Molecular Toxicology: Chemical and Biological Mechanisms

GGPS-PHRM 590

Course Director: Trevor M. Penning, Professor of Systems Pharmacology and Translational

Therapeutics

Email: penning@upenn.edu www.med.upenn.edu/ceet

Course Goals: Exposures to foreign compounds (drugs, carcinogens, and pollutants) can disrupt normal cellular processes leading to toxicity. This course will focus on the molecular mechanisms by which environmental exposures lead to end-organ injury and to diseases of environmental etiology (neurodegenerative and lung diseases, and reproduction and endocrine disruption). Students will learn the difficulties in modeling response to low-dose chronic exposures, how these exposures are influenced by metabolism and disposition, and how reactive intermediates alter the function of biomolecules. Mechanisms responsible for cellular damage, aberrant repair, and end-organ injury will be discussed. In addition, students will discuss the relationship between genetics and epigenetics and environmental exposures. Students will learn about modern predictive toxicology to classify toxicants, predict individual susceptibility and response to environmental triggers, and how to develop and validate biomarkers for diseases of environmental etiology. Students are expected to write a term paper on risk assessment on an environmental exposure using available TOXNET information. This course is required for those pursuing the Certificate Program in Environmental Health Sciences.

Lecture Course: 60 minute lectures meets twice per week on Monday's and Wednesday's and also some

Friday's

Course Unit: 1 credit unit

Proposed Days: Mon and Weds (and some Friday's)

Semester: Spring

Course Materials: Casarett & Doulls: Toxicology: The Basic Science of Poisons (7th or 8th Edition) and

relevant literature.

Pre-requisites: Undergraduate course work in biochemistry and chemistry essential. Exceptions allowed

based on past course work. Please consult with the Course Director.

Students: All 1st and 2nd year BGS students with required prerequisites; residents in Environmental

and Occupational Health, and professional masters students (MPH and MTR).

Molecular Toxicology Course Schedule

Mondays and Wednesdays: 2:00PM- 3:00 PM. (NOTE: Some classes are 2p – 4p and also some Fridays as highlighted in yellow below) The Risk Assessment Presentations and Final Exam also occur during different timeframes as noted in yellow below.

Location: 10-146 Smilow Center for Translational Research

2020 Spring Schedule

Week	Date	Location	Торіс	Lecturer		
	General Principles					
1	M, Jan 13	10-146 SCTR	Introduction /Orientation	Penning		
	W, Jan 15	10-146 SCTR	Regulatory Policy and EPA	Pepino		
	F, Jan 17	10-146 SCTR	General Principles: Dose Response &	Penning		
			Exposures			
2	M, Jan 20		No Class - Martin Luther King Day			
	W, Jan 22	10-146 SCTR	Heavy Metal Toxicity	Howarth		
	F, Jan 24	10-146 SCTR	Reactive Oxygen Species	Ischiropoulos		
3	M, Jan 27	10-146 SCTR	Metabolism Phase I	Penning		
	W, Jan 29	10-146 SCTR	Metabolism Phase II	Penning		
	F, Jan 31	10-146 SCTR	Chemical Carcinogenesis by	Penning		
			Genotoxic Agents			
4	M, Feb 3	10-146 SCTR	Chemical Carcinogenesis by Non-	Penning		
			Genotoxic			
	W, Feb 5	10-146 SCTR	Mutagenesis / Mutational Signatures	Field		
5	M, Feb 10	10-146 SCTR	DNA Adducts and their Repair	Penning		
	W, Feb 12	10-146 SCTR	Mitochondrial Dysfunction	Blair		
			nvironment Interactions			
6	M, Feb 17	10-146 SCTR	Epigenetics	Heller		
	W. Feb 19	10-146 SCTR	Transcriptome-Analysis-Technologies	Tobias		
			and Experimental Design			
	F, Feb 21	10-146 SCTR	Folate and Methylation	Whitehead		
7	M, Feb 24	10-146 SCTR	Toxicogenetics – Toxicology and	Burczynski		
	<u>2:00-4:00</u>		DNA Variation			
			Toxicogenomics- Toxicology and			
	W E 1 26	10.146 CCTD	RNA Expression	D .		
	W, Feb 26	10-146 SCTR	Risk Assessment Assignment	Penning		
	F, Feb 28	10-146 SCTR	Midterm (2:00 – 5:00 PM)			
			Exposure Science			
8	M, Mar 2	10-146 SCTR	Protein-Biomarkers-Proteomics	Mesaros		
	W, Mar 4	10-146 SCTR	Exposure and Response Biomarkers	Mesaros		
	F, Mar 6		Biosensors	Johnson		
9	M, Mar 9		Spring Break			
	W, Mar 11		Spring Break			

Week	Date	Location	Торіс	Lecturer			
		Org	an-Based Toxicology				
Lung and Airway-Disease							
10	M, Mar 16	10-146 SCTR	Mesothelioma	Moon			
	W, Mar 18	10-146 SCTR	Toxic Responses of the Respiratory	Christofido			
			System	Solomidou			
11	M. Mar 23	10-146 SCTR	Lung Cancer	Vachani			
	W. Mar 25	10-146 SCTR	Inhalation Toxicology 1: Respiratory	Joseph Jude			
	2:00 -4:00		Physiology	(Rutgers)			
			Inhalation Toxicology 2:				
	Mechanisms of Lung Injury						
	Nervous System						
12	M, Mar 30	10-146 SCTR	Mechanisms of Neurotoxicity	Ischiropoulos			
	W, Apr 1	10-146 SCTR	Overview of the Nervous System and	Robinson			
12	NA A	10.146 CCTD	Neurotoxicants	***			
13	M, Apr 6	10-146 SCTR	Sleep Disturbance and Neurodegenerative Disease	Veasey			
		Reproducti	ive & Endocrine Disruption				
	W, Apr 8	10-146 SCTR	In utero Genetic Imprinting	Bartolomei			
14	M, Apr 13	10-146 SCTR	Mechanisms of Reproductive	Gerton			
14	WI, Apr 13	10-140 501K	Disruption-Male	Gerton			
	W, Apr 15	10-146 SCTR	Environmental Reproductive	Burris			
	, 1		Epidemiology				
			tion & Predictive Toxicology				
15	M, Apr 20	10-146 SCTR	Data-Integration-Bioinformatics	Weljie			
	W, Apr 22	10-146 SCTR	Exposure Biology Informatics	Jason Moore			
	F, Apr 24	10-146 SCTR	Predictive Toxicology and TOX 21st	Penning			
16	34 4 25	10.146 CCTD	Century	T2: 11 TT (1			
16	M, Apr 27	10-146 SCTR	Risk Assessment Presentations – 9:00 am- 11:00 am	Field, Howarth,			
			am- 11:00 am	Pepino, Penning			
	W, Apr 29	10-146 SCTR	Final Examination – 9:00am -	1 chining			
	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	10 110 5011	12:00pm				

Evaluation: Mid-Term: 30% Final Exam: 40%

Risk Assessment Paper: 30% Text: Cassarett & Doull's: Toxicology: The Basic Science of Poisons

Lecturers:	Email Address
Marisa Bartolomei	BARTOLOM@PENNMEDICINE.UPENN.EDU
IAN A BLAIR, PhD	ianblair@upenn.edu
Ted Burczynski	tedburczynski@gmail.com
Melpo Christofidou-Solomidou	melpo@pennmedicine.upenn.edu
Jeff Field	jfield@upenn.edu

Lecturers:	Email Address
George Gerton	GERTON@PENNMEDICINE.UPENN.EDU
Harry Ischiropoulos	ISCHIROP@PENNMEDICINE.UPENN.EDU
Charlie Johnson	cjohnson@physics.UPENN.EDU
Joseph Jude (New)	joseph.antonyjude@rutgers.edu
Jianghong Liu	jhliu@nursing.upenn.edu
Clementina Mesaros	mesaros@pennmedicine.upenn.edu
Edmund Moon	Edmund.Moon@uphs.upenn.edu
Sam Parry	parry@pennmedicine.upenn.edu
Trevor Penning	penning@upenn.edu
Richard Pepino	rpepino@sas.upenn.edu
Michael Robinson	Robinson@pennmedicine.upenn.edu
Becky Simmons	rsimmons@pennmedicine.upenn.edu
Carsten Skarke	cskarke@pennmedicine.upenn.edu
John Tobias	jtobias@pcbi.upenn.edu
Sigrid Veasey	veasey@pennmedicine.upenn.edu
Paul Wang	wangpaul@pennmedicine.upenn.edu
Aalim Weljie	aalim@pennmedicine.upenn.edu
Steve Whitehead	aswhiteh@upenn.edu