



UNIVERSITÀ  
DEGLI STUDI DI MILANO-BICOCCA

## COURSE SYLLABUS

### Analysis, Control and Optimization of Biological Systems

2425-1-F0802Q079

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#### Aims

The course aims to train students in the analysis of biological systems in their most generic and broad sense (networks of biochemical reactions, metabolic and transcription networks, single cells or populations dynamics), to give them the basis for their control and the methods of parametric identification.

##### Knowledge and understanding:

At the end of the course, students will be able to obtain quantitative and qualitative information on the behavior of biological systems through qualitative analysis and simulations of the mathematical models that describe them. The tools for the numerical simulation in MATLAB of the studied biological systems will also be given.

##### Ability to apply knowledge and understanding:

At the end of the course, students will be able to apply the acquired methodologies to complex biological systems of various kinds, not necessarily covered in class

##### Autonomy of judgment:

Students will be able to re-elaborate and apply the most appropriate analysis methodologies learned, according to the biological contexts investigated.

##### Communication skills:

At the end of the course, students will be able to express themselves appropriately in the description of the topics addressed with properties of language and confidence in exposure.

##### Learning ability:

At the end of the course, students will be able to analyze, apply, integrate and connect the knowledge acquired - and subsequently matured with the consultation of the literature - with what they have learned in related courses, in order to solve scientific problems in Biological Sciences and Biotechnologies.

## Contents

This course provides methodologies for the analysis, identification and control of specific computational models of biological systems. The methodologies can be divided into 3 main categories: (i) qualitative analysis (what are the emergent properties of a system?); (ii) quantitative analysis (how can I simulate the computational model of a biological system?); (iii) control (what control mechanisms are apparent in nature? What kind of specifications can I meet in a control system for biological systems?)

The examples of the course will deal with different biological contexts, including transcription networks, metabolic networks, cell cycle and growth models, epidemic spreading models. Most topics will be analyzed through multidisciplinary analysis of specific case studies. One or more of the case studies will also be analyzed in the course Systems Biochemistry with a complementary biological, biochemical and molecular perspective

## Detailed program

- Input/output relationships: a biological system is characterized by the richness of laboratory experiments. State/inputs/outputs of a system. Discrete-time and continuous-time systems. Stationary systems. Linear systems. Free and forced evolution. Regime and transient response.
- Analysis of the qualitative behavior of a system. Equilibrium points, stability and multi-stability. Exponential growth. Bifurcations, oscillations and limit cycles. Chaotic behaviors. Examples on models of transcription networks, enzymatic reactions, cell growth, spread of epidemics.
- Systems identification. Estimating model parameters. Under/overdeterminate system. Deterministic approach: least squares and recursive least squares. Analytical solution for linear relationships. Case study: integration of kinetic, metabolic and proteomic data for the identification of a metabolic network.
- Simulation of deterministic models of biological systems. All simulations will be carried out in MATLAB environment.
- Control of biological systems. Feedback control mechanisms existing in nature (negative self-regulation in transcription networks, optimal expression of genes, glucose-insulin regulation system, metabolic engineering).

## Prerequisites

No specific knowledge of mathematics other than those already acquired in the basic courses of a three-year degree in Biological Sciences or Biotechnology is required.

## Teaching form

All didactic activities are conveyed by means of face-to-face lectures

- 30 hours of delivered/interactive didactics
- 1CFU (7 hours) of interactive teaching, in collaboration with the teacher of Systems Biochemistry (Prof. Marco Vanoni) to jointly address a case study. This didactic approach will allow for presentation and examination from complementary points of view pathways, functions and biological systems seen through the eye of biologist and computational modeller.
- 8 hours of in-person tutorial activities aimed at guiding and assisting students throughout hands-on applications by interactive teaching (Didattica Interattiva, DI)

## **Textbook and teaching resource**

The slides and MATLAB codes developed at lessons will be available on the e-learning page of the course. Further review articles and book chapters will be recommended in class and uploaded to the e-learning platform of the course

The following texts are recommended for appropriate further information:

- U. Alon, An introduction to systems biology: design principles of biological circuits, Chapman & Hall/CRC, Taylor & Francis Group, 2019
- E. Klipp, W. Liebermeister, C. Wierling, A. Kowald, Systems Biology – A textbook. 2nd Ed. Wiley, 2016

## **Semester**

Second semester

## **Assessment method**

The exam is divided into two parts:

- In the first part, the student will be evaluated (orally) on the in-depth analysis of a scientific article previously assigned to them
- Oral questions on the entire syllabus covered in class.

In both cases, the student will be assessed on their ability to express and summarize, as well as their understanding of the topics covered.

## **Office hours**

Students are invited to contact the teacher by email to agree upon a date (also on WebEx)

## **Sustainable Development Goals**

GOOD HEALTH AND WELL-BEING

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