## **Classifying Routes of Administration**

- Where effects are seen
  Systemic
  - > Local
- Access to blood
  - Intra-vascular
  - > Extra-vascular

#### Fundamental Reason: Bioavailability

- Location at which substance is applied
- Target of action

## Route of administration on the basis of exposure of body



## Route of administration on the basis of exposure of body



## **Oral Administration**

- Absorption mainly takes place from the intestine.
- Drugs administered orally that act locally:
  - Vancomycin, Mesalazine and Olsalazine
- Factors Affecting Absorption on Oral Administration:
  - > Physiological
  - Drug related







## **Oral Administration of Drugs**



#### ADVANTAGES

- ➢ Safe
- Convenient
- Economical
- Usually good absorption
- > No need for sterilization

#### DISADVANTAGES

- Cannot be used uncompliant patients
- Irritable and unpalatable drugs
- Local degradation
- Slow absorption and action
- First-pass effect
- Interactions

## **Bioavailability and Bioequivalence**



#### BIOAVAILABILITY

- Indicates the <u>fraction</u> (F) of an orally administered dose that reaches the systemic circulation as intact drug
- > Depends on:
  - > Drug preparation
  - > Host factors
- Bioavailability of the same preparation can vary.



## **Bioavailability and Bioequivalence**



#### BIOEQUIVALENCE

- Bioavailability + rate of absorption = Bioequivalence
- Includes:
  - $\succ$  Maximum concentration achieved  $C_{max}$
  - $\succ$  Time taken from dosing to reach maximum concentration,  $t_{max}$
  - $\blacktriangleright$  Area Under Curve AUC<sub>0-∞</sub>
- Values of each parameters must be between 80% and 125%

## **Rectal Administration of Drugs**

- > Drugs that are administered rectally as a suppository.
- Can serve as local as well as systemic route of administration of drugs.
- Unreliable absorption
- Preferable in patients who are vomiting, post-operative, difficult to establish intravenous access (child having seizures)



## **Rectal Administration of Drugs**

#### ADVANTAGES

#### Used in children

- Little or no first pass effect
- Used in vomiting or unconscious
- Higher concentrations rapidly achieved

DISADVANTAGES

- Inconvenient
- Absorption is slow and erratic
- Irritation or inflammation of rectal mucosa can occur

Sublingual and Buccal Administration of Drugs



- > When rapid response is required
- Drug unstable at gastric pH
- > Drug rapidly metabolised by the liver
- Absorbed drugs directly reach systemic circulation, bypassing portal circulation, hence escapes first pass metabolism.

## Sublingual and Buccal Administration of Drugs



#### ADVANTAGES

- Drug absorption is quick
- Quick termination
- First-pass avoided
- Can be self administered
- Economical

#### DISADVANTAGES

- Unpalatable & bitter drugs
- Irritation of oral mucosa
- Large quantities not given
- Few drugs are absorbed

#### Parenteral Route of Drug Administration

A) Intradermal

B) Subcutaneous (SC)

C) Intramuscular (IM)

D) Intravascular (IV)



# Intravenous Administration of Drugs



- Fastest route of administration of drugs
- Peak concentration reaching tissues depends on rate of administration
- Bolus dosing
- Infusion dosing





## Intravenous Administration of Drugs

#### ADVANTAGES

#### DISADVANTAGES

Most common route for drugs not absorbed orally.

Avoidsfirst-passmetabolism;absorptionbypassed

Permits a rapid effect and a maximal degree of control over the circulating levels of the drug. Titration of dose with response.

Large quantities can be given

Cannot be easily removed

May induce hemolysis or cause other adverse reactions by the too-rapid delivery of high concentrations of drug

Thrombophlebitis of vein and necrosis of adjoining tissue if extravasation occurs Intramuscular and Subcutaenous Administration of Drugs

- Large skeletal muscle are used for intra-muscular administration
- Subcutaenous Administration: drug is deposited in loose subcutaneous tissue
- Faster absorption than oral administration
- Rate limiting factors:
  - > Diffusion through the tissue
  - Removal by local blood flow



## Intramuscular Administration of Drugs

#### ADVANTAGES

- Absorption reasonably uniform
- Rapid onset of action
- Mild irritants can be given
- First pass avoided, Gastric factors avoided

#### DISADVANTAGES

- Only up to 10ml drug given
- Local pain and abscess, infection
- Expensive
- Nerve damage
- Local hematoma can occur in anticoagulant treated pt.

## Subcutaenous Administration of Drugs

#### ADVANTAGES

DISADVANTAGES

- Less blood supply: slow absorption
- Depot preparation can be used
- Risk associated with intravascular injection avoided
- Should be avoided in shock patient
- Only small volume can be injected

## Transdermal Route of Drug Administration



- Achieves systemic effects by application of drugs to the skin
- Rate of absorption determined by drug factors and site of application
- Slow effect (prolonged drug action)
- First pass effect avoided
- Absorption- increase by oily base, occlusive dressing, rubbing preparation

## Local Route of Drug Administration

Routes of Drug Administration for MBBS 1<sup>st</sup> Year

# Local Application: Application to Epithelial Surface

#### **Cutaenous Adminstration:**

Usually absorption poor over intact skin.

Advantages:

Steady rate of drug delivery

Pre Systemic metabolism avoided

Disadvantages:

Only applicable for Lipid soluble drugs Relatively Expensive





# Local Application: Application to Epithelial Surface

- > Eye/Ear Drops:
- Eye Drops: Absorption through conjunctival sac epithelium
- Ear Drops: Absorption through epithelial lining of External ear
- Relatively brief contact time with absorbing surface
- Sterile; require aseptic handling
- Desirable for local effects: lacks systemic reactions.



### **Administration By Inhalation**

#### Systemic Administration of drugs

- Large surface area and large blood flow: rapid exchange of drugs, possible to adjust plasma levels rapidly
- > Nasal Sprays
- Local Administration of drugs

## Local Application: Application to Epithelial Surface







## Intrathecal and Intravitreal Administration of Drugs

## Intrathecal administration

- Injection into subarachnoid space
- Minimises systemic adverse effects





## Intravitreal administration

- Administered byOphthalmologist
- Ranibizumab for wet age-related macular degeneration.



## **Topical Therapy: Pros and Cons**

#### **Advantages**

Disadvantages

Delivery onto the target organ at an optimal concentration

Time consuming

Rapid onset of action

Depends largely on patient's compliance and patience

Systemic effects are less

More expensive

### **Other Topical Preparations**

Medicated plaster, dressings, strips Suppository Douche

Medicated Vaginal Rings Intra-uterine devices Intra-nasal drug delivery Inhaled Drug Delivery

## **Dosage forms for Various Topical Routes**

- Mouth and pharynx: Paints, lozenges, mouth washes, gargles.
- Anal canal: As ointment, suppositories.
- Gastrointestinal tract: As non-absorbable drugs given orally e.g. aluminium hydroxide, kaolin, neomycin.
- Bronchi and lungs: As inhalations, aerosols (nebulised solution or fine powder)- e.g. salbutamol, cromolyn sodium

## **Dosage forms for Various Topical Routes**

- Eyes, ears and nose: As drops, ointments, irrigation, nasal spray.
- Urethra: As jellies e.g. lidocaine, irrigating solutions.
- Vagina: As pessaries, vaginal tablets, inserts, cream, powders, douches

## That will be all for today!

> Any Queries?

➤ Thank you!