**Instructor:** K.C. Nicolaou, BRC 363

kcn@rice.edu, 713-348-8860

**Office Hours:** Immediately after lectures or by appointment. There is a TA for this course.

**Goals:** This course is about the art and science of total synthesis of natural and designed molecules.

Through select case studies, students learn several aspects of total synthesis, including the motivation for the synthesis, retrosynthetic analysis, synthetic strategy design, method development, chemical biology studies, and drug discovery and development. Emphasis is placed on both

knowledge and creativity.

Prerequisites: CHEM 211 AND CHEM 212 AND CHEM 401 AND CHEM 442

Required Texts: 1. K.C. Nicolaou and Jason Chen, Classics in Total Synthesis III, Wiley-VCH, 2011.

ISBN: 3-527-32957-9. Available from Wiley-VCH or Amazon

(soft cover: \$83.54; also available in Rice Bookstore).

Recommended Books:

1. K.C. Nicolaou and Scott Snyder, Classics in Total Synthesis II, Wiley-VCH, 2003.

ISBN: 3-527-30684-6. (Available in Rice Bookstore.)

2. K.C. Nicolaou and E.J. Sorensen, Classics in Total Synthesis, VCH, 1996.

ISBN: 3-527-29231-4.

3. K.C. Nicolaou and Tamsyn Montagnon, *Molecules That Changed the World*, Wiley-VCH, **2008**.

ISBN: 3-527-29284-5.

4. László Kürti and Barbara Czakó, Strategic Applications of Named Reactions in Organic

*Synthesis*, Academic Press, **2005**. ISBN: 978-0-12-429785-2.

5. E.J. Corey, László Kürti and Barbara Czakó, Molecules and Medicine, Wiley, 2007.

ISBN: 978-0-470-22749-7.

6. E.J. Corey and X. Cheng, *The Logic of Chemical Synthesis*, Wiley, **1989**.

ISBN: 0-471-1159-0.

7. E.J. Corey and László Kürti, Enantioselective Chemical Synthesis, Direct Book Publishing, 2010.

ISBN: 978-0-615-39515-9.

Lectures and Tutorials:

Lectures will be held in the BioScience Research Collaborative (BRC), Room 285, on Tuesdays and Thursdays from 9:25-10:40 a.m. Tutorials will be held in BRC, Room 308, from 7:00-8:00 p.m.

on Tuesday nights.

Exams and Assignments:

Exam: A comprehensive exam will be given on Thursday, February 25 during the normal class

period. No make up exams will normally be allowed.

Assignment 1: Analysis of the total synthesis of a schindalactone A (*Angew. Chem. Int. Ed.* **2011**, 50, 7373) in the classics style (5 pages total, including structures). Point out the innovative aspects

of the work and the novel mechanisms of the key reactions involved.

Assignment 2: Proposal of a natural product total synthesis (3 pages, including schemes, excluding references). Target molecule must be cleared with instructor in advance. Emphasis should be

placed on novel and imaginative synthetic strategies and modern methods.

Reading before each class: By doing so, you will be able to participate in class more effectively.

Grades: Exam: 250 points

Assignment 1: 125 points Assignment 2: 125 points

**Disabilities:** Student disabilities will be accommodated in accordance with the guidelines put forth by Disability Support Services (DSS). Students should register with the DSS Office in Allen Center and let the

instructor know of their disabilities and needs as soon as possible.

Honor Code: The Rice University Honor Code applies to this course. Both assignments 1 and 2 are to be

exclusively the work of each individual student. Proper references must be given to the original

work when included in these home assignments.

## Schedule (Subject to Change)

Week 1	Tuesday, <b>Jan 12</b>	Lecture 1	Reserpine	Chapter 4, Classics I
				(handout will be given)
	Thursday, <b>Jan 14</b>	Lecture 2	Ginkgolide B	Chapter 25, Classics I
Week 2	marsaay, <b>Jan</b> 14	Locidio 2	Giringolide B	(handout will be given)
				(Harideat IIII 20 given)
	Tuesday, <b>Jan 19</b>	Lecture 3	Taxol	Chapter 34, Classics I
				(handout will be given)
Week 3	Thursday, <b>Jan 21</b>	Lecture 4	Tetrodotoxin	Chapter 2, Classics III
	Tuesday, <b>Jan 26</b>	Lecture 5	Littoralisone, Oseltamivir,	Chapter 7, Classics III
			Hirsutellone B	
Week 4	Thursday, Jan 28	Lecture 6	Cyanthiwigns U & F	Chapter 9, Classics III
	Tuesday, Feb 2	Lecture 7	Sporolide B	Chapter 22, Classics III
	Thursday, <b>Feb 4</b>	Lecture 8	Lomaiviticin A	J. Am. Chem. Soc. <b>2011</b> , 133, 7260
Week 5	marcady, 1 CD	Locidio	Zomarviion / t	(reprint will be given)
	Tuesday, <b>Feb 9</b>	Lecture 9	Daphmanidin E	Angew. Chem. Int. Ed. <b>2011</b> , 50, 11501
				(reprint will be given)
	Thursday, Feb 11	Lecture 10	Acetylaranotin	J. Am. Chem. Soc. <b>2012</b> , 134, 1930
Week 6				(reprint will be given)
	Tuesday, <b>Feb 16</b>	Lecture 11	Viridicatumtoxin B	Angew. Chem. Int. Ed. <b>2013</b> , 52, 8736
	. assaay, r ca	200101011	Vindioataintexiii 2	(reprint will be given)
	Thursday, <b>Feb 18</b>	Lecture 12	Ryanodol	J. Am. Chem. Soc. <b>2014</b> , 136, 5916
				(reprint will be given)
Week 7	Tuesday, <b>Feb 23</b>	Lecture 13	Taiwaniadducts B, C & D	J. Am. Chem. Soc. <b>2014</b> , 136, 8185
				(reprint will be given)
	Thursday, <b>Feb 25</b>	Exam		Assignments 1 and 2 Due
				by Tuesday, March 1