Gene Therapy
Code: 100901
ECTS Credits: 6

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<tr>
<th>Degree</th>
<th>Type</th>
<th>Year</th>
<th>Semester</th>
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<tr>
<td>2500252 Biochemistry</td>
<td>OT</td>
<td>4</td>
<td>0</td>
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</tbody>
</table>

Contact
Name: Maria Fátima Bosch Tubert
Email: Fatima.Bosh@uab.cat

Use of languages
Principal working language: catalan (cat)
Some groups entirely in English: No
Some groups entirely in Catalan: Yes
Some groups entirely in Spanish: No

Teachers
Pedro José Otaegui Goya
Miguel García Martínez
Federico Mingozzi
Ivet Elias Puigdomenech
Verónica Jiménez Cenzano

Prerequisites
There are no prerequisites to attend this course. However, to facilitate the student's understanding of the subject matter and the achievement of the learning goals proposed, it is advisable that the student has previous knowledge on Cellular Biology, Genetics, Molecular Biology and Recombinant DNA technology.

It is also advisable that the students have basic knowledge of English, so that they can use the information sources of the field, which are mostly in this language.

Objectives and Contextualisation
The objective of the subject "Gene and Cell Therapy" is to provide the students with up-to-date knowledge in the methodologies available for the genetic engineering of cells for therapeutic purposes. Thus, the content of the subject will cover the following topics: Description of in vivo and ex vivo gene therapy; Study of the different vectors used for viral and non-viral gene transfer and their respective advantages and shortcomings; study of the possible routes of administration of the vectors and the applications to the treatment of hereditary and non-hereditary human diseases. Description of cell therapy. Transplant of fully differentiated somatic cells (pancreatic islets, hepatocytes, and bone marrow). Transplant of pluripotent stem cells (embryonic and adult). Sources of cells for cell therapies. Therapeutic applications of cell therapies. Biosafety, ethics and legal aspects associated to the use of cell therapies in humans.

Skills
Learning outcomes

1. Apply general laboratory security and operational standards and specific regulations for the manipulation of different biological systems.
2. Be able to self-evaluate.
3. Collaborate with other work colleagues.
4. Combine research and and the generation of knowledge with problem-solving in one's own field, showing sensibility to ethical and social questions.
5. Describe the structural, physiological and biochemical characteristics of the different types of cells and explain how their properties fit in with their biological function.
6. Interpret experimental results and identify consistent and inconsistent elements.
7. Make an oral, written and visual presentation of one's work to a professional or non-professional audience in English and understand the language and proposals of other specialists.
8. Read specialised texts both in English and one's own language.
9. Show a capacity for leadership.
10. Show initiative and an entrepreneurial spirit.
11. Stay abreast of new knowledge of the structure, organisation, expression, regulation and evolution of genes in living beings.
12. Take responsibility for one's own learning after receiving general instructions.
13. Think in an integrated manner and approach problems from different perspectives.
14. Use ICT for communication, information searching, data processing and calculations.

Content

Knowledge on the following topics will be imparted during the theoretical classes:

**TOPIC 1**

Molecular basis of Gene Therapy. Gene therapy *in vivo* and *ex vivo*. Introduction to Cell therapy.

**TOPIC 2**

**TOPIC 3**


**TOPIC 4**


**TOPIC 5**


**TOPIC 6**


**TOPIC 7**


**TOPIC 8**

Interference RNA (siRNA). Gene therapy based on the use of siRNA. Applications.

**TOPIC 9**

Genome editing. Different types of nucleases: Meganucleases, Zinc-finger nucleases, TALENs, CRISPR Technologies. Applications in the field of Gene Therapy.

**TOPIC 10**

*Ex vivo gene therapy;* hematopoietic stem cells.

**TOPIC 11**

Introduction to Embryonic Stem Cells (*ES cells*). Differentiation of *cells*. Clinical Applications of *ES cells*.

**TOPIC 12**

Adult stem cells and their applications.

**TOPIC 13**

Reprogramming and *Induced Pluripotent Stem Cells (iPS)*.

**TOPIC 14**

TOPIC 15

TOPIC 16

TOPIC 17
Cell therapies for the regeneration of bone and cartilage.

TOPIC 18
Cell therapies for the regeneration of skin. Cell therapies for the regeneration of eye diseases.

TOPIC 19
Cell therapies for Parkinson disease and other Central Nervous System diseases.

TOPIC 20
Cell therapies for diabetes mellitus and cardiovascular diseases.

TOPIC 21

Three exercises are proposed for the laboratory practice classes. The first assignment is the design of a gene or cell therapy strategy for a given disease, from design of the gene or cell product to proof-of-concept studies and translational studies that allow a clinical trial in humans. This exercise is proposed as a review of all the concepts introduced in the theory classes. The second proposed activity allows students to acquire experience in the use of laboratory animals in gene and cell therapy studies. Finally, in the third proposed activity we show students how to perform a partial hepatectomy for gene therapy approaches directed to the liver.

Content of the laboratory practice classes:
- Design of a gene or cell therapy strategy; discussion of the different options.
- Introduction to the use of laboratory animals in gene and cell therapy studies.
- Methods and routes of administration of viral and non-viral vectors.

Methodology
The subject "Gene and Cell Therapy" consists of theory and laboratory classes, and tutored oral presentations of relevant literature. The formative activities of the subject are complementary.

Theoretical classes
The contents of the theoretical classes will be imparted by a Professor in a series of master classes supported by audio-visual material. The slides used by each professor in each class will be available to the students through the subject's Campus Virtual/Moodle. These master classes will constitute the main form of transfer of theoretical contents. Students are advised to periodically consult the books and links suggested in the Bibliography section of this document and at the Campus Virtual/Moodle to consolidate and clarify, if necessary, the contents explained in class.
### Laboratory practice classes

The laboratory practice classes have been designed to help students get familiarized with the methodologies used in the field of gene therapy and have a direct experience in the handling of laboratory animals for gene therapy studies. We expect that, during these laboratory practice classes, students will be able to experience a "real world" experimental situation. We would like students to experience the excitement associated to research in the field of gene therapy.

The laboratory practice classes are composed of 3 sessions of 4 h each (from 3PM to 7PM), during which students will work in groups of 2 people under the supervision of an experienced professor. The dates assigned to each laboratory practice group will be published in the subject's Campus Virtual/Moodle with sufficient anticipation.

Attendance to laboratory practice classes is mandatory.

The laboratory practice guide will be available through the Campus Virtual/Moodle. Students must bring their own lab coat, a waterproof marker and the Laboratory Practice Guide to each laboratory practice class.

### Oral presentations of selected papers

Students will analyse and discuss in an oral presentation in front of the whole class a selected recent scientific publication on gene or cell therapy published in a recognized international scientific journal. To this end, students will pair with a fellow classmate. During the process of analysis of the paper's content and preparation of the oral presentation, students will be tutored by researchers with experience in the field of gene and cell therapy. Students will have 10 minutes for the oral presentation, equally divided amongst the members of the group, plus 5 minutes for questions (total of 15 minutes). The objective of this evaluating activity is that students get used -under the supervision of a tutor- to the process of searching, reading and understanding of scientific literature, and if necessary, develop a critical view on the figures, tables and results described in the publication. On the other hand, with this activity students will increase their knowledge of the current applications of gene and cell therapy technologies.

### Tutoring

The oral presentations of selected papers will be tutored. In addition, upon request from the students, individualized tutoring will be available throughout the course. The objective of this sessions will be to help the student resolve doubts and review basic concepts and to provide them with advice on sources of information and the best way to discuss scientific results in public.

### Activities

<table>
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<tr>
<th>Title</th>
<th>Hours</th>
<th>ECTS</th>
<th>Learning outcomes</th>
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<tr>
<td><strong>Type: Directed</strong></td>
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<td></td>
<td></td>
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<tr>
<td>Oral presentations</td>
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<td>0.32</td>
<td>16, 3, 5, 6, 7, 10, 15, 11, 8, 14, 2, 12, 13</td>
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<tr>
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<tr>
<td>Theoretical lessons</td>
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<td>1.36</td>
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<td><strong>Type: Supervised</strong></td>
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<tr>
<td>Tutorials</td>
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<td>0.24</td>
<td>3, 7, 15, 11, 8, 14</td>
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<td><strong>Type: Autonomous</strong></td>
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<tr>
<td>Individual study time</td>
<td>74</td>
<td>2.96</td>
<td>16, 5, 6, 7, 10, 15, 11, 4, 8, 14, 2, 12, 13</td>
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</table>
Evaluation

To pass the course, students must achieve a final score of 5 points (over a total of 10 points) and must attend the laboratory practice classes. The evaluation activities are:

1.- Final examination of theoretical classes

Accounts for 50% of the final score (5 points out of 10). Assessment will consist of a written examination, under the format of a True or False test, on topics explained during the theoretical classes. A score greater than 2.5 in this examination is required to pass the course.

There will be a Second Chance/Recovery Exam, under the same format as the original exam.

2.- Examination of Laboratory classes

Accounts for 15% of the final score (1.5 points out of 10). Assessment will consist of a written examination, under the format of a True or False test, on topics explained during the practical classes. It will be held at the end of practical classes period.

Attendance to practical sessions (or field trips) is mandatory. Students missing more than 20% of programmed sessions will be graded as "No Avaluable.

3.- Self-study exercise

Accounts for 10% of the final score (1 point out of 10). Assessment will consist of an exercise that the student will have to develop on their own. Details will be posted online in the "Campus Virtual" at the end of April.

4.- Oral presentations of selected research papers

Accounts for 15% of the final score (1.5 points out of 10). Students will be evaluated individually, both on their performance during the oral presentation of the selected paper and on the audio-visual material that they prepared to support their group presentation.

5.- Attendance to the oral presentations of research papers

Accounts up to 10% of the final score (1 point out of 10). Both attendance and participation in the scientific discussions of the sessions will be evaluated, following the scale:

Attendance 90-100% = 1 point
Attendance 80-89% = 0,8 points
Attendance 70-79% = 0,7 points
Attendance 60-69% = 0,6 points
Attendance 50-59% = 0,5 points
Attendance 0-49% = 0 points

To be eligible for the retake process, the student should have been previously evaluated in a set of activities equaling at least two thirds of the final score of the course or module. Thus, the student will be graded as "No Avaluable" if the weight of all conducted evaluation activities is less than 67% of the final score.
Evaluation activities

<table>
<thead>
<tr>
<th>Title</th>
<th>Weighting</th>
<th>Hours</th>
<th>ECTS</th>
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<td>16, 1, 3, 5, 6, 7, 10, 15, 9, 11, 4, 8, 14, 2, 12, 13</td>
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<td>Oral presentations of selected research papers</td>
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<td>1</td>
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<td>16, 1, 3, 5, 6, 7, 10, 15, 11, 4, 8, 14, 2, 12, 13</td>
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<tr>
<td>Self-study exercise</td>
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<td>1</td>
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<td>16, 1, 3, 5, 6, 7, 10, 15, 9, 11, 4, 8, 14, 2, 12, 13</td>
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Bibliography


