# Guidelines for preparation and evaluation of assignments for PHGY 550 course Molecular Physiology of Bone 2017

#### **Summaries:**

Each student will prepare a one page summary for each discussion session. The summary should answer the question "In your opinion, which set of experimental data is most critical to support the conclusion of the study". The following format should be adhered to:

Maximum length: 1 page, Minimum font size: 12 point

Margins of <sup>3</sup>/<sub>4</sub>" (2 cm) around entire document

Spacing: 1.5-spaced.

Header must contain student's name, ID number, and the date of the discussion session.

The summary should contain the following information:

- 1. Authors, Title, publication details for the scientific article
- 2. Objective of the study
- 3. Description of, in your estimation, the critical experiment or data set that supports the main study conclusion
- 4. Justification of the critical nature of this experiment or data set in supporting the conclusion of the study

Summaries are to be given to the instructor in the end of the corresponding discussion session – no extensions will be granted unless a doctor's note is provided. Summaries will be evaluated on a 0-100% scale using the criteria described below. In each category partial mark can be given, so that the summary that is better than satisfactory but does not reach excellence can be given a mark between 5 and 8, for example 6.5. If all the formatting instructions are respected, the final mark would be (6.5+2)/10 or 85%. The average of all summary marks will determine 30% of student's final grade.

Criteria	excellent	satisfactory	unacceptable
Content/analysis	Student demonstrates	Student demonstrates	Student does not
	clear understanding of	general understanding of	understand the
	the paper's objective,	the paper's objective,	objective of the paper,
	correctly describes the	describes the relevant	describes the
	experimental setup and	experiment, and provides	experiment incorrectly
	rationally justifies the	some justification for its	chooses the experiment
	critical role of the	importance for the study.	not critical for the
	experiment in supporting		study, or fails to justify
	the conclusions of the		it's importance
	study		
	80	50	0
Organization/	All formatting and	All formatting	The summary is less
formatting	organization	requirements are respected,	then ½ page or more
	requirements are	some organization	then 1 page, the font is
	respected	elements are not identified	too small or too big,
			Structural elements are
	20	10	missing 0

#### **Presentations:**

Each student will participate in the presentation of 2 papers during the discussion sessions. In general, one paper will be divided between 2-4 students. Each of the presenters is expected to carefully read the paper, understand the relevant background and methodology, and participate in critical discussion of any part of the paper. The presentations are expected

- 1. To provide a specific introduction that covers salient features relevant to the study,
- 2. To use simple, clear slides that highlight major points,
- 3. To provide brief descriptions of each method used for each of the shown figures. The description could be verbal for common methods, or more detailed for newer methodologies
- 4. To summarize the significance of each figure for reaching the conclusion of the paper,
- 5. To summarize and discuss the conclusions drawn from the study

In preparation for the critical discussion of the paper the presenting students should think about the limitations of the study, additional or different experiment that may improve the certainty of the conclusions, or alternative explanations for the results. Even if the presenter finds the question which he/she is unable to answer, it is important to bring it to the discussion.

Presentations will be evaluated on a 0-100% scale using the criteria described below. The average of all presentation marks will determine 25% of student's final grade.

Criteria	excellent	good	satisfactory	unacceptable
Content	Student understands	Student understands	Student shows	Student does
	paper objective,	paper objective,	general	not
	provides extended	provides sufficient	understanding of	understand
	background that	background for	paper objective,	the objective
	significantly	article	provides some	of the paper,
	improves article	understanding by	background for	and/or the
	understanding by	the group, provides	article	logic of the
	the group,	logical description	understanding by	study
	provides clear,	of the study	the group, provides	
	logical description		description of the	
	of the study		study	
	60	53	45	0
Critical	Student identifies	Student raises	Students	Students is
analysis	important	important	contributes to	unable to
-	discussion question	discussion question	discussion	contribute to
	or finds the answer	and contributes to	moderated by the	discussion
	for a question that	the discussion	instructor	moderated by
	required serious	aimed at finding the		the instructor
	creative thought	answer		
	30	20	10	0
Organization/	Presentation is	The presentation is	The presentation is	Slides are
Style	well-organized,	well-organized,	organized, slides	badly-
	slides are clear,	slides are clear,	are clear, some	organized,
	readable, logical	readable	problems are noted	unclear,
			but were considered	barely
			minor	readable,
	10	7	5	illogical <b>0</b>

#### **Essays:**

Students will write 3 essays on selected topics. The objective of these papers is to synthesize the knowledge gained from lectures, readings and discussion and to use it to formulate an original hypothesis and suggest an experimental approach to test it. Originality of the hypothesis and suitability of the methods to answer the proposed questions are the main criteria in evaluation of your essays. The following format should be adhered to:

Maximum length: 5 pages Minimum font size: 12 point

Margins of 3/4" (2 cm) around entire document

Spacing: 2-spaced.

The essay should contain the following information:

- 1. Student name, student ID number, evaluator name, date and title of the essay. Do not use a separate title page.
- 2. Each essay should contain five sub-sections clearly separated by the subheadings:
- 1) Introduction (review existing knowledge essential for formulation of hypothesis) generally <sup>3</sup>/<sub>4</sub> of a page.
  - 2) Hypothesis (clearly formulate in 1-2 sentences).
- 3) Suggested experimental approach (explain the strategy to test the hypothesis, anticipate potential pitfalls/problems and suggest different complementary experimental approaches, consider appropriate controls and discuss alternative outcomes), 2-3 pages.
- 4) Significance (describe anticipated results and consider the importance of knowledge gained from your study).
  - 5) References (no more then 10).

Essays are to be given to Ms. Rosetta Vasile, Department of Physiology, Room 1021, McIntyre Bldg, by 3.00 PM on the corresponding deadline date – no extensions will be granted unless a doctor's note is provided.

Essays will be evaluated on a 0-100% scale using the criteria described below. The average of all essay marks will determine 40% of final grade.

Criteria	Excellent	Good	Satisfactory	Unacceptable
Hypothesis	Original,	Original	Interesting	Hypothesis is
	unambiguous	hypothesis,	hypothesis, but	obscure, or the
	hypothesis	somewhat broad	has been tested	answer represents
			before in limited	general knowledge
			number of studies	
	15	11	7	0
Content	Logical	Logical	Suggested	Suggested
	experiments are	experiments are	experiments may	experiments will
	suggested,	suggested;	provide relevant	not address the
	alternative	however, some	information, some	question, controls
	outcomes are	controls are	controls are	are not included -
	considered/	missing, or some	missing,	as a result, the
	discussed and	alternative	alternative	suggested
	appropriate controls	outcomes are not	outcomes are not	methodology will
	are suggested	discussed	discussed	not provide

				evidence for the
				hypothesis
	60	50	40	0
Support	Introduction	Introduction	Introduction	Introduction
	demonstrates	demonstrates	demonstrates	demonstrates
	excellent	good knowledge	some knowledge	limited knowledge
	knowledge of the	of the field, the	of the field, the	of the field, the
	field, the student is	student is able to	student attempts	student is not able
	able to place his	place his study	to place his study	to place his study
	study within	within broader	within broader	within broader
	broader content,	content,	content,	content,
	referencing is	referencing is	referencing is	referencing is
	accurate	accurate	accurate	inaccurate
	15	12	8	0
Style	All structural	All structural	Structural	Main structural
	elements are	elements are	elements are	element is missing
	present; text within	present but not all	present but poorly	(such as
	the sections is	are identified;	identified, the text	Introduction), the
	logically divided	Text is logical,	is hard to follow.	text is hard to
	into paragraphs,	good flow of	All formatting	follow. The length
	Good flow of ideas.	ideas, occasional	requirements are	is less then 2.5 or
	All formatting	repetitions. All	respected	more then 5 pages.
	requirements are	formatting		Formatting
	respected	requirements are		requirements are
	_	respected		not respected

#### PHGY 550 2017

# Molecular Physiology of Bone (3 credit course) SCHEDULE - LECTURES AND DISCUSSIONS

Location: Room 1101 McIntyre Medical Bldg.

LECTURE 1	September 5	Introduction to bone cells, their differentiation and function
Dr. Monzur Murshed	Tuesday	
	8:35-9:25 AM	
	Review article:	Biology of bone and how it orchestrates the form and function of the
		skeleton.
		Sommerfeldt DW, Rubin CT. Eur Spine J. 2001 Oct;10 Suppl 2:S86-95.
		Transcriptional control of skeletogenesis.
		Karsenty G. Annu Rev Genomics Hum Genet. 2008;9:183-96.
DISCUSSION 1	September 11	Smpd3 Expression in both Chondrocytes and Osteoblasts Is Required
	Monday	for Normal Endochondral Bone Development.
	9:00-10:25 AM	Li J, Manickam G, Ray S, Oh CD, Yasuda H, Moffatt P, Murshed M.
		Mol Cell Biol. 2016 Aug 12;36(17):2282-99.

INFO SESSION	September11	Course essays – requirements, evaluation criteria etc.
	Monday	
	8:35-9:00 AM	

LECTURE 2 Dr. Kerstin	September 12 Tuesday	Osteoclast differentiation, function and signaling
Tiedemann	8:35-9:25 AM	
	Review article:	Osteoclast differentiation and activation.
		Boyle WJ, Simonet WS, Lacey DL.
		Nature. 2003 May 15;423(6937):337-42.
		A Comprehensive Review of Immunoreceptor Regulation of Osteoclasts.
		Humphrey MB, Nakamura MC.
		Clin Rev Allergy Immunol. 2016 Aug;51(1):48-58. doi: 10.1007/s12016-
		015-8521-8.
DISCUSSION 2	September 18	Differential effects of alendronate and losartan therapy on osteopenia
	Monday	and aortic aneurysm in mice with severe Marfan syndrome.
	8:35-10:25 AM	Nistala H, Lee-Arteaga S, Carta L, Cook JR, Smaldone S, Siciliano G,
		Rifkin AN, Dietz HC, Rifkin DB, Ramirez F. Hum Mol Genet. 2010 Dec
		15;19(24):4790-8

LECTURE 3 Dr. Bettina Willie	September 19 Tuesday 8:35-9:25 AM	Osteocytes and skeletal tissue mechanobiology
	Review article:	Bone Structural Adaptation and Wolff's Law; BETTINA WILLIE, GEORG
		N. DUDA AND RICHARD WEINKAMER (please see the attachment)
DISCUSSION 3	September 25	Targeted ablation of osteocytes induces osteoporosis with defective
	Monday	mechanotransduction.
	8:35-10:25 AM	Tatsumi S, Ishii K, Amizuka N, Li M, Kobayashi T, Kohno K, Ito M,
		Takeshita S, Ikeda K.
		Cell Metab. 2007 Jun;5(6):464-75.

LECTURE 4	September 26	Mathematical modeling as a tool to understand complex questions
Dr. Svetlana Komarova	Tuesday	in bone biology
	8:35-9:25 AM	
	Review article:	Mathematical modeling in bone biology: from intracellular signaling
		to tissue mechanics.
		Pivonka P, Komarova SV. Bone. 2010 Aug;47(2):181-9
DISCUSSION 4	October 02	A minimal mathematical model of calcium homeostasis.
	Monday	Raposo JF, Sobrinho LG, Ferreira HG. J Clin Endocrinol Metab. 2002
	8:35-10:25 AM	Sep;87(9):4330-40

## Tuesday, October 3, the first essay is due (Murshed, Tiedemann, Willie and Komarova)

Dr. Pierre Moffat	October 03 Tuesday 8:35-9:25 AM	New signaling molecules and their potential therapeutic use in bone disease
	Review article:	Regulatory pathways revealing new approaches to the development of anabolic drugs for osteoporosis.  Martin TJ, Sims NA, Ng KW. Osteoporos Int. 2008 Aug;19(8):1125-38.
DISCUSSION 5	October 16 Monday 8:35-10:25 AM	Phosphorylation of GSK-3beta by cGMP-dependent protein kinase II promotes hypertrophic differentiation of murine chondrocytes.  Kawasaki Y, Kugimiya F, Chikuda H, Kamekura S, Ikeda T, Kawamura N, Saito T, Shinoda Y, Higashikawa A, Yano F, Ogasawara T, Ogata N, Hoshi K, Hofmann F, Woodgett JR, Nakamura K, Chung UI, Kawaguchi H. J Clin Invest. 2008 Jul;118(7):2506-15.

LECTURE 6	October 10	General discussion and techniques to analyze bone.
Dr. Monzur Murhsed	Tuesday	
	8:35-9:25 AM	

LECTURE 7	October 17	Calcium –sensing receptor and disorders of calcium metabolism
Dr. Geoffrey Hendy	Tuesday	
	8:35-9:25 AM	
	Review article:	Diseases associated with the extracellular calcium-sensing receptor.
		Thakker RV. Cell Calcium. 2004 Mar;35(3):275-82.
DISCUSSION 7	October 23	Acquired hypocalciuric hypercalcemia due to autoantibodies against
	Monday	the calcium-sensing receptor.
	8:35-10:25 AM	Pallais JC, Kifor O, Chen YB, Slovik D, Brown EM. N Engl J Med. 2004
		Jul 22;351(4):362-9.

LECTURE 8	October 24	Calcium sensing receptor as a drug target
Dr. Geoffrey Hendy	Tuesday	
	8:35-9:25 AM	
	Review article:	Calcimimetic and calcilytic drugs: just for <b>parathyroid</b> cells?
		Nemeth EF. Cell Calcium. 2004 Mar;35(3):283-9.

DISCUSSION 8	October 30	Cinacalcet HCl reduces hypercalcemia in primary
	Monday	hyperparathyroidism across a wide spectrum of disease severity.
	8:35-10:25 AM	Peacock M, Bilezikian JP, Bolognese MA, Borofsky M, Scumpia S,
		Sterling LR, Cheng S, Shoback D.
		J Clin Endocrinol Metab. 2011 Jan;96(1):E9-18.

LECTURE 9	October 31	Osteogenesis Imperfecta
Dr. Frank Rauch	Tuesday	
	8:35-9:25 AM	
	Review article:	Osteogenesis imperfecta.
		Forlino A, Marini JC.
		Lancet. 2016 Apr 16;387(10028):1657-71.
DISCUSSION 9	November 6	Attenuated BMP1 function compromises osteogenesis, leading to
	Monday	bone fragility in humans and zebrafish.
	8:35-10:25 AM	Asharani PV, Keupp K, Semler O, Wang W, Li Y, Thiele H, Yigit G, Pohl
		E, Becker J, Frommolt P, Sonntag C, Altmüller J, Zimmermann K,
		Greenspan DS, Akarsu NA, Netzer C, Schönau E, Wirth R,
		Hammerschmidt M, Nürnberg P, Wollnik B, Carney TJ.
		Am J Hum Genet. 2012 Apr 6;90(4):661-74.

## Tuesday, November 7, the second essay is due (Hendy, Moffatt and Rauch)

LECTURE 10 Dr. Juliana Marulanda/ Dr. Monzur Murshed	November 7 Tuesday 8:35-9:25 AM Review article:	Diseases associated with abnormal skeletal tissue mineralization  Molecular determinants of extracellular matrix mineralization in bone and blood vessels.  Murshed M, McKee MD.  Curr Opin Nephrol Hypertens. 2010 Jul;19(4):359-65.
DISCUSSION 10	November 13 Monday 8:35-10:25 AM	Matrix Gla protein deficiency impairs nasal septum growth, causing midface hypoplasia.  Marulanda J, Eimar H, McKee MD, Berkvens M, Nelea V, Roman H, Borrás T, Tamimi F, Ferron M, Murshed M.  J Biol Chem. 2017 Jul 7;292(27):11400-11412.

LECTURE 11	November 14	Reciprocal regulation of bone and energy metabolism
Dr. Mathieu Ferron	Tuesday	
	8:35-9:25 AM	
	Review article:	Regulation of energy metabolism by
		the skeleton: osteocalcin and beyond.
		Ferron M, Lacombe J.
		Arch Biochem Biophys. 2014 Nov 1;561:137-46.
DISCUSSION 11	November 20	Glucose Uptake and Runx2 Synergize to Orchestrate Osteoblast
	Monday	Differentiation and Bone Formation.
	8:35-10:25 AM	Wei J, Shimazu J, Makinistoglu MP, Maurizi A, Kajimura D, Zong H,
		Takarada T, Iezaki T, Pessin JE, Hinoi E, <b>Karsenty</b> G.
		Cell. 2015 Jun 18;161(7):1576-91.

LECTURE 12	November 21	Bone pain
Dr. Laura Stone	Tuesday	
	8:35-9:25 AM	
	Review article:	The neurobiology of skeletal pain.
		Mantyh PW. Eur J Neurosci. 2014 Feb;39(3):508-19
DISCUSSION 13	November 27	Morphine treatment accelerates sarcoma-induced bone pain, bone
	Monday	loss, and spontaneous fracture in a murine model of bone cancer.
	8:35-10:25 AM	King T, Vardanyan A, Majuta L, Melemedjian O, Nagle R, Cress
		AE, Vanderah TW, Lai J, Porreca F. Pain. 2007 Nov;132(1-2):154-68.

LECTURE 13	November 28	Cancer and Bone
Dr. Peter Siegel	Tuesday	
	8:35-9:25 AM	
	Review article:	Cancer to bone: a fatal attraction.
		Weilbaecher KN, Guise TA, McCauley LK. Nat Rev Cancer. 2011
		Jun;11(6):411-25
DISCUSSION 12	December 4	The osteogenic niche promotes early-stage bone colonization of
	Monday	disseminated breast cancer cells.
	8:35-10:25 AM	Wang H, Yu C, Gao X, Welte T, Muscarella AM, Tian L, Zhao H, Zhao Z,
		Du S, Tao J, Lee B, Westbrook TF, Wong ST, Jin X, Rosen JM, Osborne
		CK, Zhang XH. Cancer Cell. 2015 Feb 9;27(2):193-210

Monday, December 4, the third essay is due (Siegel, Stone, Marulanda/Murshed and Ferron)