

Page 1 / 8

#### APPROVED

at the Chair meeting of 25.08.23, minute no.1, Head of the Biochemistry and Clinical Biochemistry Chair, MD, PhD., prof., Olga TAGADIUC \_\_\_\_\_

#### THE THEMATIC PLAN

#### of courses and laboratory work in Pharmaceutical Biochemistry for students of the Faculty of Pharmacy, year III, academic year 2023-2024

|   |          | the autumn semester(5) - year III                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               |                                                                                                                                                                                                                                                                                                                                                                                                                |
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| Ν | Date     | Course, theme name                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              | Laboratory work, name of the topic                                                                                                                                                                                                                                                                                                                                                                             |
| 1 | 04-08.09 | <ol> <li>Classification and structure of lipids. Lipids of<br/>pharmaceutical interest. Essential fatty acids. Digestion and<br/>absorption of lipids in the digestive tract. Chylomicrons and<br/>Plasma Lipoproteins.</li> <li>Biosynthesis of saturated and unsaturated fatty acids.<br/>Biosynthesis of triacylglycerols and phosphoglycerides:<br/>Lipotropic substances, their role.<br/>Biosynthesis of sphingo- and glycolipids: general concepts.<br/>Cholesterol biosynthesis – stages, reactions of stage I (up to<br/>mevalonic acid), ways of use and elimination of cholesterol.<br/>Eicosanoids (prostaglandins, leukotrienes, thromboxanes) –<br/>synthesis and biomedical role. The therapeutic effect of<br/>eicosanoids.<br/>Fat-soluble vitamins - A, D, E, K - structure and biomedical<br/>role. Their use in therapy. Hypo- and avitaminoses.</li> </ol> | The biological role of lipids. Protoplasmic and reserve lipids.<br>Classification and structure of lipids. Biological membranes.<br>Digestion and absorption of lipids in the gastrointestinal tract. Plasma<br>lipoproteins. The action of pancreatic phospholipases.<br>Identification of bile acids.                                                                                                        |
| 2 | 11-15.09 |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 | <ul> <li>Biosynthesis of saturated and unsaturated fatty acids.</li> <li>Biosynthesis of triacylglycerols and phosphoglycerides: reactions and regulation. Lipotropic substances, their role.</li> <li>Biosynthesis of sphingo- and glycolipids: general concepts.</li> <li>Cholesterol biosynthesis – stages, stage I reactions (up to mevalonic acid). Ways of using and eliminating cholesterol.</li> </ul> |



| P | age 2 / 8 |
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| 3 | 18-22.09 | <ol> <li>Catabolism of triacylglycerols:<br/>Glycerol oxidation. Oxidation of fatty acids:         <ol> <li>saturated with an even number of carbon atoms;</li> <li>unsaturated with an even number of carbon atoms;</li> <li>saturated with an odd number of carbon atoms;</li> <li>Catabolism of phospho-, sphingo- and glycolipids.<br/>Metabolism of ketone bodies. Biosynthesis and catabolism -<br/>reactions, enzymes, coenzymes, regulation. The biomedical<br/>role.<br/>Regulation of lipid metabolism at cellular and<br/>neurohormonal level. The role of catecholamines, glucagon,<br/>thyroid hormones, insulin.</li> <li>Pathology of lipid metabolism: a) disorders of lipid digestion<br/>and absorption; b) fatty degeneration of the liver, obesity; c)<br/>lipid metabolism disorders in diabetes, starvation and<br/>alcoholism.</li> </ol> </li> </ol> | Eicosanoids (prostaglandins, leukotrienes, thromboxanes) –<br>synthesis and biomedical role. The therapeutic effect of eicosanoids.<br>Fat-soluble vitamins - A, D, E, K - structure and biomedical role.<br>Their use in therapy. Hypo- and avitaminoses. Determination of total<br>lipids in blood serum.<br>Catabolism of triacylglycerols:<br>Glycerol oxidation. Oxidation of fatty acids:<br>i. saturated with an even number of carbon atoms;<br>ii. unsaturated with an even number of carbon atoms;<br>iii. saturated with an even number of carbon atoms;<br>catabolism of phospho-, sphingo- and glycolipids.<br>Metabolism of ketone bodies. Biosynthesis and catabolism -<br>reactions, enzymes, coenzymes, regulation. The biomedical role.<br>Regulation of lipid metabolism at cellular and neurohormonal level.<br>The role of catecholamines, glucagon, thyroid hormones, insulin.<br>Pathology of lipid metabolism: a) disorders of lipid digestion and<br>absorption; b) fatty degeneration of the liver, obesity; c) lipid<br>metabolism disorders in diabetes, starvation and alcoholism. |
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| 4 | 25-29.09 |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              | Total test to: "Lipid metabolism"                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               |
| 5 | 02-06.10 | <ul> <li>1. The need for protein in food. The nutritional value of dietary proteins. The dynamic state of proteins. Nitrogen balance. Protein deficiency. Parenteral protein nutrition. Protein digestion and absorption. Proteolytic enzymes Zymogens and their activation. Protein digestion in the stomach. Protein digestion in the intestine. Absorption of amino acids in the intestine. Active secondary and facilitated transport of amino acids. Amino acid putrefaction in the large intestine. The products of putrefaction. The mechanisms of</li> </ul>                                                                                                                                                                                                                                                                                                         | The need for protein in food. The nutritional value of dietary proteins.<br>The dynamic state of proteins. Nitrogen balance. Protein deficiency.<br>Proteolytic enzymes Protein digestion in the stomach. Protein<br>digestion in the intestine. Absorption of amino acids in the intestine.<br>Active secondary and facilitated transport of amino acids.<br>Amino acid putrefaction in the large intestine. The products of<br>putrefaction. The mechanisms of detoxification of toxic products in<br>the liver (microsomal oxidation, conjugation). Conjugation agents,<br>enzymes.                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          |



Page 3 / 8

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|   |          | detoxification of toxic products in the liver (microsomal      | Transamination of amino acids: mechanism, enzymes, coenzymes,                                          |
|   |          | oxidation, conjugation). Conjugation agents, enzymes.          | significance of the process. The diagnostic value of determining the                                   |
|   |          | Transamination of amino acids: mechanism, enzymes,             | activity of transaminases (ALT and AST) in the blood.                                                  |
|   |          | coenzymes, significance of the process. The diagnostic value   | Deamination of amino acids. The types. Direct deamination of                                           |
|   |          | of determining the activity of transaminases (ALT and AST)     | amino acids. Oxidative deamination of glutamic acid (reaction,                                         |
|   |          | in the blood. Deamination of amino acids. The types. Direct    | enzyme, coenzymes, importance of the process). Indirect deamination                                    |
|   |          | deamination of amino acids. Oxidative deamination of           | of amino acids. The stages. Enzymes, coenzymes. Biological role.                                       |
|   |          | glutamic acid (reaction, enzyme, coenzymes, importance of      | Determination of gastric juice acidity. Identification of pathological                                 |
|   |          | the process). Indirect deamination of amino acids. The stages. | components of gastric juice.                                                                           |
|   |          | Enzymes, coenzymes. Biological role.                           |                                                                                                        |
|   |          | 1. Mechanisms of ammonia toxicity. Modes of ammonia            |                                                                                                        |
|   |          | detoxification: glutamine synthesis b) urea synthesis          |                                                                                                        |
|   |          | (ureogenetic cycle), clinical importance of urea determination |                                                                                                        |
|   |          | c) elimination of NH3 in the form of ammonium salts            |                                                                                                        |
|   |          | (NH4+).                                                        |                                                                                                        |
|   |          | Decarboxylation of amino acids. The influence of biogenic      |                                                                                                        |
| 6 | 00.12.10 | amines on the body, their detoxification.                      |                                                                                                        |
| 6 | 09-13.10 |                                                                | Mechanisms of ammonia toxicity.                                                                        |
|   |          |                                                                | Ways to detoxify ammonia:                                                                              |
|   |          |                                                                | a) glutamine synthesis b) urea synthesis (ureogenetic cycle), the                                      |
|   |          |                                                                | clinical importance of urea determination c) elimination of NH3 in                                     |
|   |          |                                                                | the form of ammonium salts (NH4+).                                                                     |
|   |          |                                                                | Decarboxylation of amino acids. The influence of biogenic amines<br>on the body, their detoxification. |
|   |          |                                                                | Determination of urea in urine.                                                                        |
| 7 | 16-20.10 | 1. Metabolism of phenylalanine and tyrosine.                   | Peculiarities of the metabolism of some amino acids.                                                   |
|   | 10 20.10 | Metabolism of glycine and serine. Metabolism of methionine     | Metabolism of phenylalanine and tyrosine. Congenital disorders of                                      |
|   |          | and cysteine. Metabolism of dicarboxylic amino acids,          | the metabolism of these amino acids (phenylketonuria, alkaptonuria,                                    |
|   |          | Reciprocal link between protein, carbohydrate and lipid        | albinism)                                                                                              |
|   |          | metabolism.                                                    | Metabolism of glycine and serine. The role of tetrahydrofolic acid in                                  |
|   |          | 2. Nucleoprotein metabolism.                                   | their metabolism. Mechanism of action of sulfanilamides.                                               |



Page 4 / 8

|   |             | a) Digestion and absorption of nucleic acids.                                            | Metabolism of methionine and cysteine. The role of methionine in                                             |
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|   |             | b) Biosynthesis of purine nucleotides - "de novo" and from                               | the synthesis of phospholipids, creatine, adrenaline. Synthesis and                                          |
|   |             | synthesized nitrogenous bases.<br>c) Degradation of purine nucleotides. Uric acid. Gout. | role of phosphocreatine.<br>Metabolism of dicarboxylic amino acids, their use as medicina                    |
|   |             | d) Biosynthesis of pyrimidine nucleotides. The role of                                   | preparations.                                                                                                |
|   |             | thioredoxin in the formation of deoxyribonucleotides.                                    | The mutual connection between protein, carbohydrate and lipid                                                |
|   |             | e) Degradation of pyrimidine nucleotides (general notions).                              | metabolism.                                                                                                  |
|   |             | Metabolism of chromoproteins.                                                            | Urinary creatinine dosage. Identification of homogentisinic acid in                                          |
|   |             | a) Digestion of chromoproteins. b) Heme biosynthesis.                                    | urine.                                                                                                       |
|   |             | Location, stages, regulation. The porphyrias. c) Heme                                    |                                                                                                              |
|   |             | catabolism. Location, stages. Jaundice.                                                  |                                                                                                              |
| 8 | 23-27.10    |                                                                                          | Nucleoprotein metabolism.                                                                                    |
|   |             |                                                                                          | a) Digestion and absorption of nucleic acids.                                                                |
|   |             |                                                                                          | b) Biosynthesis of purine nucleotides - "de novo" and from                                                   |
|   |             |                                                                                          | synthesized nitrogenous bases.                                                                               |
|   |             |                                                                                          | c) Degradation of purine nucleotides. Uric acid. Gout.                                                       |
|   |             |                                                                                          | d) Biosynthesis of pyrimidine nucleotides. The role of thioredoxin in the formation of deoxyribonucleotides. |
|   |             |                                                                                          | e) Degradation of pyrimidine nucleotides (general notions).                                                  |
|   |             |                                                                                          | Metabolism of chromoproteins.                                                                                |
|   |             |                                                                                          | a) Digestion of chromoproteins.                                                                              |
|   |             |                                                                                          | b) Heme biosynthesis. Location, stages, regulation. The porphyrias.                                          |
|   |             |                                                                                          | c) Heme catabolism. Location, stages. Jaundice.                                                              |
| 0 | 20.10.02.11 |                                                                                          | Dosage of uric acid in urine.                                                                                |
| 9 | 30.10-03.11 | 1. Notions about hormones General properties. Classification                             |                                                                                                              |
|   |             | of hormones. Mechanisms of hormone action:                                               |                                                                                                              |
|   |             | Hypothalamus hormones - liberins and statins. Vasopressin and oxytocin.                  | Test total:                                                                                                  |
|   |             | Pituitary hormones - chemical nature, mechanism of action,                               | "Metabolism of simple and complex proteins"                                                                  |
|   |             | biological effect, regulation of secretion and its deregulation,                         | metabolism of simple and complex proteins                                                                    |
|   |             | practical use as medicinal preparations.                                                 |                                                                                                              |



Page 5 / 8

| 10 06-10.11 | <ul> <li>Parathyroid hormone. Regulation of phosphorus and calcium metabolism. Parathyroid gland function disorders.</li> <li>2. Pancreatic hormones (insulin, glucagon). Mechanism of action of insulin and glucagon. Pharmaceutical preparations based on insulin synthesized in the laboratory. Thyroid gland hormones (iodothyronines and thyrocalcitonin) – their biosynthesis and regulation. Hypo-and hyperfunction of the thyroid gland. Adrenal medullary hormones (adrenaline, noradrenaline) and adrenal cortex (gluco- and mineralocorticoids). Sex hormones:</li> <li>a) androgens and their biological role, anabolic steroids as active pharmaceutical preparations.</li> <li>b) estrogens, physiological action and mechanism of action;</li> </ul> | Notions about hormones and substances with hormonal action, their<br>use as pharmaceutical preparations.<br>General properties. Classification of hormones.<br>Mechanisms of hormone action:<br>a) membrane-cytosolic (indirect), the role of G-proteins and second<br>messengers in the transmission of hormonal information;<br>b) cytosolic-nuclear (direct).<br>Hypothalamus hormones - liberins and statins. Vasopressin and<br>oxytocin.<br>Pituitary hormones - chemical nature, mechanism of action,<br>biological effect, regulation of secretion and its deregulation,<br>practical use as medicinal preparations.<br>a) the corticotropin family (ACTH, MSH, lipotropins and related<br>peptides);<br>b) the family of glycoprotein hormones (TSH, FSH, LH and placental<br>chorionic gonadotropin);<br>c) the family of somatomammotropic hormones (STH, prolactin and |
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placental lactogen); Parathyroid hormone. Regulation of phosphorus and calcium metabolism. Parathyroid gland function disorders. Dosage of inorganic phosphorus in blood serum. The reaction to identify 17-ketosteroids in urine. Calcium dosage in blood serum. Dosage of inorganic phosphorus in blood serum 1. Pancreatic hormones (insulin, glucagon). Mechanism of action of 13-17.11 1. Chemical composition of blood plasma: 11 a) Plasma proteins (albumins, globulins, enzymes, insulin and glucagon. Pharmaceutical preparations based on insulin immunoglobulins, interferon); synthesized in the laboratory. b) Nitrogenous non-protein compounds of the blood plasma. 2. Thyroid gland hormones (iodothyronines and thyrocalcitonin) -Their origin and role in plasma. Associated pathological their biosynthesis and regulation. Hypo- and hyperfunction of the conditions. thyroid gland. c) Non-nitrogenous organic compounds. Their origin and role 3. Adrenal medullary hormones (adrenaline, noradrenaline) and in plasma. Associated pathological conditions. adrenal cortex (gluco- and mineralocorticoids). d) Mineral components of blood plasma. Their role in plasma 4. Sex hormones: and body. Blood ionogram. a) androgens and their biological role, anabolic steroids as active 2. The chemical composition and peculiarities of the pharmaceutical preparations. metabolism of erythrocytes and platelets: b) estrogens, physiological action and mechanism of action; a) Chemical composition and peculiarities of erythrocyte metabolism. i. Glycolysis and the Rapoport shunt ii. The pentose phosphate pathway and glutathione iii. Hemoglobin and oxygen transport. iv. Carbonic anhydrase and CO2 transport. b) Chemical composition and peculiarities of platelet metabolism and blood coagulation. i. Platelet coagulation factors ii. Plasma coagulation factors and blood coagulation scheme. iii. The fibrinolytic system. iv. Anticoagulant factors: natural (native) anticoagulants.



| P | age 7 / 8 |
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| 12 | 20-24.11    |                                                               | Chemical composition of blood plasma:                                     |
|----|-------------|---------------------------------------------------------------|---------------------------------------------------------------------------|
|    |             |                                                               | a) Plasma proteins (albumins, globulins, enzymes, immunoglobulins,        |
|    |             |                                                               | interferon);                                                              |
|    |             |                                                               | b) Nitrogenous non-protein compounds of the blood plasma. Their           |
|    |             |                                                               | origin and role in plasma. Associated pathological conditions.            |
|    |             |                                                               | c) Non-nitrogenous organic compounds. Their origin and role in            |
|    |             |                                                               | plasma. Associated pathological conditions.                               |
|    |             |                                                               | d) Mineral components of blood plasma. Their role in plasma and           |
|    |             |                                                               | body. Blood ionogram.                                                     |
|    |             |                                                               | The chemical composition and peculiarities of the metabolism of           |
|    |             |                                                               | erythrocytes and platelets:                                               |
|    |             |                                                               | a) Chemical composition and peculiarities of erythrocyte metabolism.      |
|    |             |                                                               | i. Glycolysis and the Rapoport shunt                                      |
|    |             |                                                               | ii. The pentose phosphate pathway and glutathione                         |
|    |             |                                                               | iii. Hemoglobin and oxygen transport.                                     |
|    |             |                                                               | iv. Carbonic anhydrase and CO2 transport.                                 |
|    |             |                                                               | b) Chemical composition and peculiarities of platelet metabolism and      |
|    |             |                                                               | blood coagulation.                                                        |
|    |             |                                                               | i. Platelet coagulation factors                                           |
|    |             |                                                               | ii. Plasma coagulation factors and blood coagulation scheme.              |
|    |             |                                                               | iii. The fibrinolytic system.                                             |
|    |             |                                                               | iv. Anticoagulant factors: natural (native) anticoagulants.               |
|    |             |                                                               | Determination of hemoglobin in blood by the cyanmethemoglobin             |
| 12 | 27.11.01.12 |                                                               | method.                                                                   |
| 13 | 27.11-01.12 | 1. Biotechnology of medicinal preparations (liposomes,        | Pharmaceutical biochemistry. Biotechnology of medicinal                   |
|    |             | genetic engineering, monoclonal antibodies with targeted      | preparations (liposomes, genetic engineering, monoclonal antibodies       |
|    |             | action, application of enzymes as chemical reagents, etc.).   | with targeted action, application of enzymes as chemical reagents,        |
|    |             | Absorption (digestive, dermal, pulmonary), tissue             | etc.).                                                                    |
|    |             | distribution (volume of distribution, tissue affinity, tissue | Absorption (digestive, dermal, pulmonary), tissue distribution            |
|    |             | storage) and elimination (urinary, biliary, salivary, sweat,  | (volume of distribution, tissue affinity, tissue storage) and elimination |
|    |             | lactation) of xenobiotics (drugs). Passive transport (simple  | (urinary, biliary, salivary, sweat, lactation) of xenobiotics (drugs).    |



Page 8 / 8

|    |          | diffusion), facilitated transport, active transport. Structural-                                                   | Passive transport (simple diffusion), facilitated transport, active                                           |
|----|----------|--------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------|
|    |          | functional peculiarities of xenobiotic (drug) transporters:                                                        | transport. Structural-functional peculiarities of xenobiotic (drug)                                           |
|    |          | ABC-transporters, SLC-transporters.                                                                                | transporters: ABC-transporters, SLC-transporters.                                                             |
|    |          | Biotransformation (metabolization) of xenobiotics                                                                  | Biotransformation (metabolization) of xenobiotics (medicines).                                                |
|    |          |                                                                                                                    |                                                                                                               |
|    |          | (medicines). Location of biotransformation: cavity,<br>extracellular and tissue. The conditions that determine the | Location of biotransformation: cavity, extracellular and tissue. The                                          |
|    |          |                                                                                                                    | conditions that determine the metabolism of drugs. Dependence of                                              |
|    |          | metabolism of drugs. Dependence of drug action on their                                                            | drug action on their metabolism: deactivation, activation,                                                    |
|    |          | metabolism: deactivation, activation, modification of the main                                                     | modification of the main effect.                                                                              |
|    |          | effect.                                                                                                            | General biochemical reactions of drug biotransformation -                                                     |
|    |          | General biochemical reactions of drug biotransformation -                                                          | oxidation, reduction, hydrolysis and synthesis (conjugation).                                                 |
|    |          | oxidation, reduction, hydrolysis and synthesis (conjugation).                                                      | Two phases of biotransformation of xenobiotics (medicines):                                                   |
|    |          | 2. Two phases of biotransformation of xenobiotics                                                                  | modification and conjugation.                                                                                 |
|    |          | (medicines): modification and conjugation.                                                                         | The phase of modification of drugs and xenobiotics by microsomal                                              |
|    |          | The phase of modification of drugs and xenobiotics by                                                              | oxidation. Mechanism of microsomal oxidation and the role of                                                  |
|    |          | microsomal oxidation. Mechanism of microsomal oxidation                                                            | cytochrome P450. Monooxidase and reductase chains.                                                            |
|    |          | and the role of cytochrome P450. Monooxidase and reductase chains.                                                 | Non-microsomal drug modification phase (alcohol dehydrogenase, xanthine oxidase, mono- and diamine oxidases). |
|    |          | Non-microsomal drug modification phase (alcohol                                                                    | Synthetic phase - conjugation. Examples of type I conjugations                                                |
|    |          | dehydrogenase, xanthine oxidase, mono- and diamine                                                                 | (glucuronic, sulfate, acetyl, thiosulfate, methyl). Examples of type II                                       |
|    |          | oxidases).                                                                                                         | conjugations (glycine, glutathione).                                                                          |
|    |          | Synthetic phase - conjugation. Examples of type I                                                                  |                                                                                                               |
|    |          | conjugations (glucuronic, sulfate, acetyl, thiosulfate, methyl).                                                   |                                                                                                               |
|    |          | Examples of type II conjugations (glycine, glutathione).                                                           |                                                                                                               |
| 14 | 04-08.12 |                                                                                                                    | Test : "Hormones. The blood. Pharmaceutical biochemistry".                                                    |
| 15 | 11-15.12 |                                                                                                                    | Individual work. Admission to the session.                                                                    |

N O T E: The course is fully taught by PhD, university lecturer, Eugen Simionică;

Duration of lectures – 2 hours, practical works – 3 hours.