MANIPAL UNIVERSITY

Reg. No.

M. PHARM. PART-I DEGREE EXAMINATION - MAY/JUNE 2013

SUBJECT: MEDICINAL CHEMISTRY – I (PCH 601) (SPECIALIZATION: PHARMACEUTICAL CHEMISTRY)

Friday, May 24, 2013

Time: 10:00 - 13:00 Hrs.

Max. Marks: 100

- 1A. Explain the generation of FBDD libraries. Write any four advantages of FBDD over HTS. Give the usefulness of racemic mixture with suitable examples.
- 1B. Mention different types of functionalization reactions. Explain reductive and hydrolytic biotransformation.

(10+10 = 20 marks)

- 2A. How prodrugs are used for drug targeting? Explain with justification.
- 2B. Enlist the types of HTS assay and discuss any one of them.
- 2C. Giving suitable example, explain the binding role of carboxylic group and carbon skeleton .

(5+5+10 = 20 marks)

- 3A. Explain simplification of structure and ring fusion as drug optimizing strategies.
- 3B. Prove the structure of cardiac glycosides.
- 3C. Discuss lead discovery strategy in detail.

(10+5+5 = 20 marks)

- 4A. Mention desirable properties of radio ligands and give their limitations.
- 4B. Mention the application of fluorogenic substrate to configure peptidase screening.
- 4C. Explain the role of solubility in drug design and give BCS classification of drugs.

(5+5+10 = 20 marks)

- 5A. Discuss the mechanisms of drug interaction.
- 5B. How do you use microbes as source of drugs? Explain.
- 5C. Explain the strategies to make drugs less resistant to drug metabolism.
- 5D. Discuss about intrinsic ion channels as drug targets.

(5+5+5+5 = 20 marks)

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M. PHARM. PART-I DEGREE EXAMINATION – MAY/JUNE 2013 SUBJECT: MEDICINAL CHEMISTRY – II (PCH 602) (SPECIALIZATION: PHARMACEUTICAL CHEMISTRY)

Monday, May 27, 2013

Time: 10:00 - 13:00 Hrs.

Max. Marks: 100

- Answer ALL questions.
- 1A. Explain the usage of scavenger resin in combinatorial chemistry. Add a note on the solid supports used.
- 1B. Discuss the various steps used in recombinant DNA technology and enlist their applications.

(10+10 = 20 marks)

- 2A. Write notes on different classes of biomarkers and their applications.
- 2B. Explain the criteria for patentability in India .
- 2C. Mention the methods of peptide synthesis and explain Fmoc polyamide synthesis of a dipeptide.

(8+5+7 = 20 marks)

- 3A. Discuss the toxicity testing methods for a new chemical entitiy.
- 3B. Define integrity and purity. Mention their applications in the evaluation of a new chemical entity.
- 3C. The data for serum cholesterol levels (mg/dl) of three groups of rats is given below. Suggest an appropriate statistical test to find out if atorvastatin and NAT-1 have hypo-cholesterolemic actions. Justify your answer at an appropriate confidence interval estimate.

| Control | 220 | 245 | 230 | 310 | 240 | 255 | 235 | 215 |
|--------------|-----|-----|-----|-----|-----|-----|-----|-----|
| Atorvastatin | 215 | 240 | 225 | 295 | 235 | 240 | 225 | 290 |
| Test [NAT-1] | 195 | 210 | 185 | 175 | 165 | 190 | 180 | 175 |

 $^{(5+5+10 = 20 \}text{ marks})$

- 4A. How is the plasma protein binding (PPB) of a new chemical entity determined? Mention the factors affecting PPB.
- 4B. What do you mean by Quantitative structure activity relationship? Explain with the help of graphs and equations.
- 4C. Discuss the problems associated with CoMFA.

- 5A. Explain the stepwise bond rotation in conformational analysis. Add a note on Monte and Metropolis method.
- 5B. Explain the following:
 - i) Rigid docking by shape complementarity
 - ii) Taft steric factor
 - iii) Rigid docking by matching the hydrogen bonding groups

(8+(5+2+5) = 20 marks)

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M. PHARM. PART-I DEGREE EXAMINATION – MAY/JUNE 2013

SUBJECT: PHARMACEUTICAL PROCESS CHEMISTRY (PCH 603) (SPECIALIZATION: PHARMACEUTICAL CHEMISTRY)

Wednesday, May 29, 2013

Time: 10:00 - 13:00 Hrs.

Max. Marks: 100

& Answer ALL Questions.

1. Explain the various techniques used for modifying and controlling reagent reactivity in chemical process development.

(20 marks)

- 2A. Write the methods used for the protection and deprotection of alcohols as acetals and ethers.
- 2B. What is phase transfer catalysis? Classify PTC systems and give a note on various phase transfer catalysts.
- 2C. Discuss about the practical problems encountered in classical work-up operations and mention the solutions.

(10+5+5 = 20 marks)

- 3A. Define and classify polymorphism. Mention its importance in pharmaceutical industry.
- 3B. Write a note on salt selection strategy. Discuss the techniques for the characterization of salts.

(10+10 = 20 marks)

- 4A. Write briefly about the principles of chemical process safety.
- 4B. What are the advantages of biocatalytic transformations performed in organic media? Classify the solvent systems used for enzyme catalyzed reactions .
- 4C. Explain the following in amide hydrolysis:
 - i) Esterase method
 - ii) Amidase method
 - iii) Acylase method
 - iv) Hydantoinase method
 - v) Lactamase method

(5+10+5 = 20 marks)

(5+5+5+5 = 20 marks)

- 5A. Discuss about the preparation of ionic liquids.
- 5B. Discuss the effect of solvent on recrystallization process.
- 5C. Discuss effect of inclusion and occlusion on crystallization.
- 5D. Write a note on mechanism of nucleation.

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MANIPAL UNIVERSITY M. PHARM. PART-I DEGREE EXAMINATION – MAY/JUNE 2013 SUBJECT: ADVANCED ORGANIC CHEMISTRY (PCH 604) (SPECIALIZATION: PHARMACEUTICAL CHEMISTRY)

Friday, May 31, 2013

Time: 10:00 – 13:00 Hrs.

Max. Marks: 100

- 1A. Write two methods of preparation of quinoline. Compare the oxidations of quinoline and isoquinoline.
- 1B. What are functional group inter conversion and addition in synthetic planning? Add a note on protocol for synthetic design.
- 1C. Explain the retro synthesis and forward synthesis of Losartan.

(6+6+8 = 20 marks)

2A. What are aza - indolizines? Give a method of preparation of each one of them.

- 2B. Write the methods of preparation and reactions of pyrones.
- 2C. With example, explain the mechanism involved in Grignard reaction.

(5+10+5 = 20 marks)

- 3A. List out the reagents used in peptide synthesis and write a brief note on any two such reagents.
- 3B. Explain in detail asymmetric transfer hydrogenation.
- 3C. With example, explain the mechanism involved in Heck reaction.

(10+5+5 = 20 marks)

- 4A. Explain in detail the important chiral auxiliaries used for aldol reactions.
- 4B. Discuss Enantioselective dihydroxylation reactions of olefins.
- 4C. With example, explain the mechanism involved in Mitsunobu reaction.

(8+7+5 = 20 marks)

- 5A. What is retro Diels-Alder reaction? Explain.
- 5B. With example, explain the mechanism involved in Michael Addition reaction.
- 5C. Discuss thermodynamic resolution.
- 5D. Enumerate the uses of affinity chromatography.

(5+5+5+5 = 20 marks)

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M. PHARM. PART-I DEGREE EXAMINATION - MAY/JUNE 2013

SUBJECT: SPECTRAL AND CHROMATOGRAPHIC METHODS IN PHARMACEUTICAL CHEMISTRY (PCH 605) (SPECIALIZATION: PHARMACEUTICAL CHEMISTRY & PHARMACOGNOSY)

Time: 10:00 - 13:00 Hrs.

Monday, June 03, 2013

Max. Marks: 100

1A. Discuss the interpretation of HMBC spectrum of IPSENOL with help of a neat sketch.

- 1B. Write a short note on:
 - i) Deuterium exchange reactions in PMR
 - ii) Shift reagents and
 - iii) Anisotropic effect
- Mention the advantages of 2D NMR over 1D NMR and give an outline of the steps in 2D NMR experiment.

(5+10+5 = 20 marks)

- 2A. Discuss in detail about Fast atom bombardment ionization technique.
- 2B. Explain the principle and technique of carrying out gel electrophoresis.
- 2C. Assign the given ¹³C chemical shift values to the appropriate carbons in the following compounds.
 - i) δ : 143.6 and 109.9 in Furan,
 - ii) δ : 150.06, 148.02, 135.6, 127.9, 126.19, 120.7, 127.54, 129.114, 129.175 in Quinoline
 - iii) δ : 156.6, 29.1, 127.9, 128.4, 125.7, 144.2 in ethyl benzene iv) δ : 54.1, 160, 114.1, 129.5 and 120.8 in Anisole.

(5+5+10 = 20 marks)

- 3A. List out the applications of mass spectrometric technique.
- 3B. Explain the steps involved in the determination of molecular composition from Mass spectrum.
- 3C. Discuss the advantages and disadvantages of Gas-solid chromatography.*
- 3D. Write in detail about gases used in GC.

(5+5+5+5 = 20 marks)

4A. Calculate λ_{max} for the following:





4B. Explain the following transitions:

i) n $\neg \pi^*$ ii) $\pi \neg \pi^*$ iii) n $\neg \sigma^*$ iv) $\sigma \neg \sigma^*$

- 4C. Describe some characteristic IR absorption bands with their probable region and intensity for amides.
- 4D. In IR spectroscopy, explain about coupled vibrations and Fermi Resonance.
- 4E. Write the expected IR peaks for 4-nitro benzoic acid.

(6+4+3+4+3 = 20 m)

- 5A. What is the need of derivatization in HPLC? Explain the techniques available with examples.5B. With a neat labelled diagram explain the instrumentation of supercritical fluid chromatography.
- 5C. Discuss the effect of HPLC mobile phase strength on resolution.

(10+5+5 = 20 marks)