

BISP 194: Advanced Topics in Modern Biology
Tu 3:30-5:00pm // SCRM 1013A/B

1) Course title: "Molecular genetic basis of neural function and neurological disease"

2) One of the greatest challenges remaining in biomedical research is to understand how the brain works. Given the complexity of the central nervous system, one useful approach to deconstructing the operation of the human brain and nervous system has been to determine the basis of neurological diseases, ranging from developmental disorders such as autism and mental retardation, to neurodegenerative disorders such as Parkinson's disease and Alzheimer's disease, to neuropsychiatric disorders such as schizophrenia and depression. In this course, we will examine the recent advances in our understanding of these disorders - progress that has been driven by molecular genetic and genomic technologies in most cases. The goal of this course is that you will not only learn about these exciting discoveries and their implications for treating these diseases, but also you will learn to read, critically evaluate, and present primary data from published recent articles in the scientific literature.



PREREQUISITES: BICD100 (Genetics), BIBC100[02] (structural/metabolic biochemistry), an introductory course in Neuroscience, and their prerequisites. Additionally, it is **HIGHLY RECOMMENDED** that you have completed BIMM100 (Molecular Biology) before taking this class.

3) Enrollment is limited to 30 students.

4) Tuesday afternoons, 3:30 to 5:00 PM, ending 80 minutes after the start time.
Dates (10): 1-8; 1-15; 1-22; 1-29; 2-5; 2-12; 2-19; 2-26; 3-5; 3-12

5) No textbook is required. However, Thompson & Thompson, Genetics in Medicine, 7th edition, Saunders-Elsevier, ISBN # 9781416030805, is a valuable reference book, chapters 1-4 and 9-14.

6) Instructors:

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7) Course coordinator

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8) Requirements

Attendance is mandatory. Original papers will be selected and disseminated to students by Ms. Novak about one week prior to each class. Students will be expected to discuss

the distributed papers during class, and ask questions as to the scientific / biological significance of the work and how the study was performed. **Students are required to write and submit via email to nanovak@ucsd.edu a total of 3 SUBSTANTIVE questions relating to the research papers assigned. The 3 questions must be received by 9 AM on the day of the class – late submissions will receive no credit: NO EXCEPTIONS!** At least one question per original paper must be provided (so no questions should be submitted for review articles).

Examples of exemplary questions are given here:

1. The authors reference mGluR1 in intro but only suppress mGluR5 50%. Is this typical? How can suppression of mGluR5 be maintained when challenged? I would have liked to see proof of the 50% specificity of mGluR5 in the body of the paper in comparison to other related proteins.
2. Does the degree of severity of the polyglutamine repeats impact the expression of the different protein complexes? While 32Q is considered WT and 82Q is considered expanded, this is not all-or-none in terms of polyglutamine expansion, so does some (linear) gradient exist as well for protein complex expression?
3. In my opinion the authors have omitted an interesting and straightforward control: Take an ESC from embryo with fragile X mutation that expresses the *FMRI* gene and differentiate the cell to a somatic phenotype that has undergone transcriptional silencing for that gene. Then induce pluripotency with iPSC factors and see if the reprogramming fails in the same cell line. If the cell does not fully reprogram, there is more support for ESC as a fragile X developmental model, otherwise, light may be shed on the mechanism of differential fragile X syndrome manifestation between ESC and iPSC.

9) Grading policy

Students are expected to attend all classes. Grades will be based upon the quality of the written questions that students turn in each week; attendance; and the quality of student participation at each meeting of the class.

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SCRM 1013

DATE	NAME	MODERATOR	TOPIC
1/8/2013	La Spada	ALS	Intro / neurodegeneration - repeat dz
1/15/2013	Muotri	Muotri	Stem cell modeling of neurodevelopment disease
1/22/2013	Cleveland	ALS	From Charcot to Lou Gehrig: understanding ALS
1/29/2013	Kelsoe	Muotri	The genetics of bipolar disorder
2/5/2013	Yeo	ALS	RNA binding proteins in neurodegeneration
2/12/2013	Masliah	Muotri	Parkinson's disease pathogenesis
2/19/2013	Goldstein	Muotri	Cellular and molecular basis of Alzheimer's disease
2/26/2013	Koo	ALS	Alzheimer's disease mechanisms and pathways
3/5/2013	Sebat	Muotri	Neuropsychiatric disease genetics
3/12/2013	Mobley	ALS	Alzheimer's disease and Down syndrome