

Course timetable

Week	Date	Topic
1	5 Sep 2023	Course introduction - getting to know each other, what to expect, dos and donâ€™ts o What makes GABAergic signaling such an important drug target in the clinic?
2	12 Sep 2023	GABA_A receptors as ligand-gated ion channels - key targets for anxiolytics, sedatives, hypnotics and general anesthetics
3	19 Sep 2023	GABA_B receptors as Class C G-protein coupled receptors - targets for treatment of spasticity, addiction (nicotine, alcohol) and substance abuse
4	26 Sep 2023	Techniques used to study the synapse â€™ brief intro to electrophysiology, florescence imaging (wide-field, confocal, multiphoton), molecular biology, systems neuroscience
5	3 Oct 2023	Molecular pharmacology of GABA receptors I â€™ anxiolytics and antidepressants
6	10 Oct 2023	Constructing inhibitory synapses I â€™ what traps receptors at postsynaptic compartments? <i>Mid-term test; Paper for news and views to be given out</i>
7	17 Oct 2023	Constructing inhibitory synapses II â€™ how are synapses assembled?
8	24 Oct 2023	Anxiety I â€™ synaptic GABA receptor levels determine extent of anxiety
9	31 Oct 2023	Anxiety II â€™ brain circuit-specific coupling of GABA receptor signaling increases anxiety
10	7 Nov 2023	Phasic and tonic inhibition I â€™ phasic and tonic inhibition mediated by distinct receptor isoforms <i>News and views due; Paper for significance statement to be given out</i>
11	14 Nov 2023	Phasic and tonic inhibition II â€™ tonic inhibition and sensory processing
12	21 Nov 2023	Epilepsy I â€™ genetic epilepsy due to GABA receptor variants (mutations)
13	28 Nov 2023	Epilepsy II â€™ role of tonic inhibition in absence epilepsy <i>Significance statement due; Essay topics to be given out</i>

14	23 Jan 2024	<u>Interneurons</u> – the presynaptic viewpoint
15	30 Jan 2024	<u>Development of interneurons</u> – how do various types of interneurons find their targets?
16	6 Feb 2024	<u>Interneuron dysfunction in psychiatric disorders</u> – schizophrenia and bipolar disorder <i>Essay due; Figures for <u>legend writing assessment</u> to be given out</i>
17	13 Feb 2024	<u>– GABA in development I</u> – there is more to GABA than inhibition
18	20 Feb 2024	<u>GABA in development II</u> – critical periods during development are controlled by GABAergic signaling <i>Figure legends due; Paper for <u>graphical abstract</u> given out</i>
19	27 Feb 2024	<u>Plasticity of the inhibitory synapse I</u> – regulation of GABAergic inhibition by excitatory glutamatergic signaling
20	5 Mar 2024	<u>Plasticity of the inhibitory synapse II</u> – circuit specific inhibitory plasticity during critical periods of brain development <i>Mid-term test</i>
21	19 Mar 2024	<u>Molecular pharmacology of GABA receptors II</u> – anesthetics, neurosteroids and other ligands
22	26 Mar 2024	<u>Ischemia</u> – neuroprotective role of GABA receptors following ischemia <i>Graphical abstract due; Paper for <u>peer review</u> to be given out</i>
23	2 Apr 2024	<u>Down syndrome</u> – cognition, over-inhibition and a failed drug trial
24	9 Apr 2024	<u>Rett syndrome</u> – Rett phenotypes can arise due to loss of MECP2 in interneurons
25	16 Apr 2024	<u>Neurodegeneration and dementia</u> – GABA _B -APP interactions as therapeutic targets in Alzheimer’s disease
26	23 Apr 2024	<u>Summary of the inhibitory synapse</u> – general discussion <i>Peer review due</i>

– available for student presentation

Venue Robinson 107

Instructor Saad Hannan, Ph.D.

Research Associate

Hensch Lab

Department of Molecular and Cellular Biology

saadhannan@fas.harvard.edu

Office hours Tuesdays 5:00 – 6:00 pm or by appointment

Prerequisites

MCB/Neuro 80 (Neurobiology of Behavior) and permission of the instructor

Course goals

This overall objectives of this course are:

1. To help students to gain an in-depth understanding of the function of synapses at cellular, circuit and behavioral levels as well as the dysfunction that causes brain disorders and treatment strategies;
2. To learn key techniques used in neurosciences (e.g. molecular biology, electrophysiology, and cellular imaging) and help students critically analyze experimental results in tutor-guided small group sessions;
3. To practice reading scientific literature and communicating scientific results through presentations and brief digests.

Required reading

Reading materials will include scientific literature in the form of original research papers and reviews. This course does not require textbooks. Required readings for each week will be posted on course website at least two to three weeks in advance of tutorial sessions.

Typical tutorial format

Each tutorial will begin with a brief lecture component that will introduce the topic and paper of discussion along with addressing any unresolved issues from the previous week's tutorial. Next, the session will move onto discussing the contents of paper chosen for the week by one or more students with help from the tutor. The introductory part should take around ~15-30 min while the paper discussion should take roughly 45 min. The final few minutes will be used by the tutor to wrap up discussion and to briefly summarize how the chosen paper has extended our broader understanding of the area and what subsequent studies/ drug trials have revealed.

Typical workload

Typical workload will be approximately 3 to 5 hours a week. This will be spent on various assessments (see below) which will provide tutees multiple opportunities to get high grades and readings that will be assigned every week. While students are not expected to come up with a specific number of questions, the best tutorials are typified by high levels of student engagement and participation. Therefore, the expectation at the outset is that the students will ask questions to the tutor and each other. During the course, each tutee will pick at least one study and present it to the group. Preparation for this may require an additional 1-2 hours/ week. Pre-presentation help and feedback will all be made available for all tutees.

Assessment

This course will provide tutees with multiple opportunities to improve their grades. The types of assignments chosen are deliberately diverse to equip tutees with a wide range of skills that are necessary

not only in science but also related sectors in data analysis and presentation. Rubrics for grading for each assignment and feedback on progress will be provided at regular intervals. Feedback will also be provided on the essay assignment provided the tutee sends the essay no later than three weeks prior to the deadline.

The assessment breakdown for the course is as follows (pie chart shows summary):

Attendance – 5% of total

Assessment 1 – Mid-term quiz x 2 (10% of total; 10 Oct 2023 and 5 March 2024)

Assessment 2 – News and views (10% of total; due 7 Nov 2023)

Assessment 3 – Significance statement (10% of total; due 28 Nov 2023)

Assessment 4 – Paper presentation (15% of total; date to be chosen by tutees)

Assessment 5 – Essay (15% of total; due 30 Jan 2024)

Assessment 6 – Writing figure legends (10% of total; due 20 Feb 2024)

Assessment 7 – Graphical abstract (10% of total; due 26 Mar 2024)

Assessment 8 – Peer review (15% of total; due 23 Apr 2024)

Significance statement – tutees will write a 120-word significance statement on a seminal paper which was multiple firsts in neuroscience, molecular and cellular biology and pharmacology. So plenty to write about it.

News and views – here the tutees will write a 750 – 900 words summary of a 2015 study on an inhibitory scaffold that is important for learning and memory. Detailed guidelines will be provided to students on how to approach this piece of coursework along with where to find numerous examples - on nature's website!

Guidance on how to write significance statements can also be found at:
<https://neuronline.sfn.org/professional-development/how-to-write-an-accurate-concise-and-meaningful-significance-statement>

Course essay – three essay topics will be made available to tutees at the end of November with a deadline for submission at the end of January. Tutees will write a 3000-word essay on their topic of choice. This deadline has been chosen to give tutees ample time to work on the assignment over the winter months and to coincide with the winter recess. The essay should be a piece of original work on the core theme of inhibitory neurotransmission. Tutees are expected to read original research papers and reviews similar to those already covered in the course by that point and come up with their own assessment of their chosen topic. Feedback will be provided on the essay assignment provided the tutee sends the essay no later than three weeks prior to the deadline.

Figure legend writing – this exercise is aimed at giving students hands-on training on how to write a figure legend. Three unpublished figures from one of my scientific studies will be provided and tutees will write the figure legends, the most likely statistical test used by analyzing the bar/ box plots and include n numbers where possible. The figures will be multi-panel and will be designed in a way so that the tutees appreciate the value of keeping the text succinct. This coursework is due on the week of tutorial 18 and by then the students should have a good foundation to take on such an exercise.

Additional details on the format (from Nature): – Each figure legend should begin with a brief title for the whole figure and continue with a short description of each panel and the symbols used. legends should not contain any details of methods. Legends should be fewer than 300 words each. All error bars and statistics must be defined in the figure legend.

Graphical abstract – graphical depictions of scientific studies are rapidly gaining in popularity in various social media. Graphical abstracts are therefore powerful visual tools for reaching out to diverse audiences. This exercise is aimed at giving tutees the opportunity to gain hands-on experience of

designing a graphical abstract while nurturing their artistic talents and receiving feedback from the tutor and the group.

The abstract will be prepared according to on the Cell Press format as follows:
https://www.cell.com/pb/assets/raw/shared/figureguidelines/GA_guide.pdf

The graphical abstract is due on the week of tutorial 22 and by then the tutees will have gained multiple exposure to papers with such abstracts as exemplars.

Seminar paper presentation - These will be informal presentations where the tutee will present findings, take questions from peers and guide/ moderate class discussion. Tutees will be marked on multiple criteria including their ability to critically analyze findings, style of presenting, approach to taking questions from peers etc. Help and guidance will be provided in preparation stages should there be a need.

Peer review - peer review of scientific papers and grants is an integral part of the scientific journey and this exercise will give tutees an opportunity to become an expert reviewer for a journal. An unpublished paper from the pre-print server bioRxiv containing materials covered on the course will be chosen. Tutees will be provided with guidelines on writing peer reviews and pointed to details of journals who publish peer review reports which can be used as examples. The final peer-review report does not have a word limit and will be excellent for honing critical thinking skills while analysing a real-world unpublished scientific paper.

Grading

Attendance, completion of writing and oral assignments, and thoughtful participation in class discussions will lead to an 'A' grade. Letter grades will decrease due failure to complete assignments and low attendance.

Course Policies

Confidentiality - while most of the materials in the course are already in public domain, occasionally tutees may be given real-world unpublished results eg - figure legend writing assignment etc. Moreover, the peer-review assignment will be on an unpublished manuscript deposited in a pre-print server. I encourage all tutees to discuss their ideas and opinions with their peers and expect them to engage in these academic exercises with full vigour. However, due to the unpublished nature of some results, tutees are asked to refrain from disseminating contents of these assignments in the public domain. If you have any questions regarding this, then please feel free to get in touch.

Classroom environment - Tutorials provide an excellent opportunity for students to iron out creases in their conceptual understanding of topics. Moreover, small group learning is ideal for student participation. Therefore, it is my expectation that the tutees will come to class prepared and actively participate in class discussions with respectful dialogue with classmates. Our classroom will be an inclusive environment that values diverse ideas which exists to support all students as they learn and grow throughout the year.

Attendance/Participation Policy - Students are expected to attend all classes and be an active participant in class. Three unexcused absences over the course of the school year will result in the drop of an entire letter grade. If you know that you are unable to attend class on a specific day, please contact the instructor as early as possible.

Feedback - Students will be continually provided feedback when sought. In addition, tutees will be provided with some opportunities to provide feedback at the end of tutorials to write down their views on content, format, or any other matter. These will be collected, reviewed and factored in to tailor the course to suit more to the needs of our pupils.

Late Work - Students are expected to turn in all work on time. Weekly assignments that are not turned in by 5 pm. on the day they are due will be assessed a 10% penalty for each day that it is late. If you have extenuating circumstances during the course, please speak with the instructor in advance of the assignment due date.

Academic integrity - Written assignments that are submitted for evaluation are expected to be your own, independent work. Students are required to correctly cite literature (only books and peer reviewed journals allowed). Training will be provided during the course on how to cite correctly in scientific writing.

Students must also acknowledge any assistance they received with their writing (e.g. feedback on drafts, etc.).

Course Schedule - The policies, schedule, or assignments described within this syllabus are subject to change during the course at the discretion of the instructor. Students will be notified of any changes to the schedule or readings/assignments at least one week ahead of time.

Use of artificial intelligence (AI) - We expect that all work students submit for this course will be their own. Students are discouraged from using ChatGPT or any other generative AI tools at all stages of the work process, including preliminary ones. We draw your attention to the fact that different classes at Harvard could implement different AI policies, and it is the student's responsibility to conform to expectations for each course.