



**Regulatory Affairs
(Department of Pharmaceutics)
National Institute of Pharmaceutical Education and Research
(NIPER) - Raebareli
Bijnor-Sisendi Road, Sarojini Nagar,
Near CRPF Base Camp, Lucknow (U.P.) – 226002**

M.S. (Pharm.) Regulatory Affairs

Course No.	Course Name	Credits
Semester – I		
RA-510	Good Regulatory Practices	03*
RA-520	Regulatory Aspects of Drugs, Biologicals and Medical Devices	03*
PE-540	Regulatory Consideration for Pharm Development	01
GE-520	Fundamentals of Intellectual Property (IP) & Technology Management	01
GE-510	Biostatistics	02
RT-550	Introduction to Regulatory Toxicology	02
GE-511	Seminar	01
LG-510	Regulatory Aspects of Drugs, Phytopharmaceuticals, Biologicals and Medical Devices - Lab	07
Total Credits		20
Semester – II		
RA-610	Documentation and Regulatory Writing	02
RA-620	Clinical Research Regulations	02
PA-610	Pharmacopoeial Methods of Analysis	02
PA-630	Stability Testing	01
PA-640	Quality Control and Quality Assurance	02
RT-650	Good Regulatory Practice in Regulatory Toxicology	02
GE-611	Seminar	01
LS-610	Regulatory Aspects of Drug Formulations and Medical Devices	08
Total Credits		20
Semester – III		
TH-598	Synopsis	15
TH-599	Presentation	05
Total Credits		20
Semester – IV		
TH-698	Thesis	15
TH-699	Defence of Thesis	05
Total Credits		20
Grand Credits (I to IV Semesters)		80

*The course includes 2 credits for theory classes and 1 credit assigned for tutorial

M.S. (Pharm.) Regulatory Affairs

SEMESTER - I

RA 510 - Good Regulatory Practices (3* Credits)

1. **Introduction to good regulatory practices (GRP) and its importance, Coverage of GRP**
Good Manufacturing Practices (GMP): GMP principles focusing on US cGMP regulations outlined in Part 210 and Part 211, EC principles of GMP (Directive 91/356/EEC), WHO cGMP guidelines, GAMP-5, and IMDRF guidance documents. Quality systems in GMP. Schedule M. Data integrity
2. **Good Distribution Practices:** Introduction to GDP, Legal GDP requirements put worldwide, Principles, Personnel, Documentation, Premises, and Equipment, Deliveries to Customers, Returns, Self-Inspection, Provision of information, WHO GDP, USP GDP (Supply chain integrity), relevant CDSCO guidance and ISO standards. New Drug Approval Guidelines in India. Audit systems
3. **Clinical Research Related Guidelines:** Good Clinical Practice Guidelines (ICH GCP E6), Indian GCP Guidelines, ICMR Ethical Guidelines for Biomedical Research, CDSCO guidelines.
4. **Good Laboratory Practices (GLP):** Introduction to GLP, USFDA GLP Regulations, controlling the GLP inspection process, documentation, audit, goals of Laboratory Quality Audit, audit tools, and the future of GLP regulations along with relevant ISO and QCI Standards.
5. **Good Automated Laboratory Practices (GALP):** Introduction to GALP, principles, requirements, SOPs, training documentation, 21 CFR Part 11 compliance, general checklists, spreadsheets used in GLP, software evaluation checklists. Relevant ISO and QCI standards. Management of laboratory standards. Deviation management in GLP. Computer system validation
6. **Quality Management Systems:** Overview of ICH Q9 and Q10. Concept of Quality, Total Quality Management, Six Sigma concept, Out of Specifications (OOS) and out of trend (OOT), Deviations, change control. Validation: Types of validation, Types of qualification, Validation master plan (VMP).

Recommended Books:

1. Weinberg S. (Ed.). (2007). Good Laboratory Practice Regulations, NY: Marcel Dekker, (4th ed.). CRC Press.
2. Sharp J. (2004). Good Pharmaceutical Manufacturing Practice: Rationale and Compliance (1st ed.). CRC Press.
3. Willig SH, Stoker JR, Stockdale JM. (2008). Pharmaceutical Manufacturing Handbook, Regulations and Quality by. Wiley-Interscience, A John Wiley and Sons, Inc., Publications.
4. Berry IR, Martin RP. (Eds.). (2008). The Pharmaceutical Regulatory Process (2nd ed.). CRC Press.
5. Pisano DJ, Mantus DS. (Eds.). (2008). FDA Regulatory Affairs: A Guide for Prescription Drugs, Medical Devices, and Biologics (2nd ed.). CRC Press.
6. Sharp J. (2004). Good Pharmaceutical Manufacturing Practice: Rationale and Compliance (1st ed.). CRC Press.

7. Singer DC, Stefan RI, Van Staden JF. (2005). Laboratory Auditing for Quality and Regulatory Compliance (1st ed.). Taylor & Francis Publisher.
8. Anderson M. (2002). GLP Essentials: A Concise Guide to Good Laboratory Practice, 2nd Ed., CRC Press.

RA 520 - Regulatory Aspects of Drugs, Biologicals and Medical Devices (3* Credits)

1. Organizational structure and functions of FDA and EMA. Regulatory framework of ROW countries
Code of Federal Regulations; Federal, Food, Drug and Cosmetic Act (FFDCA), Manufacturing, packaging, and labelling requirements for drugs and biologicals in regulated markets.
Regulatory requirements for the approval and registration of API (including orphan drugs) and their products (single as well as fixed dose combinations), biologics and novel formulations/therapies; Registration of generic drugs (including their BA-BE assessments)
2. **Regulations for clinical studies:** Investigator brochure, Conducting clinical trials. Role of institutional review board/ ethics committees, informed consent process, study process, pharmacovigilance and monitoring of safety.
Post approval regulatory affairs: Nonclinical development; CTD formats; Dossier preparation; Liaison between the industry and regulatory agencies.
3. Brief overview of ICH guidelines (Q, S, E, M) in context to API/product development. Marketing authorization procedure in USA and EU; Mutual recognition procedure. Regulatory considerations for manufacturing, packaging, distribution and labelling of pharmaceuticals and biologicals.
4. Introduction to biologics and biosimilars, different biologic products, difference between biologic and small molecular drugs/products. Bio-similarity assessment
Laws, regulations, and guidelines for the development of biosimilar/biologic and approval of biologics and biosimilars (IND, PMA, BLA, NDA). Pre-clinical and clinical development
Principles for the development, manufacturing, distribution, commercialization of biologics in USA, EU and India; Post-marketing surveillance and pharmacovigilance on biologics, Case studies.
5. **Medical devices in diagnosis and therapeutics:** Definition, classification, global nomenclatures, essential principles and life cycle of medical devices. Drug-device combination and other hybrids
History of medical device regulation and regulatory approval of medical devices in US and European markets. Role of International Medical Device Regulators Forum (IMDRF). Existing regulations on registration of medical devices in India. An overview of chemical tests on medical device components
6. Good clinical practice in context to the investigation of medical devices; Safety and risks; Quality management system for medical devices, ISO 13485
International and national standards for medical devices; Future of medical devices in India
Assessment of quality, effectiveness and risks; Reporting of adverse events; Post-marketing surveillance

Recommended Books:

1. The Pharmaceutical Regulatory Process, Edited by Ira R. Berry and Robert P. Martin
2. Guidebook for drug regulatory submissions by Sandy Weinberg, John Wiley & Sons
3. Clinical Trials and Human Research: A Practical Guide to Regulatory Compliance by Fay A. Rozovsky and Rodney K. Adams
4. Preparation and maintenance of the IND application in eCTD Format by William K. Sietsema
5. FDA Regulatory Affairs: A Guide for Prescription Drugs, Medical Devices, and Biologics, Second edition by Douglas J. Pisano and David S. Mantus
6. Biological Drug Products: Development and Strategies; Wei Wang and Manmohan Singh; By Wiley, 2013
7. Guidelines on Similar Biologics: Regulatory Requirements for Marketing Authorization by the Department of Biotechnology, CDSCO, Ministry of Health and Family Welfare, Govt. of India
8. www.ema.europa.eu
9. Medical Product Regulatory Affairs: Pharmaceuticals, Diagnostics, Medical Devices, Second Edition by John J. Tobin and Gary Walsh
10. Compliance Handbook for Pharmaceuticals, Medical Devices and Biologics by Carmen
11. Medical Device Regulations: Global overview and guiding principles, World Health Organization.
12. Quality System Information - Guidance documents released by regulatory agencies
13. Leachables and Extractables Handbook: Safety Evaluation, Qualification, and Best Practices Applied to Inhalation Drug Products, Douglas J. Ball, Daniel L. Norwood, Cheryl L. M. Stults and Lee M. Nagao
14. Indian regulations on registration of medical devices.
15. Scientific journals

PE 540 - Regulatory considerations for Pharmaceutical Development - II (1 Credit)

1. **International regulatory trends in pharmaceutical industry.**
2. **Role of regulatory affairs department in pharmaceutical organization:** regulatory audits, interactions with various other departments, single point contact with regulatory agencies.
3. **Types of regulatory filings for pharmaceutical products:** goals of regulatory registration procedures investigational new drug applications, introduction to various type of regulatory filings.
4. **New drug applications:** stages involved in NDA, different phases of clinical trials, purpose of IND, types, and categories of IND applications information to be given in IND applications.
5. **Chemistry, manufacturing, and control (CMC) information in NDA:** information related to drug substance like manufacturing process, specifications, description of tests methods. Information related to drug product: description of method of manufacturing, specifications, and acceptable limits. Information related to placebo.
6. **Hybrid NDA:** a difference from NDA, historical background, literature-based hybrid NDAs and other sources of information for hybrid NDA, examples of types of products considered under hybrid NDA.
7. **Abbreviated New Drug applications (ANDAs):** historical developments leading to creation of ANDA process, Hatch Waxman Act, patent term restoration, criteria for patent term extension, various types of Hatch Waxman Exclusivities, concept of therapeutic equivalence, ANDA review process.

8. **Paragraph IV certification ANDAs:** different ANDA patent certification options, Medicare Modernization Act, implications of this act on 30 months stay period and 180-day exclusivity, triggering and forfeiture of 180-day exclusivity, shared exclusivity.
9. **ANDA with suitability petition:** case studies of drug products considered appropriate for filing under suitability petition.

GE 520 - Fundamentals of Intellectual Property (IP) and Technology Management (1 Credit)

1. **Intellectual property:** Concepts and fundamentals; Concepts regarding intellectual property (IP), intellectual property protection (IPP) and intellectual property rights (IPR); Economic importance, mechanisms for protection of intellectual property-patents, copyrights, trademark; Factors effecting choice of IP protection; Penalties for violation; Role of IP in pharmaceutical industry; Global ramifications and financial implications.
2. **Trade related aspects of intellectual property rights:** Intellectual property and international trade; Concept behind WTO (World Trade Organization), WIPO (World Intellectual Property Organization), GATT (General Agreement on Tariff and Trade), TRIPs (Trade Related Intellectual Property Rights), TRIMS (Trade Related Investment Measures), and GATS (General Agreement on Trade in Services); Protection of plant and animal genetic resources; Biological materials; Gene patenting; Biotechnology / drug related IPR issues; Status in India and other developing countries; Case studies and examples; TRIPS issues on herbal drugs.
3. **Nuts and bolts of patenting, copyright and trademark protection criteria for patentability, types of patents; Indian Patent Act, 1970; WTO and modifications under TRIPS:** Filing of a patent application; Precautions before patenting-disclosures / nondisclosures, publication-article / thesis; Prior art search-published patents, internet search patent sites, specialized services-search requests, costs; Patent application-forms and guidelines, fee structure, time frames, jurisdiction aspects; Types of patent applications provisional, non-provisional, PCT and convention patent applications; International patenting requirement procedures and costs; Financial assistance for patenting- introduction to schemes by NRDC and TIFAC; Publication of patents-gazette of India, status in Europe and US; Patent annuity; Patent attorneys technical aspects, criteria for selection, addresses, fee, rights and responsibilities of a patentee; Practical aspects regarding maintaining of a PATENT FILE; Patent infringement- meaning, scope, litigation, case studies and examples; Patenting by research students, lecturers and scientists-University / organizational rules in India and abroad; Thesis research paper publication, credit sharing by workers, financial incentives; Useful information sources for patents related information-internet sites, brochures, periodicals, CD roms; Significance of copyright protection for researchers; Indian Copyright Law and digital technologies-Berne convention, WIPO copyright treaty (WCT), WIPO performance and Phonogram Treaty (WPPT); Protection for computer data bases, multimedia works; Trademarks legislation and registration system in India-an introduction, meaning of trademark criteria for eligibility; filling application for trademark registration; Trade secrets-scope modalities and protection; Case studies-drug related patents infringements.
4. **Technology development / transfer / commercialization related aspects:** Technology development-meaning; Drug related technology development; Toxicological studies, bioequivalence (BU), clinical trials-phase-I, phase-II and phase-

III; Approved bodies and agencies; Scale-up, semi-commercialization and commercialization-practical aspects and problems; Significance of transfer of technology (TOT), bottlenecks; Managing technology transfer-guidelines for research students, scientists and related personal; TOT agencies in India-APCTD, NRDC, TIFAC, BCIL, TBSE/SIDBI; TOT related documentation-confidentiality agreements, licensing, MOUs, legal issues; Compulsory licensing excess to medicine issues; DOHA declaration, POST WTO product patent regime from 2005; Challenges for Indian pharmaceutical industry in the context of globalization of IP; Drug registration and licensing issues-national and global; Drug master file submissions, SOPs; Related registration and marketing issues; Case studies-antiretroviral drugs and others.

5. Funding sources for commercialization of technology: Preparation of a project report, financial appraisal, business models; GOI schemes and incentives; NRDC, TePP, HGT, TDB schemes. PATSER; Venture capitalists, banks. Incubator concept-Case studies with respect to IIT, CCMB, IMTECH, and NIPER. Documentation and related aspects.

6. Ethics and values in IP: IP and ethics-positive and negative aspects of IPP; Societal responsibility; Avoiding unethical practices; Echo-responsibility-economic, social, and environmental benefits of modern biotechnology; Voluntary adoption of pollution control strategies.

Recommended Books:

1. Law Relating to Intellectual Property by B.L. Wadhera
2. IPR Handbook for Pharma Students and Researchers by P.Bansal
3. The Patents Act, 1970 (Bare Act with Short Notes) (New Delhi: Universal Law Publishing Company Pvt. Ltd. 2012)
4. Patent Agent Examination by Sheetal Chopra and Akash Taneja
5. Making Innovation Happen- A simple and Effective Guide to Turning Ideas into Reality by Michael Morgan
6. Making Breakthrough Innovation Happen by Porus Munshi
7. Innovation X- Why a Company's Toughest Problems are its Greatest Advantage by Adam Richardson
8. Legal Drafting for the Layman by Nabhi Kumar Jain
9. How to Write and Publish a Scientific Paper by Rober A Day
10. Concise Law Dictionary-with Legal Maxims, Latin Terms and Words and Phrases by Justice Y.V. Chandrachud
11. Biomedical Research- From Ideation to Publication by G. Jagadeesh and others

RT 550 - Introduction to Regulatory Toxicology (2 Credits)

- 1. Drug discovery and development:** Drug Laws, Food and Drug Administration (FDA), Environmental Protection Agency (EPA), Occupational Safety and Health Administration (OSHA), Organization for Economic Co-operation and Development (OECD), International Council for Harmonization of Technical Requirements for Pharmaceuticals for Human Use (ICH), Central Drugs Standard Control Organization (CDSCO).
- 2. New Drugs and Clinical Trials Rules, 2019:** Salient features including definitions, Ethics Committees (EC), registration of clinical studies and biomedical and health research, academic clinical trials, and role of ECs in compensation.
- 3. Overview of guidelines:** OECD Guidelines for the Testing of Chemicals (Section 4 - Health Effects), ICH guidelines (ICH Q3A to ICH Q3D, ICH S1 to ICH S12, ICH M3,

ICH M4, ICH M7), ISO 10993 guidelines brief overview, Introduction to ISO 10993 standards that govern biocompatibility and toxicity testing of medical devices, Medical Devices Rules, 2017 – CDSCO, Guidelines on Umbilical Cord Blood Banking, Guidelines for Stem Cell Research, Framework for the Regulation of Regenerative Medicine - stem cell and exosome based therapeutics.

4. **Drug discovery and registration:** Regulatory affairs, WTO, patent regime, accreditation, and harmonization process.
5. **Threshold limitations:** Hormesis and dose-response relationship, lower dose extrapolation.
6. **Concepts in Risk assessment:** Threshold, Determining the Point of departure (i.e., NOAEL, NOEL, LOAEL, LOEL, BMD, LD50, TD50, TTC, MABEL), MTD, Biocompatibility, Margin of safety, Margin of exposure, Weight-of-Evidence (WOE) Approach, Clinical risk/benefit analysis.
7. **Animal to human dose extrapolation:** NOAEL determination, human equivalent dose calculation, appropriate species selection and application of Uncertainty/Safety factors.
8. **Flow chart:** “Case by Case” basis in non-clinical development and its influences on safety assessment, usefulness and limitations Models and bioassay: Methods in toxicity testing, dose-response characterization.
9. **Regulations of human pharmaceuticals:** FDA approval pathways and preclinical safety evaluation of biotechnology-derived pharmaceuticals.
10. **Influence of new technologies:** Discovery development gap, future of drug safety.

Recommended Books and Websites:

1. Regulatory Toxicology by Shayne C. Gad Taylor & Francis
2. Principles and Methods of Toxicology by A. Wallace Hayes
3. <https://www.fda.gov/regulatory-information/search-fda-guidance-documents>
4. <https://cdsco.gov.in/>
5. <https://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfStandards/search.cfm>
6. <https://www.ich.org/>
7. <https://www.aami.org/standards/>
8. <https://www.fda.gov/Medical-Devices>
9. <https://www.iso.org/standard/68936.html>
10. <https://www.fda.gov/vaccines-blood-biologics/cellular-gene-therapy-products>
11. <https://main.icmr.nic.in/content/draft-guidelines-umbilical-cord-blood-banking-202>

GE 510 - Biostatistics (2 Credits)

1. **Statistics:** Introduction, its role and uses. Collection; Organization; Graphics and pictorial representation of data; Measures of central tendencies and dispersion. Coefficient of variation.
2. **Probability:** Basic concepts; Common probability distributions and probability distributions related to normal distribution.
3. **Sampling:** Simple random and other sampling procedures. Distribution of sample mean and proportion.
4. **Estimation and Hypothesis Testing:** Point and interval estimation including fiducial limits. Concepts of hypothesis testing and types of errors. Student- t and Chi square tests. Sample size and power.

5. **Experimental design and analysis of variance:** Completely randomized, randomized blocks. Latin square and factorial designs. Post- hoc procedures.
6. **Correlation and regression:** Graphical presentation of two continuous variables; Pearson's product moment correlation coefficient, its statistical significance. Multiple and partial correlations. Linear regression; Regression line, coefficient of determination, interval estimation and hypothesis testing for population slope. Introduction to multiple linear regression model. Probit and logit transformations.
7. **Non-parametric tests:** Sign; Mann-Whitney U; Wilcoxon matched pair; Kruskal Wallis and Friedman two-way ANOVA tests. Spearman rank correlation.
8. **Statistical techniques in pharmaceuticals:** Experimental design in clinical trials; Parallel and crossover designs. Statistical test for bioequivalence. Dose response studies; Statistical quality control.

Recommended Books:

1. Fundamentals of Biostatistics by Bernard Rosner
2. Pharmaceutical Statistics: Practical and Clinical Applications by Bolton and Bon
3. Statistical Misconceptions by Huck

LG 510 - Regulatory Aspects of Drugs, Herbals, Biologicals and Medical Devices - Lab (7 Credits)

1. Presentation and assignments on rules, guidelines and Indian standards related to herbal products
2. Case studies on regulatory procedures for herbal and biological products (type of tests/trials, assembling the data, overall content, format, stability requirements, ethical guidelines for pre-clinical and clinical studies, etc.)

GE 511 - Seminar (1 Credit)

1. Introduction, information retrieval systems
2. Writing term papers and reports
3. Organization of scientific material, thesis, dissertation and references
4. Reading research papers and skill in oral presentation

M.S. (Pharm.) Regulatory Affairs

SEMESTER - II

RA 610 – Documentation and Regulatory Writing (2 Credits)

- 1. Documentation in pharmaceutical industry:** Exploratory Product Development Brief (EPDB) for Drug substance and Drug product, Product Development Plan (PDP), Product Development Report (PDR). packaging development report, Trade dress, Dissolution method development report, Analytical method development and validation, Genotoxic and nitrosamine risk assessment report, ICH Q8.
Relevant guidelines of US-FDA and EU guidelines and differences therein.
- 2.** Master Formula Record, Batch Manufacturing Record and its calculations, Batch Reconciliation, Batch Packaging Records, Print pack specifications, Distribution records, Product specification. Certificate of Analysis (CoA).
- 3. Description of 505(b)(2) terminologies**
Dossier preparation and submission: Introduction and overview of dossiers, contents and organization of dossier and sections, compilation and review of dossier. Paper submissions, overview and modules of CTD, electronic CTD submissions;
Electronic submission: Planning electronic submission, requirements for submission, regulatory bindings and requirements, Tool and Technologies, electronic dossier submission process and validating the submission, Electronic Submission Gateway (ESG). None CTD electronic submissions (NeeS), Asian CTD formats (ACTD) submission.
Organizing, process and validation of submission. Submission in Sugam system of CDSCO.
- 4. Audits:** Introduction, Definition, Summary, Types of audits, GMP compliance audit, Audit policy, Internal and External Audits, Second Party Audits, External third-party audits, Auditing strategies, Preparation and conducting audit, Audit analysis, audit report and compliance, audit follow up. Auditing/inspection of manufacturing facilities by regulatory agencies. Timelines for audits/inspection. GHTF study group 4 guidance document. ISO 13485.
- 5. Inspections:** Pre-approval inspections, Inspection of pharmaceutical manufacturers, Inspection of drug distribution channels, Quality systems requirements for national good manufacturing practice inspectorates, inspection report, model certificate of good manufacturing practices, Root cause analysis, Corrective and Preventive action (CAPA).
- 6. Product life cycle management:** Prior Approval Supplement (PAS), Post Approval Changes [SUPAC] for immediate and modified release dosage forms, Changes Being Affected in 30 Days (CBE-30), Annual Report, Post marketing Reporting Requirements, Post approval Labelling Changes, Lifecycle Management, FDA Inspection and Enforcement, Establishment Inspection Report (EIR), Warning Letters, Recalls, Seizure and Injunctions. ISO Risk Management Standard. ICH Q12

Recommended Books:

1. Compliance auditing for Pharmaceutical Manufacturers. Karen Ginsbury and Gil Bismuth, Interpharm/CRC, Boca Raton, London New York, Washington D.C.
2. Pharmaceutical Manufacturing Handbook, Regulations and Quality by Shayne Cox Gad. Wiley- Interscience, A John Wiley and sons, Inc., Publications.
3. Handbook of microbiological Quality control. Rosamund M. Baird, Norman A. Hodges, Stephen P. Denyar. CRC Press. 2000.
4. Laboratory auditing for quality and regulatory compliance. Donald C. Singer, Raluca-loana Stefan, Jacobus F. Van Staden. Taylor and Francis (2005).
5. Implementing Juran's Road Map for Quality Leadership: Benchmarks and Results, By Al Endres, Wiley, 2000
6. Understanding, Managing and Implementing Quality: Frameworks, Techniques and Cases, By Jiju Antony; David Preece, Routledge, 2002
7. Organizing for High Performance: Employee Involvement, TQM, Reengineering, and Knowledge Management in the Fortune 1000: The CEO Report By Edward E. Lawler; Susan Albers Mohrman; George Benson, Jossey-Bass, 2001
8. Corporate Culture and the Quality Organization By James W. Fairfield- Sonn, Quorum Books, 2001
9. The Quality Management Sourcebook: An International Guide to Materials and Resources By Christine Avery; Diane Zabel, Routledge, 1997
10. The Quality Toolbox, Second Edition, Nancy R. Tague, ASQ Publications
11. Juran's Quality Handbook, Sixth Edition, Joseph M. Juran and Joseph A. De Feo, ASQ Publications
12. Root Cause Analysis, The Core of Problem Solving and Corrective Action, Duke Okes, 2009, ASQ Publications
13. International Medical Device Regulators Forum (IMDRF) Medical Device Single Audit Program (MDSAP)

RA 620 – Clinical Research Regulations (2 Credits)

1. **Clinical drug development process:** Types of clinical studies, Clinical trial protocol, Phases of clinical trial, Phase 0 studies, Phase I and subtype studies (single ascending, multiple ascending, dose escalation, methods, food effect studies, drug – drug interaction, PK end points), Phase II studies (proof of concept or principle studies to establish efficacy), Phase III studies (Multi ethnicity, multinational, registration studies), Phase IV studies (Post marketing authorization studies; pits and practices)
2. **Ethics in clinical research:** Historical Perspectives (Nuremberg Code study, Thalidomide study, Nazis Trials, Tuskegee Syphilis Study, The Belmont Report, The declaration of Helsinki, Origin and Principles of International Conference on Harmonization – Good Clinical Practice (ICH-GCP) guidelines, The ethics of randomized clinical trials, The role of placebo in clinical trials, Ethics of clinical research in special population, Institutional Review Board/Independent Ethics Committee/Ethics Committee, Data safety monitoring boards, Responsibilities of sponsor, CRO, and investigator in ethical, Conduct of clinical research, Informed Consent Process (Ethical principles governing informed consent process, Structure and content of Patient Information Sheet and Informed Consent Form, The informed consent process and documentation
3. **Regulations governing Clinical Trials**
India: Clinical Research regulations in India – Schedule Y & Medical Device Guidance
USA: Regulations to conduct drug studies in USA (FDA)
EU: Clinical Research regulations in European Union (EMA)

4. **Clinical Research Related Guidelines:** Good Clinical Practice Guidelines (ICH GCP E6), Indian GCP Guidelines, ICMR Ethical Guidelines for Biomedical Research, CDSCO guidelines
5. **Types and Designs used in Clinical Research**
Types of research designs based on Controlling Method (Experimental, Quasi experimental, and Observational methods) Randomization techniques (Simple randomization, restricted randomization, blocking method and stratification), Time Sequences (Prospective and Retrospective), Sampling methods (Cohort study, case Control study and cross-sectional study), Health outcome measures (Clinical & Physiological, Humanistic, and economic)
6. **Regulations for the Development and Clinical Trials of Biosimilars and Medical Devices:** Biosimilars (Guidelines and Regulations, Development and Quality aspects, Safety and Efficacy, Clinical Development, Principles for Development of Biosimilars, Data requirements for Preclinical studies and Clinical trial application); Medical Devices (Regulations and Research Phases in the lifespan of a medical devices, Conducting clinical trials on medical devices, Medical devices regulations across the world)

Recommended Books:

1. Handbook of clinical research. Julia Lloyd and Ann Raven Ed. Churchill Livingstone
2. Principles of Clinical Research edited by Giovanna di Ignazio, Di Giovanna and Haynes
3. Recent Central Drugs Standard Control Organization. Good Clinical Practices-Guidelines for Clinical Trials on Pharmaceutical Products in India. New Delhi: Ministry of Health;2013, 2017.
4. International Conference on Harmonization of Technical requirements for registration of Pharmaceuticals for human use. ICH Harmonized Tripartite Guideline. Guideline for Good Clinical Practice.E6; May 1996.
5. Ethical Guidelines for Biomedical Research on Human Subjects 2000, 2014, 2017. Indian Council of Medical Research, New Delhi. 4. Textbook of Clinical Trials edited by David Machin, Simon Day and Sylvan Green, March 2005, John Wiley and Sons.
6. Clinical Data Management edited by R K Rondels, S A Varley, C F Webbs. Second Edition, Jan 2000, Wiley Publications
7. Drug Discovery and Clinical Research by SK GUPTA, Jaypee Brothers Medical Publishers (p) Ltd. Second Edition New Delhi

PA 610 – Pharmacopoeial Methods of Analysis (2 Credits)

1. Detailing of ICH Q3 and correlation with Pharmacopeia.
2. Detailing of ICH Q4: Pharmacopoeial Harmonization; Evaluation and recommendation of Pharmacopoeial texts for use in the ICH regions.
3. Study of various pharmacopeias - IP, BP, EP, and USP
4. Pharmacopoeia General notices and their usage.
5. **Pharmacopeia General Chapters:** Chromatography, Spectroscopy, Elemental Impurities, Packaging, Residual solvents, Water determination, LOD.
6. CDSCO guideline for phytopharmaceutical, Ayurvedic Pharmacopoeia of India.
7. Detailing of dietary supplementary-general chapters.
8. Introduction to the pharmacopoeia forum
9. **Microbiological tests and assays as per Pharmacopeia:** Antimicrobial (preservative) effectiveness testing, microbial limit tests, sterility test, vitamins assay (zone of

exhibition), antibiotics assays, bacterial endotoxin test. Pharmacopeia General Chapters of microbiology.

10. **Reference standards:** Regulatory and pharmaceutical perspectives along standardization procedures.

Recommended Books:

1. The Indian Pharmacopoeia, Indian Pharmacopoeia Commission, Ghaziabad.
2. The British Pharmacopoeia, Stationary Office British Pharmacopoeia Commission, London
3. The United States Pharmacopeia-National Formulary, Board of Trustees, Rockville
4. The European Pharmacopoeia, Directorate for Quality of Medicines of the Council of Europe.

PA 630 – Stability Testing (1 Credit)

1. **Drug development cycles and stability testing:** Role and types of stability studies during different stages of drug and product development.
2. **Drug stability testing guidelines:** International, Regional, and National drug stability guidelines.
3. **WHO vs. ICH drug stability testing guidelines:** Comparison of different aspects in WHO guideline, and critical comparison with ICH parent guideline Q1A(R2).
4. **Specific discussion on following ICH guidelines:** Q1B, Q1C, Q1D, Q1E and Q5C.
5. **Additional topics:**

Stress testing and stability-indicating method development: Role, regulatory aspects, protocols/approaches, practical considerations

Stability testing of phytopharmaceuticals: Regulatory requirements

Stability test equipment: Types of stability chambers (walk-in, stand-alone), design considerations, qualification, and other critical issues.

Stability testing for Shipping & Distribution: Stability testing during transport

Stability testing of drug delivery systems

Recommended Books:

1. ICH (www.ich.org) and WHO (www.who.int) guidelines
2. Pharmaceutical Stress Testing (Predicting Drug Degradation) by Steven Baertschi and Karen Alsante
3. Drug Stability (Principles and Practices) by S. James, J.T. Carstensen
4. Stability-indicating HPLC Methods for Drug Analysis by Quanyun A. Xu, Lawrence A. Trissel
5. Stability of Drugs and Dosage Forms by Sumic Yoshioka, Valentino Stella
6. Physical Pharmacy and Pharmaceutical Sciences by Patrick Sinko, Alfred Martin
7. New Drug Approval Process (Chapter 7) by Richard Guarino
8. Handbook of Stability Testing in Pharmaceutical Development: Regulations, Methodologies, and Best Practices by Kim Huynh-Ba
9. Stability and Characterization of Protein & Peptide of Drugs by Y. John Wang
10. Peptide and Protein Drug Analysis by Ronald Reid

PA 640 – Quality Control and Quality Assurance (2 Credits)

1. **Good manufacturing practices [Schedule M] and Good laboratory practices [Schedule L-I]:** Their applications to the pharmaceutical industry.

2. **Basic principles and concepts of quality management:** Quality control, quality assurance, quality auditing, ISO system, electronic quality management system (eQMS).
3. Control of raw & packaging material and labelling, sampling, testing, release and distribution of finished products.
4. **Document control:** Preparation, review, approval, issuance, storage and retrieval (e.g., master manufacturing and packaging records, site master file, etc.), electronic document management system (e-DMS).
5. **Standard operating procedures:** SOP on SOPs, Change control procedure, annual product review/product quality review, handling of deviations & non-conformity, corrective & preventive actions (CAPA), handling of laboratory incidents and OOS and OTT test results.
6. **Qualification of facility and utilities:** Concepts of facility validation, qualification of HVAC and water systems.
7. Process validation, product change over, basic requirements of cleaning and its validation.
8. Technology transfer from R&D to manufacturing, including product life-cycle approach.
9. Handling of market complaints, recalls and returned goods.
10. Quality risk management in production area, data integrity management.
11. Introduction to concepts of QbD, PAT and continuous manufacturing.

Recommended Books:

1. Quality Assurance of Pharmaceuticals: A Compendium of Guidelines and Related Materials, Volume 1.
2. Quality Assurance of Pharmaceuticals: A Compendium of Guidelines and Related Materials, Volume 2.
3. Q.A. Manual by D. H. Shah.
4. Good Pharmaceutical Manufacturing Practice: Rationale and Compliance by John Sharp.
5. WHO Expert Committee on Specifications for Pharmaceutical Preparations.
6. Handbook of Pharmaceutical Quality Assurance by Dr. Premnath Shenoy.

LS 610 – Regulatory Aspects of Drug Formulations and Medical Devices (8 Credits)

1. Presentation and assignments on rules, guidelines and Indian standards on nutraceuticals and phytopharmaceuticals, GxP
2. Presentation and assignments on rules, guidelines and Indian standards on OTC products
3. Regulatory procedures for IP on phytopharmaceuticals, cosmeceuticals and nutraceuticals (type of tests, overall content, format, stability tests and other requirements)
4. Guidelines related to the advertising and sale of nutraceuticals and supplements
5. Certification requirements for the manufacturers of nutraceuticals
6. Assignment on preparation of documents as per guidelines
7. Preparation of SOPs and reports (covering the stability profile, validation, etc.)
8. Preparation of CTD
9. Addressing the warning letters and letter of concerns
10. Corrective and preventive actions, and their documentations
11. Clinical trial procedures

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| <ol style="list-style-type: none">12. Any other topic of industrial interest13. Industrial visit and report preparation |
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GE 611 – Seminar (1 Credit)

Students are expected to deliver a Seminar on the project to be pursued in Semesters III & IV. The seminar should include the purpose and basis of the project stating the aims, objectives and probable outcomes, literature review towards the project and method of its execution.

RT 650 – Good Laboratory Practice in Regulatory Toxicology (2 Credits)

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| <ol style="list-style-type: none">1. Good Laboratory Practices (GLP).2. Organogram, Management, Quality control and Quality Assurance.3. SOP writing and implementation: GLP Establishment.4. Historical control data: Importance of the generation of quality data, background lesions, and Use of suitable animal models in toxicity evaluation.5. Study plans: Study protocols.6. Master schedule: Responsibility of study directors.7. Multisite studies and principles investigators responsibility.8. Reporting of study results.9. Storage and retention of records and materials.10. GLP audits and inspections.11. Cost-benefit comparisons in regulatory setups |
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Recommended Books:

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| <ol style="list-style-type: none">1. Good Laboratory Practice, 2nd Edition, by Jurg P Seiler, Springer2. WHO/TDR Manual for Good Laboratory Practice, WHO/TDR, Geneva, Switzerland |
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