

BIOCHEMISTRY 2 [1203253]

The University of Jordan
Faculty: Pharmacy
Department: Biopharmaceutics and Clinical Pharmacy
Program: Pharmacy
Academic Year/ Fall Semester: 2014/15

Credit hours	3	Level	2 nd year	Pre-requisite	-
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Course website	-	E-mail		Place	Pharmacy

Course Objectives:

This course is the second course in a two-semester sequence in biochemistry. The students are expected to:

- 1- Use the knowledge gained in Biochemistry I to understand the basic concepts of metabolism
- 2- Understand the metabolic pathways of the major biomolecules; carbohydrate, lipids, proteins and nucleotides.
- 3- Understand the main issues regarding the storage and expression of genetic information

Learning Outcomes:

A. Knowledge and Understanding of:

A1. DNA, RNA, AND THE FLOW OF GENETIC INFORMATION

A1.1 Nucleotide Biosynthesis

- ❑ Purine Bases Can Be Synthesized de Novo or Recycled by Salvage Pathways
- ❑ Deoxyribonucleotides Synthesized by the Reduction of Ribonucleotides Through a Radical Mechanism
- ❑ Key Steps in Nucleotide Biosynthesis Are Regulated by Feedback Inhibition
- ❑ In de Novo Synthesis, the Pyrimidine Ring Is Assembled from Bicarbonate, Aspartate, and Glutamine
- ❑ Nucleoside Monophosphate Kinases: Catalyzing Phosphoryl Group Exchange between Nucleotides Without Promoting Hydrolysis
- ❑ Disruptions in Nucleotide Metabolism Can Cause Pathological Conditions

A1.2 DNA Structure, Replication, Recombination, and Repair

- ❑ A Nucleic Acid Consists of Four Kinds of Bases Linked to a Sugar-Phosphate Backbone
- ❑ A Pair of Nucleic Acid Chains with Complementary Sequences Can Form a Double-Helical Structure
- ❑ DNA Is Replicated by Polymerases that Take Instructions from Templates
- ❑ DNA Can Assume a Variety of Structural Forms
- ❑ DNA Polymerases Require a Template and a Primer
- ❑ Double-Stranded DNA Can Wrap Around Itself to Form Supercoiled Structures
- ❑ DNA Replication of Both Strands Proceeds Rapidly from Specific Start Sites
- ❑ Double-Stranded DNA Molecules with Similar Sequences Sometimes Recombine

A1.3 RNA Synthesis and Splicing

- ❑ Transcription Is Catalyzed by RNA Polymerase
- ❑ Eukaryotic Transcription and Translation Are Separated in Space and Time
- ❑ The Transcription Products of All Three Eukaryotic Polymerases Are further subjected to downstream processing
- ❑ Most Eukaryotic Genes Are Mosaics of Introns and Exons

A1.4 Protein Synthesis

- ❑ Gene Expression Is the Transformation of DNA Information Into Functional Molecules
- ❑ Amino Acids Are Encoded by Groups of Three Bases Starting from a Fixed Point
- ❑ Protein Synthesis Requires the Translation of Nucleotide Sequences Into Amino Acid Sequences
- ❑ Mutations Involve Changes in the Base Sequence of DNA
- ❑ Aminoacyl-Transfer RNA Synthetases Read the Genetic Code
- ❑ A Ribosome Is a Ribonucleoprotein Particle (70S) Made of a Small (30S) and a Large (50S) Subunit
- ❑ Protein Factors Play Key Roles in Protein Synthesis
- ❑ Eukaryotic Protein Synthesis Differs from Prokaryotic Protein Synthesis Primarily in Translation Initiation and subsequent coupling reactions.

A1.5 The Gene Expression Regulation

- ❑ Prokaryotic DNA-Binding Proteins Bind Specifically to Regulatory Sites in Operons
- ❑ The Greater Complexity of Eukaryotic Genomes Requires Elaborate Mechanisms for Gene Regulation
- ❑ Transcriptional Activation and Repression Are Mediated by Protein-Protein Interactions
- ❑ Some Receptors Dimerize in Response to Ligand Binding and Signal by Cross-phosphorylation
- ❑ Gene Expression Can Be Controlled at Posttranscriptional Levels

A1.6 Molecular basis of inherited diseases

- ❑ Restriction Enzymes: Performing Highly Specific DNA-Cleavage Reactions
- ❑ DNA recombinations are helpful in establishing genomic as well as cDNA libraries
- ❑ Antibiotic resistance genes can select for the transfected cloning vector.
- ❑ The utility of Sanger dideoxy method is basically for purified DNA sequencing.
- ❑ Restriction Fragment Length Polymorphism analysis is a direct diagnostic tool of sickle cell disease.
- ❑ Polymerase chain reaction is highly advantageous in detecting/tracing low abundance nucleic acid sequences
- ❑ Microarray technique is quite a handy Analytical means of determinations of the gene expression products.
- ❑ ELISA and Western blots can be Important Techniques to investigate specific proteins.

A2. TRANSDUCING & STORING OF ENERGY INTERMEDIARY METABOLISM

A2. 1 Glycolysis and Gluconeogenesis

- ❑ Metabolism Is Composed of Many Coupled, Interconnecting Reactions
- ❑ The Oxidation of Carbon Fuels Is an Important Source of Cellular Energy
- ❑ Glycolysis Is an Energy-Conversion Pathway in Many Organisms
- ❑ The Glycolytic Pathway Is Tightly Controlled
- ❑ Glucose Can Be Synthesized from Noncarbohydrate Precursors
- ❑ Gluconeogenesis and Glycolysis Are Reciprocally Regulated

A2. 2 Citric Acid Cycle

- ❑ The Citric Acid Cycle Oxidizes Two-Carbon Units
- ❑ Entry to the Citric Acid Cycle and Metabolism Through It Are Controlled

- ✘ The Citric Acid Cycle Is a Source of Biosynthetic Precursors. Amino Acids Are Made from Intermediates of the Citric Acid Cycle and Other Major Pathways

A2. 3 The Pentose Phosphate Pathway

- ✘ the Pentose Phosphate Pathway Generates NADPH and Synthesizes Five-Carbon Sugars
- ✘ The Metabolism of Glucose 6-Phosphate by the Pentose Phosphate Pathway Is Coordinated with Glycolysis
- ✘ Glucose 6-Phosphate Dehydrogenase Plays a Key Role in Protection Against Reactive Oxygen Species

A2. 4 Glycogen, hexoses and disaccharides Metabolism

- ✘ Glycogen Breakdown Requires the Interplay of Several Enzymes
- ✘ Phosphorylase Is Regulated by Allosteric Interactions and Reversible Phosphorylation
- ✘ Epinephrine and Glucagon Signal the Need for Glycogen Breakdown
- ✘ Glycogen Is Synthesized and Degraded by Different Pathways
- ✘ Glycogen Breakdown and Synthesis Are Reciprocally Regulated
- ✘ All hexoses are to be phosphorylated before they are any further metabolized
- ✘ Hexose epimerase can substitute for lacking dietary sources of galactose
- ✘ Lactose synthesis is mainly mediated by galactosyltransferases

A3. LIPID METABOLISM

A3. 1 Fatty Acid Metabolism

- ✘ Triacylglycerols Are Highly Concentrated Energy Stores
- ✘ The Utilization of Fatty Acids as Fuel Requires Three Stages of Processing
- ✘ Certain Fatty Acids Require Additional Steps for Degradation
- ✘ Fatty Acids Are Synthesized and Degraded by Different Pathways
- ✘ Acetyl Coenzyme A Carboxylase Plays a Key Role in Controlling Fatty Acid Metabolism via Carnitine shuttle modulation
- ✘ Elongation and Unsaturation of Fatty Acids Are Accomplished by Accessory
- ✘ Ketogenesis is strictly hepatic and ketone bodies can be consumed by brain as well as muscle cells

A3. 2 The Biosynthesis of Membrane Lipids and Steroids

- ✘ Phosphatidic acid Is a Common Intermediate in the Synthesis of Phospholipids and Triacylglycerols
- ✘ Cholesterol Is Synthesized from Acetyl Coenzyme A in Three Stages
- ✘ The Complex Regulation of Cholesterol Biosynthesis Takes Place at Several Levels
- ✘ Important Derivatives of Cholesterol Include Bile Salts and Steroid Hormones

A4. PROTEIN TURNOVER AND AMINO ACID CATABOLISM

- ✘ Proteins Are Degraded to Amino Acids
- ✘ Protein Turnover Is Tightly Regulated
- ✘ Many Enzymes Are Activated by Specific Proteolytic Cleavage
- ✘ The First Step in Amino Acid Degradation Is the Removal of Nitrogen
- ✘ Ammonium Ion Is Converted Into Urea in Most Terrestrial Vertebrates
- ✘ Carbon Atoms of Degraded Amino Acids Emerge as Major Metabolic Intermediates
- ✘ Inborn Errors of Metabolism Can Disrupt Amino Acid Degradation
- ✘ Amino Acids Are Precursors of Many Biomolecules

A5. THE INTEGRATION OF METABOLISM

- ✘ Metabolic Pathways Contain Many Recurring Motifs
- ✘ Metabolism Consist of Highly Interconnected Pathway
- ✘ Each Organ Has a Unique Metabolic Profile

- ☞ Food Intake and Starvation Induce Metabolic Changes
- ☞ Ethanol Alters Energy Metabolism in the Liver
- ☞ Peptide hormones, namely insulin and glucagon, are actively involved in reciprocal regulation of metabolism during absorptive and postabsorptive phases

B. Intellectual skills (cognitive and analytical):

- ☞ Integrate metabolic pathways, and analyze the complete integrated metabolic map.
- ☞ Interpret metabolic abnormalities and relate them to possible causes and mechanisms.
- ☞ Relate the biochemical events at the cellular level to the physiological processes occurring in the whole animal.
- ☞ Follow up the flow of genetic information; DNA→RNA→Protein

C. Subject specific skills

D. Transferable Skills

- ☞ The development of problem solving and critical thinking skills.
- ☞ Use oral communication to effectively transmit ideas and conclusions to a scientific audience.

References:

ISBN	Title	Author	Year
716712261	BIOCHEMISTRY 4TH EDITION	STRYER, LUBERT	1995C
781769604	BIOCHEMISTRY LIPPINCOTT'S ILLUSTRATED REVIEWS, 4TH EDITION	CHAMPE, PAMELA; HARVEY, RICHARD; FERRIER, DENISE; COOPER, MICHAEL	2008C
7167743396	LEHNINGER PRINCIPLES OF BIOCHEMISTRY	LEHNINGER, ALBERT	2005C
9780071765763	HARPER'S ILLUSTRATED BIOCHEMISTRY-27ED.	MURRAY, ROBERT K. (ROBERT KINCAID)	2012
0272797138	ESSENTIALS OF HUMAN BIOCHEMISTRY	PATERSON, COLIN RALSTON	1983

Lecture notes are available on

<http://blackboard.ju.edu.jo>

User name: pharm_std

Password: pharm_std

Course Contents and Schedule

<u>Subject</u>	<u>No. of lectures</u>
Introduction	1
Storage and expression of genetic information.	
• Nucleotide metabolism.	3
• DNA structure and replication	3
• RNA structure and synthesis	2
• Protein Synthesis	3
• Regulation of gene expression	3
• Molecular basis of inherited disease	5
Intermediary metabolism.	
• Glycolysis.	2
• Gluconeogenesis.	2
• Hexose Monophosphate pathway	2
• Citric acid cycle.	2
MIDTERM	
Carbohydrate metabolism	
• Glycogen metabolism.	2
• Metabolism of monosaccharides and disaccharides	
Lipid metabolism.	
• Metabolism of dietary lipids.	2
• Fatty acid and triacylglycerol metabolism.	2
• Phospholipid metabolism	1
• Cholesterol and steroid metabolism.	3
Nitrogen metabolism.	
• Disposal of Nitrogen.	2
• Metabolism of carbon skeleton.	1
• Conversion of amino acids to specialized products	1
Integration of metabolism.	
• Metabolic effects of insulin and glucagon.	1
• Metabolism in the well-fed state.	1
• Metabolism in starvation and diabetes mellitus	1
FINAL EXAM	