PHARMACEUTICALCHEMISTRY(MPC)

MODERN PHARMACEUTICAL ANALYTICAL TECHNIQUES (MPC 101T)

Scope

This subject deals with various advanced analytical instrumental techniques for identification, characterization and quantification of drugs. Instruments dealt are NMR, Mass spectrometer, IR, HPLC, GC etc.

Objectives

After completion of course student is able to know about chemicals and excipients

- The analysis of various drugs in single and combination dosage forms
- Theoretical and practical skills of the instruments

THEORY 60 Hrs

- 1. a. UV-Visible spectroscopy: Introduction, Theory, Laws, 10 Instrumentation associated with UV-Visible spectroscopy, Choice Hrs of solvents and solvent effect and Applications of UV-Visible spectroscopy, Difference/ Derivative spectroscopy.
 - b. IR spectroscopy: Theory, Modes of Molecular vibrations, Sample handling, Instrumentation of Dispersive and Fourier Transform IR Spectrometer, Factors affecting vibrational frequencies and Applications of IR spectroscopy, Data Interpretation.
 - c. Spectroflourimetry: Theory of Fluorescence, Factors affecting fluorescence (Characterestics of drugs that can be analysed by flourimetry), Quenchers, Instrumentation and Applications of fluorescence spectrophotometer.
 - d. Flame emission spectroscopy and Atomic absorption spectroscopy: Principle, Instrumentation, Interferences and Applications.
- 2 NMR spectroscopy: Quantum numbers and their role in NMR, Principle, Instrumentation, Solvent requirement in NMR, Relaxation process, NMR signals in various compounds, Chemical shift, Factors influencing chemical shift, Spin-Spin coupling, Coupling constant, Nuclear magnetic double resonance, Brief outline of principles of FT-NMR and 13C NMR. Applications of NMR spectroscopy.

- 3 Mass Spectroscopy: Principle, Theory, Instrumentation of Mass 10 Spectroscopy, Different types of ionization like electron impact, Hrs chemical, field, FAB and MALDI, APCI, ESI, APPI Analyzers of Quadrupole and Time of Flight, Mass fragmentation and its rules, Meta stable ions, Isotopic peaks and Applications of Mass spectroscopy.
- 4 Chromatography: Principle, apparatus, instrumentation, 10 chromatographic parameters, factors affecting resolution, isolation Hrs of drug from excipients, data interpretation and applications of the following:
 - a) Thin Layer chromatography
 - b) High Performance Thin Layer Chromatography
 - c) Ion exchange chromatography
 - d) Column chromatography
 - e) Gas chromatography
 - f) High Performance Liquid chromatography
 - g) Ultra High Performance Liquid chromatography
 - h) Affinity chromatography
 - i) Gel Chromatography
- 5 a.Electrophoresis: Principle, Instrumentation, Working 10 conditions, factors affecting separation and applications of the Hrs following:
 - a) Paper electrophoresis b) Gel electrophoresis c) Capillary electrophoresis d) Zone electrophoresis e) Moving boundary electrophoresis f) Iso electric focusing
 - b.X ray Crystallography: Production of X rays, Different X ray methods, Bragg's law, Rotating crystal technique, X ray powder technique, Types of crystals and applications of X-ray diffraction.
- 6 a. Potentiometry: Principle, working, Ion selective Electrodes 10 and Application of potentiometry.
 - b. Thermal Techniques: Principle, thermal transitions and Instrumentation (Heat flux and power-compensation and designs), Modulated DSC, Hyper DSC, experimental parameters (sample preparation, experimental conditions, calibration, heating and cooling rates, resolution, source of errors) and their influence, advantage and disadvantages, pharmaceutical applications. Differential Thermal Analysis (DTA): Principle, instrumentation

and advantage and disadvantages, pharmaceutical applications, derivative differential thermal analysis (DDTA). TGA: Principle, instrumentation, factors affecting results, advantage and disadvantages, pharmaceutical applications.

- 1. Spectrometric Identification of Organic compounds Robert M Silverstein, Sixth edition, John Wiley & Sons, 2004.
- 2. Principles of Instrumental Analysis Doglas A Skoog, F. James Holler, Timothy A. Nieman, 5th edition, Eastern press, Bangalore, 1998.
- 3. Instrumental methods of analysis Willards, 7th edition, CBS publishers.
- 4. Practical Pharmaceutical Chemistry Beckett and Stenlake, Vol II, 4th edition, CBS Publishers, New Delhi, 1997.
- 5. Organic Spectroscopy William Kemp, 3rd edition, ELBS, 1991.
- 6. Quantitative Analysis of Drugs in Pharmaceutical formulation P D Sethi, 3rd Edition, CBS Publishers, New Delhi, 1997.
- 7. Pharmaceutical Analysis Modern Methods Part B J W Munson, Vol 11, Marcel. Dekker Series
- 8. Spectroscopy of Organic Compounds, 2nd edn., P.S/Kalsi, Wiley estern Ltd., Delhi.
- 9. Textbook of Pharmaceutical Analysis, KA.Connors, 3rd Edition, John Wiley & Sons, 1982.

ADVANCED ORGANIC CHEMISTRY - I (MPC 102T)

Scope

The subject is designed to provide in-depth knowledge about advances in organic chemistry, different techniques of organic synthesis and their applications to process chemistry as well as drug discovery.

Objectives

Upon completion of course, the student shall be to understand

- The principles and applications of reterosynthesis
- The mechanism & applications of various named reactions
- The concept of disconnection to develop synthetic routes for small target molecule.
- The various catalysts used in organic reactions
- The chemistry of heterocyclic compounds

THEORY 60 Hrs

- 1. Basic Aspects of Organic Chemistry:
- 12 Organic intermediates: Carbocations, carbanions, free Hrs radicals, carbenes and nitrenes. Their method of formation, stability and synthetic applications.
 - 2. Types of reaction mechanisms and methods of determining them,
 - regarding 3. Detailed knowledge the reactions. mechanisms and their relative reactivity and orientations.

Addition reactions

- Nucleophilic uni- and bimolecular reactions (SN1 and a) SN2)
- b) Elimination reactions (E1 & E2; Hoffman & Saytzeff's rule)

12

Hrs

- Rearrangement reaction
- 2 Study of mechanism and synthetic applications of following named Reactions:

Ugi reaction, Brook rearrangement, Ullmann coupling reactions, Dieckmann Reaction, Doebner-Miller Reaction, Sandmever Reaction, Mitsunobu reaction, Mannich reaction, Vilsmeyer-Haack Reaction, Sharpless asymmetric epoxidation, Baeyer-Villiger oxidation, Shapiro & Suzuki reaction, Ozonolysis and Michael addition reaction

3 Synthetic Reagents & Applications:
Aluminiumisopropoxide, N-bromosuccinamide, diazomethane, dicyclohexylcarbodimide, Wilkinson reagent, Witting reagent.
Osmium tetroxide, titanium chloride, diazopropane, diethyl azodicarboxylate, Triphenylphosphine, Benzotriazol-1-yloxy) tris (dimethylamino) phosphonium hexafluoro-phosphate (BOP).

12 Hrs

Protecting groups

- a. Role of protection in organic synthesis
- b. Protection for the hydroxyl group, including 1,2-and1,3-diols: ethers, esters, carbonates, cyclic acetals & ketals
- c. Protection for the Carbonyl Group: Acetals and Ketals
- d. Protection for the Carboxyl Group: amides and hydrazides, esters
- e. Protection for the Amino Group and Amino acids: carbamates and amides
- 4 Heterocyclic Chemistry:

12 Hrs

Organic Name reactions with their respective mechanism and application involved in synthesis of drugs containing five, six membered and fused hetrocyclics such as Debus-Radziszewski imidazole synthesis, Knorr Pyrazole Synthesis Pinner Pyrimidine Synthesis, Combes Quinoline Synthesis, Bernthsen Acridine Synthesis, Smiles rearrangement and Traube purine synthesis.

Synthesis of few representative drugs containing these hetrocyclic nucleus such as Ketoconazole. Metronidazole. Miconazole, celecoxib, antipyrin, Metamizole sodium. Terconazole. Alprazolam. Triamterene. Sulfamerazine. Hydroxychloroguine, Ouinine, Trimethoprim, Chloroquine, Prochlorpherazine. Ouinacrine. Amsacrine. Promazine. Chlorpromazine, Theophylline, Mercaptopurine and Thioguanine.

5 Synthon approach and retrosynthesis applications

- i. Basic principles, terminologies and advantages of retrosynthesis; guidelines for dissection of molecules. Functional group interconvertion and addition (FGI and FGA)
- ii. C-X disconnections; C-C disconnections alcohols and carbonyl compounds; 1,2-, 1,3-,1,4-, 1,5-, 1,6-difunctionalized compounds
- iii. Strategies for synthesis of three, four, five and six-membered ring.

- 1. "Advanced Organic chemistry, Reaction, Mechanisms and Structure", J. March, John Wiley and Sons, New York.
- 2. "Mechanism and Structure in Organic Chemistry", ES Gould, Hold Rinchart and Winston, New York.
- 3. "Organic Chemistry" Clayden, Greeves, Warren and Woihers., Oxford University Press 2001.
- 4. "Organic Chemistry" Vol I and II. I.L. Finar. ELBS, Pearson Education Lts, Dorling Kindersley 9India) Pvt. Ltd.,.
- 5. A guide to mechanisms in Organic Chemistry, Peter Skyes (Orient Longman, New Delhi).
- 6. Reactive Intermediates in Organic Chemistry, Tandom and Gowel, Oxford & IBH Publishers.
- 7. Combinational Chemistry Synthesis and applications Stephen R Wilson & Anthony W Czarnik, Wiley Blackwell.
- 8. Carey, Organic Chemistry, 5th Edition (Viva Books Pvt. Ltd.)
- 9. Organic Synthesis The Disconnection Approach, S. Warren, Wily India
- 10. Principles of Organic Synthesis, ROC Norman and JM Coxan, Nelson Thorns.
- 11. Organic Synthesis Special Techniques. VK Ahluwalia and R Agarwal, Narosa Publishers.
- 12. Organic Reaction Mechanisms IVth Edtn, VK Ahluwalia and RK Parashar, Narosa Publishers.

ADVANCED MEDICINAL CHEMISTRY (MPC 103T)

Scope

The subject is designed to impart knowledge about recent advances in the field of medicinal chemistry at the molecular level including different techniques for the rational drug design.

Objectives

At completion of this course it is expected that students will be able to understand

- Different stages of drug discovery
- Role of medicinal chemistry in drug research
- Different techniques for drug discovery
- Various strategies to design and develop new drug like molecules for biological targets
- Peptidomimetics

THEORY 60 Hrs

Drug discovery: Stages of drug discovery, lead discovery; 12 identification, validation and diversity of drug targets.

Biological drug targets: Receptors, types, binding and activation, theories of drug receptor interaction, drug receptor interactions, agonists vs antagonists, artificial enzymes.

2 Prodrug Design and Analog design:

- a) Prodrug design: Basic concept, Carrier linked prodrugs/ Bioprecursors, Prodrugs of functional group, Prodrugs to improve patient acceptability, Drug solubility, Drug absorption and distribution, site specific drug delivery and sustained drug action. Rationale of prodrug design and practical consideration of prodrug design.
- b) Combating drug resistance: Causes for drug resistance, strategies to combat drug resistance in antibiotics and anticancer therapy, Genetic principles of drug resistance.
- c) Analog Design: Introduction, Classical & Non classical, Bioisosteric replacement strategies, rigid analogs,

alteration of chain branching, changes in ring size, ring position isomers, design of stereo isomers and geometric isomers, fragments of a lead molecule, variation in inter atomic distance.

a) Medicinal chemistry aspects of the following class of drugs

12 Hrs

Systematic study, SAR, Mechanism of action and synthesis of new generation molecules of following class of drugs:

- a) Anti-hypertensive drugs, Psychoactive drugs, Anticonvulsant drugs, H1 & H2 receptor antagonist, COX1 & COX2 inhibitors, Adrenergic & Cholinergic agents, Antineoplastic and Antiviral agents.
- b) Stereochemistry and Drug action: Realization that stereo selectivity is a pre-requisite for evolution. Role of chirality in selective and specific therapeutic agents. Case studies, Enantio selectivity in drug adsorption, metabolism, distribution and elimination.
- 4 Rational Design of Enzyme Inhibitors 12
 Enzyme kinetics & Principles of Enzyme inhibitors, Enzyme Hrs
 inhibitors in medicine, Enzyme inhibitors in basic research,
 rational design of non-covalently and covalently binding enzyme
 inhibitors.
- Peptidomimetics 12
 Therapeutic values of Peptidomimetics, design of Hrs peptidomimetics by manipulation of the amino acids, modification of the peptide backbone, incorporating conformational constraints locally or globally. Chemistry of prostaglandins, leukotrienes and thromboxones.

- 1. Medicinal Chemistry by Burger, Vol I –VI.
- Wilson and Gisvold's Text book of Organic Medicinal and Pharmaceutical Chemistry, 12th Edition, Lppincott Williams & Wilkins, Woltess Kluwer (India) Pvt.Ltd, New Delhi.
- 3. Comprehensive Medicinal Chemistry Corwin and Hansch.
- 4. Computational and structural approaches to drug design edited by Robert M Stroud and Janet. F Moore

- 5. Introduction to Quantitative Drug Design by Y.C. Martin.
- 6. Principles of Medicinal Chemistry by William Foye, 7th Edition, Ippincott Williams & Wilkins, Woltess Kluwer (India) Pvt.Ltd, New Delhi.
- 7. Drug Design Volumes by Arienes, Academic Press, Elsevier Publishers, Noida, Uttar Pradesh..
- 8. Principles of Drug Design by Smith.
- 9. The Organic Chemistry of the Drug Design and Drug action by Richard B.Silverman, II Edition, Elsevier Publishers, New Delhi.
- 10. An Introduction to Medicinal Chemistry, Graham L.Patrick, III Edition, Oxford University Press, USA.
- 11. Biopharmaceutics and pharmacokinetics, DM.Brahmankar, Sunil B. Jaiswal II Edition, 2014, Vallabh Prakashan, New Delhi.
- 12. Peptidomimetics in Organic and Medicinal Chemistry by Antonio Guarna and Andrea Trabocchi, First edition, Wiley publishers.

CHEMISTRY OF NATURAL PRODUCTS (MPC 104T)

Scope

The subject is designed to provide detail knowledge about chemistry of medicinal compounds from natural origin and general methods of structural elucidation of such compounds. It also emphasizes on isolation, purification and characterization of medicinal compounds from natural origin.

Objectives

At completion of this course it is expected that students will be able to understand-

- Different types of natural compounds and their chemistry and medicinal importance
- The importance of natural compounds as lead molecules for new drug discovery
- The concept of rDNA technology tool for new drug discovery
- General methods of structural elucidation of compounds of natural origin
- Isolation, purification and characterization of simple chemical constituents from natural source

THEORY 60 Hrs

- Study of Natural products as leads for new pharmaceuticals
 for the following class of drugs
 Hrs
 - a) Drugs Affecting the Central Nervous System: Morphine Alkaloids
 - b) Anticancer Drugs: Paclitaxel and Docetaxel, Etoposide, and Teniposide
 - c) Cardiovascular Drugs: Lovastatin, Teprotide and Dicoumarol
 - d) Neuromuscular Blocking Drugs: Curare alkaloids
 - e) Anti-malarial drugs and Analogues

emetine and reserpine.

- f) Chemistry of macrolid antibiotics (Erythromycin, Azithromycin, Roxithromycin, and Clarithromycin) and β Lactam antibiotics (Cephalosporins and Carbapenem)
- 2 a) Alkaloids 12
 General introduction, classification, isolation, purification, Hrs
 molecular modification and biological activity of alkaloids, general
 methods of structural determination of alkaloids, structural
 elucidation and stereochemistry of ephedrine, morphine, ergot,

b) Flavonoids

Introduction, isolation and purification of flavonoids, General methods of structural determination of flavonoids; Structural elucidation of guercetin.

c) Steroids

General introduction, chemistry of sterols, sapogenin and cardiac glycosides. Stereochemistry and nomenclature of steroids, chemistry of contraceptive agents male & female sex hormones (Testosterone, Estradiol, Progesterone), adrenocorticoids (Cortisone), contraceptive agents and steroids (Vit - D).

3 a) Terpenoids

12 Hrs

12

Hrs

Classification, isolation, isoprene rule and general methods of structural elucidation of Terpenoids; Structural elucidation of drugs belonging to mono (citral, menthol, camphor), di(retinol, Phytol, taxol) and tri terpenoids (Squalene, Ginsenoside) carotinoids (β carotene).

b) Vitamins

Chemistry and Physiological significance of Vitamin A, B1, B2, B12, C, E, Folic acid and Niacin.

- 4 a). Recombinant DNA technology and drug discovery rDNA technology, hybridoma technology, New pharmaceuticals derived from biotechnology; Oligonucleotide therapy. Gene therapy: Introduction, Clinical application and recent advances in gene therapy, principles of RNA & DNA estimation
 - b). Active constituent of certain crude drugs used in Indigenous system Diabetic therapy Gymnema sylvestre, Salacia reticulate, Pterocarpus marsupiam, Swertia chirata,
- Structural Characterization of natural compounds
 Structural characterization of natural compounds using IR,
 1HNMR, 13CNMR and MS Spectroscopy of specific drugs e.g.,
 Penicillin, Morphine, Camphor, Vit-D, Quercetin and Digitalis
 glycosides.

Antitumor - Curcuma longa Linn.

Trigonella foenum graccum: Liver dysfunction - Phyllanthus niruri:

- 1. Modern Methods of Plant Analysis, Peech and M.V.Tracey, Springer Verlag, Berlin, Heidelberg.
- 2. Phytochemistry Vol. I and II by Miller, Jan Nostrant Rein Hld.
- 3. Recent advances in Phytochemistry Vol. I to IV Scikel Runeckles, Springer Science & Business Media.
- 4. Chemistry of natural products Vol I onwards IWPAC.
- Natural Product Chemistry Nakanishi Gggolo, University Science Books, California.
- 6. Natural Product Chemistry "A laboratory guide" Rapheal Khan.
- 7. The Alkaloid Chemistry and Physiology by RHF Manske, Academic Press.
- 8. Introduction to molecular Phytochemistry CHJ Wells, Chapmannstall.
- 9. Organic Chemistry of Natural Products Vol I and II by Gurdeep and Chatwall, Himalaya Publishing House.
- Organic Chemistry of Natural Products Vol I and II by O.P. Agarwal, Krishan Prakashan.
- 11. Organic Chemistry Vol I and II by I.L. Finar, Pearson education.
- 12. Elements of Biotechnology by P.K. Gupta, Rastogi Publishers.
- 13. Pharmaceutical Biotechnology by S.P.Vyas and V.K.Dixit, CBS Publishers.
- 14. Biotechnology by Purohit and Mathur, Agro-Bios, 13th edition.
- 15. Phytochemical methods of Harborne, Springer, Netherlands.
- 16. Burger's Medicinal Chemistry.

PHARMACEUTICAL CHEMISTRY PRACTICAL - I (MPC 105P)

- Analysis of Pharmacopoeial compounds and their formulations by UV Vis spectrophotometer, RNA & DNA estimation
- Simultaneous estimation of multi component containing formulations by UV spectrophotometry
- 3. Experiments based on Column chromatography
- 4. Experiments based on HPLC
- 5. Experiments based on Gas Chromatography
- 6. Estimation of riboflavin/quinine sulphate by fluorimetry
- 7. Estimation of sodium/potassium by flame photometry

To perform the following reactions of synthetic importance

- 1. Purification of organic solvents, column chromatography
- Claisen-schimidt reaction.
- 3. Benzyllic acid rearrangement.
- 4. Beckmann rearrangement.
- 5. Hoffmann rearrangement
- 6. Mannich reaction
- 7. Synthesis of medicinally important compounds involving more than one step along with purification and Characterization using TLC, melting point and IR spectroscopy (4 experiments)
- 8. Estimation of elements and functional groups in organic natural compounds
- Isolation, characterization like melting point, mixed melting point, molecular weight determination, functional group analysis, co-chromatographic technique for identification of isolated compounds and interpretation of UV and IR data.
- 10. Some typical degradation reactions to be carried on selected plant constituents

ADVANCED SPECTRAL ANALYSIS (MPC 201T)

Scope

This subject deals with various hyphenated analytical instrumental techniques for identification, characterization and quantification of drugs. Instruments dealt are LC-MS, GC-MS, ATR-IR, DSC etc.

Objectives

THEORY

At completion of this course it is expected that students will be able to understand-

60Hrs

Hrs

- Interpretation of the NMR, Mass and IR spectra of various organic compounds
- Theoretical and practical skills of the hyphenated instruments
- Identification of organic compounds

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1.	UV and IR spectroscopy: Wood ward – Fieser rule for 1,3- butadienes, cyclic dienes and α , β -carbonyl compounds and interpretation compounds of enones. ATR-IR, IR Interpretation of organic compounds.	12 Hrs
2	NMR spectroscopy: 1-D and 2-D NMR, NOESY and COSY, HECTOR, INADEQUATE techniques, Interpretation of organic compounds.	12 Hrs
3	Mass Spectroscopy Mass fragmentation and its rules, Fragmentation of important functional groups like alcohols, amines, carbonyl groups and alkanes, Meta stable ions, Mc Lafferty rearrangement, Ring rule, Isotopic peaks, Interpretation of organic compounds.	12 Hrs
4	Chromatography:	12

Principle, Instrumentation and Applications of the following:

Exclusion Chromatography) k) Flash chromatography

a) GC-MS b) GC-AAS c) LC-MS d) LC-FTIR e) LC-NMR f) CE-MS g) High Performance Thin Layer chromatography h) Super critical fluid chromatography i) I-DC (Ion-

- 5 a). Thermal methods of analysis 12 Introduction, principle, instrumentation and application of DSC, Hrs DTA and TGA.
 - b). Raman Spectroscopy Introduction, Principle, Instrumentation and Applications.
 - c). Radio immuno assay Biological standardization, bioassay, ELISA, Radioimmuno assay of digitalis and insulin.

- 1. Spectrometric Identification of Organic compounds Robert M Silverstein, Sixth edition, John Wiley & Sons, 2004.
- 2. Principles of Instrumental Analysis Doglas A Skoog, F. James Holler, Timothy A. Nieman, 5th edition, Eastern press, Bangalore, 1998.

 3. Instrumental methods of analysis – Willards, 7th edition, CBS publishers.
- 4. Organic Spectroscopy William Kemp, 3rd edition, ELBS, 1991.
- 5. Quantitative analysis of Pharmaceutical formulations by HPTLC P D Sethi, CBS Publishers, New Delhi,
- 6. Quantitative Analysis of Drugs in Pharmaceutical formulation P D Sethi, 3rd Edition, CBS Publishers, New Delhi, 1997.
- 7. Pharmaceutical Analysis- Modern methods Part B J W Munson, Volume 11, Marcel Dekker Series

ADVANCED ORGANIC CHEMISTRY - II (MPC 202T)

Scope

The subject is designed to provide in-depth knowledge about advances in organic chemistry, different techniques of organic synthesis and their applications to process chemistry as well as drug discovery.

Objectives

Upon completion of course, the student shall able to understand

- The principles and applications of Green chemistry
- The concept of peptide chemistry.
- The various catalysts used in organic reactions
- The concept of stereochemistry and asymmetric synthesis.

THEORY 60 Hrs

1. Green Chemistry: 12

a. Introduction, principles of green chemistry

- Hrs
- b. Microwave assisted reactions: Merit and demerits of its use, increased reaction rates, mechanism, superheating effects of microwave, effects of solvents in microwave assisted synthesis, microwave technology in process optimization, its applications in various organic reactions and heterocycles synthesis
- c. Ultrasound assisted reactions: Types of sonochemical reactions, homogenous, heterogeneous liquid-liquid and liquid-solid reactions, synthetic applications
- d. Continuous flow reactors: Working principle, advantages and synthetic applications.
- 2 Chemistry of peptides

- a. Coupling reactions in peptide synthesis
- b. Principles of solid phase peptide synthesis, t-BOC and FMOC protocols, various solid supports and linkers: Activation procedures. peptide bond formation, deprotection cleavage from resin, low and high HF cleavage protocols, formation of free peptides and peptide amides, purification and case studies, site-specific chemical modifications of peptides
- c. Segment and sequential strategies for solution phase peptide synthesis with any two case studies
- d. Side reactions in peptide synthesis: Deletion peptides, side

reactions initiated by proton abstraction, protonation, overactivation and side reactions of individual amino acids.

Photochemical Reactions

12 Hrs

Basic principles of photochemical reactions. Photo-oxidation, photo-addition and photo-fragmentation.

Pericyclic reactions

Mechanism, Types of pericyclic reactions such as cyclo addition, electrocyclic reaction and sigmatrophic rearrangement reactions with examples

4 Catalysis:

12

- a. Types of catalysis, heterogeneous and homogenous catalysis, Hrs advantages and disadvantages
- Heterogeneous catalysis preparation, characterization, kinetics, supported catalysts, catalyst deactivation and regeneration, some examples of heterogeneous catalysis used in synthesis of drugs.
- Homogenous catalysis, hydrogenation, hydroformylation, hydrocyanation, Wilkinson catalysts, chiral ligands and chiral induction, Ziegler-Natta catalysts, some examples of homogenous catalysis used in synthesis of drugs
- d. Transition-metal and Organo-catalysis in organic synthesis: Metal-catalyzed reactions
- e. Biocatalysis: Use of enzymes in organic synthesis, immobilized enzymes/cells in organic reaction.
- f. Phase transfer catalysis theory and applications

5 Stereochemistry & Asymmetric Synthesis

- a. Basic concepts in stereochemistry optical activity, specific rotation, racemates and resolution of racemates, the Cahn, Ingold, Prelog (CIP) sequence rule, meso compounds, pseudo asymmetric centres, axes of symmetry, Fischers D and L notation, cis-trans isomerism, E and Z notation.
- b. Methods of asymmetric synthesis using chiral pool, chiral auxiliaries and catalytic asymmetric synthesis, enantiopure separation and Stereo selective synthesis with examples.

- 1. "Advanced Organic chemistry, Reaction, mechanisms and structure", J March, John Wiley and sons, New York.
- "Mechanism and structure in organic chemistry", ES Gould, Hold Rinchart and Winston, NewYork.
- 3. "Organic Chemistry" Clayden, Greeves, Warren and Woihers., Oxford University Press 2001.
- 4. "Organic Chemistry" Vol I and II. I.L. Finar. ELBS, Sixth ed., 1995.
- 5. Carey, Organic chemistry, 5th edition (Viva Books Pvt. Ltd.)
- 6. Organic synthesis-the disconnection approach, S. Warren, Wily India
- 7. Principles of organic synthesis, ROCNorman and JMCoxan, Nelson thorns
- 8. Organic synthesis- Special techniques VK Ahluwalia and R Aggarwal, Narosa Publishers.
- Organic reaction mechanisms IV edtn, VK Ahluwalia and RK Parashar, Narosa Publishers.

COMPUTER AIDED DRUG DESIGN (MPC 203T)

Scope

The subject is designed to impart knowledge on the current state of the art techniques involved in computer assisted drug design.

Objectives

At completion of this course it is expected that students will be able to understand

- Role of CADD in drug discovery
- Different CADD techniques and their applications
- Various strategies to design and develop new drug like molecules.
- Working with molecular modeling softwares to design new drug molecules
- The in silico virtual screening protocols

Theory 60 Hrs

Introduction to Computer Aided Drug Design (CADD)
 Hrs

History, different techniques and applications.

Quantitative Structure Activity Relationships: Basics

History and development of QSAR: Physicochemical parameters and methods to calculate physicochemical parameters: Hammett equation and electronic parameters (sigma), lipophilicity effects and parameters (log P, pi-substituent constant), steric effects (Taft steric and MR parameters) Experimental and theoretical approaches for the determination of these physicochemical parameters.

- Quantitative Structure Activity Relationships: Applications Hansch analysis, Free Wilson analysis and relationship between them, Advantages and disadvantages; Deriving 2D-QSAR equations.
 - 3D-QSAR approaches and contour map analysis.

Statistical methods used in QSAR analysis and importance of statistical parameters.

3 Molecular Modeling and Docking

12

- a) Molecular and Quantum Mechanics in drug design.
- Hrs

12 Hrs

b) Energy Minimization Methods: comparison between global

- minimum conformation and bioactive conformation
- c) Molecular docking and drug receptor interactions: Rigid docking, flexible docking and extra-precision docking. Agents acting on enzymes such as DHFR, HMG-CoA reductase and HIV protease, choline esterase (AchE & BchE)
- 4 Molecular Properties and Drug Design

12

12

- a) Prediction and analysis of ADMET properties of new Hrs molecules and its importance in drug design.
- b) De novo drug design: Receptor/enzyme-interaction and its analysis, Receptor/enzyme cavity size prediction, predicting the functional components of cavities, Fragment based drug
- c) Homology modeling and generation of 3D-structure of protein.
- 5 Pharmacophore Mapping and Virtual Screening of pharmacophore, pharmacophore Hrs Concept mapping, identification of Pharmacophore features and Pharmacophore modeling; Conformational search used in pharmacophore mapping.

In Silico Drug Design and Virtual Screening Techniques Similarity based methods and Pharmacophore based screening, structure based In-silico virtual screening protocols.

- 1. Computational and structural approaches to drug discovery, Robert M Stroud and Janet. F Moore, RCS Publishers.
- 2. Introduction to Quantitative Drug Design by Y.C. Martin, CRC Press, Taylor & Francis group...
- 3. Drug Design by Ariens Volume 1 to 10, Academic Press, 1975, Elsevier Publishers.
- 4. Principles of Drug Design by Smith and Williams, CRC Press, Taylor & Francis.
- 5. The Organic Chemistry of the Drug Design and Drug action by Richard B. Silverman, Elsevier Publishers,
- 6. Medicinal Chemistry by Burger, Wiley Publishing Co.

- 7. An Introduction to Medicinal Chemistry -Graham L. Patrick, Oxford University Press.
- 8. Wilson and Gisvold's Text book of Organic Medicinal and Pharmaceutical Chemistry, Ippincott Williams & Wilkins.
- 9. Comprehensive Medicinal Chemistry Corwin and Hansch, Pergamon Publishers.
- 10. Computational and structural approaches to drug design edited by Robert M Stroud and Janet. F Moore

PHARMACEUTICAL PROCESS CHEMISTRY (MPC 204T)

Scope

Process chemistry is often described as scale up reactions, taking them from small quantities created in the research lab to the larger quantities that are needed for further testing and then to even larger quantities required for commercial production. The goal of a process chemist is to develop synthetic routes that are safe, cost-effective, environmentally friendly, and efficient. The subject is designed to impart knowledge on the development and optimization of a synthetic route/s and the pilot plant procedure for the manufacture of Active Pharmaceutical Ingredients (APIs) and new chemical entities (NCEs) for the drug development phase.

Objectives

At completion of this course it is expected that students will be able to understand

- The strategies of scale up process of apis and intermediates
- The various unit operations and various reactions in process chemistry

THEORY 60 Hrs

1. Process chemistry
Introduction, Synthetic strategy
Stages of scale up process: Bench, pilot and large scale process.
In-process control and validation of large scale process.
Case studies of some scale up process of APIs.
Impurities in API, types and their sources including genotoxic impurities

2 Unit operations

12

- a) Extraction: Liquid equilibria, extraction with reflux, Hrs extraction with agitation, counter current extraction.
- b) Filtration: Theory of filtration, pressure and vacuum filtration, centrifugal filtration,
- c) Distillation: azeotropic and steam distillation
- d) Evaporation: Types of evaporators, factors affecting evaporation.
- e) Crystallization: Crystallization from aqueous, nonaqueous solutions factors affecting crystallization, nucleation. Principle and general methods of Preparation of polymorphs, hydrates, solvates and amorphous APIs.

2	Unit Processes - I	12
	 Nitration: Nitrating agents, Aromatic nitration, kinetics and mechanism of aromatic nitration, process equipment for technical nitration, mixed acid for nitration, 	Hrs
	b) Halogenation: Kinetics of halogenations, types of halogenations, catalytic halogenations. Case study on industrial halogenation process.	
	 c) Oxidation: Introduction, types of oxidative reactions, Liquid phase oxidation with oxidizing agents. Nonmetallic Oxidizing agents such as H₂O₂, sodium hypochlorite, Oxygen gas, ozonolysis. 	
4	Unit Processes - II	12
	 Reduction: Catalytic hydrogenation, Heterogeneous and homogeneous catalyst; Hydrogen transfer reactions, Metal hydrides. Case study on industrial reduction process. 	Hrs
	b) Fermentation: Aerobic and anaerobic fermentation. Production of	
	i. Antibiotics; Penicillin and Streptomycin,ii. Vitamins: B2 and B12	
	iii. Statins: Lovastatin, Simvastatin	
	c) Reaction progress kinetic analysis	
	i. Streamlining reaction steps, route selection,	
	 ii. Characteristics of expedient routes, characteristics of cost-effective routes, reagent selection, families of reagents useful for scale-up. 	
5	Industrial Safety	12
	a) MSDS (Material Safety Data Sheet), hazard labels of chemicals and Personal Protection Equipment (PPE)	Hrs
	b) Fire hazards, types of fire & fire extinguishers	
	c) Occupational Health & Safety Assessment Series 1800	
	(OHSAS-1800) and ISO-14001(Environmental	

Management System), Effluents and its management

- 1. Process Chemistry in the Pharmaceutical Industry: Challenges in an Ever-Changing Climate-An Overview; K. Gadamasetti, CRC Press.
- 2. Pharmaceutical Manufacturing Encyclopedia, 3rd edition, Volume 2.
- 3. Medicinal Chemistry by Burger, 6th edition, Volume 1-8.
- 4. W.L. McCabe, J.C Smith, Peter Harriott. Unit operations of chemical engineering, 7th edition, McGraw Hill
- 5. Polymorphism in Pharmaceutical Solids .Dekker Series Volume 95 Ed: H G Brittain (1999)
- Regina M. Murphy: Introduction to Chemical Processes: Principles, Analysis, Synthesis
- 7. Peter J. Harrington: Pharmaceutical Process Chemistry for Synthesis: Rethinking the Routes to Scale-Up
- 8. P.H.Groggins: Unit processes in organic synthesis (MGH)
- 9. F.A.Henglein: Chemical Technology (Pergamon)
- M.Gopal: Dryden's Outlines of Chemical Technology, WEP East-West Press
- 11. Clausen, Mattson: Principle of Industrial Chemistry, Wiley Publishing Co.,
- 12. Lowenheim & M.K. Moran: Industrial Chemicals
- 13. S.D. Shukla & G.N. Pandey: A text book of Chemical Technology Vol. II, Vikas Publishing House
- 14. J.K. Stille: Industrial Organic Chemistry (PH)
- 15. Shreve: Chemical Process, Mc Grawhill.
- 16. B.K.Sharma: Industrial Chemistry, Goel Publishing House
- 17. ICH Guidelines
- 18. United States Food and Drug Administration official website www.fda.gov

PHARMACEUTICAL CHEMISTRY PRACTICALS – II (MPC 205P)

- 1. Synthesis of organic compounds by adapting different approaches involving (3 experiments)
 - a) Oxidation
 - b) Reduction/hydrogenation
 - c) Nitration
- Comparative study of synthesis of APIs/intermediates by different synthetic routes (2 experiments)
- 3. Assignments on regulatory requirements in API (2 experiments)
- 4. Comparison of absorption spectra by UV and Wood ward Fieser rule
- 5. Interpretation of organic compounds by FT-IR
- 6. Interpretation of organic compounds by NMR
- 7. Interpretation of organic compounds by MS
- 8. Determination of purity by DSC in pharmaceuticals
- 9. Identification of organic compounds using FT-IR, NMR, CNMR and Mass spectra
- 10. To carry out the preparation of following organic compounds
- 11. Preparation of 4-chlorobenzhydrylpiperazine. (an intermediate for cetirizine HCl).
- 12. Preparation of 4-iodotolene from p-toluidine.
- 13. NaBH₄ reduction of vanillin to vanilly alcohol
- 14. Preparation of umbelliferone by Pechhman reaction
- 15. Preparation of triphenyl imidazole
- 16. To perform the Microwave irradiated reactions of synthetic importance (Any two)
- 17. Determination of log P, MR, hydrogen bond donors and acceptors of selected drugs using softwares
- 18. Calculation of ADMET properties of drug molecules and its analysis using softwares
 - Pharmacophore modeling
- 19. 2D-QSAR based experiments
- 20. 3D-QSAR based experiments
- 21. Docking study based experiment
- 22. Virtual screening based experiment

(Pharmaceutical Chemistry-MPC)

(Pharmaceutical Chemistry-MPC)								
		In	Internal Assessment			End Semester Exams		
Course Code	Course	Cont inuo	inuo Exams		Tot	Mar	Du	Total Marks
		us Mod e	Mar ks	Durati on	al	ks	rati on	
			SEMEST	ΓER I				
MPC101T	Modern Pharmaceutic al Analytical Techniques	10	15	1 Hr	25	75	3 Hrs	100
MPC102T	Advanced Organic Chemistry -I	10	15	1 Hr	25	75	3 Hrs	100
MPC103T	Advanced Medicinal chemistry	10	15	1 Hr	25	75	3 Hrs	100
MPC104T	Chemistry of Natural Products	10	15	1 Hr	25	75	3 Hrs	100
MPC105P	Pharmaceutic al Chemistry Practical I	20	30	6 Hrs	50	100	6 Hrs	150
-	Seminar /Assignment	-	-	-	-	-	-	100
			otal					650
			SEMEST	TER II				
MPC201T	Advanced Spectral Analysis	10	15	1 Hr	25	75	3 Hrs	100
MPC202T	Advanced Organic Chemistry -II	10	15	1 Hr	25	75	3 Hrs	100
MPC203T	Computer Aided Drug Design	10	15	1 Hr	25	75	3 Hrs	100
MPC204T	Pharmaceutic al Process Chemistry	10	15	1 Hr	25	75	3 Hrs	100
MPC205P	Pharmaceutic	20	30	6 Hrs	50	100	6	150

al Chemistry Practical II						Hrs	
Seminar - /Assignment	-	-	-	-	-	-	100
Total					650		

Tables - 19: Schemes for internal assessments and end semester examinations (Pharmaceutical Analysis-MPA)

Course		Internal Assessment				End Semester Exams		Total
Code	Course	Contin uous Mode	Ex Mark	sional ams Durati	Tot al	Mark s	Dura tion	Marks
			s SEMEST	On FFR I				
	Modern	'	SEIVIES	LICI				
MPA101T	Pharmaceuti cal Analysis	10	15	1 Hr	25	75	3 Hrs	100
MPA102T	Advanced Pharmaceuti cal Analysis	10	15	1 Hr	25	75	3 Hrs	100
MPA103T	Pharmaceuti cal Validation	10	15	1 Hr	25	75	3 Hrs	100
MPA104T	Food Analysis	10	15	1 Hr	25	75	3 Hrs	100
MPA105P	Pharmaceuti cal Analysis- I	20	30	6 Hrs	50	100	6 Hrs	150
-	Seminar /Assignment	-	-	-	-	-	-	100
			otal					650
		,	SEMEST	ER II				
MPA201T	Advanced Instrumental Analysis	10	15	1 Hr	25	75	3 Hrs	100
MPA202T	Modern Bio- Analytical Techniques	10	15	1 Hr	25	75	3 Hrs	100
MPA203T	Quality Control and Quality	10	15	1 Hr	25	75	3 Hrs	100

Table - 4: Course of study for M. Pharm. (Pharmaceutical Chemistry)

Course Code	Course	Credit Hours	Credit Points	Hrs./w k	Marks
	Sem	ester I			
MPC101T	Modern Pharmaceutical Analytical Techniques	4	4	4	100
MPC1012T	Advanced Organic Chemistry -I	4	4	4	100
MPC103T	Advanced Medicinal chemistry	4	4	4	100
MPC104T	Chemistry of Natural Products	4	4	4	100
MPC105P	Pharmaceutical Chemistry Practical I	12	6	12	150
-	Seminar/Assignment	7	4	7	100
	Total	35	26	35	650
	Seme	ester II			
MPC201T	Advanced Spectral Analysis	4	4	4	100
MPC202T	Advanced Organic Chemistry -II	4	4	4	100
MPC203T	Computer Aided Drug Design	4	4	4	100
MPC204T	Pharmaceutical Process Chemistry	4	4	4	100
MPC205P	Pharmaceutical Chemistry Practical II	12	6	12	150
-	Seminar/Assignment	7	4	7	100
	Total	35	26	35	650

Tables - 26: Schemes for internal assessments and end semester examinations (Semester III&IV)

		Int	Internal Assessment			End Semester Exams		Tota
Course Code	Course	Conti nuou		sional ams Tot		Γot Mark	Durati	l Mark s
		s Mode	Mark s	Durati on	al	s	on	3
			SEMES	ΓER III				
MRM30 1T	Research Methodology and Biostatistics*	10	15	1 Hr	25	75	3 Hrs	100
-	Journal club	-	-	-	25	-	-	25
-	Discussion / Presentation (Proposal Presentation)	-	-	-	50	-	-	50
-	Research work*	-	-	-	-	350	1 Hr	350
			Total					525
		ı	SEMEST	TER IV				
-	Journal club	-	-	-	25	-	-	25
-	Discussion / Presentation (Proposal Presentation)	-	-	-	75	-	-	75
-	Research work and Colloquium	-	-	-	-	400	1 Hr	400
Total								500

^{*}Non University Examination

11.2. Internal assessment: Continuous mode

The marks allocated for Continuous mode of Internal Assessment shall be awarded as per the scheme given below.

Table - 27: Scheme for awarding internal assessment: Continuous mode

Theory				
Criteria	Maximum Marks			
Attendance (Refer Table – 28)	8			
Student – Teacher interaction	2			
Total	10			
Practical				
Attendance (Refer Table – 28	10			
Based on Practical Records, Regular viva voce, etc.	10			
Total	20			

Table - 28: Guidelines for the allotment of marks for attendance

Percentage of Attendance	Theory	Practical
95 – 100	8	10
90 – 94	6	7.5
85 – 89	4	5
80 – 84	2	2.5
Less than 80	0	0

11.2.1. Sessional Exams

Two sessional exams shall be conducted for each theory / practical course as per the schedule fixed by the college(s). The scheme of question paper for theory and practical sessional examinations is given in the table. The average marks of two sessional exams shall be computed for internal assessment as per the requirements given in tables.

12. Promotion and award of grades

A student shall be declared PASS and eligible for getting grade in a course of M.Pharm.programme if he/she secures at least 50% marks in that particular courseincluding internal assessment.

13. Carry forward of marks

In case a student fails to secure the minimum 50% in any Theory or Practical course as specified in 12, then he/she shall reappear for the end semester examination of that course. However his/her marks of the Internal Assessment shall be carried over and he/she shall be entitled for grade obtained by him/her on passing.

14. Improvement of internal assessment

A student shall have the opportunity to improve his/her performance only once in the sessional exam component of the internal assessment. The re-conduct of the sessional exam shall be completed before the commencement of next end semester theory examinations.

15. Reexamination of end semester examinations

Reexamination of end semester examination shall be conducted as per the schedule given in table 29. The exact dates of examinations shall be notified from time to time.

Table - 29: Tentative schedule of end semester examinations

Semester	For Regular Candidates	For Failed Candidates
I and III	November / December	May / June
II and IV	May / June	November / December

16. Allowed to keep terms (ATKT):

No student shall be admitted to any examination unless he/she fulfills the norms given in 6. ATKT rules are applicable as follows:

A student shall be eligible to carry forward all the courses of I and IIsemesters till the III semester examinations. However, he/she shall not be eligible to attend the courses of IV semester until all the courses of I, II and III semesters are successfully completed.

A student shall be eligible to get his/her CGPA upon successful completion of the courses of I to IV semesters within the stipulated time period as per the norms.

Note: Grade AB should be considered as failed and treated as one head for deciding ATKT. Such rules are also applicable for those students who fail to register for examination(s) of any course in any semester.

17. Grading of performances

17.1. Letter grades and grade points allocations:

Based on the performances, each student shall be awarded a final letter grade at the end of the semester for each course. The letter grades and their corresponding grade points are given in Table – 30.

Table – 30: Letter grades and grade points equivalent to Percentage of marks and performances

Percentage of Marks Obtained	Letter Grade	Grade Point	Performance
90.00 - 100	0	10	Outstanding
80.00 - 89.99	Α	9	Excellent
70.00 – 79.99	В	8	Good
60.00 - 69.99	С	7	Fair
50.00 - 59.99	D	6	Average
Less than 50	F	0	Fail
Absent	AB	0	Fail

A learner who remains absent for any end semester examination shall be assigned a letter grade of AB and a corresponding grade point of zero. He/she should reappear for the said evaluation/examination in due course.

18. The Semester grade point average (SGPA)

The performance of a student in a semester is indicated by a number called 'Semester Grade Point Average' (SGPA). The SGPA is the weighted average of the grade points obtained in all the courses by the student during the semester. For example, if a student takes five courses (Theory/Practical) in a semester with credits C1, C2, C3 and C4 and the student's grade points in these courses are G1, G2, G3 and G4, respectively, and then students' SGPA is equal to:

$$SGPA = C_1G_1 + C_2G_2 + C_3G_3 + C_4G_4$$

$$C_1 + C_2 + C_3 + C_4$$

The SGPA is calculated to two decimal points. It should be noted that, the SGPA for any semester shall take into consideration the F and ABS grade awarded in that semester. For example if a learner has a F or ABS grade in course 4, the SGPA shall then be computed as:

$$SGPA = \begin{array}{c} C_1G_1 + C_2G_2 + C_3G_3 + C_4* ZERO \\ \\ C_1 + C_2 + C_3 + C_4 \end{array}$$

19. Cumulative Grade Point Average (CGPA)

The CGPA is calculated with the SGPA of all the IV semesters to two decimal points and is indicated in final grade report card/final transcript showing the grades of all IV semesters and their courses. The CGPA shall reflect the failed statusin case of F grade(s), till the course(s) is/are passed. When the course(s) is/are passedby obtaining a pass grade on subsequent examination(s) the CGPA

shall only reflect the new grade and not the fail grades earned earlier. The CGPA is calculated as:

$$CGPA = \begin{array}{c} C_1S_1 + C_2S_2 + C_3S_3 + C_4S_4 \\ \\ C_1 + C_2 + C_3 + C_4 \end{array}$$

where C_1 , C_2 , C_3 ,... is the total number of credits for semester I,II,III,... and S_1,S_2 , S_3 , is the SGPA of semester I,II,III,.....

20. Declaration of class

The class shall be awarded on the basis of CGPA as follows:

First Class with Distinction = CGPA of. 7.50 and above First Class = CGPA of 6.00 to 7.49 Second Class = CGPA of 5.00 to 5.99

21. Project work

All the students shall undertake a project under the supervision of a teacher in Semester III to IV and submit a report. 4 copies of the project report shall be submitted (typed & bound copy not less than 75 pages).

The internal and external examiner appointed by the University shall evaluate the project at the time of the Practical examinations of other semester(s). The projects shall be evaluated as per the criteria given below.

Evaluation of Dissertation Book:

Objective(s) of the work done	50 Marks
Methodology adopted	150 Marks
Results and Discussions	250 Marks
Conclusions and Outcomes	50 Marks

Total 500 Marks

Evaluation of Presentation:

Presentation of work	100 Marks
Communication skills	50 Marks
Question and answer skills	100 Marks

Total 250 Marks

22. Award of Ranks

Ranks and Medals shall be awarded on the basis of final CGPA. However, candidates who fail in one or more courses during the M.Pharm program shall not be eligible for award of ranks. Moreover, the candidates should have completed the M. Pharm program in minimum prescribed number of years, (two years) for the award of Ranks.

23. Award of degree

Candidates who fulfill the requirements mentioned above shall be eligible for award of degree during the ensuing convocation.

24. Duration for completion of the program of study

The duration for the completion of the program shall be fixed as double the actual duration of the program and the students have to pass within the said period, otherwise they have to get fresh Registration.

25. Revaluation I Retotaling of answer papers

There is no provision for revaluation of the answer papers in any examination. However, the candidates can apply for retotaling by paying prescribed fee.

26. Re-admission after break of study

Candidate who seeks re-admission to the program after break of study has to get the approval from the university by paying a condonation fee.