

Introduction to Enzyme Mechanisms

CHM 6301

Spring 2012

Instructor: Steven Bruner

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Office hours: Mondays 1:30-2:30, Wednesdays 10-11 and Thursdays 2-3 or by appointment (email).

Course overview: This course will survey the chemical mechanism and function of enzymes

Prerequisites: CHM 3218, BCH4024

Course Requirements:	Exam #1	2/10/2012	20%
	Exam #2	3/19/2012	20%
	Exam #3	4/16/2012	20%
	Problem sets		10%
	Presentation		20%
	Participation		10%

Course objectives: Understand the chemical mechanisms behind enzyme catalyzed reactions, experimental methodology used to decipher enzyme chemistry and mechanisms of action of common pharmaceuticals.

Recommended textbook: Organic Chemistry of Enzyme-Catalyzed Reactions, Richard B. Silverman, revised edition, 2002.

Handouts, journal articles and additional course material will be posted online (<https://lss.at.ufl.edu>). We will also make extensive use of the primary literature and references will be provided in class.

Attendance: Successful performance in the course requires regular attendance, though no formal monitoring of attendance will occur. Exams that are missed due to an excused absence can be taken at an arranged time. An absence is considered excused if there is an acceptable reason according to UF policy (<http://www.registrar.ufl.edu/catalog/policies/regulationattendance.html>). Otherwise, the absence will be considered unexcused and will result no credit.

Academic Honesty: Any act of academic dishonesty will be reported to the Dean of Students, and may result in failure of the assignment in question and/or the course. For University of Florida's honor code, see <http://www.dso.ufl.edu/sccr/honorcodes/honorcode.php>.

Accommodations for Students with Disabilities Students requesting classroom accommodation must first register with the Dean of Students Office. The Dean of Students Office will provide documentation to the student who must then provide this documentation to the Instructor when requesting accommodation. Contact the Disability Resources Center (<http://www.dso.ufl.edu/drc/>) for information about available resources for students with disabilities.

Grading scheme: At the end of the semester, grades will be curved so the median for the class is approximately a "B+".

Selection of topics to be covered in the course:

- Introduction to enzyme catalysis
- General themes of how enzymes work
- The perfect enzyme: triosephosphate isomerase

- Mechanism of proteases, especially HIV protease
- Design and development of HIV protease inhibitors
- Aminotransferases and other enzymes utilizing small diffusable reactants
- Chemistry of phosphatases/kinases
- Regulation and role in systems biology
- Development of kinase inhibitors and chemical genetics
- Cofactor chemistry: redox chemistry of NADH
- DHQ synthase
- Cofactor chemistry of flavins.
- Radical clocks as mechanistic probes
- Baeyer-Villiger monooxygenase
- Cofactor chemistry: PLP
- Diversity of PLP chemistry
- Cofactor chemistry: thiamin
- Benzoin condensation / Baylis Hillman, pyruvate dehydrogenase
- Cofactor chemistry: methylations
- Metalloenzymes
 - Introduction to heme and nonheme iron/cobalt
 - Cytochrome P450s
 - Vitamin C and non-heme iron enzymes
 - Proline hydrolase and scurvy
- Cobalamin / vitamin B12
- Complex non-heme metalloenzymes
 - Methane monooxygenase, Isopenicillin synthase, ACC oxidase
- Prostaglandins and the enzymology of pain killers
- Biosynthesis and function of alkaloid opiates
- Enzymology of Viagra
 - NO synthase
- Multicomponents, complex metallocluster enzymes.