Annexure VIII



UNIVERSITY OF KERALA

POST GRADUATE DIPLOMA IN MOLECULAR DIAGNOSTICS

Scheme and Syllabus w.e.f. 2015 admission

Post Graduate Diploma in Molecular Diagnostics

2015

About the course

Molecular Diagnostics are the tools based on the principles of Molecular Diagnosis. It is the process of identifying a disease by understanding the molecules, such as proteins, DNA, and RNA, in a tissue or fluid, which forms the markers of the diseases directly or indirectly. Molecular diagnostics is a new discipline that captures genomic and proteomic expression patterns and uses the information to distinguish between two or more conditions at the molecular level. The conditions under investigation can be human genetic disease or infectious diseases. Molecular diagnostics is not confined to human diseases but can be used in animals and plants also. Molecular diagnostics can also be used to identify foodstuffs, vegetables, meet types and food processing methods etc., and can be also used in environmental monitoring, detect the presence of specific microorganisms in various samples including food materials. The course will provides the theory and use of molecular techniques in the diagnostic techniques, with more importance on nucleic acids based techniques. Common analytical techniques and molecular techniques related to the development and use of diagnostics such as polymerase chain reaction (PCR), quantitative real time PCR (qRT-PCR), microarray analysis, chromosomal techniques, hybridization techniques and DNA bioinformatic tools will be emphasized. The laboratory exercises are designed to provide a hands-on context for the topics being presented in the syllabus.

Objectives of the Course

The course focuses on learning and understanding how the various molecular techniques that were studied in other classes can be developed and utilized in diagnosis and sold in diagnostic kits. Many of the diagnostic techniques are developed and marketed as Diagnostic Kits by biotechnology companies. The main source of information is web sites of such companies that develop and market the molecular diagnostic kits, which they continuously develop.

Eligibility for admission

A first class with 60% M. Sc., Degree in Biotechnology, Botany, Zoology, Biochemistry, Microbiology, Genomic Sciences, Agriculture, Veterinary Sciences, Fisheries Sciences, Pharmacy or M.Tech., in Biotechnology, or Post graduate degree in a branch of Life Sciences, Medical Sciences or equivalent. A minimum of 50 % marks or equivalent for SC/ST candidates.

Mode of Selection

Selection will be made based on the basis of an National entrance test conducted by the University and Reservation Policy stipulated by the Government from time to time. A total of 10 students will be selected each year.

Course structure and Duration

The course consists of two semesters and shall extend over a period of 12 months, including the examinations.

5. Examination and Evaluation

The conductance of the examinations and evaluations will be as per the general regulations of the University adopted for the PG Diploma Courses. Each Theory course carries 3 credits and practical course carries 5 credits.

For each course, 40% of the total marks will be allotted to continuous assessment or internal assessment, and 60% for End semester examination by the University. The internal assessment (CA) for each course will be carried out by the concerned faculty member or the course instructor.

The break up for Continuous Assignment will be as follows:

a) For Theory Paper

Mid Sem Examination 15 Marks
Seminar / Tests / Quiz 10 Marks
Assignment 10 Marks
Attendance 05 Marks
Total 40 Marks

b) For Practical Course

Mid Sem Examination 15 Marks

Performance of experiments And Results 10 Marks

Viva 10 Marks

Maintenance of records 05 Marks

Total 40 Marks

Project Work evaluation

a) Project Presentation and Viva 100 Marks (50+ 50)

c) Dissertation 100 Marks

Total 200 Marks

Project Presentation, Viva and evaluation of the Dissertation will be done by a committee composed of all Faculty members of the course Evaluation and External Examiner.

Declaration of final results

- 1. The marks obtained in each course, CA (Internal) and ESA (Final University Examination) will be entered separately as general procedures of the University.
- 2. A candidate shall be declared to have passed in each course if he/she secures not less than 40% marks in the ESA (University Examination) and 40% marks in the CA (Internal Assessment) and not less than 50% in the aggregate, taking CA and ESA marks together.
- 3. The final results will be based on total marks obtained by adding CA and ESA together for each Candidates. Successful candidates passing the examinations and earning 80 % or above shall be declared to have passed in First Class with distinction and those who earned marks 60 % or above will be declared to have passed in First class. Those who have scored below 60 % shall be declared to have passed in Second Class. Candidates earning Marks below 50 will be declared Failed.

Course Structure and Scheme of evaluation

S1.	Course	Course	Credits		Marks	
No.	Code	Title		CA	ESA	Total
1	MD 511	Introduction to Molecular Diagnostics	3	40	60	100
2	MD 512	Human Genetics and Genomics	3	40	60	100
3	MD 513	Biochemistry, Molecular Biology and Immunology in Diagnosis	3	40	60	100
4	MD 514 (MD 521)	Molecular diagnostics- I (Nucleic acid and Protein based diagnostics)	5	40	60	100
5	MD 515 (MD 522)	Molecular diagnostic II (Genetic & Cytologic techniques)	5	40	60	100
6	MD 523	Project Dissertation	6			100
7	MD 524	Viva Voce				100
		Total	25			700

Detailed Syllabus

Semester I

MD 511 Introduction to Molecular Diagnostics

Module 1

- · Introduction and History of diagnostics, Diseases- infectious, physiological and metabolic errors, genetic basis of diseases, inherited diseases. Infection mode of transmission in infections, factors predisposing to microbial pathogenicity, types of infectious diseases- bacterial, viral, fungal, protozoans and other parasites.
- Philosophy and general approach to clinical specimens, Sample collection- method of collection, transport and processing of samples, Interpretation of results, Normal microbial flora of the human body, Host - Parasite relationships.

Module 2

- Traditional disease diagnosis methods and tools diagnosis of infection caused by *Streptococcus*, *Coliforms*, *Salmonella*, *Shigella*, *Vibrio*, *andMycobacterium*., Diagnosis of fungal infections. Major fungal diseases: Dermetophytoses, Candidiosis and Aspergillosis.
- · Diagnosis of DNA and RNA viruses- Pox viruses, Adenoviruses, Rhabdo Viruses, Hepatitis Viruses and
- Retroviruses.
- Diagnosis of Protozoan diseases: Amoebiosis, Malaria, Trypnosomiosis, Leishmaniasis. Study of helminthic diseases- *Fasciola hepatica* and *Ascaris lumbricoides*. Filariasis and Schistosomiasis.

Module 3

· Major Metabolic disorders and its causes. Traditional methods for the diagnosis of metabolic errors. Disease due to genetic disorders - Identifying human disease genes.

- · Cancer- different types of cancers, genetics of cancer- oncogenes, tumour suppressor genes. Methods available for the diagnosis of genetic diseases and metabolic disorders.
- · Genetic disorders- Sickle cell anemia, Duchenne muscular Dystrophy, Retinoblastoma, Cystic Fibrosis and Sex linked inherited disorders.
- Neonatal and Prenatal disease diagnostics. Gender identification using amelogenin gene locus. Amplification of Y chromosome specific Short Tandem Repeats (Y-STR). Analysis of mitochondrial DNA for maternal inheritance. Molecular diagnosis for early detection of cerebral palsy, Down syndrome etc.
- Blood (formation, composition, function and pathology of blood disorders (haemoglobinopathies, sickle cell anemia, hemophilia), Muscle disorders (Duchene muscular dystrophy-DMD, Becker's muscular dystrophy-BMD, spinal muscular atrophy-SMA), Bone disorders (Osteogenesis imperfecta, Rheumatoid arthritis), Skin disorder (Albinism), Eye disorder (Retinitis pigmentosa)

References

- 1. Medical Microbiology (1997), Edited by Greenwood, D, Slack, R and Peutherer, J, ELST Publishers.
- 2. Parasitology (1997), Chatterjee K.D, Chatterjee Medical Publishers.
- 3. Bailey & Scott's Diagnostic Microbiology (2002), Betty A. Forbes , Daniel F. Sahm, Alice S. Weissfeld , Ernest A. Trevino, Published by C.V. Mosby
- 4. Jawetz, Melnick, & Adelberg's Medical Microbiology (2004), Geo F. Brooks, Stephen A. Morse, Janet S. Butel.
- 5. Fundamentals of Molecular Diagnostics (2007). David E. Bruns, Edward R. Ashwood, Carl A. Burtis. Saunders Group.
- 6. Henry's Clinical Diagnosis And Management By Laboratory Methods (2007) Mcpherson
- 7. Molecular Diagnostics: Fundamentals, Methods & Clinical applications (2007). Lele Buckingham and Maribeth L. Flaws
- 8. Molecular Diagnostics for the Clinical Laboratorian 2Ed. 2006, W.B. Coleman. Humana Press.
- 9. Molecular Pathology in Clinical Practice (2007). D. G. B. Leonard.
- 10. Microbial Functional Genomics (2004) by J.Zhou, D.K. Thomson. Y.Xu. J.M. Tiedje. J.Wiley & Sons Publishers.
- 11. Expert Review of Molecular Diagnostics

Semester 1

MD 512 Human Genetics and Genomics

Module 1

- Principles of inheritance- Mendelian and Non-Mendelian inheritance, Polygenic inheritance, Multifactorial trait, Threshold trait, Genetic Susceptibility & risk factors, Mitochondrial genome and disorders.
- Medical Genetics: Human genome Project, Genome Organization, Genome Annotations and databases, Identifying human disease genes. Genetic markers for diseases. (microsatellites, SNPs), Pharmacogenomics, Ecogenomics, Metabolomics, Teratogenetics

Mapping and identification of disease genes (linkage analysis, LOD score, association study)
 SNPs in diagnostics

Module 2

- · Human Cytogenetics Chromosomal basis of inheritance, sex chromosome, X-chromosome inactivation, Basics of cell culture, Techniques of cell cultures (short term lymphocyte, primary and secondary cell cultures, maintenance of cell lines) Techniques of chromosome analysis
- · Chromosome preparation from cultured lymphocytes, cell lines and solid tumors, Karyotyping, C-,G-banding and fluorescence banding, nomenclatures of bandings, *In-situ* hybridization techniques FISH and GISH,
- Meiotic chromosomes- Chromosomal anomalies and disorders, Numerical variations (polyploidy, aneuploidy, autosomal, sex-chromosomal) Structural variations (deletion, duplication, translocation, inversion, isochromosome, ring chromosome) Chromosomal abnormalities in cancer.

Module 3

- · Genetic disorders: Classification of genetic disorders, Single gene Disorders Sickle cell anaemia, Duchenne muscular Dystrophy, Retinoblastoma, Cystic Fibrosis and Marfan's syndrome Multifactorial disorders - Diabetes, Atherosclerosis, Schizophrenia
- · Sex linked inherited disorders.
- · Cancer genetics- Molecular basis of cancer, oncogenes, tumour suppressor genes. Gene therapy and other molecular based therapeutic approaches.. Genetic Counseling, Ethical and legal issues in genetic counseling

Module 4

- Molecular Techniques for diagnosis PCR- RFLP, ARMS-PCR, ELISA, Multiplex-PCR, SSCP, CSGE, DGGE, DHPLC, MALDI-TOF, DNA Sequencing
- Disease identification and Genetic tests for following disorders- Thalassemia, Fanconi anemia, Sickle Cell anemia, Fragile-X syndrome, Alzheimer's disease, Duchenne Muscular Dystrophy/ Becker's Muscular Dystrophy, Huntington's disease
- · Allelic susceptibility test for multifactorial disorders (Neural Tube Defect, Cleft Lip and Palate, Cardio Vascular Disorder, Male infertility)
- Risk evaluation (Mendelian risk, empirical risk), Prenatal and pre-implantation diagnosis. Non-invasive: Triple test, Ultrsonography (USG), Invasive: Amniocentesis (AC), chorionic villi sampling (CVS), Fetal blood sampling (FBS), Population screening for genetic disorders,
- · Treatment and management of genetic disorders.

References

- 1. Basic Concepts of Molecular Pathology Series: Molecular Pathology Library, Vol. 2; Cagle, Philip T. Allen, Timothy C. (Eds.); Springer 2009, Softcover ISBN: 9780387896250 \$79.95
- 2. Molecular Pathology: The Molecular Basis of Human Disease; William B. Coleman, Gregory J. Tsongalis (Eds.); Academic Press; 1 edition 2009 ISBN 10: 0123744199 ISBN 13: 978-0123744197

- 3. Genomics and Personalized Medicine (2 volumes); Huntington F. Willard, Geoffry S. Ginsburg; Elsevier 2009 ISBN: (set) 97801236942011 \$319.96
- 4. Medical Genetics, 4th Edition; Lynn B. Jorde, John C. Carey, and Michael J. Bamshad, Mosby ISBN: 9780323053730 \$67.95
- 5. DNA from A to Z & Back Again; Carol A. Holland and Daniel H. Farkas; AACC Press 2008 159425088X \$30.00
- 6. Molecular Genetic Pathology, 1st ed.; Liang Cheng and David Zhang; Humana Press 2008 ISBN: 1588299740
- 7. Genomics and Clinical Medicine; Dhavendra Kumar and David Weatherall; Oxford University Press 2008 ISBN 13: 978019518834
- 8. Fundamentals of Molecular Diagnostics; David E Bruns, Edward R Ashwood, and Carl A Burtis; Saunders 2007 ISBN: 1416037373 \$59.95
- 9. Molecular Diagnostics: Fundamentals, Methods, & Clinical Applications; Lela Buckingham and Maribeth Flaws; F. A. Davis Company 2007 0803616597
- 10. Thompson & Thompson Genetics in Medicine, 7th ed.; Robert L. Nussbaum, Roderick R. McInnes, and Huntington F. Willard; WB Saunders 2007 9781416030805 \$62.65
- 11. Medical Genetics at a Glance; Dorian J. Pritchard, Bruce R. Korf; Wiley-Blackwell 2007 ISBN13: 9781405148467
- 12. Molecular Pathology in Clinical Practice; Debra GB Leonard; Springer 2006 038733226X \$164.00; New Edition Springer 2009 Softcover ISBN: 978038787373
- 13. Molecular Diagnostics; George P. Patrinos and Wilhelm Ansorge, (Eds.); Academic Press 2005 ISBN: 0125466617 \$104.00
- 14. Molecular Diagnostics for the Clinical Laboratorian, 2nd ed.; William B. Coleman and Gregory J. Tsongalis, (Eds.); Humana Press 2005 ISBN: 1588293564 \$135.00
- 15. Smith's Recognizable Patterns of Human Malformation Sixth Edition; Kenneth Jones; Saunders 2005 ISBN 10: 0721606156 ISBN 13: 9780721606156
- 16. Principles of Molecular Pathology; Anthony A. Killeen; Humana Press 2003 ISBN: 1588290859 \$120.00
- 17. Molecular Testing in Laboratory Medicine: Selections from Clinical Chemistry, 1998-2001; David E. Bruns, Y.M. Dennis Lo, and Carl T. Wittwer; AACC Press 2003 ISBN: 1890883603 \$62.00 (\$50.00 for AACC members)
- 18. Human Molecular Genetics 3; Tom Strachan and Andrew P. Read; Garland Science 2003 ISBN: 0815341822 \$104.95
- 19. Human Molecular Biology: An Introduction to the Molecular Basis of Health and Disease; Richard J. Epstein; Cambridge University Press 2003 ISBN: 052164481X \$65.00
- 20. Inborn Errors of Development: The Molecular Basis of Clinical Disorders of Morphogenesis; Charles J. Epstein, Robert Erickson, Anthony Wynshaw- Boris; Oxford University Press 2003 ISBN: 9780195145021
- 21. Molecular Microbiology: Diagnostic Principles and Practice David H. Persing, Fred C. Tenover, James Versalovic, Yi-Wei Tang, Elizabeth R. Unger, David A.; Relman, and Thomas J. White, (Eds.) ASM Press 2003 ISBN: 155581221X \$124.95
- 22. Manual of Clinical Microbiology, 9th ed. Patrick R. Murray, Ellen Jo Baron, James H. Jorgensen, Marie Louise Landry, Michael A. Pfaller, (Eds.); ASM Press 2007 ISBN:9781555813710 \$209.95
- 23. Molecular Pathology in Clinical Practice: Infectious Diseases Debra G.B. Leonard; Springer, 2008 ISBN 10: 0387873678 ISBN 13: 978-0387873671

- 24. Molecular Protocols in Transfusion Medicine Gregory A. Denomme, Maria Rios, and Marion E. Reid; Academic Press 2000 ISBN: 0122093704 \$96.95
- 25. Genetics and Molecular Aspects of Gastrointestinal Disease, An issue of Clinics in Laboratory Medicine David C. Whitcomb and Antonia Sepulveda; Saunders Title 2005 ISBN: 9781416027027 \$74.95.
- 26. The Molecular and Genetic Basis of Neurologic and Psychiatric Disease, 4th Edition Roger N. Rosenberg, Salvatore DiMauro, Henry L Paulson, Louis Ptacek, and Eric J. Nestler (Eds.); Lippincott Williams & Wilkins 2007 ISBN: 9780781769563
- 27. Molecular Basis of Cardiovascular Disease, 2nd Edition A Companion to Braunwald's Heart Disease Kenneth R.Chien: Saunders 2004 ISBN: 9780721694283 \$172.
- 28. Inborn Metabolic Diseases: Diagnosis and Treatment 2nd ed. John Fernandes, Jean-Marie Saudubray, Georges van den Berghe, John H. Walter (Eds.); Springer 2006 ISBN: 9783540287834

Semester 1

MD 513. Biochemistry, Molecular Biology and immunology in Diagnosis

Module 1

- · Biochemical tests for detection and quantification of sugar, albumin, urea, protein, globulin, vitamin
- Biochemistry and diagnostic tests of following diseases Duchenne Muscular Dystrophy (DMD) (Creatine phosphokinase-CPK), Phenylketonuria-PKU (phenylketone), G6PD deficiency syndrome (G6PD), Mucopolysaccharidosis, Endocrine disorders related to thyroid and reproduction (TSH, T3, T4, Estradiol, Testosterone, LH, FSH)
- Isolation and Purification of Nucleic acids- Principles and Methods. Molecular cloning, labeling of nucleic acids, hybridization.

Module 2

- Nucleic acid amplification methods and types of PCR: Reverse Transcriptase-PCR, Real-Time PCR, Inverse PCR, Multiplex PCR, Nested PCR, Alu-PCR, Hot-start, *In situ* PCR, Long-PCR, PCR-ELISA, Arbitrarily primed PCR, Ligase Chain Reaction.
- Proteins and Amino acids, Qualitative and quantitative techniques: Protein stability, denaturation; amino acid sequence analysis,
- · Hybridization techniques Southern, Northern, in-situ (including FISH), microarrays types and applications; Protein extraction and analysis (including PAGE and its variations); Western Blot

Module 3

- Applications of PCR- PCR based microbial typing: Eubacterial identification based on 16S rRNA sequences- Amplified Ribosomal DNA Restriction analysis (ARDRA)-Culture independent analysis of bacteria- DGGE and TRFLP. Molecular diagnosis of fungal pathogens based on 18S rRNA sequences- Detection of viral pathogens through PCR. RAPD for animal and plants.
- PCR in forensic science- AmpFLP, STR, Multiplex PCR- Determination of Paternity- Human identification and sex determination.

Module 4

- · Automated DNA sequencing- Principles, Methods and Instrumentation- Advances in DNA sequencing- New Generation sequencing Methods, Pyrosequencing
- · Microarrays- Personalised Medicine- Pharmacogenomics.
- · Levels for rDNA experiments. Biosafety aspects of transgenic plants and germplasm.

Module 5

- · Proteomics- Clinical Proteomics.
- · Overview of immune system, Antigens and antibodies, Antigen-antibody interactions, Major Histocompatibility Complex (MHC), HLA typing, Immunotherapy and immunodiagnostics
- · Immunodiagnostics Introduction, antigen-antibody binding interactions and assays; antibodies-polyclonal and monoclonal antibodies,
- · Immunoassays types [RIA, ELISA, ChemiluminescentIA, FIA] and specific applications; Immunohistochemistry principle and techniques. Good Laboratory Practices.
- · Different Levels of Biosafety, Containment,

References

- 1. Genes IX by B. Lewin, Oxford University Press.
- 2. An Introduction to Genetic Analysis (2000) by A.J.F. Griffiths, J.H. Miller, D.T. Suzuki, R.C. Lewontin and W.M. Gelbart, W.H. Freeman, New York.
- 3. Molecular Biology of the Gene (2004) by J.D.Watson, Tania A baker, Stephen P. Bell, Alexander Gann, Michael Levine, Richard Losick, Pearson Education Pte. Ltd. (Singapore).
- 4. Essentials of Molecular Biology (1998) by G. M. Malacinski and D. Friefelder, Jones & Bartlett Publishers.
- 5. rDNA safety guidelines- Government of India, Ministry of Science and Technology, Dept.of Biotechnology, New Delhi.
- 6. rDNA safety guidelines & regulations-Government of India, Ministry of Science and Technology, Dept. of Biotechnology, New Delhi.
- 7. An Introduction to Forensic DNA Analysis (2002) Rudin, N and Inman, K.CRC Press.
- 8. Forensic DNA Typing. Biology, Technology and Genetics of STR markers (2005) John M. Butler, Elsevier Academic Press, Amsterdam.
- 9. Molecular Diagnostics: Fundamentals, Methods & Clinical applications (2007). Lele Buckingham and Maribeth L. Flaws.
- 10. Fundamentals of Molecular Diagnostics (2007). David E. Bruns, Edward R. Ashwood, Carl A. Burtis. Saunders Group.
- 11. Expert Review of Proteomics and Molecular Diagnostics (Journals) Basic Concepts of Molecular Pathology Series: Molecular Pathology Library, Vol. 2; Cagle, Philip T. Allen, Timothy C. (Eds.); Springer 2009, Softcover ISBN: 9780387896250 \$79.95
- 12. Molecular Pathology: The Molecular Basis of Human Disease; William B. Coleman, Gregory J. Tsongalis (Eds.); Academic Press; 1 edition 2009 ISBN 10: 0123744199 ISBN 13: 978-0123744197
- 13. Genomics and Personalized Medicine (2 volumes); Huntington F. Willard, Geoffry S. Ginsburg; Elsevier 2009 ISBN: (set) 97801236942011 \$319.96

- 14. Medical Genetics, 4th Edition; Lynn B. Jorde, John C. Carey, and Michael J. Bamshad, Mosby ISBN: 9780323053730 \$67.95
- 15. DNA from A to Z & Back Again; Carol A. Holland and Daniel H. Farkas; AACC Press 2008 159425088X \$30.00
- 16. Molecular Genetic Pathology, 1st ed.; Liang Cheng and David Zhang; Humana Press 2008 ISBN: 1588299740
- 17. Genomics and Clinical Medicine; Dhavendra Kumar and David Weatherall; Oxford University Press 2008 ISBN 13: 978019518834
- 18. Fundamentals of Molecular Diagnostics; David E Bruns, Edward R Ashwood, and Carl A Burtis; Saunders 2007 ISBN: 1416037373 \$59.95
- 19. Molecular Diagnostics: Fundamentals, Methods, & Clinical Applications; Lela Buckingham and Maribeth Flaws; F. A. Davis Company 2007 0803616597
- 20. Thompson & Thompson Genetics in Medicine, 7th ed.; Robert L. Nussbaum, Roderick R. McInnes, and Huntington F. Willard; WB Saunders 2007 9781416030805 \$62.65
- 21. Medical Genetics at a Glance; Dorian J. Pritchard, Bruce R. Korf; Wiley-Blackwell 2007 ISBN13: 9781405148467
- 22. Molecular Pathology in Clinical Practice; Debra GB Leonard; Springer 2006 038733226X \$164.00; New Edition Springer 2009 Softcover ISBN: 978038787373
- 23. Molecular Diagnostics; George P. Patrinos and Wilhelm Ansorge, (Eds.); Academic Press 2005 ISBN: 0125466617 \$104.00
- 24. Molecular Diagnostics for the Clinical Laboratorian, 2nd ed.; William B. Coleman and Gregory J. Tsongalis, (Eds.); Humana Press 2005 ISBN: 1588293564 \$135.00
- 25. Smith's Recognizable Patterns of Human Malformation Sixth Edition; Kenneth Jones; Saunders 2005 ISBN 10: 0721606156 ISBN 13: 9780721606156
- 26. Principles of Molecular Pathology; Anthony A. Killeen; Humana Press 2003 ISBN: 1588290859 \$120.00
- 27. Molecular Testing in Laboratory Medicine: Selections from Clinical Chemistry, 1998-2001; David E. Bruns, Y.M. Dennis Lo, and Carl T. Wittwer; AACC Press 2003 ISBN: 1890883603 \$62.00 (\$50.00 for AACC members)
- 28. Human Molecular Genetics 3; Tom Strachan and Andrew P. Read; Garland Science 2003 ISBN: 0815341822 \$104.95
- 29. Human Molecular Biology: An Introduction to the Molecular Basis of Health and Disease; Richard J. Epstein; Cambridge University Press 2003 ISBN: 052164481X \$65.00
- 30. Inborn Errors of Development: The Molecular Basis of Clinical Disorders of Morphogenesis; Charles J. Epstein, Robert Erickson, Anthony Wynshaw- Boris; Oxford University Press 2003 ISBN: 9780195145021
- 31. Molecular Microbiology: Diagnostic Principles and Practice David H. Persing, Fred C. Tenover, James Versalovic, Yi-Wei Tang, Elizabeth R. Unger, David A.; Relman, and Thomas J. White, (Eds.) ASM Press 2003 ISBN: 155581221X \$124.95
- 32. Manual of Clinical Microbiology, 9th ed. Patrick R. Murray, Ellen Jo Baron, James H. Jorgensen, Marie Louise Landry, Michael A. Pfaller, (Eds.); ASM Press 2007 ISBN:9781555813710 \$209.95
- 33. Molecular Pathology in Clinical Practice: Infectious Diseases Debra G.B. Leonard; Springer, 2008 ISBN 10: 0387873678 ISBN 13: 978-0387873671
- 34. Molecular Protocols in Transfusion Medicine Gregory A. Denomme, Maria Rios, and Marion E. Reid; Academic Press 2000 ISBN: 0122093704 \$96.95

- 35. Genetics and Molecular Aspects of Gastrointestinal Disease, An issue of Clinics in Laboratory Medicine David C. Whitcomb and Antonia Sepulveda; Saunders Title 2005 ISBN: 9781416027027 \$74.95.
- 36. The Molecular and Genetic Basis of Neurologic and Psychiatric Disease, 4th Edition Roger N. Rosenberg, Salvatore DiMauro, Henry L Paulson, Louis Ptacek, and Eric J. Nestler (Eds.); Lippincott Williams & Wilkins 2007 ISBN: 9780781769563
- 37. Molecular Basis of Cardiovascular Disease, 2nd Edition A Companion to Braunwald's Heart Disease Kenneth R.Chien; Saunders 2004 ISBN: 9780721694283 \$172.
- 38. Inborn Metabolic Diseases: Diagnosis and Treatment 2nd ed. John Fernandes, Jean-Marie Saudubray, Georges van den Berghe, John H. Walter (Eds.); Springer 2006 ISBN: 9783540287834

Practical Course

MD 514 Molecular diagnostics- I (Nucleic acid and Protein based Diagnostics)

List of experiments

- 1. Hormone assay for thyroid (TSH, T3, T4) and
- 2. Sexual disorders (testosterone, dihydrotestosteron, estradiol, FSH, LH by RIA)
- 3. Isolation of Genomic DNA from microbe (*E.coli*), Plant (*Bacopa monnieri*), Animal (*C. elegans*) and Human (*Peripheral Blood*).
- 4. Isolation of Metagenome (sediment/soil).
- 5. Plasmid DNA isolation by Alkaline lysis and Boiling method.
- 6. Quality / Quantity checking of Nucleic acids by *a*) UV Spectrophotometer *and* b) Agarose Gel Electrophoresis.
- 7. Molecular Cloning- ligation
- 8. Competent cell preparation
- 9. Transformation- selection of recombinants (Blue white selection)
- 10. Confirmation of recombinants by gel electrophoresis.
- 11. Nucleic acid labeling and Southern Hybridization
- 12. Automated DNA sequencing (Demo).
- 13. RNA isolation (from rat liver) and Pulsed Field Gel Electrophoresis
- 14. PAGE Denaturing gradient gels, Temperature gradient gels
- 15. and Western Blot
- 16. DNA sequencing, Automated DNA Sequencing PCR
- 17. Identification of human bacterial pathogens by Polymerase chain reaction,
- 18. PCR based diagnosis of plant bacterial pathogen
- 19. Culture independent analysis of microbes by DGGE (Denatured Gradient Gel Electrophoresis) and T-RFLP (Terminal Restriction Fragment Length Polymorphism).
- 20. Molecular diagnosis of parasitic disease
- 21. Detection of transgenes in GMOs
- 22. PCR amplification using primers that amplify the regions consisting of dinucleotide repeats, trinucleotide repeats and that detects restriction enzyme polymorphic sites, Alu ins/del sites.

- 23. Amplification of Short Tandem Repeats (STR)/Microsatellites.
- 24. Multiplex STR PCR (Demo).
- 25. Single strand conformation polymorphism (SSCP) analysis.
- 26. HLA typing and tissue transplantation matching (Demo)
- 27. Microarrays for pathogen detection and SNP studies (Demo).
- 28. Bioinformatic tools for genome and proteome analysis.

Practical II

MD 515 Molecular diagnostic II (Genetic & Cytologic techniques)

List of Experiments

- 1. Metaphase chromosome preparations from bone marrow of mouse, rat, cultured Lymphocytes
- 2. Chromosome preparation from lymphocyte culture / mouse bone marrow
- 3. Karyotype using Human lymphocyte culture
 - a. Q-banding
 - b. G-banding

Fluorescence *in-situ* Hybridization (FISH)

- 4. Automated Karyotyping
- 5. Sex chromatin (buccal mucosa, hair bud)
- 6. Comet assay
- 7. Micronucleus assay bone marrow of mouse and cultured lymphocytes
- 8. Chromosome preparation from chorionic villi, stem cells, cell line
- 9. Sister Chromatid Exchange (SCE)
- 10. Study of Chromosomal Aberrations in Mice
- 11. Meiotic Chromosome preparations from testis
- 12. Flowcytometry.

Proteome analysis

- 15. Characterization of Proteins by SDS-PAGE and 2D Gel Electrophoresis
- 16. Purification of proteins by HPLC
- 17. Immunological methods- Agglutination (ABO/Bacterial),
- 18. Precipitation, Immunodiffusion, Immunoelectrophoresis

Molecular Diagnosis of human genetic disorders

- 19. Beta thalassemia
- 20. Checking of p53 gene polymorphism for susceptibility to cancer
- 21. Down's Syndrome
- 22. Retinitis pigmentosa
- 23. Human identification and paternity determination (simulated) by VNTR Probes
- 24. Mutation detection methods
- 25. Functional protein identification- 2D gels
- 26. Forensic Application Paternity Testing, victim Identification, crime detection,

MD 521 Project Dissertations

Each of the students has to undertake a project during the first semester itself and should be comp0leted in the end of second semester. They have to submit the dissertation at the end of 2^{nd} semester for evaluation. Each student has to present their work before a committee consisting of the faculty members and an external expert as an examiner.

MD 522 General Viva voce

The committee constituted for evaluating the project dissertation also has to conduct the General viva voce examination along with the Project presentation. They have to submit the mark list to the chairman of the examination Board, who will the Department head, or the Director of the Centre or Institute.



UNIVERSITY OF KERALA

POST GRADUATE DIPLOMA IN MOLECULAR DIAGNOSTICS

Scheme and Syllabus w.e.f. 2015 admission