

# **Neurofunctional Techniques**

**Lesson 14**

13, 18 November 2024

**Paper presentations**

# Calendar

- **M 30 Sept:** Course introduction
- **W 2 Oct:** Functional imaging
- **F 4 Oct:** Statistics (Cesca)
- **M 7 Oct:** Functional imaging
- **W 9 Oct:** Biophysics of diffusion
- **F 11 Oct:** Statistics (Cesca)
- **M 14 Oct:** Functional imaging
- **W 16 Oct:** General introduction to the papers for the presentations
- **F 18 Oct:** Statistics (Cesca)
- **M 21 Oct:** Modeling in neuroscience
- **W 23 Oct:** Molecular approaches in neuroscience
- **F 25 Oct:** Statistics (Cesca)
- **F 25 Oct:** Laboratory (14:00- 18:00)
- **M 28 Oct:** Practical exercises on the first part of the course
- **W 30 Oct:** Genome editing in neuroscience (Dr. Jaudon)
- **M 4 Nov:** Optogenetics
- **W 6 Nov:** Papers assignment to the groups; introductions to the specific papers
- **T 12 Nov:** X-genetics + Practical exercises on the second part of the course
- **W 13 Nov:** Introductions to the specific papers
- **M 18 Nov:** Introductions to the specific papers
  
- **9, 10, 11 Dic:** Paper presentation 15:00-19:00)
- **Tue 17 Dic:** Test (14:00 - 16:00 Room 3A, Building H2bis)

# Structure of the exam

1. Paper presentation in small groups (5 students 1 hour) **Maximum score 20 (+1):**

a) Presentation (30 min)

- What was known
- What is the gap
- What are the main findings
- Are the techniques appropriate
- What are the broader implications

b) Questioning on individual figures and techniques used (30 min)  
**(9, 10, 11 December)**

2. Questionnaire on the Moodle platform. 10 multiple choice questions **(also on statistics)**. Each question has only one correct answer, and each correct answer is awarded 1 point. **Maximum score: 10.**

**(17 December Room 3A Building H2bis)**

**First turn: 14:00 - 15:00**

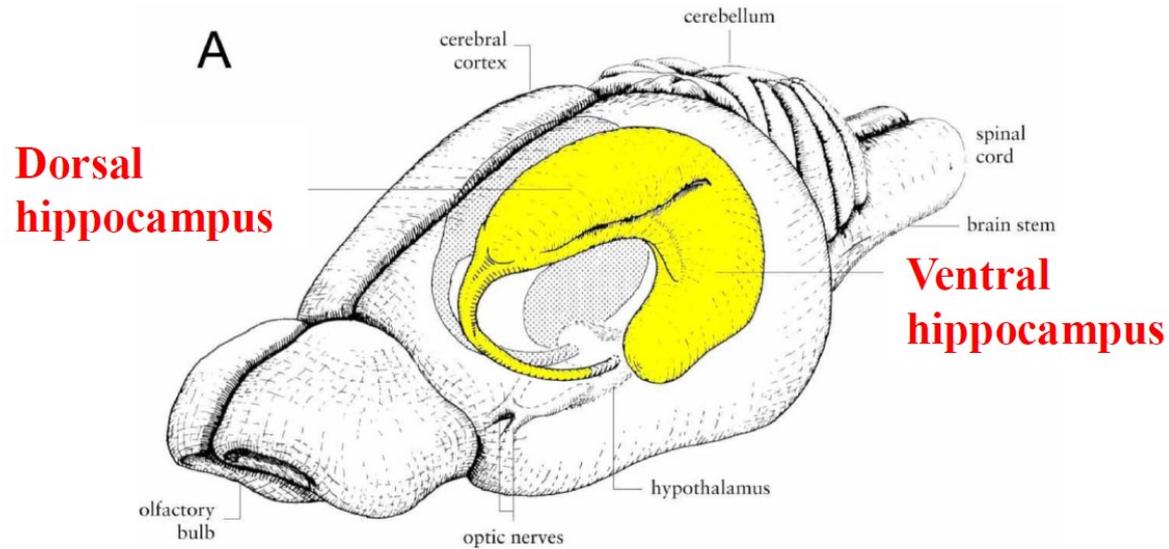
**Second turn: 15:00 - 16:00**

Monday 9.12.24		Tuesday 10.12.24		Wednesday 11.12.24		Thursday 12.12.24	
Hall 2A-Bld D (14:00-16:00)		Hall exCla-Bld C1 (16:00-18:00)		Hall F-Bld C1 (14:00-16:00)		Hall exCla-Bld C1 (16:00-18:00)	
Hall exCla-Bld C1 (16:00-18:00)				Hall exCla-Bld C1 (16:00-18:00)			
<b>Group 3 - Paper 1</b>		<b>Group 6 - Paper 1</b>		<b>Group 7 - Paper 1</b>		<b>Group 5 - Paper 1</b>	
Mayia My		Rossella Di Pompeo		Cristina Baio		Sofia Mosconi	
Giuseppina Russo		Elena Lunardon		Fabiana Cusimano		Maria Sole Faeti	
Diomira Elettra Lenti		Beatrice Buso		Angela Marchiano		Valeria clai	
Saadet Alkan		AnnaMaria Benetti		Rimoun Kaldas		Alessandra guida	
Matteo Theodule		Tobia De Rosso				Maryamsadat Seyedi	
<b>Group 2 - Paper 2</b>		<b>Group 8 - Paper 2</b>		<b>Group 4 - Paper 2</b>		<b>Group 10</b>	
Martina Merga		Francesca Ronchi		GAMAL AHMED MAHMOUD ABDELAZIM		Corinna Perone	
Irene Giovani		Zahra Lashkari		MOHAMED AL SIYABI		Davide Pontiggia	
Carlotta Tiranzoni		Lorenzo sieni		Amanuel Bekaffa		Marianna Cis	
Nosiba Yaseen		Silvia Cassani		Emmanuel		Sara Abbasigharaei	
Jiulija Vodenik		Virginia Camporesi		Yabets Woldegiorgis		Delaram Forouzeh	
<b>Group 1 - Paper 3</b>				<b>Group 9 - Paper 3</b>			
Canonero Marida				Teo Zakula			
Di Filippo Dalila				Danilo Zanghi			
Nicchiotti Francesca				Diana Dall'Olio			
Pau Alessandra				Noemi Valero			
Russo Martina				Gulnur Asci			
				azziza haddad			

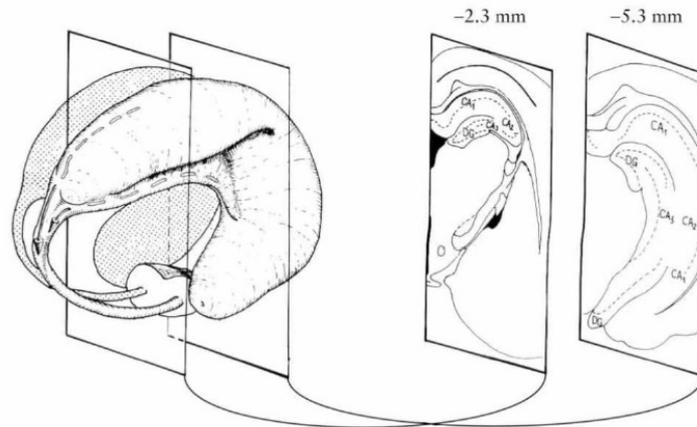
# **Hippocampus & entorhinal cortex anatomy**



# Hippocampus in rodents



**Coronal sections**



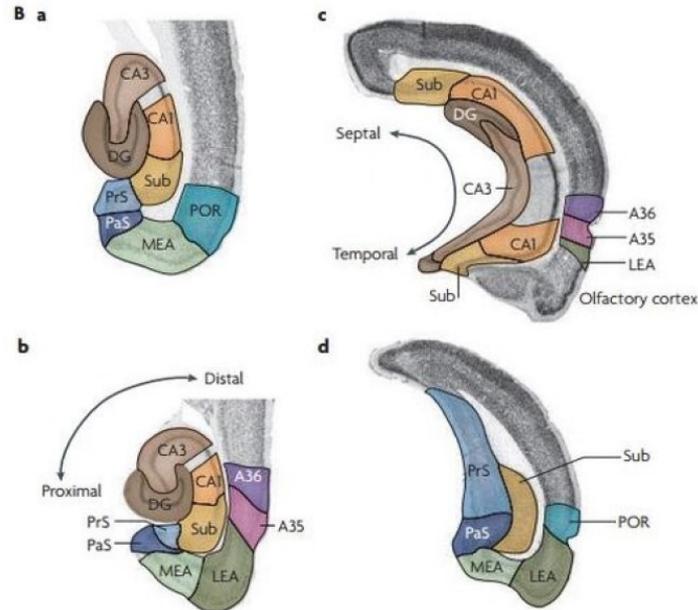
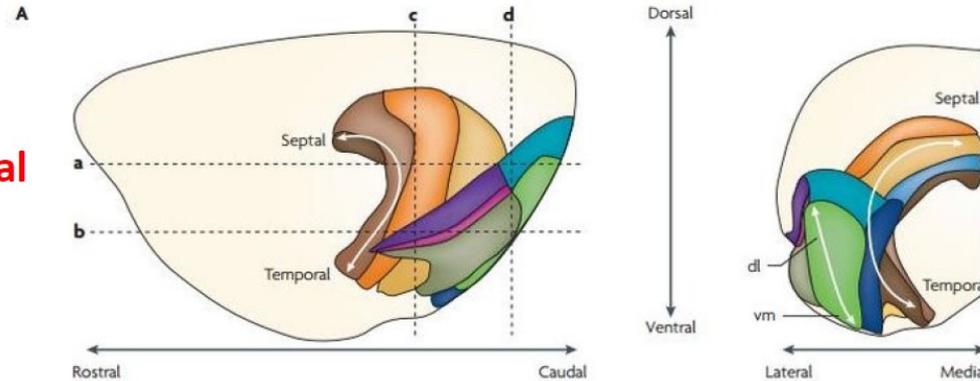
**Dorsal hippocampus**

**Ventral hippocampus**

# Hippocampus in rodents

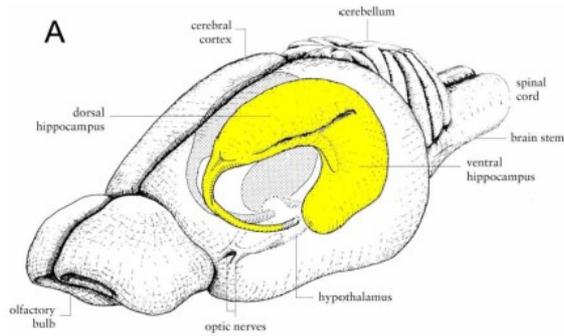
## Coronal sections

## Horizontal sections

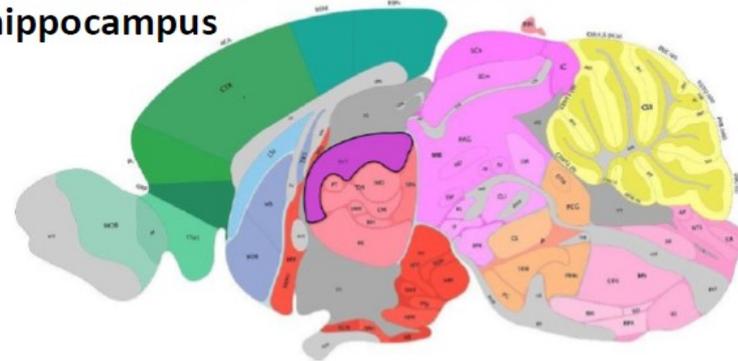


## Ventral hippocampus

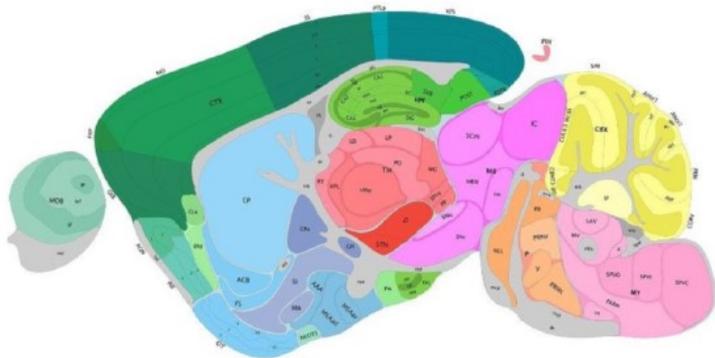
# Hippocampus in rodents - sagittal sections



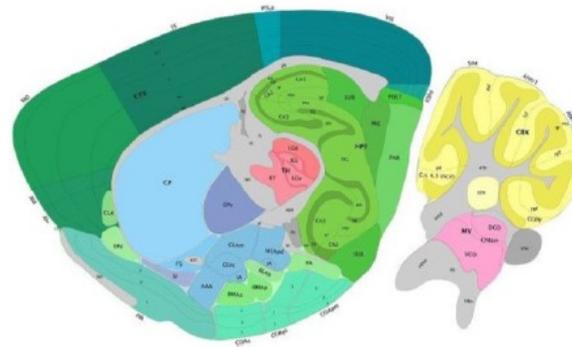
Sagittal section through the midline:  
no hippocampus



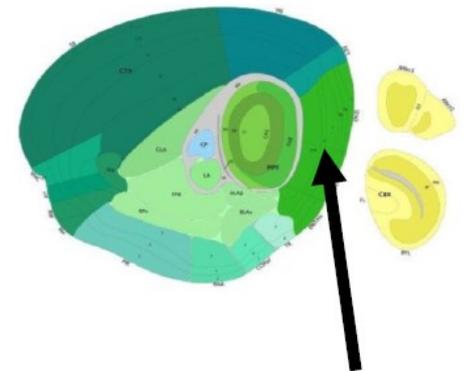
Sagittal section immediately to the right/left of the middle line:  
dorsal hippocampus + dorsal subiculum



More lateral sagittal section:  
Dorsal and ventral hippocampus



Very lateral sagittal section:  
Lateral view of the ventral hippocampus

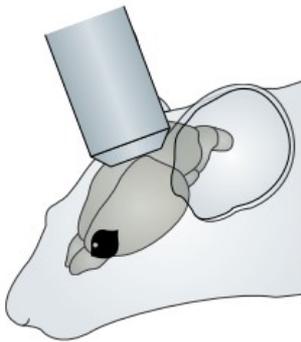


**Medial entorhinal cortex**

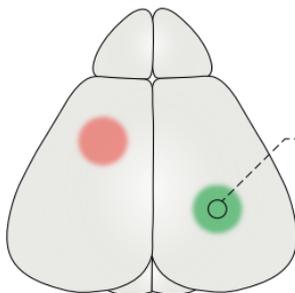
***In vivo* Ca<sup>2+</sup> imaging in freely moving mice**

# *In vivo* experimental setup configurations for different levels of functional imaging, from macroscopic to subcellular level analyses

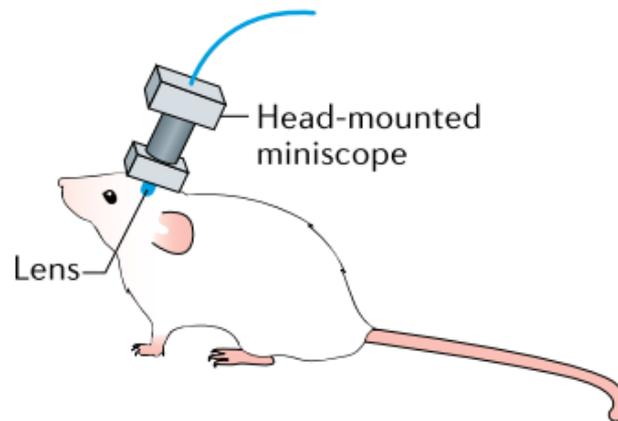
**Fiber photometry**  
in freely moving animals  
Macroscopic - mesoscopic



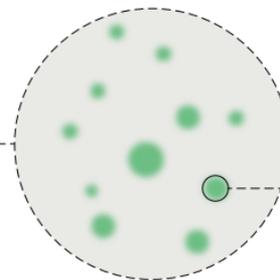
Dorsal surface of cortex



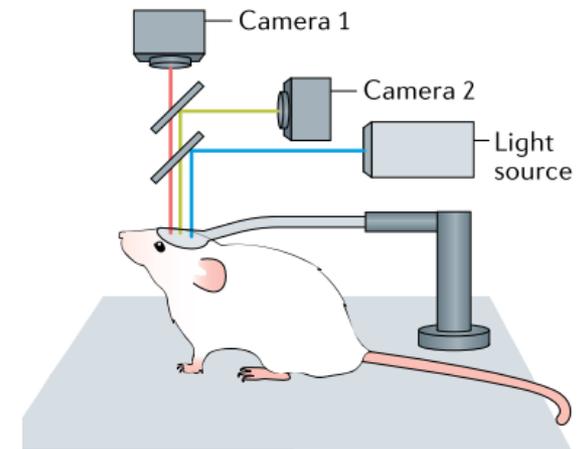
**Miniscope**  
in freely moving animals  
Circuit - cellular



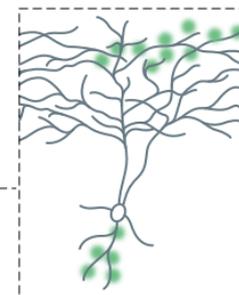
Cellular resolution



**2-foton microscopy**  
in head-fixed animals  
Cellular - subcellular



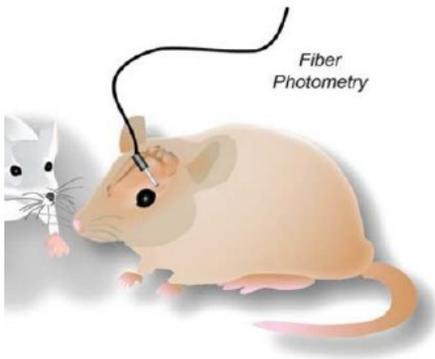
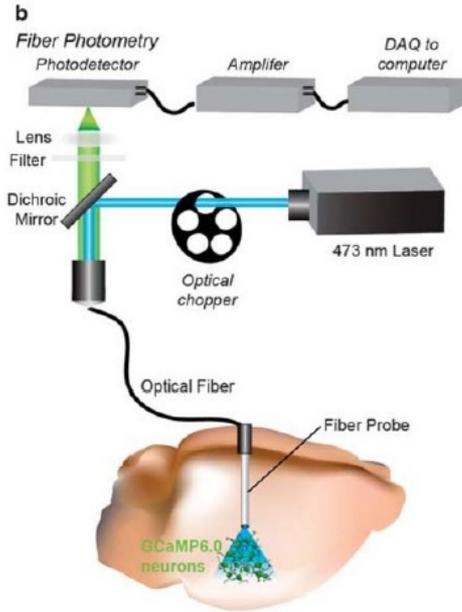
Subcellular resolution



# Fiber photometry

## in freely moving animals

### Macroscopic - mesoscopic



# Neuron

## A direct lateral entorhinal cortex to hippocampal CA2 circuit conveys social information required for social memory

### Highlights

- Lateral entorhinal cortex (LEC) inputs to hippocampal CA2 underlie social memory

Medial entorhinal cortex (MEC) CA2 input is weak and not involved in social memory

Social memory requires the direct but not indirect LEC inputs to CA2

LEC CA2 inputs are selectively activated by social over non-social exploration

### Authors

Jeffrey Lopez-Rojas,  
Christopher A. de Solis, Felix Leroy,  
Eric R. Kandel, Steven A. Siegelbaum

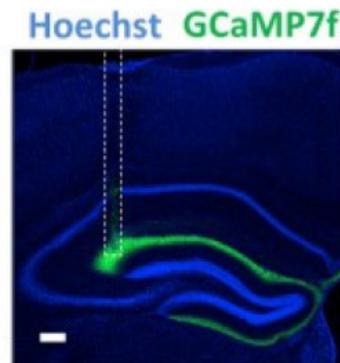
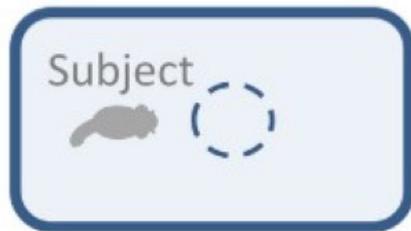
### Correspondence

jl5545@columbia.edu (J.L.-R.),  
sas8@columbia.edu (S.A.S.)

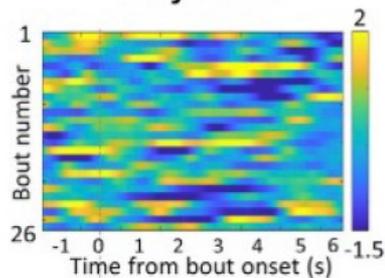
### In brief

Although the CA2 hippocampal region is essential for social memory and detection of social novelty, the inputs that provide social information to CA2 are unknown. We found that social memory depends on the direct inputs CA2 receives from the lateral entorhinal cortex. As this input is activated similarly by novel and familiar individuals, CA2 itself may compute novelty.

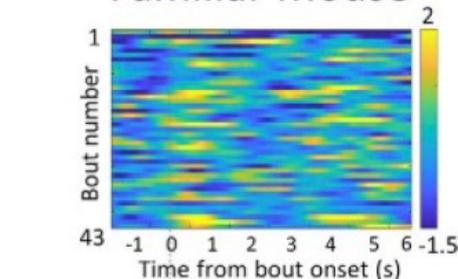
# Fiber photometry in freely moving animals Macroscopic - mesoscopic



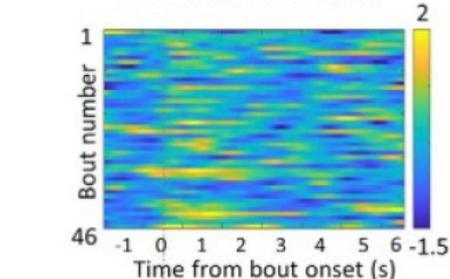
### Object 2



### Familiar Mouse



### Novel Mouse

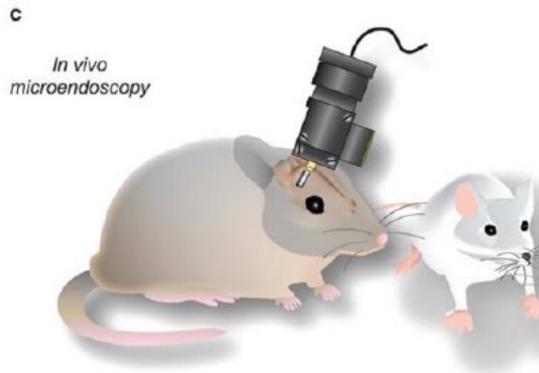
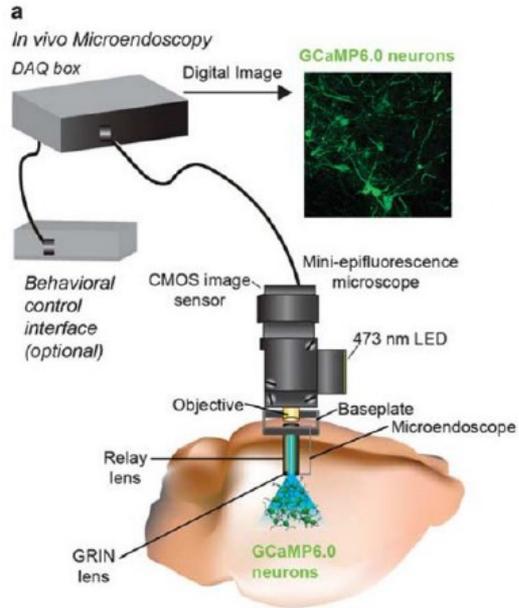


## Figure 6

# Miniscope = miniature microscope

## in freely moving animals

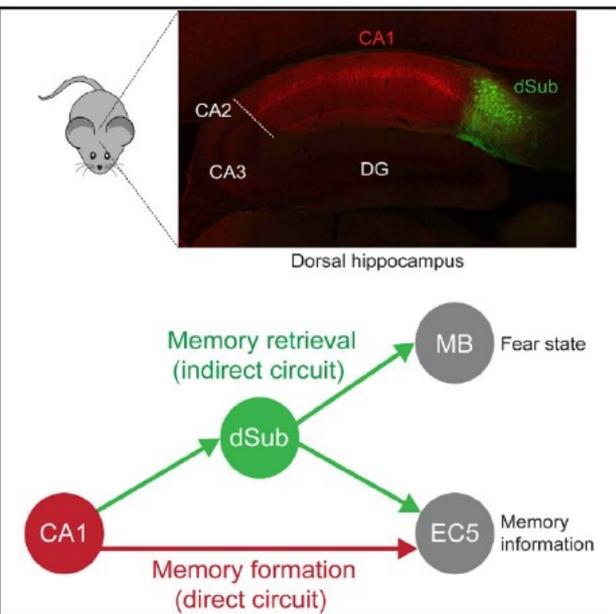
### Circuit - cellular



# Distinct Neural Circuits for the Formation and Retrieval of Episodic Memories

# in freely moving animals Circuit - cellular

Graphical Abstract



Authors

Dheeraj S. Roy, Takashi Kitamura, Teruhiro Okuyama, ..., Yuichi Obata, Atsushi Yoshiki, Susumu Tonegawa

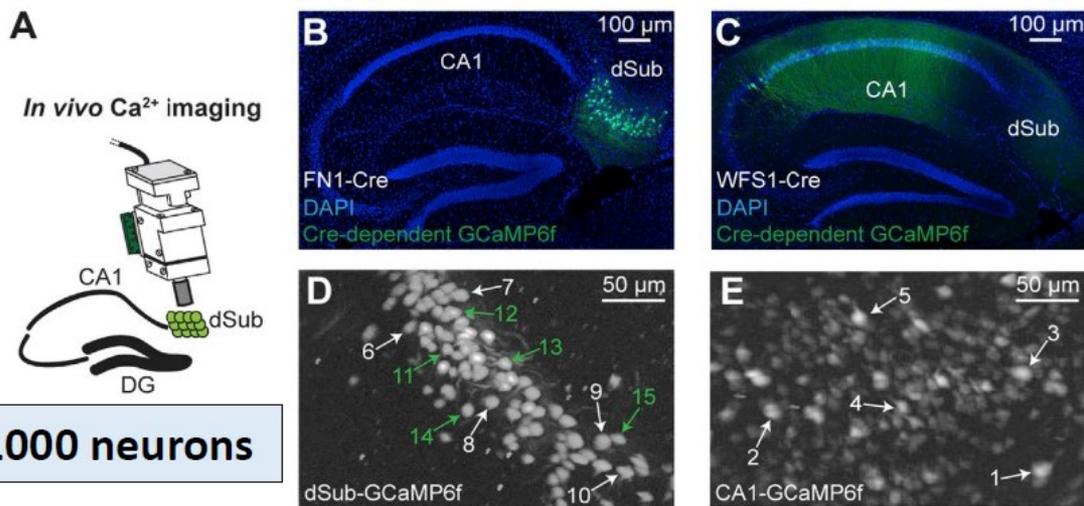
Correspondence

tonegawa@mit.edu

In Brief

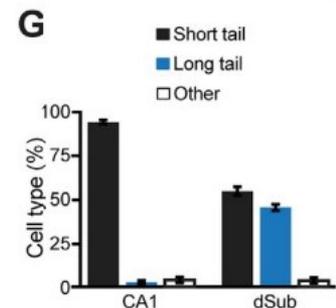
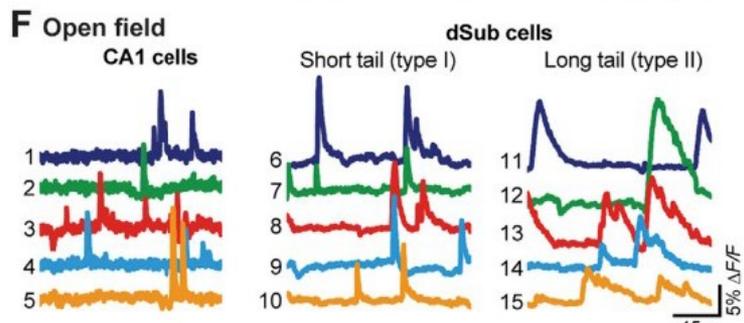
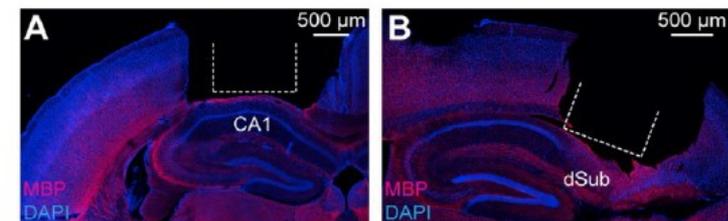
Episodic memories are formed and retrieved through distinct hippocampal pathways.

Figure 6



Possibility to image up to 1000 neurons

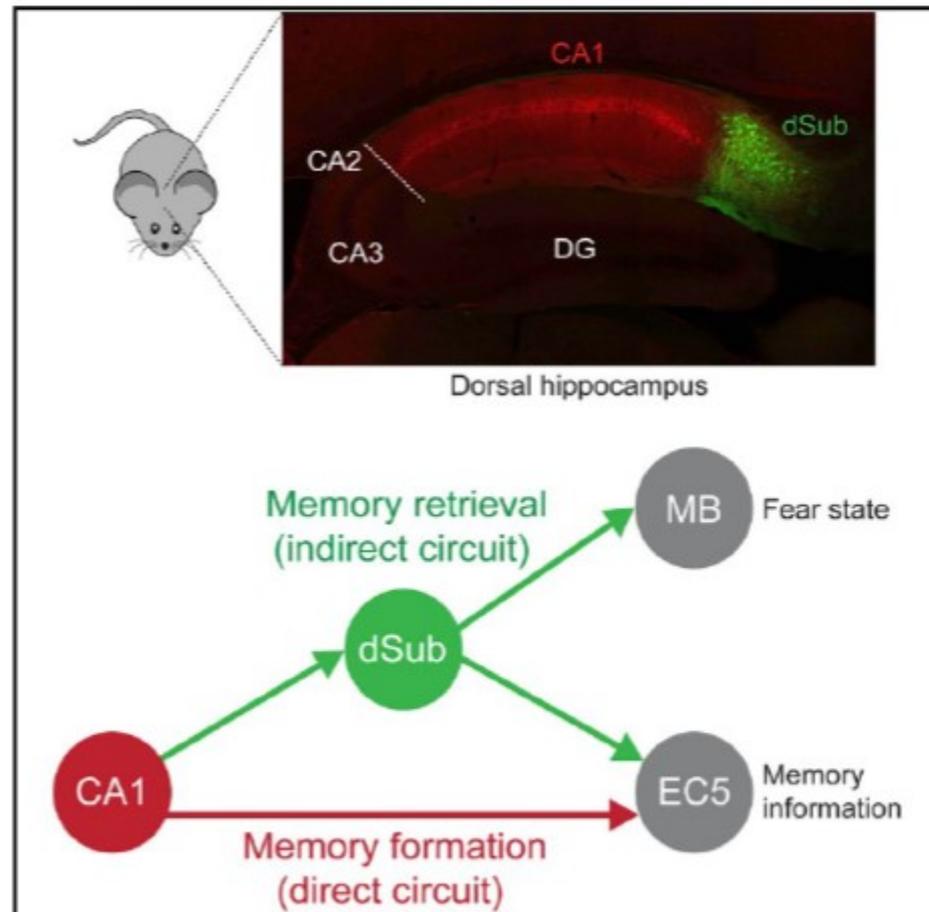
Figure S6



# **Behavioral tests**

# Distinct Neural Circuits for the Formation and Retrieval of Episodic Memories

## Graphical Abstract



## Authors

Dheeraj S. Roy, Takashi Kitamura, Teruhiro Okuyama, ..., Yuichi Obata, Atsushi Yoshiki, Susumu Tonegawa

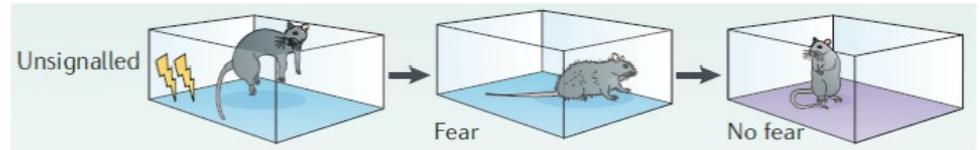
## Correspondence

tonegawa@mit.edu

## In Brief

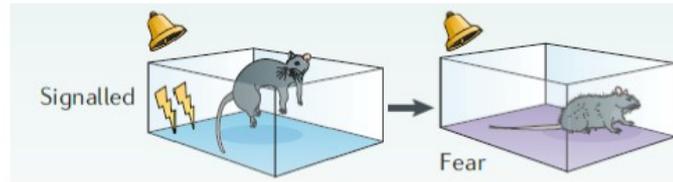
Episodic memories are formed and retrieved through distinct hippocampal pathways.

1. Contextual fear conditioning (CFC)

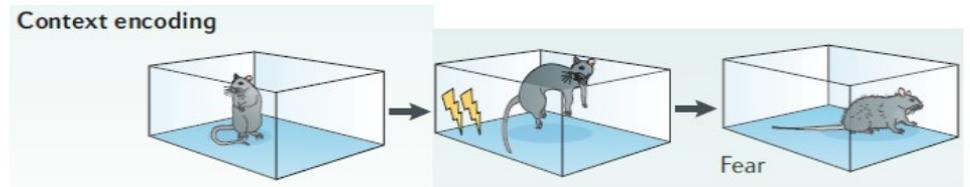


2. Trace fear conditioning (TFC)

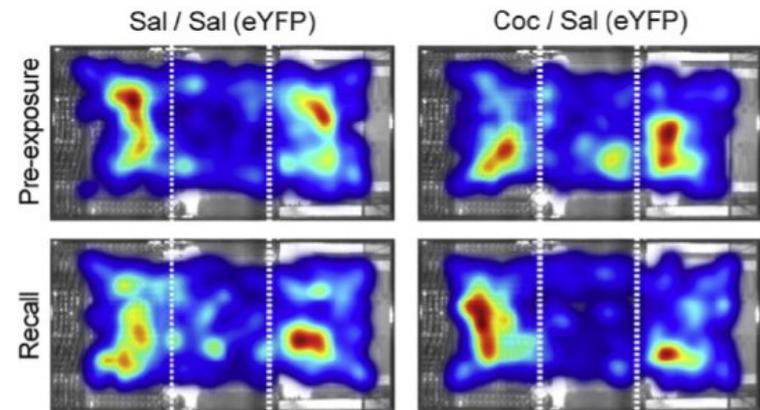
3. Delay fear conditioning (DFC)



4. Memory updating



5. Conditioned place preference (CPP)



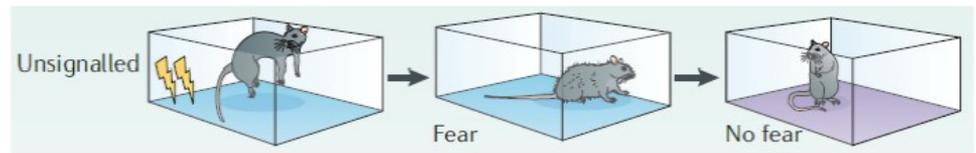
## NEUROSCIENCE

# A shift in the mechanisms controlling hippocampal engram formation during brain maturation

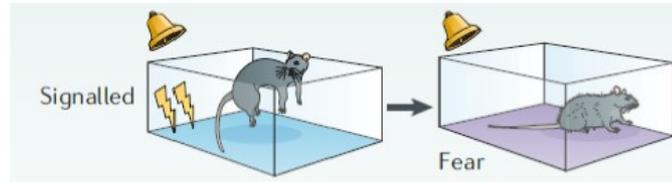
Adam I. Ramsaran<sup>1,2</sup>, Ying Wang<sup>1,3</sup>, Ali Golbabaei<sup>1,4</sup>, Stepan Aleshin<sup>5</sup>, Mitchell L. de Snoo<sup>1,4</sup>, Bi-ru Amy Yeung<sup>1,3</sup>, Asim J. Rashid<sup>1</sup>, Ankit Awasthi<sup>1</sup>, Jocelyn Lau<sup>1,3</sup>, Lina M. Tran<sup>1,3,6</sup>, Sangyoon Y. Ko<sup>1,3</sup>, Andrin Abegg<sup>1,7</sup>†, Lana Chunan Duan<sup>1,4</sup>, Cory McKenzie<sup>1,2</sup>, Julia Gallucci<sup>1</sup>‡, Moriam Ahmed<sup>1</sup>, Rahul Kaushik<sup>5,8</sup>, Alexander Dityatev<sup>5,8,9</sup>, Sheena A. Josselyn<sup>1,2,3,4,10</sup>, Paul W. Frankland<sup>1,2,3,4,11</sup>\*

The ability to form precise, episodic memories develops with age, with young children only able to form gist-like memories that lack precision. The cellular and molecular events in the developing hippocampus that underlie the emergence of precise, episodic-like memory are unclear. In mice, the absence of a competitive neuronal engram allocation process in the immature hippocampus precluded the formation of sparse engrams and precise memories until the fourth postnatal week, when inhibitory circuits in the hippocampus mature. This age-dependent shift in precision of episodic-like memories involved the functional maturation of parvalbumin-expressing interneurons in subfield CA1 through assembly of extracellular perineuronal nets, which is necessary and sufficient for the onset of competitive neuronal allocation, sparse engram formation, and memory precision.

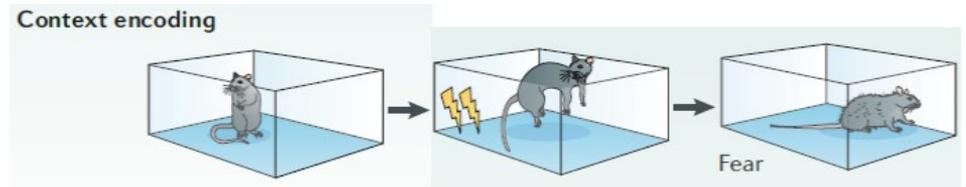
# 1. Contextual fear conditioning (CFC)



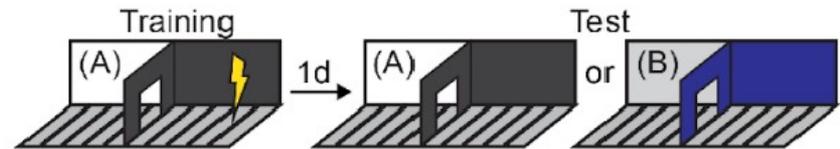
# 2. Delay fear conditioning (DFC)



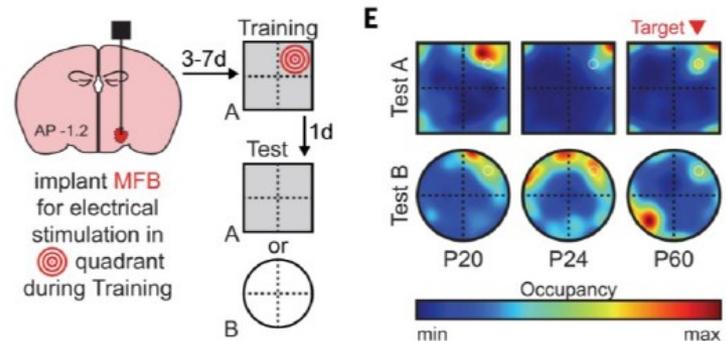
# 3. Memory updating



# 4. Inhibitory avoidance (IA)

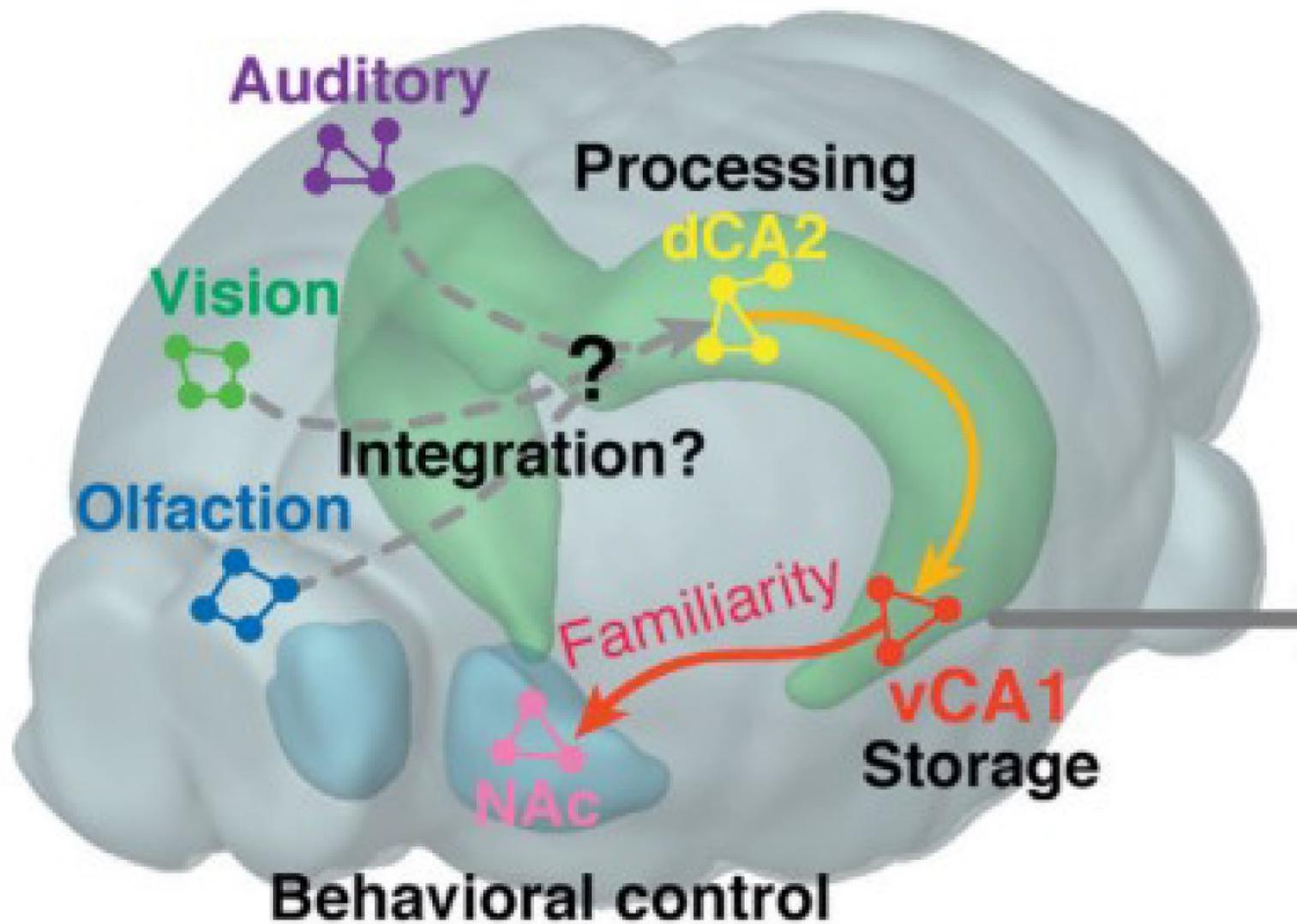


# 5. Spatial foraging Task (Conditioned place preference)



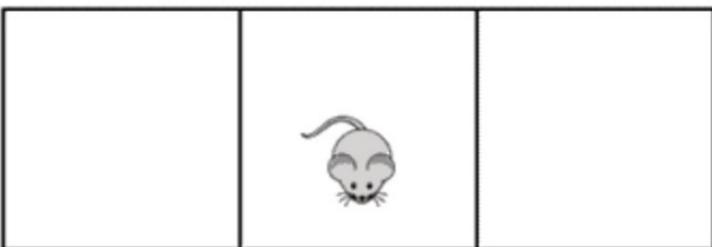
# Social memory



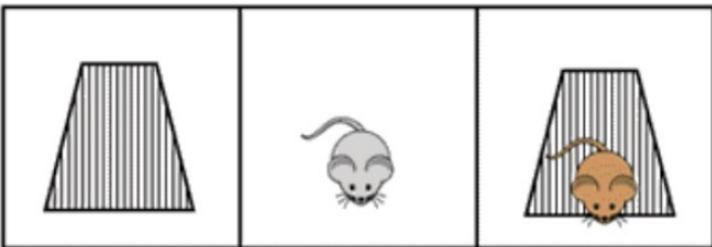




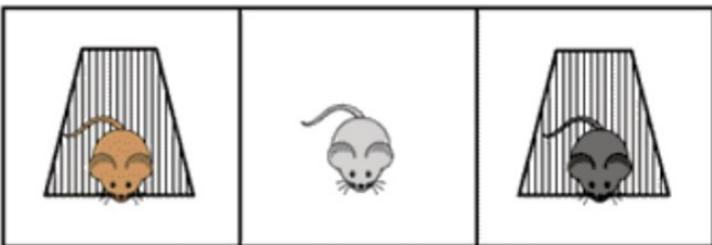
## 3-Chamber Sociability and Social Novelty Test



Habituation: Empty Apparatus

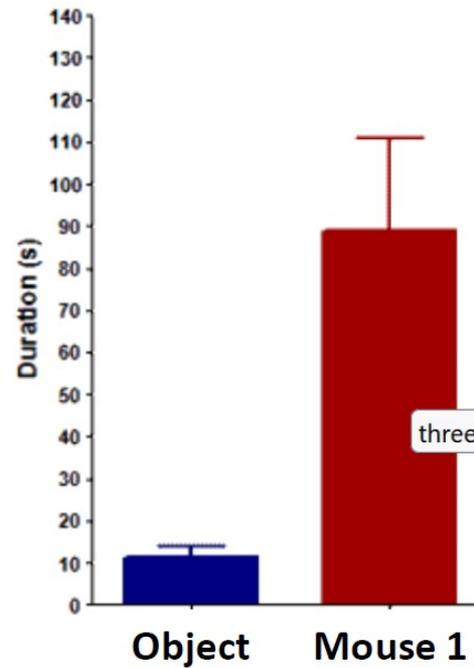


Sociability: Novel Object; Mouse 1

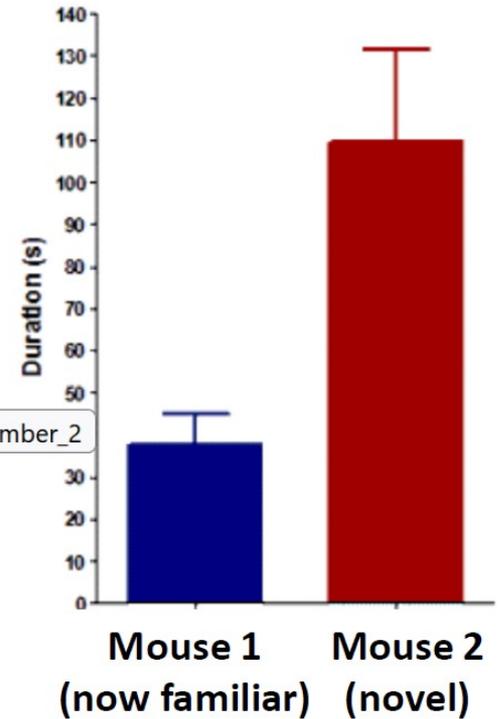


Social Novelty: Mouse 1; Mouse 2

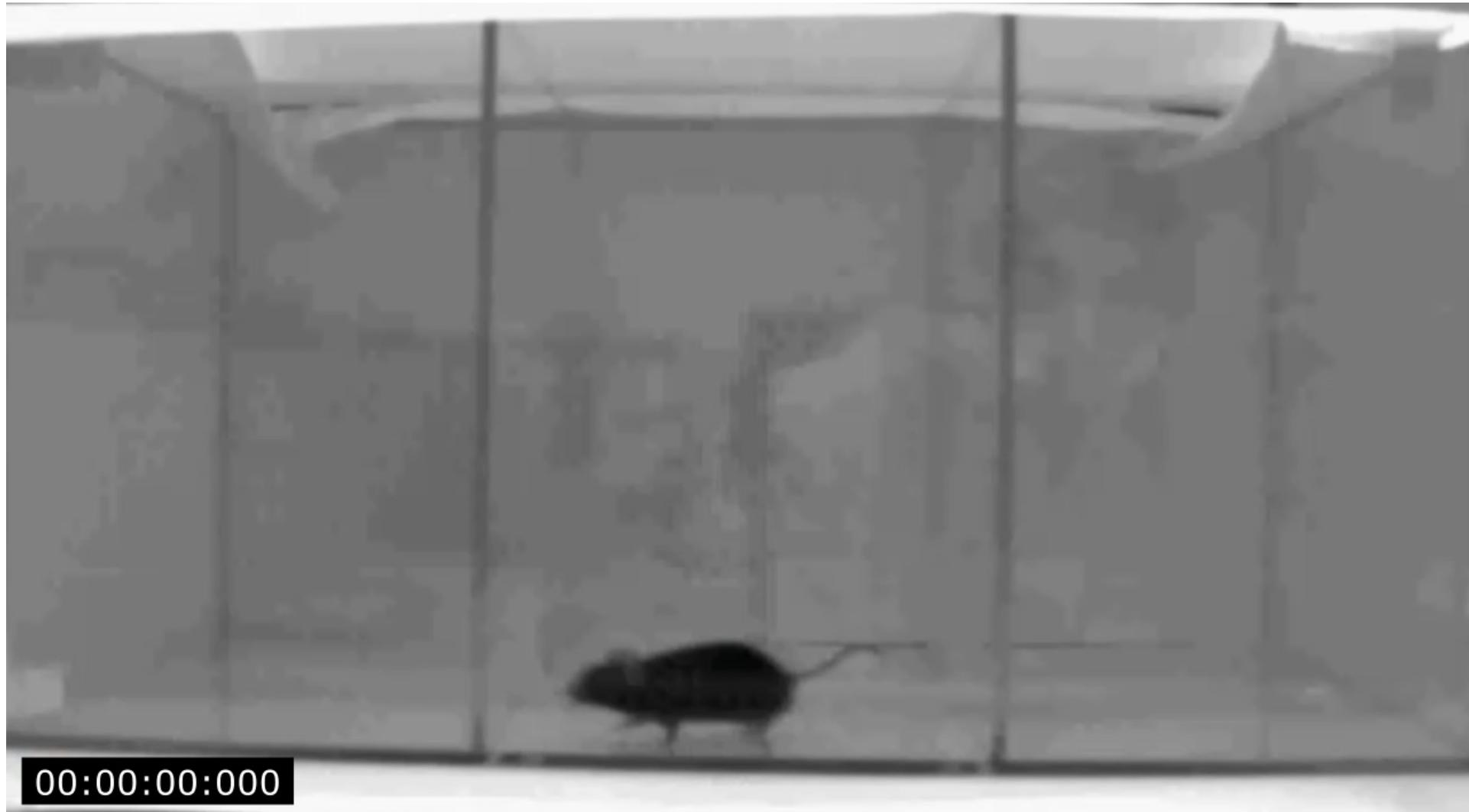
Sniffing Time: Sociability  
(C57BI6/J)



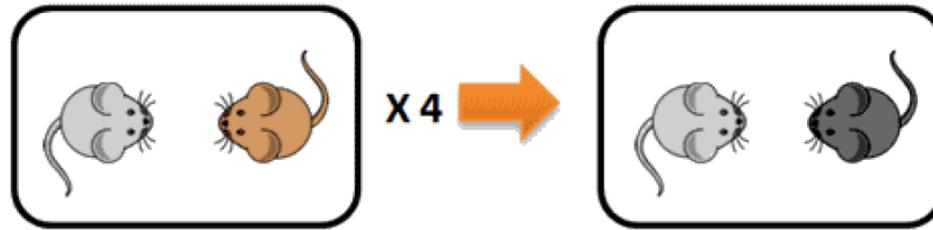
Sniffing Time: Social Novelty  
(C57BI6/J)



# 3-Chamber Sociability and Social Novelty Test



# 5-Trial Social Memory Test



Trial 1-4:  
Intruder 1

Trial 5:  
Intruder 2

## Constitutive *Itgb3* KO mice

