Molecular Toxicology: Chemical and Biological Mechanisms GGPS-PHRM 590

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

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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 (7 th – 9 th Edition) and
	relevant literature.
<u>Pre-requisites:</u>	Undergraduate course work in biochemistry and chemistry essential. Exceptions allowed based on past course work. Please consult with the Course Director.
Students:	All 1 st and 2 nd year BGS students with required prerequisites; residents in Environmental and Occupational Health, and professional masters students (e.g. MPH and MTR).

Molecular Toxicology Course Schedule- <u>IMPORTANT PLEASE READ</u>

Mondays, Wednesdays and Fridays: <u>2:00PM-3:00 PM</u> and other classes are <u>2PM-4PM</u> as highlighted in grey on the schedule. As noted, some classes are live via bluejeans and others are prerecorded and viewable on mediasite. All live bluejeans classes are highlighted in <u>yellow</u> on the schedule. The Risk Assessment Presentations occur during different timeframes as noted in <u>yellow</u> below and both the midterm and final exams are in a take home format.

There are also optional live Bluejeans tutorials with Dr. Trevor Penning every Friday from 3p - 4p beginning on Friday, January 29^{th} . The purpose of these tutorials is to provide the students with an opportunity to address any questions or issues that they are having with the course or course materials.

BlueJeans link for all live sessions is <u>https://upenn.bluejeans.com/5348332877</u>. Mediasite link for all recordings is <u>https://mediasite.med.upenn.edu/Mediasite/Catalog/Full/bcaaf0fbc98a4f02abdc82d760877fef21</u>

Location: As mentioned above, due to the COVID-19 University Restrictions, <u>ALL</u> classes will be held online. Some classes are prerecorded (see 'online recording') and others will be held live via BlueJeans (see 'BlueJeans Session' highlighted in yellow).

Week	Date	Location	Торіс	Lecturer
	General Principles			
1	W, Jan 20	BlueJeans Session	Introduction /Orientation Live Session	Penning
	F, Jan 22	Online Recording	General Principles: Dose Response & Exposures	Penning
2	M, Jan 25	Online Recording	Regulatory Policy and EPA	Pepino
	W, Jan 27	Online Recording	Heavy Metal Toxicity	Osterhoudt
	F, Jan 29	Online Recording and Optional Live BlueJeans	Reactive Oxygen Species Recording (2p – 3p) w/Ischiropolous Optional Live Tutorial Session (3p – 4p) w/ Penning	Ischiropolous Penning
3	M, Feb 1	Online Recording	Metabolism Phase I	Penning
	W, Feb 3	Online Recording	Metabolism Phase II	Penning
	F, Feb 5	Online Recording and Optional Live BlueJeans	Chemical Carcinogenesis by Genotoxic Agents Recording (2p – 3p) w/Penning Optional Live Tutorial Session (3p – 4p) w/Penning	Penning
4	M, Feb 8 2:00 – 4:00	Online Recording & Mandatory Live BlueJeans Session	Chemical Carcinogenesis by Non- Genotoxic Recording (2p – 3p) w/Penning Live Problem Set Metabolism (3p – 4p) w/Penning	Penning
	W, Feb 10	BlueJeans Session	Mutagenesis / Mutational Signatures Live BlueJeans Session	Field
	F, Feb 12	Online Recording & Optional Live BlueJeans Session	DNA Adducts and their Repair Recording (2p- 3p) w/Penning Optional Live Tutorial Session (3p – 4p) w/Penning	Penning
5	M, Feb 15 2:00-4:00	BlueJeans Sessions	Mitochondrial Dysfunction Live Session (2p – 3p) w/Blair Live Problem Set Chemical Carcinogenesis (3p – 4p) w/Penning	Blair Penning

2021 Spring Schedule

Week	Date	Location	Торіс	Lecturer
			vironment Interactions	
	W, Feb 17	BlueJeans Session	Toxicogenetics – Toxicology and	Burczynski
	2:00 - 4:00		DNA Variation Live Session	
			Toxicogenomics- Toxicology and	
			RNA Expression Live Session	
	F, Feb 19	BlueJeans Sessions	Transcriptome-Analysis-Technologies	Tobias
			and Experimental Design Live Session	Penning
			(2p - 3p) w/Tobias	
			Optional Live Tutorial Session	
			(3p – 4p) w/Penning	
6	M, Feb 22	BlueJeans Session	Epigenetics Live Session	Heller
	W, Feb 24	Online Recording	Folate and Methylation	Whitehead
	F, Feb 26	BlueJeans Sessions	Risk Assessment Assignment	Penning
			(2p–3p) w/Penning	
			Optional Live Tutorial Session	
			(3p-4p) w/Penning	
7	M, Mar 1	Take Home Exam	Midterm (Due March 8 th)	
	XV M 2		Exposure Science	М
	W, Mar 3	Online Recording	Protein-Biomarkers-Proteomics	Mesaros
	F, Mar 5	Online Recording &	Exposure and Response Biomarkers	Mesaros
		Optional Live BlueJeans	Recording (2p-3p) w/Mesaros	Penning
		BlueJeans	Optional Live Tutorial Session $(2p, 4p)$ w/ Penning	
8	M, Mar 8	BlueJeans Session	(3p-4p) w/ Penning Biosensors Live Session	Johnson
0	W, Mar 10	Diucjeans Session	Spring Break	JOHNSON
		Orga	m-Based Toxicology	
			and Airway-Disease	
	F, Mar 12	Online Recording	Inhalation Toxicology 1: Respiratory	Panettieri
	(2p - 4p)	NO LIVE	Physiology	
		TUTORIAL THIS	Inhalation Toxicology 2: Mechanisms	
		WEEK	of Lung Injury	
9	M, Mar 15	Online Recording	Toxic Responses of the Respiratory	Christofido
			System	Solomidou
	W, Mar 17	BlueJeans Session	Lung Cancer Live Session	Vachani
	F, Mar 19	Online Decording 0-	Mesothelioma Recorded Session	Moon
	1', Maf 19	Online Recording & Optional Live	(2p-3p) w/Moon	Penning
		BlueJeans Session	Optional Live Tutorial Session	1 chining
		DideJeans Session	(3p-4p) w/Penning	
			Nervous System	
10	M Mc= 22	1	•	Dehingen
10	M, Mar 22	Online Recording	Overview of the Nervous System and Neurotoxicants	Robinson
	W, Mar 24	Online Pecording	Mechanisms of Neurotoxicity	Ischiropoulos
		Online Recording		Ischiropoulos
	F, Mar 26	BlueJeans Sessions	Sleep Disturbance and	Veasey
			Neurodegenerative Disease Live	Penning
			Session (2p-3p) w/Veasey	
			Optional Live Tutorial Session $(3p, 4p)$ w/Popping	
			(3p-4p) w/Penning	

Week	Date	Location	Торіс	Lecturer	
Reproductive & Endocrine Disruption					
11	M, Mar 29	Online Recording	In utero Genetic Imprinting	Bartolomei	
	W, Mar 31	Online Recording	Mechanisms of Reproductive	Gerton	
			Disruption-Male		
	F, Apr 2	BlueJeans Sessions	Environmental Reproductive	Burris	
			Epidemiology Live Session	Penning	
			(2p-3p) w/Burris		
			Optional Live Tutorial Session		
			(3p-4p) w/Penning		
	Data Integration & Predictive Toxicology				
12	M, Apr 5	Online Recording	Data-Integration-Bioinformatics	Weljie	
	W, Apr 7	BlueJeans Session	Exposure Biology Informatics Live	Moore	
			Session		
	F, Apr 9	BlueJeans Sessions	Predictive Toxicology and TOX 21st	Penning	
			Century Live Session		
			(2p-3p) w/Penning		
			Optional Live Tutorial Session		
			(3p-4p) w/ Penning		
13	M, Apr 12	BlueJeans Session	Risk Assessment Live Presentations –	Penning, Field,	
	(9a-11a)		<mark>9:00 am- 11:00 am</mark>	Gerton, Howarth,	
				Pepino	
	W, Apr 14	Take Home	Final Examination (Due April 21 st)		

Evaluation:

Mid-Term: 30% Final Exam: 40% Risk Assessment Paper: 30% Text: Cassarett & Doull's: Toxicology: The Basic Science of Poisons Questions on Individual Lectures Can be sent by Email to:

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