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FACULTY OF PHARMACY STUDY PROGRAM 0916.1 PHARMACY CHAIR OF GENERAL CHEMISTRY

APPROVED

at the meeting of the Commission for Quality Assurance and Evaluation of the Curriculum faculty of Pharmacy

Minutes No. 4 of 06.06. 201

Chairman, PhD, assocrate professo

Uncu Livia

APPROVED
at the Council meeting of the Faculty of
Pharmacy
Minutes No. 4 of 07, 06, 2019

Dean of Faculty, PhD, associate profes

Ciobanu Nicolae

APPROVED

approved at the meeting of the chair of

General chemistry
Minutes No 15 of 04 June 2019
Head of chair, PhD, associate professor

Cheptanaru Constantin Constantin

SYLLABUS

DISCIPLINE STEREOISOMERISM AND ACTION OF MEDICINES

Integrated studies

Type of course: Optional discipline



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I. INTRODUCTION

• General presentation of the discipline: place and role of the discipline in the formation of the specific competences of the professional / specialty training program

The curriculum of *Stereoisomerism and action of medicines* as pharmacist qualification is a normative pedagogical document and a didactic tool for the efficient organization of the educational process, elaborated based on the Framework Program for Pharmaceutical Higher Education in the Republic of Moldova. Is based on the Charter of the State University of Medicine and Pharmacy "NicolaeTestemitanu", Organization rules of studies in higher education based on the National Credit Studies System, no. 1/8 of 06.04.2017. Organization rules for evaluation and academic performance in State University of Medicine and Pharmacy "NicolaeTestemitanu", no. 5/4 of 12.10.2016, coordinated with the curriculum of pharmaceutical subjects (pharmaceutical chemistry, pharmaceutical biochemistry, drug technology, pharmacology and clinical pharmacy).

• Mission of the curriculum (aim) in professional training

The aim of the discipline *Stereoisomerism and the action of the medicines* is the foundation of the notions of stereoisomerism, the acquisition of knowledge about the importance of the pure enantiomers of the pharmaceutical substances, compared to the racemic mixtures in the treatment of various diseases, knowledge that completes the professional training of future pharmacists.

- Languages of the course: Romanian, English
- Beneficiaries: students of the 3rdyear, faculty of Pharmacy



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II. MANAGEMENT OF THE DISCIPLINE

Code of discipline		S.06.A.059	
Name of the discipline		Stereoisomerism and the action of the medicines	
Person(s) in charge of the discipline		Ph.D in chem., assistant prof. Constantin Cheptanaru, Globa Elena	
Year	III	Semester/Semesters	V
Total number of hours, including:			30
Lectures	15	Practical/laboratory hours	-
Seminars	15	Self-training	-
Clinical internship			
Form of assessment	DC	Number of credits	1

III. TRAINING AIMS WITHIN THE DISCIPLINE

At the end of the discipline study the student will be able to:

• at the level of knowledge and understanding:

- to know the theoretical bases of stereoisomerism of organic compounds.
- to know the specification of the relative configuration and absolute configuration of the stereoisomers.
- to understand the concept of chirality and its importance in the field of medicine.
- to understand the importance of pure enantiomers of drug substances, relative to racemic mixtures, on their pharmacological activity.

• at the application level:

- to determine the belonging of the stereoisomers to the stereochemical series D and L, or R and S.
 - to apply the characterization of optical isomers according to optical activity, relative configuration and absolute configuration of the center of chirality.

• at the integration level:

• to appreciate the importance of stereoisomerism in the context of integration with the disciplines (pharmacology, pharmaceutical chemistry, toxicology, drug technology, pharmacognosis, etc.).



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IV. PROVISIONAL TERMS AND CONDITIONS

Curriculum - organic chemistry.

Skills - for the good grasp of the course, students must have the ability to understand, learn and apply theoretical notions. Students should have the ability to make correlations between taught notions, between course and both practical and interdisciplinary. Thorough knowledge of the organic chemistry: structural isomerism, configurational isomerism and conformational isomerism of organic compounds. Student of the III ^{rd.} year should possess: • knowledge of the language of instruction; • digital competences (use of the Internet, document processing, electronic tables and presentations, use of graphic programs); • ability to communicate and work in team; • qualities - tolerance, compassion, autonomy.

V. THEMES AND ESTIMATE ALLOCATION OF HOURS

Lectures, practical hours/laboratory hours/seminars and self-training

No. Number of ho			r of hours
d/o	THEME		Practical hours
1.	Brief history of molecular chirality. Classification and definition of terminology used in drug stereochemistry (isomerism, enantiomerism, diastereoisomerism, epimer, racemates and racemization, "meso" forms, chiral inversion, distometer, eutomer, enantioselective synthesis). The importance of chirality in pharmacology and current therapeutics.	3	3
2.	Ways of characterization of optical isomers (according to optical activity, relative configuration and absolute configuration of the chirality. <i>Cahn</i> - <i>Ingold</i> - <i>Prelog</i> convention (the order of priority of chirality center substitutes, examples). Fischer projection for representing the relative configuration. Relationship between the attribution of the S-R character (according to the <i>Cahn-Ingold-Prelog</i> convention) and D-L (according to the Fischer projection) (examples).	3	3
3.	The implications of stereochemistry in different therapeutic classes: hypnotic sedatives, general and local anesthetics, opioid analgesics, antidepressants, antiparkinsonian, non-steroidal anti-inflammatory drugs, bronchodilators and antihistamines, antiulcer drugs, preparations used in cardiology, andrenergic, antimycotic.	7	3
4.	Presentation of the "case" of Talidomide and its metabolites. Biochemical route of metabolism of Talidomide. In vivo transformation / racemization of the (R) -Ibuprofen (distomer) isomer into (S) -Ibuprofen (eutomer).		3
5.	Aspects about the discovery of new chiral drugs. Enantioselective enantiomeric preparation of chiral drugs and / or enantiomer separation. Enumeration of enantioselective chemical synthesis and transformation methods.	1	3
	Total	15	15



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VI. REFERENCE OBJECTIVES OF CONTENT UNITS

Objectives Content units Chapter 1. Theoretical basis of the stereoisomerism of organic compounds. 1. The Importance of Chirality in define isomerism. enantiomer, diastereoisomerism. epimer, Pharmacy Pharmacology. racemates and and racemization, "meso" forms, chiral inversion, **Types** of isomers. Getting distomer, eutomer. Chirality. to know the methods of characterization of optic 2. Stereoisomers: enantiomers and (after optical activity, relative diastereomers. The relative isomers and configuration and absolute configuration of the absolute nomenclature. The chirality center. properties of the enantiomers. relationship demonstrate 3. Ways of characterization of optical the between assigning the S-R character (according to the isomers (after optical activity, Cahn-Ingold-Prelog convention) and D-L relative configuration and absolute (according to the Fischer projection). configuration of the chirality to apply the classification and definition of the center. The Cahn - Ingold - Prelog terminology used in the stereochemistry of drugs Convention (the order of priority of to integrate the knowledge gained in the field chirality center substitutes, with the needs of other disciplines in the field of examples) The Fischer projection drug chemistry the representation of the relative configuration. The relation between the attribution of the character SR (according to the Cahn - Ingold - Prelog convention) and DL (according to the Fischer projection)

Chapter 2. The implications of stereochemistry in different therapeutic classes

- To know the role of chirality in the therapeutic action of drugs, metabolic biotransformation.
- to demonstrate stereochemical implications in different therapeutic classes.
- to apply stereochemical notions in the development of new drugs.
- to integrate the knowledge gained in the field with the needs of other disciplines in the field of drug chemistry and pharmacology.
- 1. The implications of stereochemistry in different therapeutic classes.
- 2. The importance of the pure enantiomers of the drug substances, compared to the racemic mixtures, on their pharmacological activity.
- 3. Aspects about the discovery of new chiral drugs. Enantioselective preparation of chiral drugs.



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VII. PROFESSIONAL (SPECIFIC (SC)) AND TRANSVERSAL (TC) COMPETENCES AND STUDY OUTCOMES

✓ Professional (specific) (SC) competences

- PC1.Knowledge of the theoretical bases of the disciplines included in the curriculum of the faculty, general principles in the design, formulation, preparation and conditioning of pharmaceutical and para-pharmaceutical products.
- PC2. Knowing the notions of stereochemistry related to different classes of drugs. Knowing the importance of drug chirality. Knowledge of the importance of stereochemistry in the process of obtaining new pharmaceutical substances. Knowledge of enantiodiscriminatory action of drugs and enantioselective toxicity of chiral drugs.
- PC3. Adoption of messages in various socio-cultural environments, including through multi-language communication, use of problem solving capabilities through interdisciplinary correlation with other fundamental and specialized subjects: pharmaceutical chemistry, pharmacognosis, pharmacology, biochemistry, etc. the bibliographic documentation capacities, the synthesis of the obtained information.

✓ Transversal competences (TC)

- TC1. Obtaining moral markers, forming professional and civic attitudes, allowing students to be honest, honest, nonconflict, cooperative, available to help people interested in community development;
- TC2. Teamwork skills.
- TC3.Using knowledge and skills in new contexts. Openness for continuous education, autonomy and responsibility, observance of professional ethics.

• .Study outcomes

- Understanding the concept of chirality and its importance in the field of medicine.
- Knowledge of aspects specific to the stereochemistry of drugs.
- Knowledge of enantiodiscriminatory action of drugs and enantioselective toxicity of chiral drugs.
- Knowledge of enantioselective methods of chiral drugs.



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VIII. METHODOLOGICAL SUGGESTIONS FOR TEACHING-LEARNING-ASSESSMENT

Teaching and learning methods used

The discipline of *Stereoisomerism and the action of the medicines*taught is taught in classical ways: lectures, practical works. At the lectures, the theoretical course will be read by the course holders. At practical work, students will broaden their knowledge in the importance of pure enantiomers of pharmaceutical substances, as compared to racemic mixtures in the treatment of various diseases.

• Applied teaching strategies / technologies (specific to the discipline)

To succeed in the discipline of *Stereoisomerism and the action of the medicines*the student should actively work both in courses and seminars, as well as in his own right, and the teacher should use the didactic technologies specific to the discipline. The most important methods in teaching organic chemistry are *problematization and brainstorming*.

Brainstorming is a technique of group creativity designed to generate a large number of ideas to solve a problem.

Problematization called and teaching through problem solving or, more specifically, teaching through productive problem solving. A didactic method consisting in putting in the minds of the students some deliberately created difficulties in overcoming which, by their own effort, the student learns something new.

• *Methods of assessment* (including the method of final mark calculation)

Current: front and / or individual control via

- 1. solving problems / exercises,
- 2. analysis of case studies
- 3. current quiz.

Final:

Differentiated colloquium (semester V).

The final grade at the differentiated colloquium in the fifth semester will be made up of the average grade from one current quiz, the grid test and the oral test.

The topics for the differential colloquium are approved at the department meeting and are brought to the attention of the students with at least one month until the session.



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Method of mark rounding at different assessment stages

Intermediate marks scale (annual average, marks from the examination stages)	National Assessment System	ECTSEquivalent	
1,00-3,00	2	F	
3,01-4,99	4	FX	
5,00	5		
5,01-5,50	5,5	E	
5,51-6,0	6		
6,01-6,50	6,5	D	
6,51-7,00	7		
7,01-7,50	7,5	C	
7,51-8,00	8		
8,01-8,50	8,5	В	
8,51-8,00	9		
9,01-9,50	9,5	- A	
9,51-10,0	10		

The average annual mark and the marks of all stages of final examination (computer assisted, test, oral) - are expressed in numbers according to the mark scale (according to the table), and the final mark obtained is expressed in number with two decimals, which is transferred to student's record-book.

Absence on examination without good reason is recorded as "absent" and is equivalent to 0 (zero). The student has the right to have two re-examinations.



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IX. RECOMMENDED LITERATURE:

A. Compulsory:

- 1. ZURABYAN S.E. Fundamentals of bioorganic chemistry. GEOTAR-Media publishing group, 2017.
- 2. STEVEN S. ZUMDAHL. Chemistry. Lexington, Massachusetts, Toronto, 1986.
- 3. FRANCIS MARION MILLER. *Chemistry, Structure and dynamics*. McGraw-Hill book company, USA., 1984.

B. Additional

- 1. NENIȚESCU C. D. Chimie organică. B.: "Regia Autonomă Monitorul Oficial", 2015.
- 2. ВОРОНКОВ Л. Г. Клиническое использование хиральных молекул как новое направление в кардиоваскулярной фармакотерапии. Киев: Национальный научный центр «Институт кардиологии им. Н.Д. Стражеско» АМН Украины.
- 3. НЕСУКАЙ Е. Г. Левовращающий переворот в кардиологии, Артериальная гипертензия. № 1(15), 2011, с.14-18.