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# UNIVERSITY OF RUHUNA

## **FACULTY OF MEDICINE**

ALLIED HEALTH SCIENCES DEGREE PROGRAMME FIRST BPHARM PART II EXAMINATION – AUGUST 2014 PH 1213: PHARMACEUTICAL CHEMISTRY II

TIME: 03 HOURS

#### INSTRUCTIONS

- Answer all six (06) questions.
- Do not use any correction fluid.
- Use illustrations where necessary.

## 01. Answer all parts

131 Draw the structural formulae for the possible products of the following reactions.

[30 marks]

- 1.2 Giving necessary **reducing** reagents and reaction conditions, draw the possible structures for the products.
  - 1.2.1 COOH

1.2.3 CH<sub>3</sub>-CH<sub>2</sub>-CHO

1.2.4 
$$C_6H_5$$
  $C_7 = C_2H_9$ 

[40 marks]

1.3 Give a suitable mechanism for the following Claisen-Schimdt reaction.

[30 marks]

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02. Answer all parts

2.1

2.1.1 Of the following molecules /ions which are more stable?



[10 marks]

2.1.2 Giving necessary structures explain your answer above in 2.1.1.

[15 marks]

2.2 Consider the two reactions given below:

2.2.1 Of the above two reactions, which one is faster?

[08 marks]

2.2.2 Draw the structure of the product(s) formed in the fast reaction according to your answer above.

[12 marks]

2.2.3 Giving appropriate mechanism, explain why the above mentioned reaction is faster.

[10 marks]

2.3 Giving necessary structures, explain the formation of the two products in the reaction below:

$$H_2O$$
  $H_2O$   $HO$   $+$   $OH$ 

[25 marks]

2.4

2.4.1 What it meant by resonance energy of benzene?

[08 marks] •

2.4.2 Using approprate structures, explain why cycloheptatrienyl cation is more stable than cycloheptatriene.

[12 marks]

03. Answer all parts

3.1

3.1.1 Explain briefly why aromatic compounds generally undergo electrophilic substitution reactions.

[10 marks]

3.1.2 Write down the complete mechanism for the following conversion.

[15 marks]

3.2 Consider the following reaction of nitration of bromobenzene:

Using electronic effects explain why:

- 3.2.1 bromobenzene needs strong conditions for the nitration.
- 3.2.2 only 2-nitrobromobenzene and 4-nitrobromobenzene are formed.

[25 marks]

3.3 Draw the structure of the product(s) formed in the following reactions:

[20 marks]

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3.4 Indicating reagents, reactions conditions and intermediates, show how you would carry out

[30 marks]

- 04. Answer all parts
  - 4.1 Consider the reaction scheme below:

- 4.1.1 Give the reactant and reagent needed in step 1.
- 4.1.2 Show the mechanism for step 2.
- 4.1.3 In step 2 an alternative product to B may be formed in theory. Draw this product and

[30 marks]

4.2 In the Fischer indole synthesis acetone and phenylhydrazine (PhNHNH2) react to give 2-methylindole (A). Using this example predict the structures of the products B, and D.

4.2.1 
$$\longrightarrow$$
 NHNH<sub>2</sub> + H<sub>3</sub>C  $\longrightarrow$  CH<sub>3</sub>  $\longrightarrow$  CH<sub>3</sub> A

4.2.2  $\longrightarrow$  NHNH<sub>2</sub> +  $\bigcirc$  C<sub>12</sub>H<sub>12</sub>BrN

4.2.3  $\bigcirc$  O<sub>2</sub>N  $\longrightarrow$  NHNH<sub>2</sub> +  $\bigcirc$  C<sub>11</sub>H<sub>10</sub>N<sub>2</sub>O<sub>4</sub>

[20 marks]

4.3 Given that in the Doebner-von Miller quinoline synthesis *trans*-but-2-enal and aniline react to give 2-methylquinoline (E), predict the products **F** and **G**.

4.3.1 
$$\downarrow_{NH_2}$$
  $\downarrow_{CH_3}$   $\downarrow_{CH_3}$ 

[20 marks]

4.4 Give the structure of the unknown products B, C, and D of the following reactions:

4.4.2 
$$NH_3$$
 C  $C_{10}H_{15}N$ 

4.4.3 Ph 
$$\frac{\text{PhCH}_2\text{NH}_2}{\text{H}^+ \text{ cat.}}$$
 D  $\frac{\text{C}_{26}\text{H}_{31}\text{NO}_2}{\text{C}_{26}\text{H}_{31}\text{NO}_2}$ 

[30 marks]

- 05. Answer all parts
  - 5.1
    - 5.1.1 Giving structures, explain the three and erythro forms of sugars.

[04marks]

5.1.2 What is meant by mutarotation pertaining to sugars?

[04marks]

- 5.1.3 Write down a chemical reaction which can be used to protect the anomeric OH of glucose.

  [04marks]
- 5.1.4 Sorbitol has a high commercial value in food industry as well as in manufacturing of toothpaste. Give a chemical method to prepare sorbitol from glucose.

[04marks]

5.1.5 "Although cellulose belongs to the family of carbohydrates, it is a very hard material present in plants". Explain this statement.

[04marks]

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5.2

5.2.1 Name two  $\alpha$ -amino acids having more than one amino group.

[02marks]

5.2.2 Give their three-letter and one-letter codes.

[04marks]

5.2.3 Draw the chemical structures of the above two amino acids.

[06marks]

5.2.4 Explain briefly the amphoteric nature of amino acids.

[04marks]

5.2.5 Define the term isoelectric point of proteins.

[04marks]

- 5.3 The octapeptide **P** with the amino acid sequence of Ala-Lys-Phe-Glu-Gly-Tyr-Arg-Leu was subjected to the following treatments. Derive the product(s) obtained after each treatment with:
  - 5.3.1 2,4 DNP followed by acid hydrolysis;
  - 5.3.2 chymotrypsin;
  - 5.3.3 carboxypeptidase;
  - 5.3.4 trypsin;

[20 marks]

5.4 Draw the structures of the organic products formed when dAMP is hydrolyzed with acid.

[10 marks]

5.5 The diagram below represents part of a DNA strand

- 5.5.1 Identify the atoms represented by **A** and **B**
- 5.5.2 Indicate the anomeric carbon atom(s)
- 5.5.3 Indicate the N-glycosidic and the phophodiester bonds
- 5.5.4 Number the both purine ring and the sugar unit
- 5.5.5 Draw structure and name the base that is complementary to the base shown.

[30 marks]

### 06. Answer all parts

- 6.1 Explain the following observations:
  - 6.1.1 p-Nitrophenol (pKa=7.15) is a stronger acid than m-nitrophenol (pKa=8.28).
  - 6.1.2 cyclopetadiene (pKa=15) is more acidic than cyclopentene (pKa=40).
  - 6.1.3 Pyridine (pKa=5.2) is a weaker base than piperidine (pKa=11.2).

[15 marks]

- 6.2 Complete each of the following acid-base reactions and indicate whether the equilibrium is favored to the right or left.
  - 6.2.1 CH<sub>3</sub>SH + NaOH
  - 6.2.2  $CH_3CH_2ONa + CH \equiv CH =$
  - 6.2.3 CH<sub>3</sub>OH + KOtBu
  - 6.2.4 HF + NH<sub>3</sub>

[15 marks]

6.3

6.3.1 Explain why imidazole is a stronger base than pyrazole



pyrazole



6.3.2 The two nitrogen of atoms of imidazole are quite different. Which nitrogen is protonated in acidic medium?

[20 marks]

6.4 Identify physical and chemical factors affecting the biodegradation of polymers.

[10 marks]

6.5 Identify the primary mechanisms, by which active agents can be released from a polymer drug delivery system.

[10 marks]

6.6 Draw a rough sketch to show in a macroscopic scale the release of the active agent in the polymer matrix.

[10 marks]

6.7 Show the typical variation of (i) melting point(s) and (ii) solubility in water with the chain length of fatty acids in a plot.

[05 marks]

6.8 Show a typical variation of melting points with the number of double bonds present in fatty acids in a typical plot.

[05 marks]

6.9 Briefly discuss the significance of omega fatty acids in diets.

[10 marks]