Title: Application of molecular biology and molecular brain imaging techniques in neurodegenerative disorders

Course number: 2315

Credits: 3.0

Date: 20160509 - 20160520

Language: English Level: Doctoral level

Responsible KI department: Department of Neurobiology, Care Sciences and Society

Specific entry requirements: No

**Learning outcomes:** After the course the students will understand the main pathophysiological features of neurodegenerative diseases, with a focus on dementia disorders. After the course a doctoral student shall be able to:

Account for recent advances in CSF and in vivo brain imaging biomarkers

- Account for the theoretical basis of main techniques used in molecular imaging and biochemical analyses used on living subjects and biological materials
- Develop, evaluate and validate in vivo brain and CSF markers
- Perform key in vitro and in vivo techniques
- Collect, process, analyze and interpret data of key techniques
- Apply what they have learned to their own research work

Contents of the course: This course will focus on scientific approaches used to study neurodegenerative disorders. Topics for the practical experiments will include the following: demonstrations of molecular imaging techniques that are used to study function and pathological changes in the brain of living patients/animal models including PET/microPET, demonstrations of MRI techniques for structural brain analysis, histopathological examination of brain lesions in human postmortem brain tissue and autoradiography and radioligand binding techniques to study localization of receptors, binding potential, changes in neurotransmitter function and to correlate these with other pathological changes in the brain, quantitative assays of biomarkers and proteins involved in signaling pathways in biological fluids and in the brain using brain homogenate, human CSF and/or plasma etc. The course will also discuss, provide a deep understanding, some hands-on training on how to setup and use common in-house sensitive assays, their advantages, applicability and caveats.

**Teaching and learning activities:** The course is full time (10 days) and will be organised as an integration of lectures, combined with practical exercises, demonstrations and literature studies.

**Examination:** All the intended learning outcomes will be assessed by a combination of written examination, written reports for the practical workshops, and oral presentation.

**Compulsory elements:** All parts of the course are mandatory. Absence from any of these will be compensated for by extra individual assignments provided by the course organizers.

Number of students: 10 - 20

**Selection of students:** Selection will be based on 1) the relevance of the course syllabus for the applicant's doctoral project (according to written motivation), 2) date for registration as a doctoral student (priority given to earlier registration date)

## More information:

<u>Course responsible</u>: Taher Darreh-Shori Institutionen för neurobiologi, vårdvetenskap och samhälle Taher.Darreh-Shori@ki.se

## Contact person:

Elena Rodriguez-Vieitez Institutionen för neurobiologi, vårdvetenskap och samhälle elena.rodriguez-vieitez@ki.se

## Application of molecular biology and molecular imaging techniques in neurodegenerative disorders (course 2315) May 9th to May 20th, 2016: WEEK 1

Monday (9/5)	Tuesday (10/5)	Wednesday (11/5)	Thursday (12/5)	Friday (13/5)
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9:00 - 10:00 Lecture	9:00 - 10:00 Lecture	9:00 - 10:00 Lecture	9:00 - 10:00 Lecture	9:00 - 10:00 Lecture
Introduction to neurodegenerative	Magnetic resonance imaging (MRI) physics	Introduction to PET imaging and radiotracer		PET imaging: focus on microPET
disorders: focus on AD and other dementia	Iwagnetic resonance imaging (wiki) physics	development	imaging of PET data	FET Imaging, locus of microfet
disorders. Todas of AD and other dementia		development	innaging of 1 L1 data	
Agneta Nordberg, KI	Farshad Falahati, KI	Bengt Långström, Uppsala U.	Zsolt Cselenyi, KI	Miklós Tóth, KI
<u> </u>	, and the second		* '	,
10:00 - 11:00	10:00 - 11:00	10:00 - 11:00	10:00 - 11:00	10:00 - 10:45
Lecture Neuropathology of neurodegenerative	Lecture Visual rating scales: clinical diagnosis and	Lecture Molecular PET imaging in Alzheimer's	Lecture PET imaging analysis: Practical procedures	Lecture/Discussion
. 0,	research	disease and related dementia disorders in a		animal models
diseases	research	clinical perspective		animai modeis
		·		
Irina Alafuzoff, Uppsala U.	Daniel Ferreira, KI	Agneta Nordberg, KI	Laure Saint-Aubert, KI	Laetitia Lemoine, KI
11:00 - 12:00	11:00 - 12:00	11:00 - 12:00	11:00 - 12:00	11:00 - 12:00
Lecture	Lecture	Lecture	Cáudu timo	Lecture
Neuropathology of neurodegenerative diseases (cont'd)	Voxel-based morphometry (VBM): Theory and applications	Principles of PET imaging methodology and semi-quantitative approaches	Study time	MicroPET image analysis in AD mouse model: Introdution to practical workshop
Irina Alafuzoff, Uppsala U.	Joana Pereira, KI	Akihiro Takano, KI		Elena Rodriguez-Vieitez, KI
* **	12:00 - 13:00	12:00 - 13:00	12:00 - 13:00	•
12:00 - 13:00 LUNCH	LUNCH	LUNCH	LUNCH	12:00 - 13:00 LUNCH
13:00 - 14:00	13:00 - 14:00	13:00 - 14:00	13:00 - 14:00	13:00 - 14:00
Lecture	Lecture	10.00	Lecture	Lecture
Functional and pathological correlates of	Introduction to MRI practical workshops:	Study time	Introduction to PET practical workshop	CSF and PET imaging biomarkers in
subjective memory complaints in Aging and	' '	·	' '	preclinical and prodromal Alzheimer's
Alzheimer's disease				disease
Patrizia Vannini,	MRI sequences, Farshad Falahati		Laure Saint-Aubert / Antoine Leuzy /	Oskar Hansson, Lund U.
Brigham and Women's Hospital, Boston,	Visual rating scales for brain atrophy and		Konstantinos Chiotis / Elena Rodriguez-	
USA	white matter lesions, Daniel Ferreira		Vieitez, KI	
	VBM analysis, <i>Joana Pereira</i>			
14:00 - 14:15 COFFEE				
14:15 - 14:45	14:00 - 17:00	14:00 - 16:00	14:00 - 17:00	14:00 - 17:00
Introduction to task assignment	Imaging workshop:		Imaging workshop:	Imaging workshop:
	MRI	14:40 Comparing CSF and amyloid PET	PET	MicroPET
	Computer lab, Huddinge library, Room	biomarkers in AD	Computer lab, Huddinge library, Room	Computer lab, Huddinge library, Room
	S311	Antoine Leuzy, KI	S311	S311
14:45 - 15:30	Joana Pereira / Daniel Ferreira / Farshad	<b>14:40 - 15:20</b> Tau PET imaging in	Laure Saint-Aubert / Antoine Leuzy /	Laetitia Lemoine / Elena Rodriguez-Vieitez,
Lecture	Falahati, KI	prodromal and dementia stages of AD	Konstantinos Chiotis / Elena Rodriguez-	KI
		Konstantinos Chiotis, KI	Vieitez	
Clinical assessment of dementia,	Div. Clinical Geriatrics, KI	<b>15:20 - 16:00</b> Using the early frames of	Div. Translational Alzheimer Neurobiology,	Div. Translational Alzheimer Neurobiology,
Vesna Jelic, KI		PET tracers as a proxy for brain perfusion	KI	KI
		Elena Rodriguez-Vieitez, KI		
		<b>16:00 - 17:00</b> Visit to PET scanners,		
		Huddinge Hospital,		
		Irina Savitcheva		

## Application of molecular biology and molecular imaging techniques in neurodegenerative disorders (course 2315) May 9th to May 20th, 2016: WEEK 2

Monday (16/5)	Tuesday (17/5)	Wednesday (18/5)	Thursday (19/5)	Friday (20/5)
9:00 - 11:00 Lecture:	9:00 - 10:00 Lecture and Overview of Lab-	9:00 - 12:00 Lab:	9:00 - 10:30 Overview of Lab-workshop: Lab:	9:00 - 12:00 Own study time
In vitro molecular imaging techniques and their application in studies of neurodegenerative diseases	Tracer development and overview of in vitro screening techniques (EA, SPA, ITC)	Enzymological analyses and IC50 and/or Ki calculation of the Lead	Laboratory on in vitro imaging studies in postmortem brain	Preparation of oral exam and written lab report
Per-Göran Gillberg, KI	Taher Darreh-Shori, KI		Laetitia Lemoine, KI Lakshman Puli, KI	
11:00 - 12:00 Lecture:	10:00 - 12:00 Lab:		10:30 - 12:00 Demonstration:	
Fundamentals on astrogliopathology: from astrodegeneration to astroglial reactivity	Hits screening by enzymological analyses		Autoradiography technique	
Prof. Alexei Verkhratsky, Faculty of Life Sciences, University of Manchester, UK	Taher Darreh-Shori, KI Rajnish Kumar, KI Erica Lana, KI	Taher Darreh-Shori, KI Rajnish Kumar, KI Erica Lana, KI	Laetitia Lemoine, KI Lakshman Puli, KI	
12:00 - 13:00 LUNCH	12:00 - 13:00 LUNCH	12:00 - 13:00 LUNCH	12:00 - 13:00 LUNCH	12:00 - 13:00 LUNCH
13:00 - 14:00	13:00 - 14:00	13:00 - 14:00	13:00 - 15:00	13:00 - 16:00
Lecture:	Lab:	Lab:		
Tracer development	Hits screening by enzymological	IC50 and/or Ki calculation of the Lead	Tutorial on lamination	EXAM
Computer-aided hits identification	analyses			
Rajnish Kumar, KI			Laetitia Lemoine, KI Lakshman Puli, KI	
14:00 - 14:15 COFFEE	14:00 - 14:15 COFFEE	14:00 - 14:15 COFFEE	14:00 - 14:15 COFFEE	
14:15 - 15:00	14:15 - 15:30	14:15 - 17:00	15:00 - 17:00	
Lecture:	Lab:	Lab:/Data analyses:	Demonstration:	
Impact of epigenetic variation in the			Immunohistochemistry technique	
study of neurodegenerative disorders				
Erica Lana, KI				CAKE AND COFFEE
15:00 - 16:45	15:30 - 16:45			
Lecture:				
Radiation safety at the lab	Preparation of written lab report / Individual consultations			
Annie Olsson, KI				
16:45 - 17:00				
Diamoria	Taher Darreh-Shori, KI Rajnish Kumar, KI	Taher Darreh-Shori, KI Rajnish Kumar, KI	Laetitia Lemoine, KI	
Discussion	Erica Lana, KI	Erica Lana, KI	Laetitia Lemoine, Ki Lakshman Puli, Kl	
JISCUSSION		The state of the s	The state of the s	

Thurs	sday (19/5)			
	oup 1 & 2			
9:0	0 - 10:00			
Overview of workshop: studies in p	Laboratory on in vitro binding postmortem brain			
Laetit	ia Lemoine			
10:	00-10:30			
introduction to autoradiog	raphy/immunostaining workflow			
	30-12:00			
Tutorial on lamination anaysis				
Laetitia Lemoine	Lakshman Puli			
12:0	00 - 13:00			
13:00 - 15:00	13:00-15:00			
demonstration of immunostaining technique Groupe 1	demonstration of autoradiography technique and binding assay Groupe 2			
Lakshman Puli	Laetitia Lemoine			
14:00 - 14:15 <i>COFFEE</i>				
15:00-17:00	15:00-17:00			
demonstration of immunostaining technique Groupe 2	demonstration of autoradiography technique and binding assay Groupe 1			
Lakshman Puli	Laetitia Lemoine			
17:	00-17:10			
Wrap ι	up of the day			
Laetitia Lemoine	Lakshman Puli			