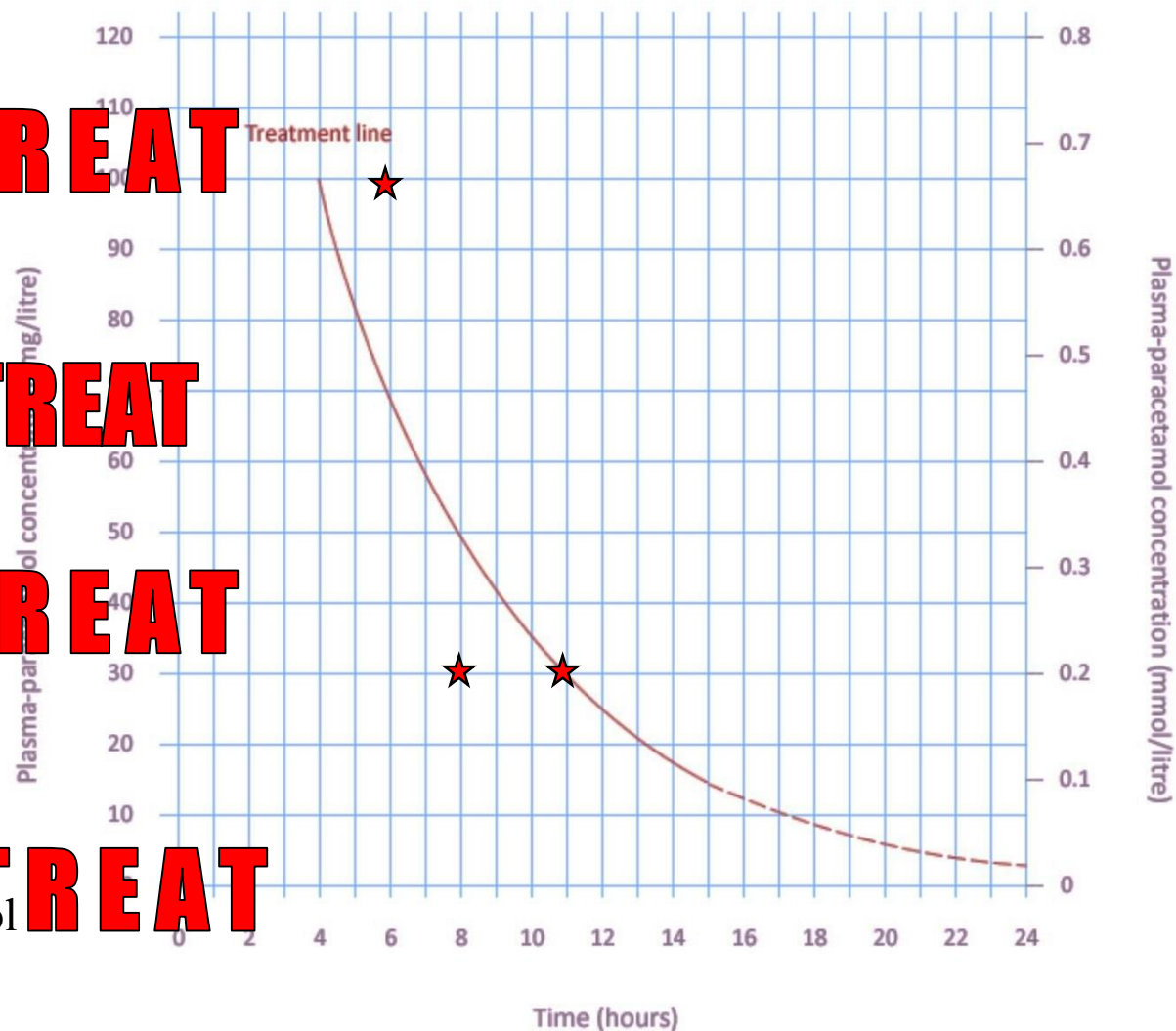
## What would you do ?

4. Staggered OD between 0900 and  
1600. Admits taking 50 Paracetamol  
tablets. Presents at 1630.

**TREAT**

5. OD at 0800. Admits taking 4  
tablets with ETOH XS. Presents at  
1100.

**WAIT!**



# Case

**54yo female, PMH- depression (previous suicide attempts), chronic ETOH abuse, malnourished. Found unconscious with multiple packets of meds.**

**1.Do they need treatment**

**2.Dose of treatment**



## 2. Dose of treatment

Total dose (300 mg/kg in 20 hours)

150 mg/kg in 200ml glucose 5% over first 15 minutes

50 mg/kg over next 4 hours in 500ml

100 mg/kg over next 16 hours in 1 litre

**For a 50kg patient:**

1. What is the total dose of NAC required?

- A. 300 mg
- B. 1500 mg
- C. 15 g
- D. 30 g
- E. 150 g

2. What is the rate (mls/hr) for the first 15 minutes of the regime?

- A. 15 mls/hr
- B. 150 mls/hr
- C. 200 mls/hr
- D. 600 mls/hr
- E. 800 mls/hr



# Liver Transplant

Criteria for possible transplant:

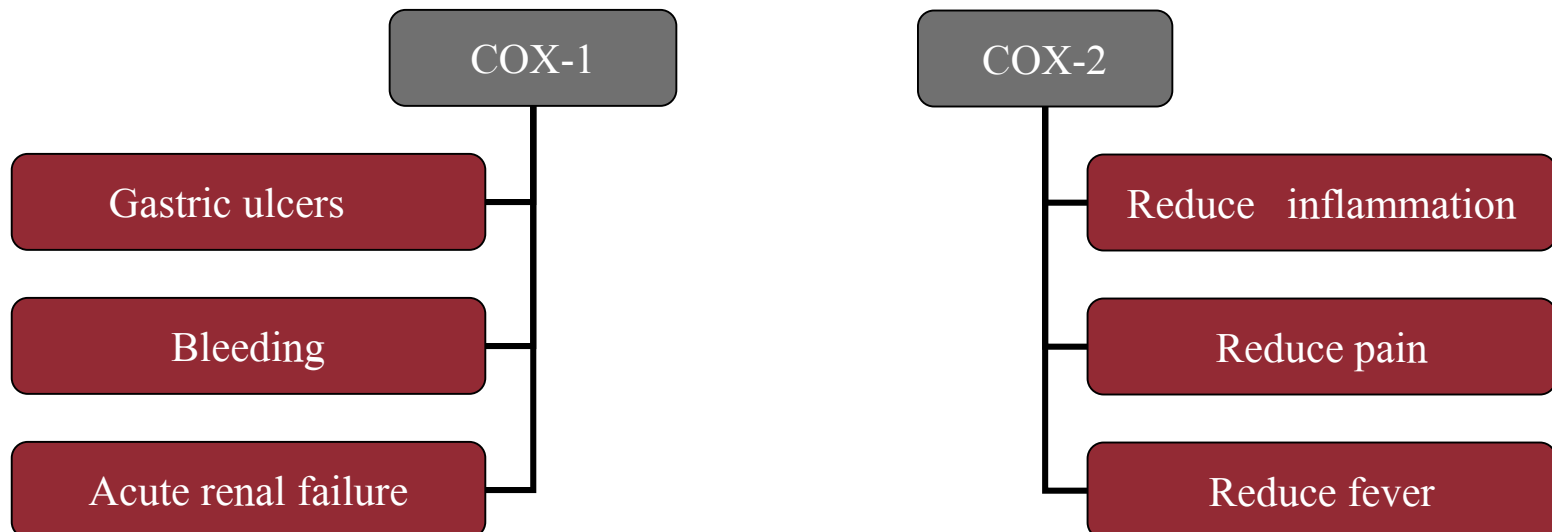
- Metabolic Acidosis - Arterial pH less than 7.30
- Hepatic encephalopathy grade III/IV **and** serum creatinine concentration  $>300 \mu\text{mol/L}$  **and** prothrombin time  $>100$  seconds
- Arterial lactate concentration  $>3.5\text{mmol/L}$  at 4hrs post or  $>3.0\text{mmol/L}$  24 hours post paracetamol ingestion
- Hyperphosphataemia - 48-96hr phosphate  $>3.7\text{ml/dl}$

Discuss with liver transplant unit as soon as the *possible* need is identified



# Non-Steroidal Anti-Inflammatory Drugs

| Drug name   | Dose               | Anti-inflammatory | Side effect risk |
|-------------|--------------------|-------------------|------------------|
| Ibuprofen   | 300-400mg TDS      | +                 | +                |
| Naproxen    | 500mg BD           | ++                | ++               |
| Diclofenac  | 75-150mg total/day | ++                | ++               |
| Indometacin | 50-200mg total/day | +++               | +++              |



# Non-Steroidal Anti-Inflammatory Drugs

| Drug name   | Dose               | Anti-inflammatory | Side effect risk |
|-------------|--------------------|-------------------|------------------|
| Ibuprofen   | 300-400mg TDS      | +                 | +                |
| Naproxen    | 500mg BD           | ++                | ++               |
| Diclofenac  | 75-150mg total/day | ++                | ++               |
| Indometacin | 50-200mg total/day | +++               | +++              |

## Side Effects:

COMMON... especially in elderly

Gastric- indigestion/nausea and gastric erosions (+/- UGI Bleed).

*Co-prescribe proton pump inhibitor if patient also on anti-coagulant*

Bronchospasm- type 1 hypersensitivity reaction. C/I in those with asthma

*Less common; fluid retention, hypertension, acute kidney injury*

# COX-2 inhibitors

Celecoxib / Etoricoxib

- + Reduce GI side effects by 50%
- Expensive

Contraindicated

- CV Disease or cerebrovascular disease
- Relatively with CV risk factors

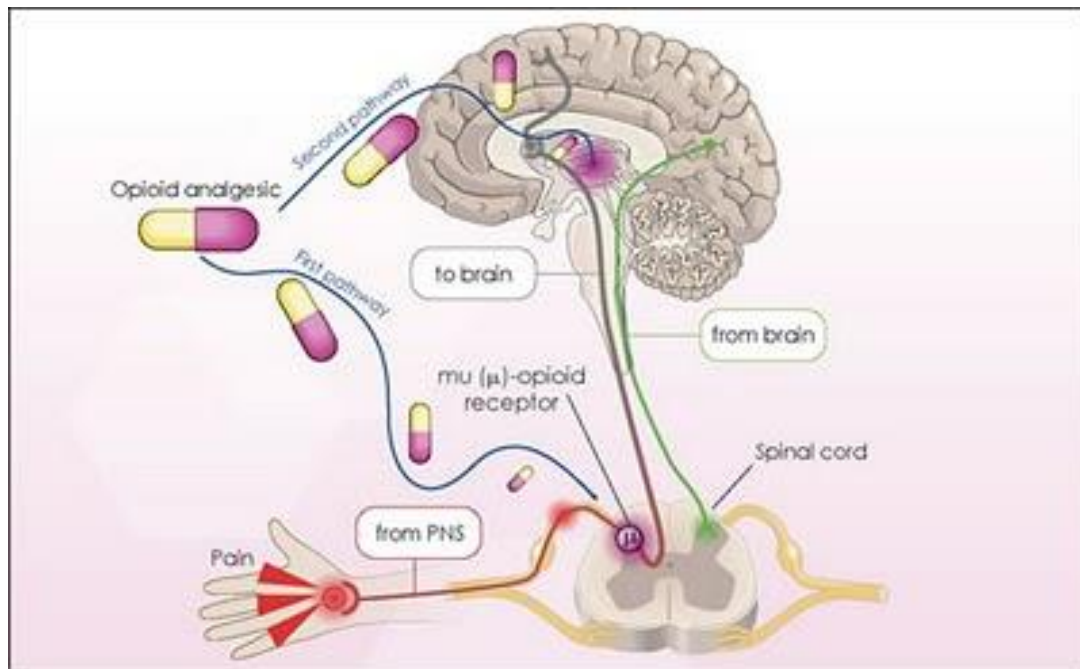
Try NSAIDs first





# Opioids

- Act on opioid receptors within CNS (mainly  $\mu$ -1 & kappa)



# Opioids

## Morphine

SC/IM/PO

2.5 - 20mg, titrate to symptoms/response



### Side effects

*(as non-specific to mu-1, actions on mu-2 receptors)*

Nausea and vomiting → co-prescribe antiemetic (e.g. Metoclopramide)

Constipation → co-prescribe laxative (e.g. lactulose PRN)

Respiratory depression

### Overdose

Accidental / Iatrogenic / Intentional

Reversal with NALOXONE 100mcg-400mcg IV (increased by 100mcg/ 2 mins PRN)

Naloxone half life < Morphine half life (so may need multiple doses)



# Opioids

## Codeine

1/12<sup>th</sup> analgesic effect of morphine

30-60mg 4 hourly (max 240mg/24hrs)

Orally

Analgesia (normally used in combination with above e.g. co-codamol/co-dydramol)

## Tramadol

50-100mg 4 hourly

Orally (rarely IV)

Opioid action + enhancement of the serotonergic and adrenergic pathways

## Fentanyl

Patches 25 / 100 ug/hr

Change every 72 hours



# Patient Controlled Analgesia

**Allows self administration of pre-determined dose of medication e.g. morphine**

**Can determine:**

- Dose (usually start with 1mg morphine)
- Maximum dose (over 24hours)
- Lock-out period (usually 5 mins)



# Palliative Symptom Control

## *Palliative Care;*

Active & total care of incurable disease aim to improve QOL within wide MDT

Symptoms can be caused by disease or treatment

Detailed history and examination required to determine cause/best treatment options

*Main symptom groups:*

- **GI Symptoms**

e.g. nausea/vomiting, anorexia, constipation, bowel obstruction

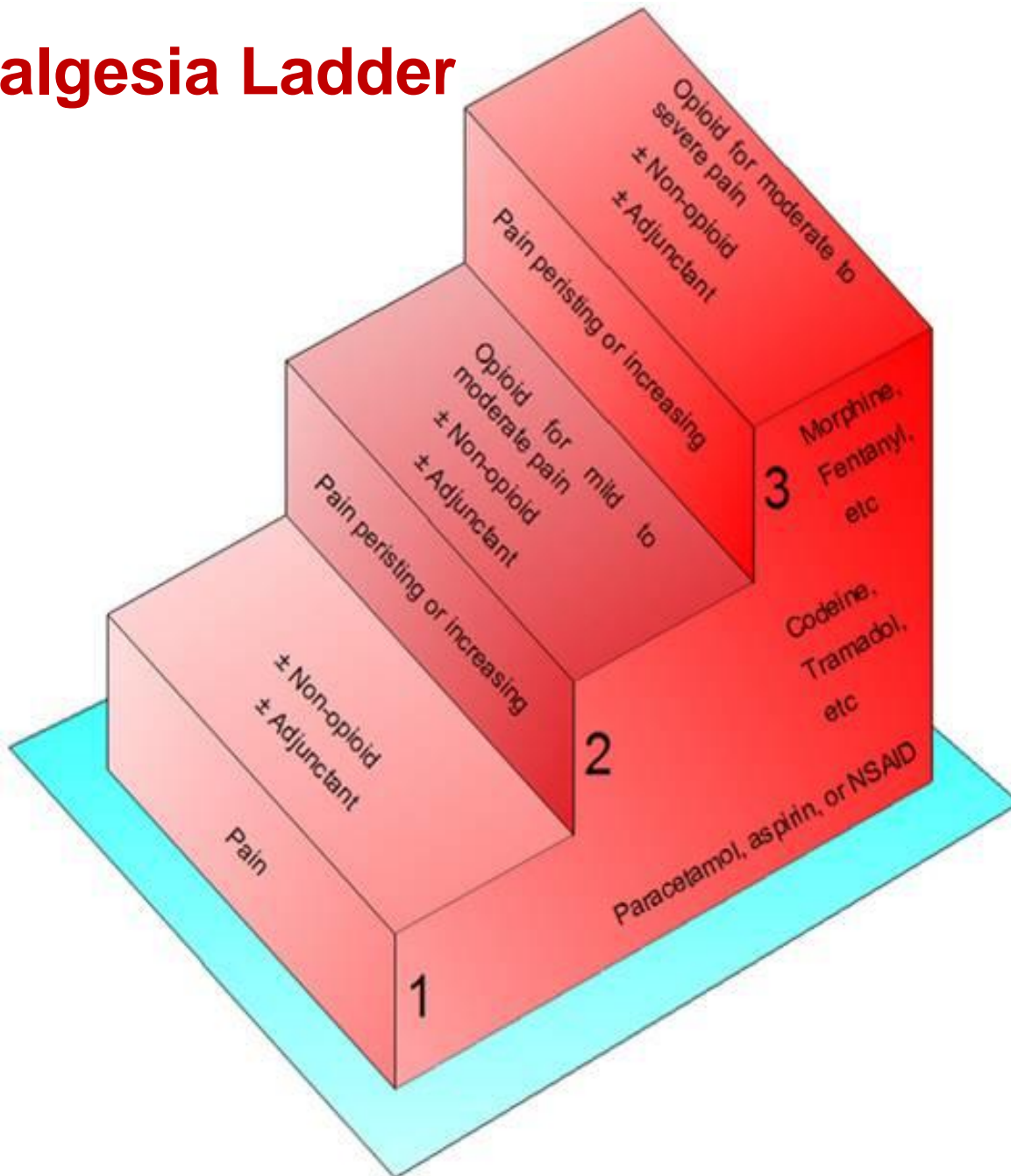
- **Respiratory Symptoms**

e.g. Breathlessness, secretions

- **Pain Symptoms... 70% of cancer patient experience severe pain**



# Analgesia Ladder



# Case...

**Mrs CA is a 80yo has known inoperable metastatic breast cancer. She is complaining of significant pain which has previously been managed with non-opioid analgesia.**

**What dose of morphine would you prescribe?**

**Morphine Sulphate 2.5-10mg every 4 hours**

**With Morphine Sulphate 5-20mg PRN 2-4hrly**





# Case...

Mrs CA's pain is relieved by Morphine Sulphate 10mg but the **relief is not sustained until the next dose 4 hours later.**

**How would you change the regular prescription?**

**Increase dose by 50%**

**So now; Morphine Sulphate 15mg every 4 hours**

**With Morphine Sulphate 5-20mg PRN 2-4hrly**



# Case...

**Mrs CA is eventually controlled over a 24 hour period using morphine sulphate as prescribed below:**

|      |                   |      |      |                   |      |
|------|-------------------|------|------|-------------------|------|
| 0800 | Morphine Sulphate | 15mg | 2100 | Morphine Sulphate | 5mg  |
| 1200 | Morphine Sulphate | 15mg | 2200 | Morphine Sulphate | 10mg |
| 1400 | Morphine Sulphate | 5mg  | 2400 | Morphine Sulphate | 15mg |
| 1600 | Morphine Sulphate | 15mg | 0400 | Morphine Sulphate | 15mg |
| 2000 | Morphine Sulphate | 15mg | 0600 | Morphine Sulphate | 10mg |

**Would you change the prescription?  
If so, how?**



# Case...

To change regular immediate acting morphine prescription into regular prescription

Work out total 24 hour analgesia requirement to control pain...

|      |                   |      |      |                   |      |
|------|-------------------|------|------|-------------------|------|
| 0800 | Morphine Sulphate | 15mg | 2100 | Morphine Sulphate | 5mg  |
| 1200 | Morphine Sulphate | 15mg | 2200 | Morphine Sulphate | 10mg |
| 1400 | Morphine Sulphate | 5mg  | 2400 | Morphine Sulphate | 15mg |
| 1600 | Morphine Sulphate | 15mg | 0400 | Morphine Sulphate | 15mg |
| 2000 | Morphine Sulphate | 15mg | 0600 | Morphine Sulphate | 10mg |

Total = 120mg

## REMEMBER SIDE EFFECTS

1. Change to Controlled Release morphine preparation

Either : Morphine Sulphate **MR** / MST 120mg OD or 60mg BD

2. Plus Breakthrough Pain prescription PRN

**One-sixth** of total dose

Morphine Sulphate 20mg PRN



# Spinal vs. Epidural

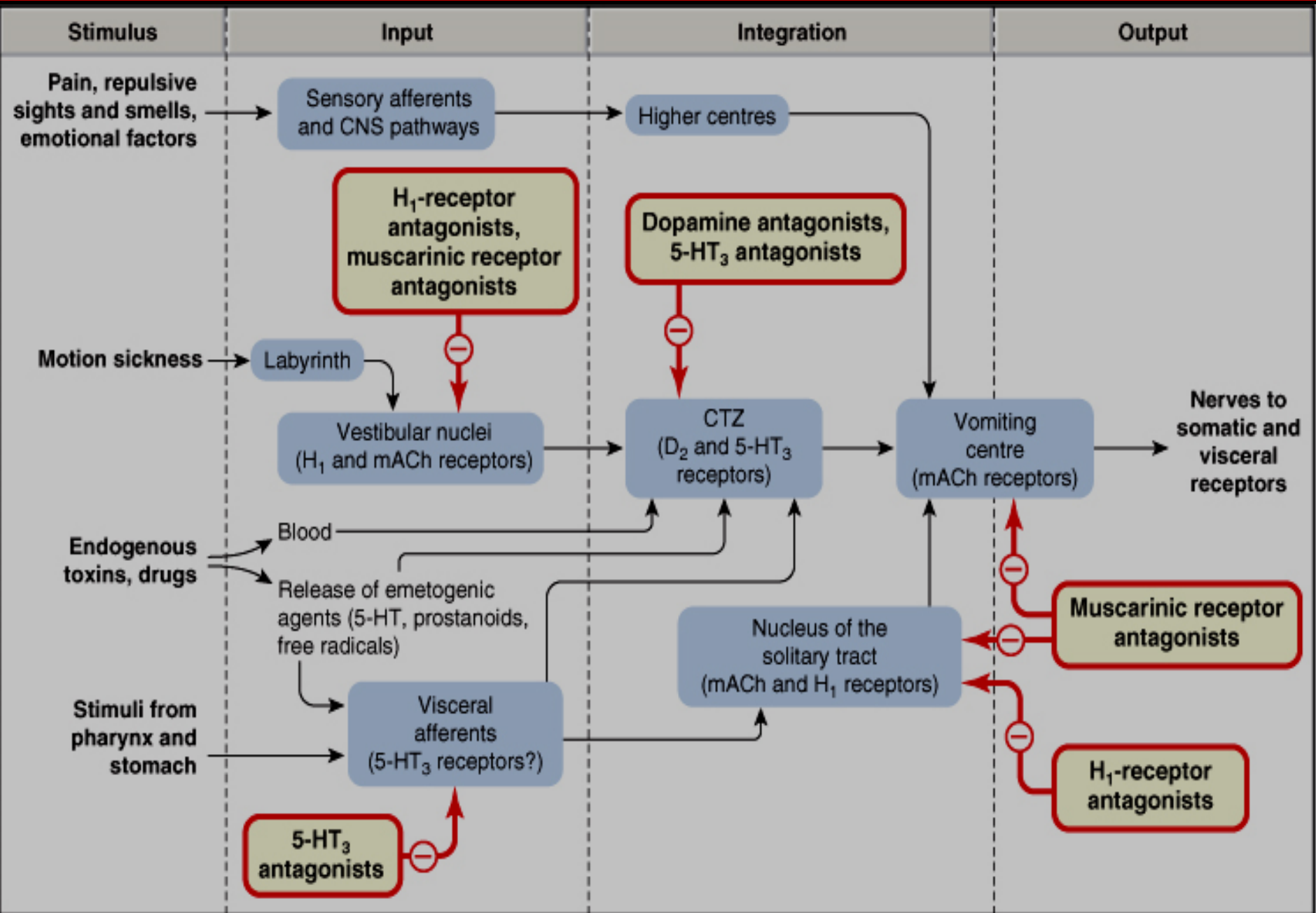


# Anti-emetics

Vomit Centre can be triggered by:

- stimulation of Chemoreceptor Trigger Zone (CTZ) in medulla (opioid, dopamine and serotonin receptors) e.g. by opioids
- Disturbance of Vestibular system (muscarinic, histamine receptors) e.g. motion sickness
- Central Nervous System trigger e.g. smells / emotion
- Peripheral Trigger e.g. GIT containing serotonin receptors





# Anti-emetics

Ondansetron                      *4-8mg BD PO/IM/IV*  
Directed towards CTZ & Peripheral Triggers  
Used in chemo/radiotherapy related vomiting

**5-HT<sub>3</sub> antagonist**

Metoclopramide                *10-20mg TDS PO/SC/IM/IV*  
Does pass blood brain barrier

**Dopamine antagonist**

S/e include extra-pyramidal e.g. dyskinesia

Domperidone                    *10-20mg TDS PO*

**Dopamine antagonist**

Does not pass blood brain barrier (nb. CTZ is outside BBB)

Cyclizine                         *50mg TDS PO/IM/IV*  
For hyperemesis or motion sickness

**Anti-histamine**

Hyoscine hydrobromide        *PO/Patch/IV*  
For motion sickness, no role in chemo related

**Anti-muscarinic**



## Anti-Emetic Receptor Sites

Avoid prescribing anti-emetics that act on the same receptor site, see table below for details

|                              | D <sub>2</sub> -<br>receptor<br>antagonist | H <sub>1</sub> -<br>rec<br>ept<br>or<br>ant<br>ago<br>nist | Muscarinic<br>receptor<br>antagonist | 5HT <sub>3</sub> -<br>receptor<br>antagonist | 5HT <sub>2</sub> -<br>receptor<br>antagonist | 5HT <sub>4</sub> -<br>receptor<br>agonist | Neurokinin -1-<br>Inhibitor |
|------------------------------|--|--|--------------------------------------|--|--|---|-----------------------------|
| Metoclopramide               | ++   | 0  | 0                                    | +  | 0  | ++  | 0                           |
| Domperidone <sup>a</sup>     | ++ <sup>a</sup>                            | 0  | 0                                    | 0  | 0  | 0   | 0                           |
| Haloperidol                  | +++  | 0  | 0                                    | 0  | 0  | 0   | 0                           |
| Prochlorperazine             | ++   | +  | 0                                    | 0  | 0  | 0   | 0                           |
| Cyclizine                    | 0  | ++   | ++                                   | 0  | 0  | 0   | 0                           |
| Hyoscine<br>hydrobromide     | 0  | 0  | +++                                  | 0  | 0  | 0   | 0                           |
| Ondansetron                  | 0  | 0  | 0                                    | +++  | 0  | 0   | 0                           |
| Dexamethasone <sup>b,c</sup> | ?  | ?  | ?                                    | ?  | ?  | ?   | ?                           |
| Levomepromazine              | ++   | +++  | ++                                   | 0  | +++  | 0   | 0                           |
| Aprepitant                   | 0  | 0  | 0                                    | 0  | 0  | 0   | +++                         |

### Pharmacological activity:

0 = none or insignificant

+ = slight

++ = moderate

+++ = marked

a domperidone does not cross the blood-brain barrier so rarely causes extra-pyramidal effects

b mode of action unknown, possibly within the cerebral cortex or other higher centres in the brain

c please half the dose of Dexamethasone when using Aprepitant



# Laxatives

## Osmotic

Non-absorbable salts which increase water retained in large bowel

e.g. **Lactulose, macrogols**

## Bulk-forming

Increase faecal mass which stimulates peristalsis. For those with small hard stools.

e.g. **Dietary fibre, Ispaghula Husk**

## Stimulant

Increase intestinal motility. s/e abdo cramping

e.g. **Senna, docusate sodium**

## Suppositories

**Glycerin**: Absorbs water into colon

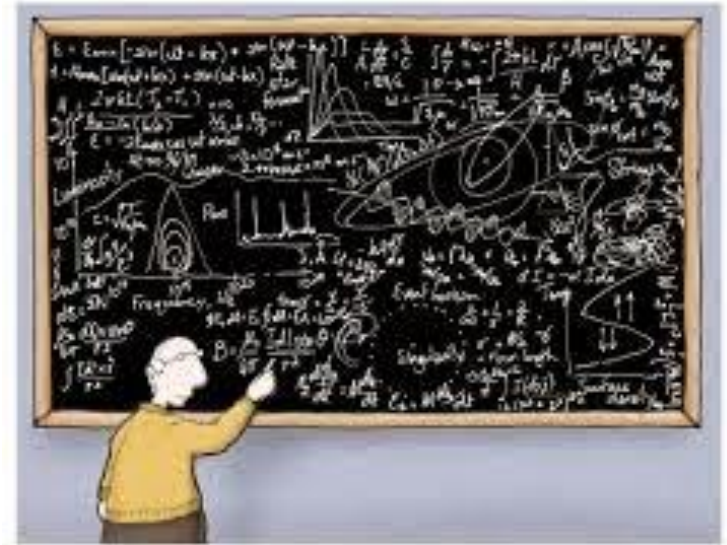
**Bisacodyl**: Induces peristalsis

## Enemas: Hyperosmotic

e.g. **Fleet enema**: Osmotically draws water into colon



# Calculations



# Calculations

1.

**0.01% Adrenaline. How much adrenaline in 1 litre of saline?**

**Same as 1:10,000 (0.01%) adrenaline**

**So 1 unit adrenaline in 10,000 units saline**

**Or 1g Adrenaline in 10,000ml saline**

**So 0.1g in 1 litre**

**(Simple Method: Move one decimal place to the right)**

2.

**What should the minimum urine output for a 80kg patient over 4 hours be?**

**UO > 0.5 ml / kg / hr**

**So at least  $0.5 \times 80 \times 4 = 160\text{ml}$**



# Calculations

3. Prescribe fluid for 14kg child in ml/hr

| Weight Range | Required <u>Daily</u> Fluid              |
|--------------|--|
| 0-10 kg      | 100 mL per kg                            |
| 10-20 kg     | 1,000 mL + 50 mL per each kg above 10 kg |
| 20-70 kg     | 1,500 mL + 20 mL per each kg above 20 kg |
| Over 70 kg   | 2,500 mL                                 |

$$(100 \times 10) + (50 \times 4) = 1200 \text{ mL/day}$$

$$1200 / 24 = \underline{50 \text{ mL/hour}}$$



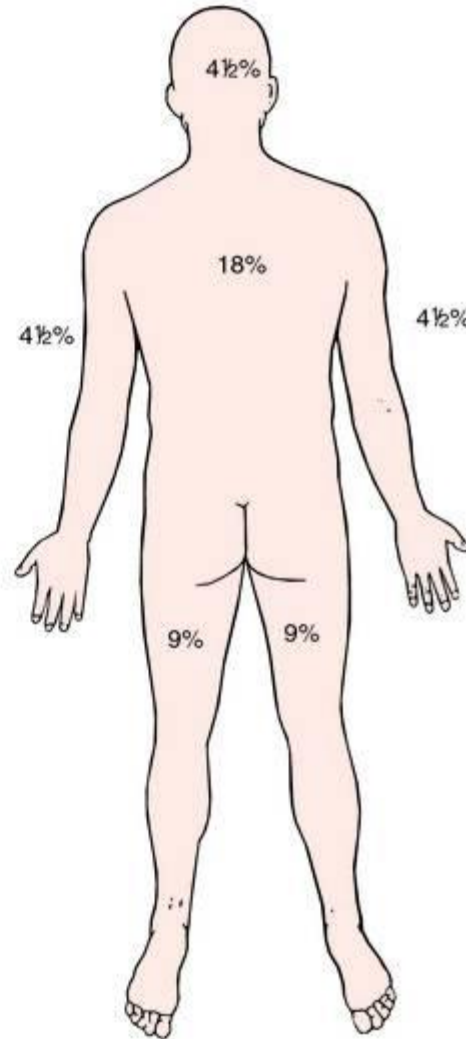
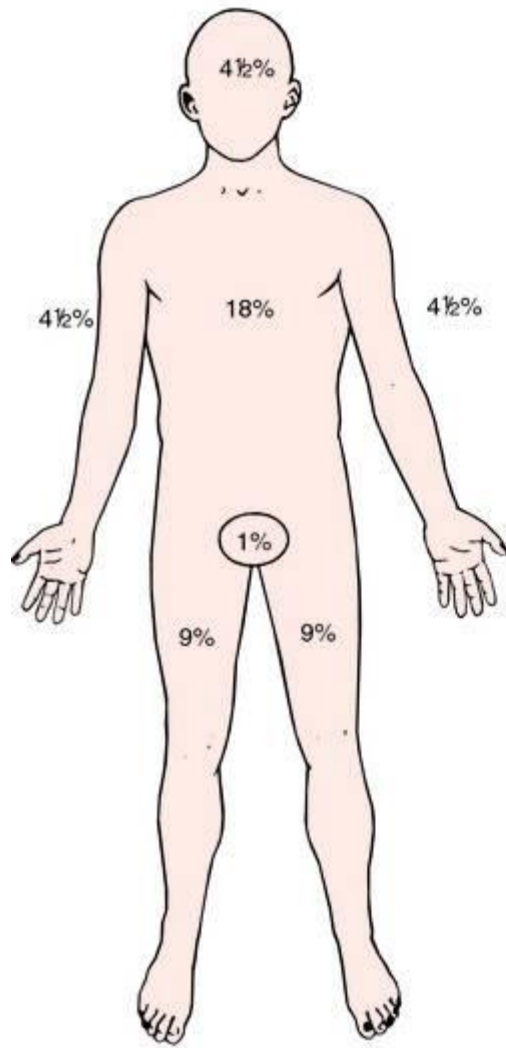
# Calculations

4. Patient presents with burns from fire.  
Effecting both his arms, his face and head.

Approximately what percentage body area has been effected?



# Herndon Rule of 9s



|                 |     |
|-----------------|-----|
| Arm             | 9%  |
| Head            | 9%  |
| Neck            | 1%  |
| Leg             | 18% |
| Anterior trunk  | 18% |
| Posterior trunk | 18% |

Patient:

Arms x2 = 18%

Head / Face = 9%

Total = 27%

# Summary

Take your time and think about the problem

Is your answer sensible?

Practice!

## Any Questions?

