

## 1 D.1 Pharmaceutical products and drug action

### 1.1 Type 1: $LD_{50}$ , $TD_{50}$ , $ED_{50}$ , Therapeutic Index, and Therapeutic Window

Could be asked as Distinguish LD and TD; Calculate TI; Define LD, TD, and ED

#### 1.1.1 Type 1.1: Define or Distinguish LD and TD

Lethal Dose 50: amount/dose that kills 50 percent of the population

Toxic Dose 50: amount/dose that negatively affects/produces toxic effects in 50 percent of the population

#### 1.1.2 Type 1.2: Distinguish TI for animals and humans

Therapeutic Index for Animal:  $\frac{LD_{50}}{ED_{50}}$

Therapeutic Index for Humans:  $\frac{TD_{50}}{ED_{50}}$

#### 1.1.3 Type 1.3: Define or Outline the significance of Therapeutic Window

Use definition of Therapeutic Window: range of doses that produce a therapeutic effect without causing toxic effects

Wide TW: Small ED and Large LD/TD

Narrow TW requires small doses because **small LD/TD** and easily reach toxic level

### 1.2 Type 2: Drug Administration

#### 1.2.1 Type 2.1: General Types of Drug Administration

- Oral
- Inhalation
- Topical
- Rectal
- Injection (Parenteral)

#### 1.2.2 Type 2.2: Types of Injection

- Intravenous
- Intramuscular
- Subcutaneous

#### 1.2.3 Type 2.3: Discuss Oral-taken Drug Administration

Advantage: Convenient to take

Disadvantage: Stomach acid reacts with drugs; Slow Effects...

### 1.3 Type 3: Bioavailability

#### 1.3.1 Type 3.1: Define Bioavailability

**fraction** of administrated dosage that reaches **target part** of the body

#### 1.3.2 Type 3.2: The method to increase/maximize bioavailability

Intravenous Injection: 100 percent

### 1.4 Type 4: Side Effects, Tolerance, and Ethical Issues

#### 1.4.1 Type 4.1: Define Side Effects

an effect produced in addition to the one intended effect (unintended effect)

#### 1.4.2 Type 4.2: Define Tolerance

person needs to take ever larger quantities of a drug to gain the original effect

#### 1.4.3 Type 4.3: Ethical Consideration

Usually choose **2 of them** to recite

- side-effects of medication on patient
- effects on environment
- potential for abuse
- drugs may be developed that are contrary to some religious doctrines
- animal testing

### 1.5 Type 5: Development of Drugs and Effects of Medicine

#### 1.5.1 Type 5.1: Stages of Development of Drugs

1. drug is isolated from existing species
2. tested on animals to establish LD<sub>50</sub>
3. tested on humans and half is given a **placebo**

#### 1.5.2 Type 5.2: General Effects of Medicine on Body

- alter physiological state
- alter emotions or mood
- alter incoming sensory sensations

## 2 D.2 Aspirin and Penicillin

### 2.1 Type 1: Characteristics of Aspirin and Mild Analgesics

#### 2.1.1 Type 1.1: Characteristics of Aspirin

Why Could not be Stored in **hot, humid** location?

- Humid → Water, Aspirin **react with water** (hydrolysis)
- Hot → Heat, Increase **rate of reaction** with water

How to identify aspirin? (memorize 2 ways)

- melting point
- mass spectrometry
- NMR

#### 2.1.2 Type 1.2: Mild Analgesics

How mild analgesics function?

- Prevent the production of **prostaglandins**
- at the site of injury

How prostaglandins function?

Prostaglandins are involved in the **transmission of pain impulses**

### 2.2 Type 2: Structure, synthesis, and conversion of Aspirin

#### 2.2.1 Type 2.1: Synthesis of Aspirin

Aspirin could be synthesized from **salicylic acid** and **acetic anhydride**

Type of reaction: esterification

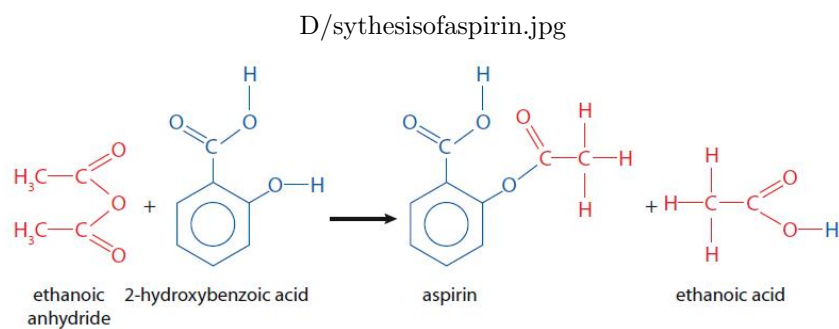


Figure 1: Synthesis of aspirin

### 2.2.2 Type 2.2: Structure of Aspirin

Partially Soluble because:

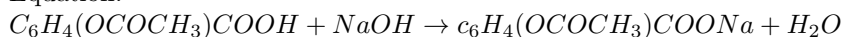
- Carboxyl: OH bond makes it soluble
- Ester: Non-polar makes it unsoluble

### 2.2.3 Type 2.3: Make Aspirin more Soluble

Could be asked with: ion salt; more soluble in water; **How to increase bioavailability** etc.

Key point: react with **NaOH**

Equation:



## 2.3 Type 3: Appropriate Use of Aspirin

### 2.3.1 Type 3.1: Use of Aspirin

Could be asked with: state the **other use** of aspirin (memorize 2 points)

- lower risk of heart attack
- prevent recurrence of heart attack
- prevent cancer of stomach/colon

### 2.3.2 Type 3.2: Misuse of Aspirin with Alcohol

Key point: It will lead to **stomach bleeding**

## 2.4 Type 4: Functions and Structures of Penicillin

### 2.4.1 Type 4.1: Penicillin work in general and as Anti-bacterial

Could be asked with: How penicillin work; how penicillin combat infection etc.

How Penicillin combat infection?

- Inhibits **enzyme** that produces cell walls
- Cells cannot reproduce

How Penicillin work as anti-bacterial?

- prevent the formation of bacterial cell walls
- causes water to enter bacterial cell and cell bursts

### 2.4.2 Type 4.2: The Beta-Lactam Ring on Penicillin

New concept: "ring strain", means bond angle smaller

Why Beta-lactam ring important

1. ring is **strained**
2. ring breaks up easily
3. binds to enzyme responsible for **cell wall formation**

Functional Group present in Beta-lactam ring: **amide**

### 2.4.3 Type 4.3 How to modify Penicillin

Could be asked with: Why necessary; How to;

How to modify?

Modify the **side chain**

Why necessary?

Modifying overcomes the resistance of bacteria

## 2.5 Type 5: Development and limitations of Penicillin

### 2.5.1 Type 5.1: Development of Penicillin

Could be asked with: Why develop?

- to overcome the resistance that bacteria develop to existing antibiotics
- prevents penicillinase enzyme from destroying penicillin

### 2.5.2 Type 5.2: Limitations of Penicillin

Could be asked with: Overprescription of penicillin

- Might cause allergic reactions
- Might wipe out beneficial bacteria
- increase the proportion of resistant bacteria

## 3 D.3 Opiates

### 3.1 Type 1: Morphine and strong analgesics

#### 3.1.1 Type 1.1 Strong Analgesics

Temporarily bind to the **receptor sites** in brain

### 3.1.2 Type 1.2: Opiates as painkillers in general

1. work directly on pain receptors
2. suppress pain impulses
3. resemble endorphins

## 3.2 Type 2: Diamorphine from Morphine

### 3.2.1 Type 2.1: Reaction from Morphine to Diamorphine

Reactant:  $CH_3COOH$

By-product:  $H_2O$

### 3.2.2 Type 2.2: Comparison on the Structure between Morphine, Diamorphine, and Codeine

Similarities:

- Benzene Ring
- Amino
- Ether

Difference:

- Two OH in morphine, one in codeine, and none in diamorphine
- Ester only in diamorphine

### 3.2.3 Type 2.3: Comparison on the effects between Morphine and Diamorphine

Could be asked with: Potent/potency; Advantage/disadvantage etc.

Why Diamorphine is more potent than morphine?

1. Diamorphine is **non-polar**
2. Can cross the **blood-brain** barrier more easily

Advantage and Disadvantage of Diamorphine

Advantage: Strong pain killer

Disadvantage: More addictive

## 3.3 Type 3: Codeine from Morphine

### 3.3.1 Type 3.1: Reaction from Morphine to Codeine

React with **methyl iodide** ( $CH_3I$ ) in **alkaline solution**

Type of reaction: Nucleophilic Substitution

### 3.3.2 Type 3.2: Comparison on the effects between Morphine and Codeine

Could be asked with: Why widely used; why without prescription etc. Codeine has a wider **therapeutic window**

### 3.4 Type 4: Addiction, Use of Morphine

#### 3.4.1 Type 4.1: Addiction of Morphine

Why opiates addictive?

- interact with opioid receptors in the brain
- withdrawal symptoms
- cause euphoria

#### 3.4.2 Type 4.2: Advantage and Disadvantage of Using Morphine

Advantage:

- Strong pain relief
- Relieve coughing

Disadvantage:

- addiction
- tolerance
- dependence

## 4 D.4 pH Regulation of the Stomach

### 4.1 Type 1: Action of Omeprazole

General Function: **Increase** the stomach pH

#### 4.1.1 Type 1.1: How does omeprazole regulates stomach pH

1. Bind to receptors of **proton pump**
2. Inhibits the secretion of stomach acid

Site of Action: Proton Pump

### 4.2 Type 2: Action of Ranitidine

General function: Increase the stomach pH

#### 4.2.1 Type 2.1: How does ranitidine reduces stomach acid production?

1. Binds to H<sub>2</sub> receptor in cells of stomach lining
2. Prevent **parental cells** from release acid

Notice that: Originally **histamine** binds to the H<sub>2</sub> receptor

### 4.3 Type 3: Buffer Solution and pKa Calculation

#### 4.3.1 Type 3.1: Calculation of Buffer Solution

$$\text{pH} = \text{pKa} + \log \frac{[\text{base}]}{[\text{acid}]}$$

### 4.4 Type 4: Antacids, neutralization, and Change in pH of Stomach

#### 4.4.1 Type 4.1: Acids present in gastric juice

Hydrochloric Acid (HCl)

#### 4.4.2 Type 4.2: Common Antacids

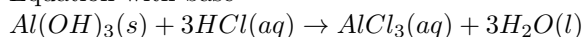
In form of **tablets** (solid)

- 1 mole:  $\text{NaHCO}_3$
- 2 mole:  $\text{CaCO}_3$  and  $\text{MgCO}_3$
- 3 mole:  $\text{Al}(\text{OH})_3$

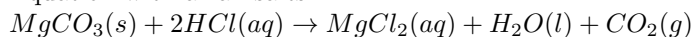
#### 4.4.3 Type 4.3: Neutralization

All about the equations.

Equation with base



Equation with alkali salts



The  $\text{CO}_2$  produced could lead to **belching**

#### 4.4.4 Type 4.4: Change in pH of Blood

Could be asked with: effect of a large amount of aspirin

1.  $\text{H}^+$  from Aspirin reacts with  $\text{HCO}_3^-$  to form  $\text{CO}_2$  and  $\text{H}_2\text{O}$
2. Causing the pH to decrease

## 5 D.5 Anti-viral Medicine

### 5.1 Type 1: Difference between Bacteria and Virus

Could be phrased as: State the difference; distinguish; Outline the difference, etc.

Answer:

- Bacteria performs living functions while Viruses is non-living.
- Bacteria have cell walls while viruses don't.
- Bacteria have capsid while viruses don't
- Bacteria are larger than viruses
- Bacteria are single cells while viruses are not cellular



## 5.2 Type 2: How Antiviral Drugs Work

Could be phrased as: x ways in which antiviral drugs work

Answer:

- alter cell's genetic material
- block enzyme activity within host cell
- inhibits virus entry/bonding to cell
- prevents virus from leaving cell
- becomes part of DNA of virus
- prevents virus from using cell to replicate

### 5.2.1 Type 2.1 How oseltamivir/zanamivir works

Answer:

- inhibits viral enzyme
- prevent virus from leaving host cells

### 5.2.2 Type 2.2: Reason for Zanamivir taken by inhalation

Answer:

Oral bioavailability is low for zanamivir

## 5.3 Type 3: Structures of Oseltimivir and Zanamivir

### 5.3.1 The functional groups they both have

- ether
- carbonyl
- amido

### 5.3.2 Functional groups: Zanamivir Only

- Carboxyl
- Hydroxyl

## 5.4 Type 4: Why difficult to treat AIDS/HIV

- HIV retrovirus attacks immune system
- virus has ability to mutate
- virus makes people vulnerable to other infections
- Metabolism of virus is linked closely to metabolism of the (host) cell
- antiretroviral agents are expensive
- Stigma of diagnosis leads to not getting treatment

## 6 D.6: Environmental impact of some medications

### 6.1 Type 1: Environmental Impact of Nuclear Waste

Explain the low environmental impact of most medical nuclear waste.

- Low radioactivity
- Short Lives

Examples and Treatment of Nuclear Wastes

- Low Level Waste: Storage until the isotope has decayed
- Medium Level Waste: Storage underground

Ethical Implications of Nuclear Waste

- Security concerns if nuclear radioactive material ended up with terrorists
- Proper disposal of nuclear wastes
- Exposure of workers to radioactivity

### 6.2 Type 2: Organic Solvents and Green Chemistry

#### 6.2.1 Type 2.1: Hazardous Solvents

- Benzene
- Methanol
- dichloromethane (chlorinated solvent)

#### 6.2.2 Type 2.2: Green Solvent

- Water
- Carbon dioxide
- Ethanol (when replacing a hazardous solvent)
- propanone (acetone) when replacing a hazardous solvent

#### 6.2.3 Type 2.3: Problem of Chlorinated Solvent and Green chemistry

Problem of Chlorinated Solvent

- incomplete combustion can produce toxic products
- ozone depletion
- contribute to formation of smog
- costs of disposal

Green Chemistry

- use organic solvent-free synthetic methods
- use a water as a solvent
- based on atom economy

### **6.3 Type 3: Antibiotics**

#### **6.3.1 Type 3.1: Consequences of Prescribing Antibiotics Unnecessarily**

- Bacterial Resistance
- Destroy beneficial bacteria
- Damage to ecosystems

#### **6.3.2 Type 3.2: Overuse of Antibiotics**

Same as 3.1.