

**PHOL 410/NTRN410 - Basic Oxygen and Physiological Function**

Schedule: lecture videos will be posted on Mondays and Wednesdays

Course Director/Professor: Joseph LaManna, PhD, 368-1112; JCL4@case.edu

Co-Instructors: Kui Xu, MD/PhD, 368-5950, kxx@case.edu, and Michelle Puchowicz, PhD, map10@case.edu

Course TA: AIREZA ABDOLLAHIFAR <axa860@case.edu>

Class: 02/01/2021 - 05/19/2021

Lecture : online only

Office Hour (Zoom) for questions and discussion. Thursday 2-3pm

Suggested readings:

Pathway for Oxygen: Weibel E. R., *Harvard Univ Press* 1984; ISBN 0-674-65790-X

Physiology of Oxygen Radicals Aubrey E. Taylor et al.; *Waverly Press, Inc* 1986; ISBN 0-683-08104-7

Atom: L. Krauss, *Library of Congress* 2001 ISBN 0-316-49946-3

Oxygen: Nick Lane; *Oxford University Press* 2002; ISBN 0-19-850803-4

Symmorphosis: E. Weibel; *Harvard Univ Press* 2000; ISBN 0-674-00068-4

Bioenergetics: David G. Nicholls, Stuart J. Ferguson, 1999; *Academic Press* ISBN 0-12-518121-3

Scaling: K. Schmidt-Nielsen, *Cambridge Univ Press*, 1984; ISBN 0-521-31987-0

High Altitude and Man: J.B. West, S. Lahiri; *Amer Physiol Soc. The Williams & Wilkins Co.* 1984; ISBN 0-683-08945-5

**Grading: A: 85 and above; B: 70 -84; C: <70**

**Weekly quizzes: 15%, Block 1: 25%, Block 2: 30%, Block 3: 30%; Extra credit book report up to 10 points.**

<u>SESSION</u>	<u>DATE</u>	<u>TOPIC</u>
1	1-Feb	<b>Origin of Oxygen - JCL</b>
2	3-Feb	<b>Discovery of Oxygen - JCL</b> history / discovery; chemistry and physics of oxygen <i>"Oxygen: a play in 2 acts, by Carl Djerassi and Roald Hoffmann"</i>
3	8-Feb	<b>Life before Oxygen - JCL</b>
4	10-Feb	<b>Photosynthesis - JCL</b>
5	15-Feb	<b>Oxygen Toxicity, Hyperoxia and Hyperbaric Oxygen Therapy - JCL</b> chemistry of free radicals; mammalian defense mechanisms
6	17-Feb	<b>Measurement of Oxygen: Methods of Detection - JCL</b> O2 trodes, Optodes, mass spec, fluorescence/phosphorescence, etc <u>Selected readings</u> 1) <i>Imaging of Phosphorescence: A Novel Method for Measuring Oxygen Distribution in Perfused Tissue</i> ; Rumsey et al. <i>Science</i> 1988 2) <i>Oxygen in mammalian tissue: methods of measurement and affinities of various reactions</i> ; Vanderkooi, J.M. et al; <i>Amer J. Physiol</i> 1991 3) <i>The Redox State of Cytochrome Oxidase in Brain in Vivo: An historical perspective</i> ; Joseph C. LaManna 4) <i>Brain tissue oxygen concentration measurements</i> . Ndubuizu O, LaManna JC. <i>Antioxid Redox Signal</i> . 2007 Aug;9(8):1207-19. Review.
7	22-Feb	<b>Block Exam 1</b>
8	24-Feb	<b>Evolution of Lung - JCL</b>
9	1-Mar	<b>Structure and Function of the Respiratory System - KX</b> structure and function in the mammalian respiratory system compartmentation; distribution of lung volume; lung capacity; gas exchange; <u>Selected readings</u> : Weibel, <i>Pathway for Oxygen</i>
10	3-Mar	<b>Blood and Hemoglobin, Heart and Circulation - KX</b> Hb-O2 dissociation theory; characteristics of blood in various species; oxygen carriage by blood cardiac output, CaO2, etc
11	8-Mar	<b>Microcirculation - JCL</b> Historical: Krogh cylinder model (red vs white muscle; brain); Fick's Law, diffusion fields Capillary Distribution / Stewart Hamilton- mean transit time / oxygen delivery-capacity Intro / history- transporters; clinical applications <u>Suggested reading</u> L. Sokoloff, <i>Historical Review of Developments in the field of Cerebral Blood Flow and Metabolism in History of CBF and Metabolism</i>
12	10-Mar	<b>Blood Brain Barrier and Gas Channels, Transporters - JCL</b> <u>Selected readings</u> 1) Seider G et al, <i>GLUT-1 deficiency syndrome caused by haploinsufficiency of the blood-brain barrier hexose carrier</i> ; 1998; <i>Nature Genetics</i>

- 2) *De Vivo, D et al, Defective Glucose Transport Across the Blood-Brain Barrier; 1991; NEJM*  
 glucose, MCT; diffusion vs carrier mediated;  
 substrate uptake from blood to brain- Michaelis-Menten kinetics  
 hemodynamics; convection, diffusion; measuring O2 consumption; blood gas content
- 13 **15-Mar Energy Expenditure - JCL**  
 total body energy expenditure
- 14 **17-Mar Temperature, Hibernation- JCL**  
Selected Reading: *LaManna et al, Temperature Coefficients for the Oxidative Metabolic Responses to Electrical Stimulation in Cerebral Cortex, J Neurochem 1980*
- 15 **22-Mar Symmorphosis - JCL**  
Selected Reading: *Porter & Brand, Cellular oxygen consumption depends on body mass, 1995, AJP*
- 24-Mar No-class day**
- 16 **29-Mar Scaling - JCL**  
Selected Reading: *Porter & Brand, Cellular oxygen consumption depends on body mass, 1995, AJP*
- 17 **31-Mar Induction to Mitochondria - JCL**  
 Introduction/Origin of Mitochondria; Discovery of respiratory chain; chemiosmotic energy and bioenergetics (theory and biological applications); prevention of mitochondrial oxidative damage
- 18 **5-Apr Discovery of Cytochrome - JCL**
- 19 **7-Apr Mitochondrial Metabolism - MAP**  
Selected Readings: 1) *Green K et al, Prevention of Mitochondrial Oxidative Damage as a Therapeutic Strategy in Diabetes, 2004, Diabetes* 2) *Wallace D, A Mitochondrial Paradigm of Metabolic and Degenerative Diseases, Aging, and Cancer: A Dawn for Evolutionary Medicine, 2005; Annu Rev Genet.*  
 Oxidative phosphorylation and mitochondrial transport systems; maintenance of redox  
Selected Reading: *Sato, K. et al, Insulin, ketone bodies, and mitochondrial energy transduction, 1995; FASEB Journal.*
- 20 **12-Apr Reading day**
- 21 **14-Apr Block Exam 3**
- 22 **19-Apr Hypoxia: Adaptation/acclimatization to Altitude - JCL**  
Selected reading *J.C. LaManna et al, Review: Structural and functional adaptation*  
*Beall CM: Tibetan and Andean patterns of adaptation to high-altitude hypoxia*
- 23 **21-Apr Hypoxic Adaptation: Angioplasticity - JCL**
- 24 **26-Apr Anoxia - JCL**
- 25 **28-Apr Metabolic State of Ketosis, and its Protective Properties - MAP**
- 26 **3-May Brain Evolution - JCL**
- 27 **5-May Brain Functional Imaging - JCL**  
 measuring cerebral energy metabolism: fMRI, BOLD, NMR spectroscopy  
 1) *Fox PT, et al, Nonoxidative glucose consumption during focal physiologic neural activity, 1988; Science.*  
 2) *Raichle, M, Behind the scenes of functional brain imaging: A historical and physiological perspective; 1998; Proc Natl. Acad Sci*  
 3) *Magnetic Resonance Imaging: From Atomic Physics to Visualization, Understanding and Treatment of Brain Disorders, Breakthroughs in Bioscience www.faseb.org*
- 28 **10-May Ischemia: Focal vs Global, Experimental Studies - KX**  
 Blood flow, autoregulation, edema  
 Circle of Willis; cardiac arrest and resuscitation; edema  
Selected reading *J.C. LaManna and W.D. Lust, Intrinsic and Extrinsic Optical Probes of Cerebrovascular and Metabolic function in Cerebrovascular Disease: Pathology, Diagnosis, and Management, 1998; Ginsberg, ed; Blackwell Science*
- 29 **12-May Reading Day**
- 30 **17-May Block Exam 3**
- 31 **20-May Extra credit paper due 5pm EST**