

# Neuronal plasticity in the olfactory bulb during simple and complex perceptual learning



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September 2017

# Olfactory system

Food  
search

Danger  
avoidance

Social  
interactions

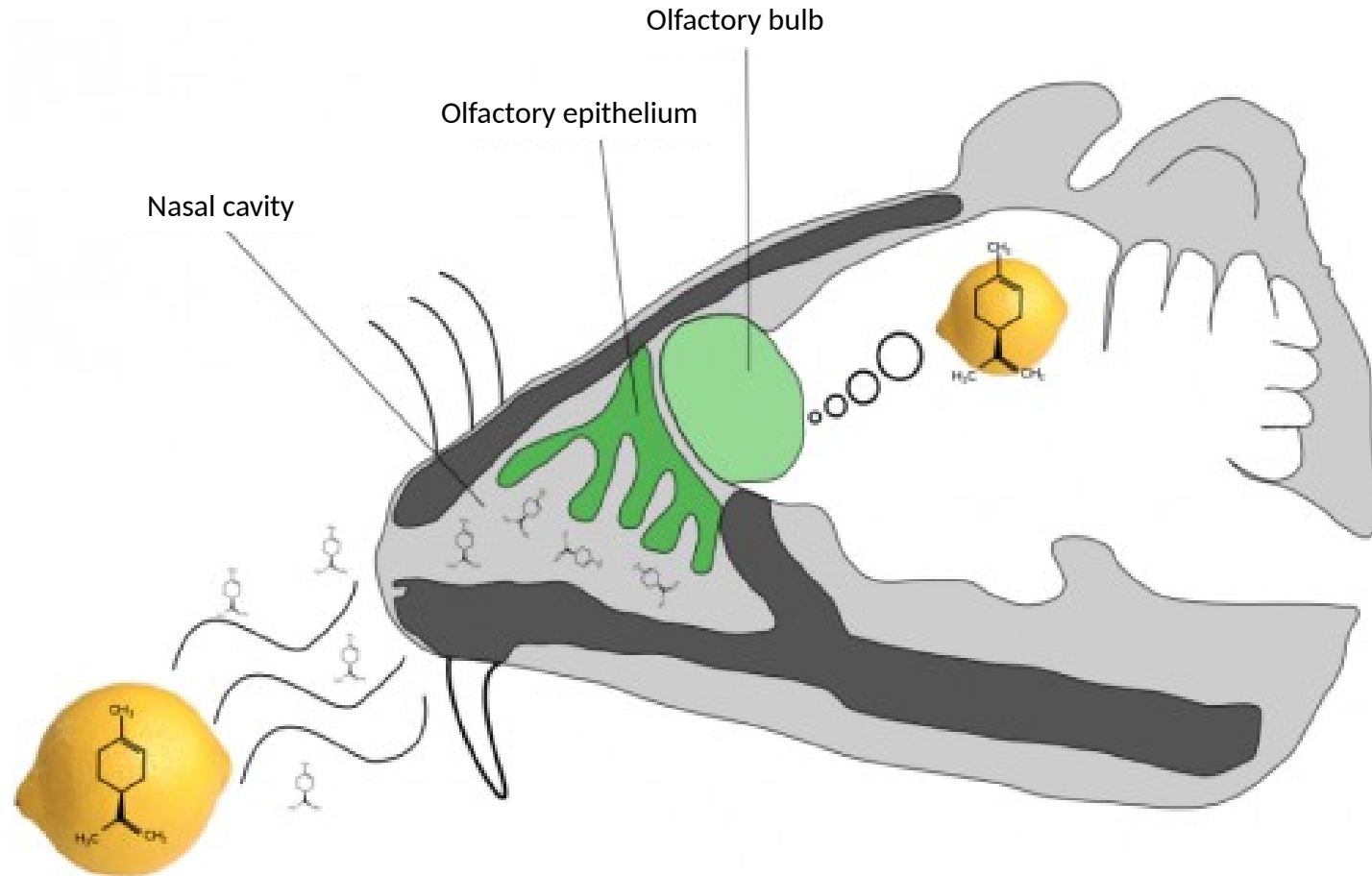
Human



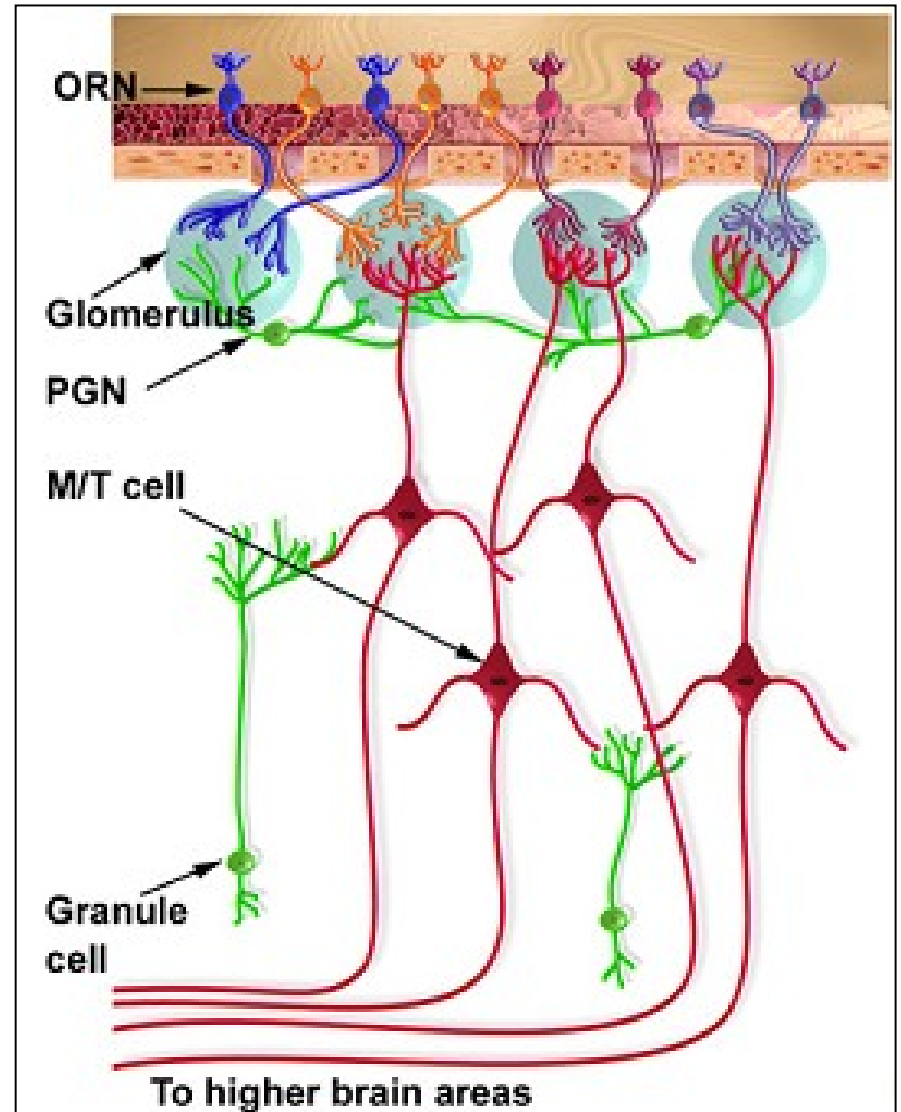
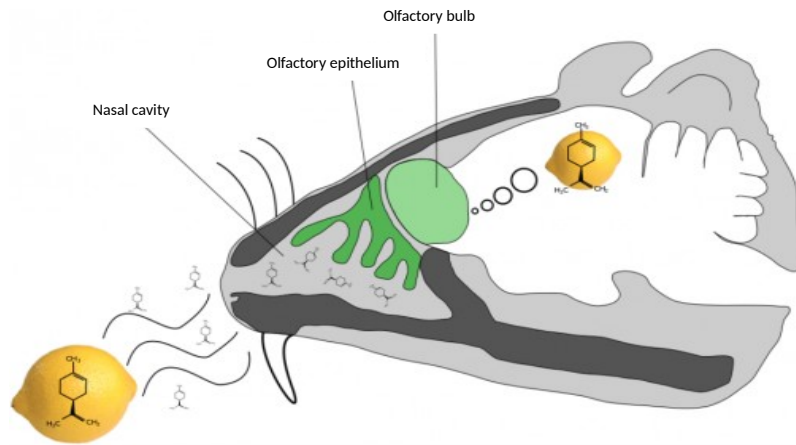
Mice



# Olfactory system

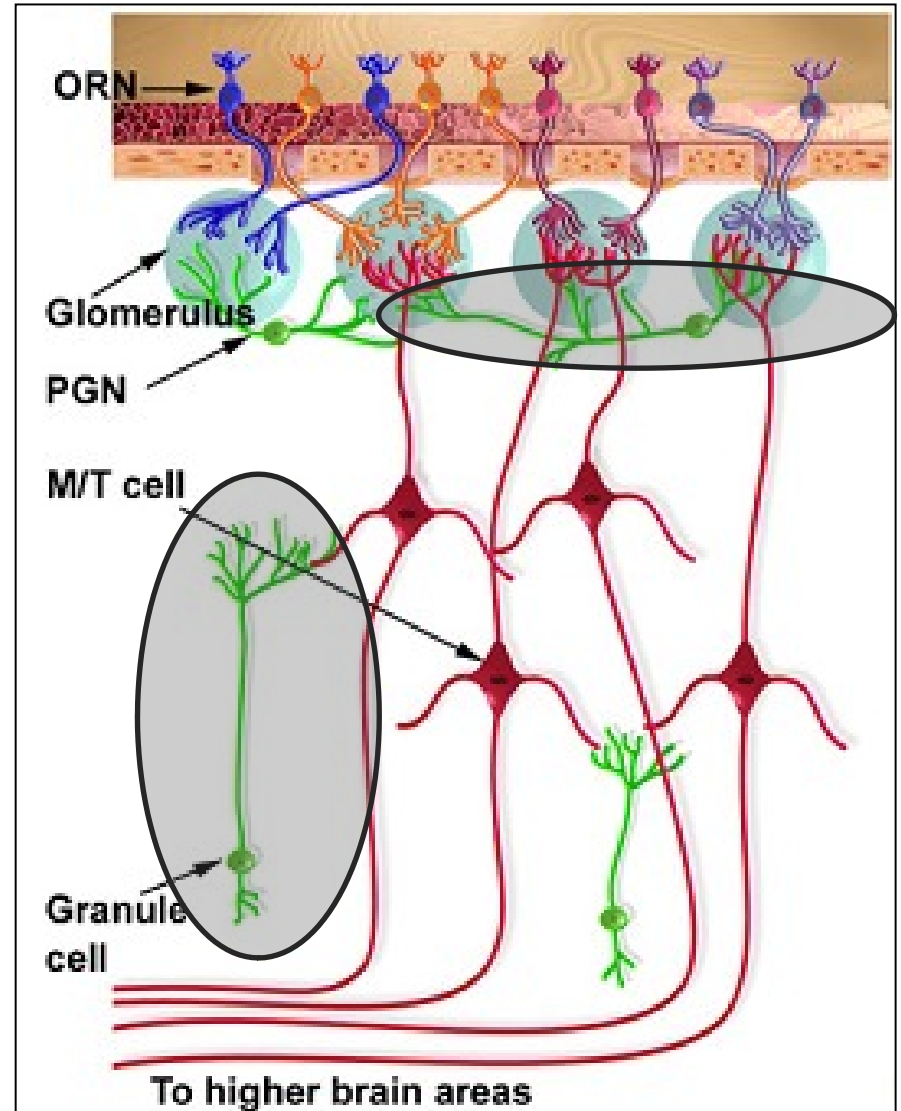
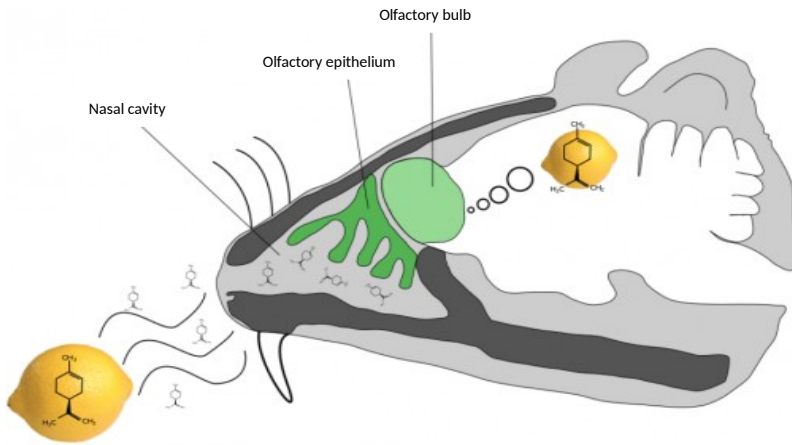


# Olfactory system



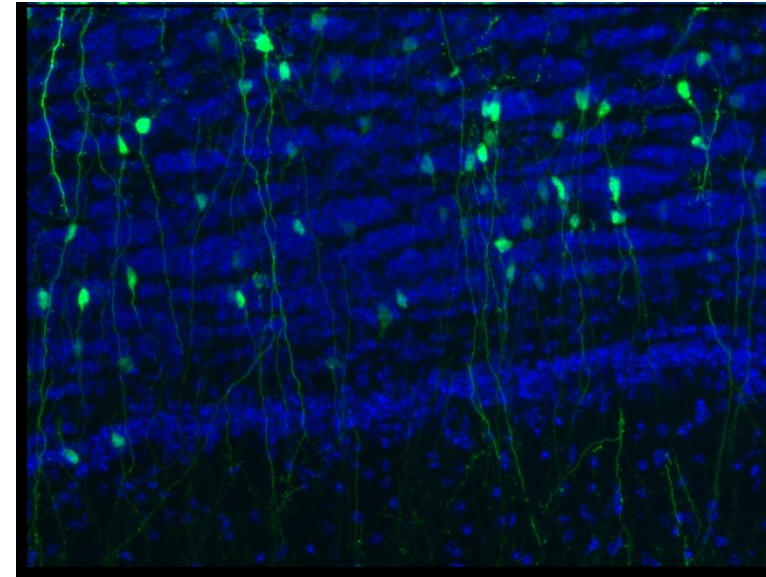
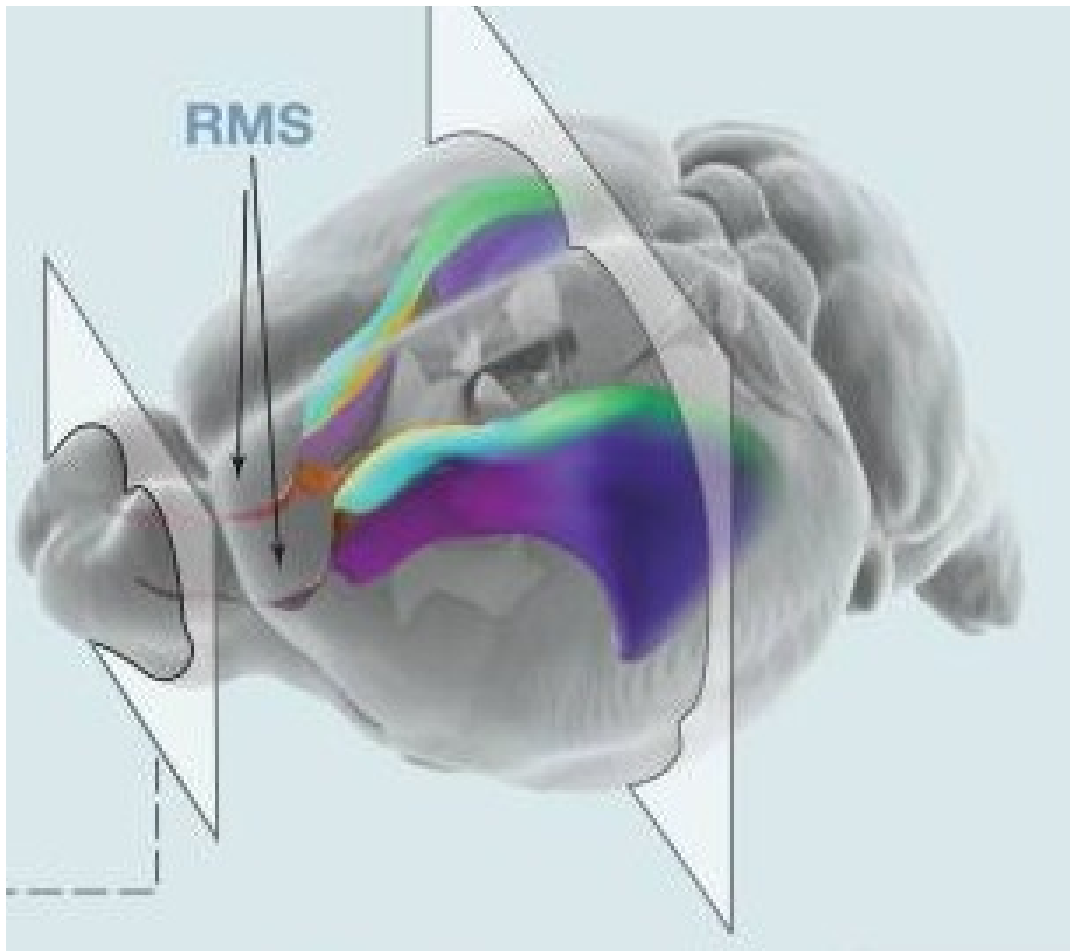
Adapted from Adam and Mizrahi 2010 Curr Opin Neurobiol

# Olfactory system



Adapted from Adam and Mizrahi 2010 Curr Opin Neurobiol

# Adult neurogenesis

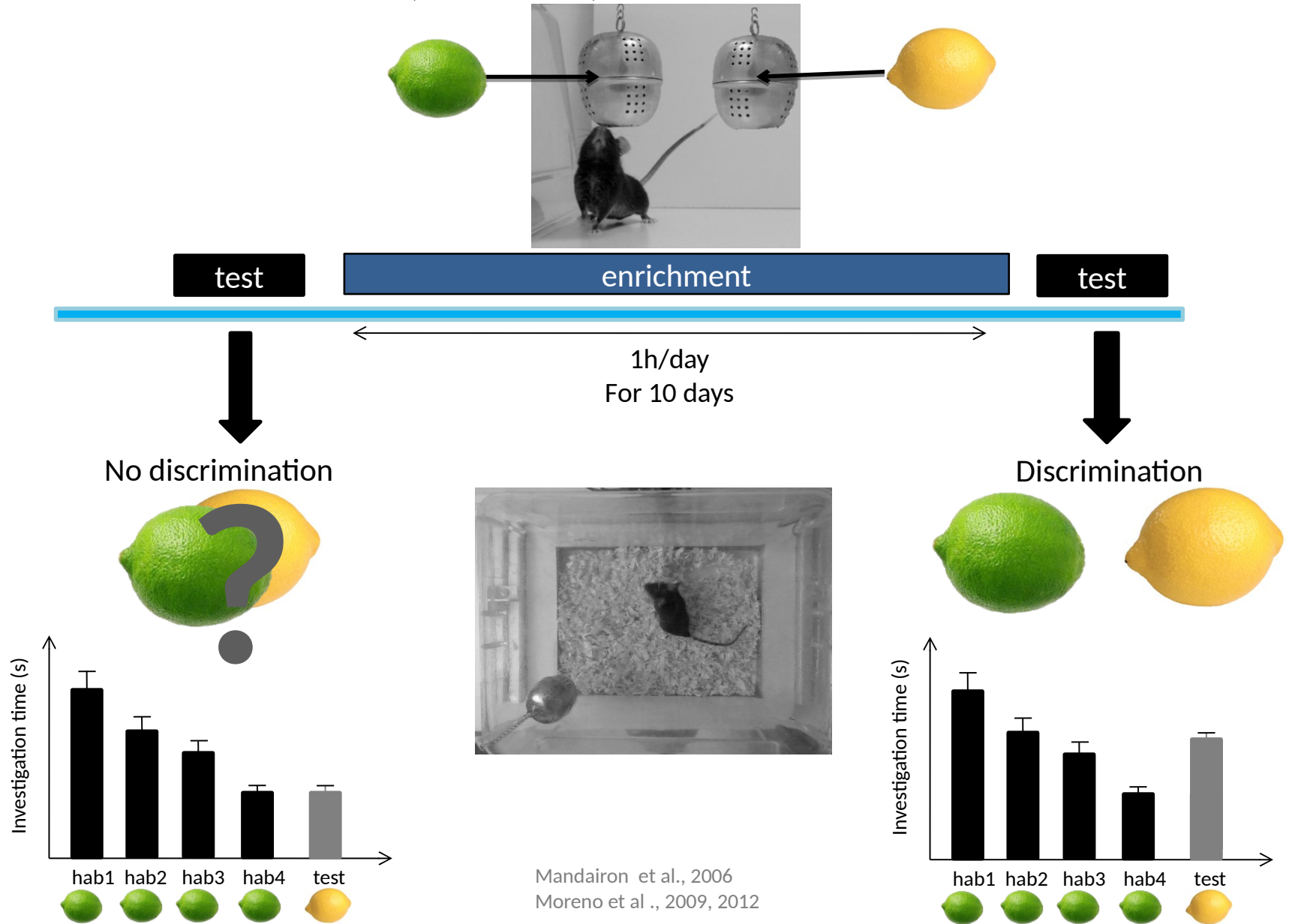


From Tong and Alvarez-Buylla, 2014, Neuron

Lledo et al. Nat Neuro 2006 ; Rocherfort et al. J Neuro 2002 ; Mandairon et al. Behav Neuro, 2006;  
Mandairon et al., J Neuro, 2011 ; Alonso et al., J Neuro, 2006 ; Moreno et al., PNAS, 2009

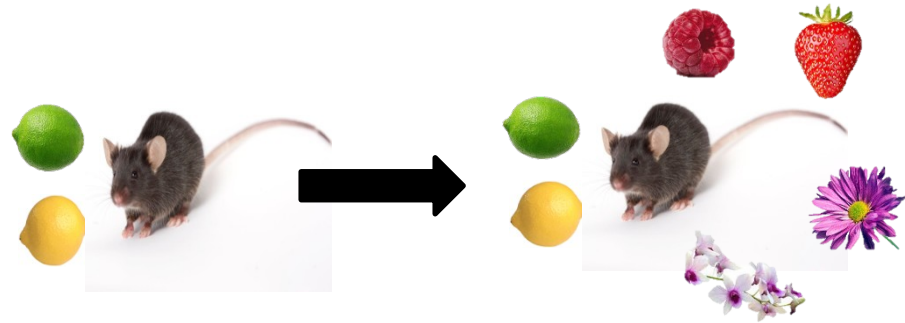
# Perceptual learning

Significant improvement of the discrimination abilities of perceptually close odorants after repeated exposition to these same odorants(= enrichment).

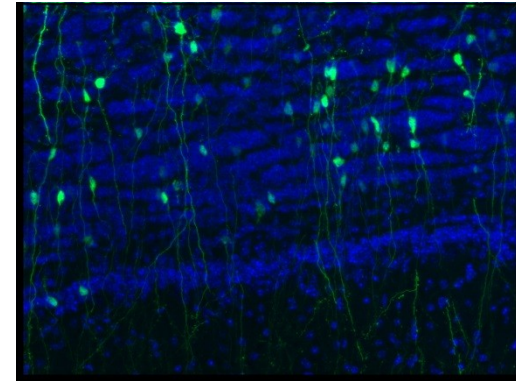


# Perceptual learning

- **Simple perceptual learning** paradigm = 1 pair of odorants.  
Real olfactory environment is **more complex** = several pairs of odorants



- There are **adult-born neurons** in the olfactory bulb but also **preexisting neurons**, born during ontogenesis

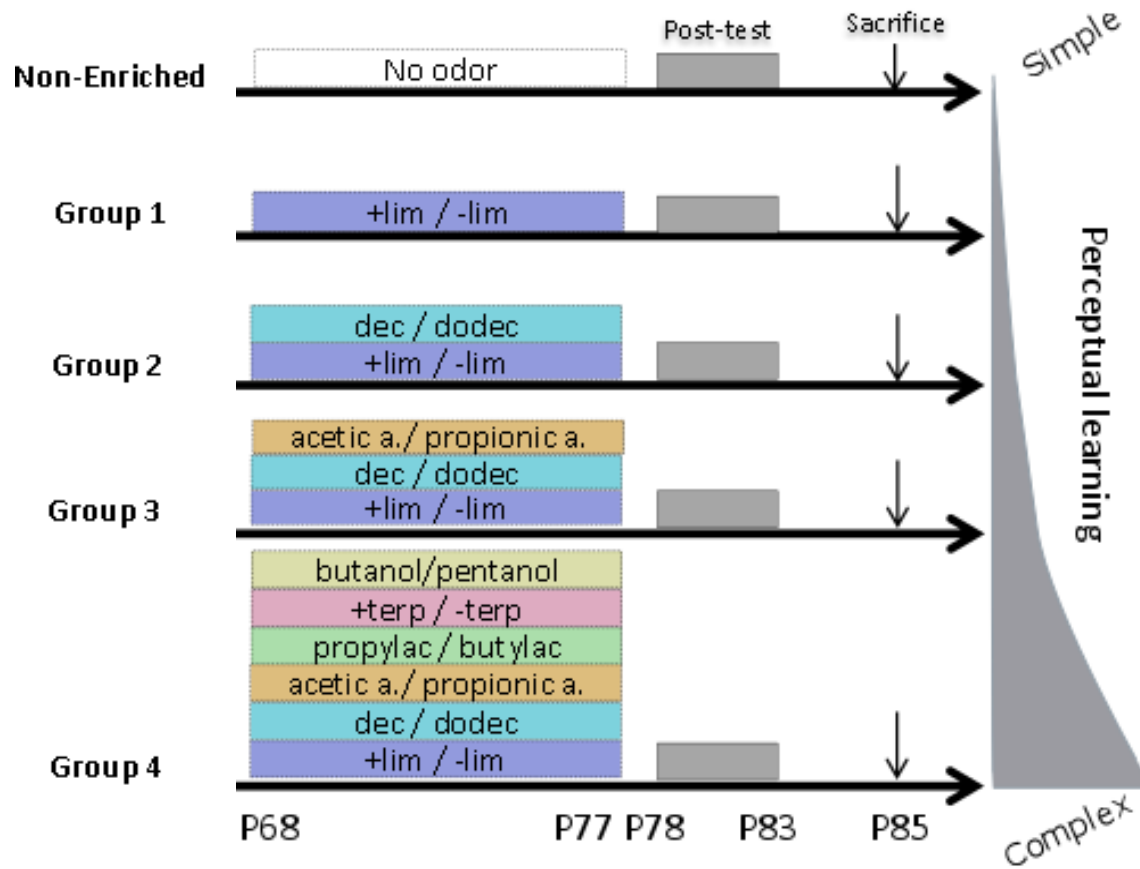


Up to which point can we push neurogenesis?

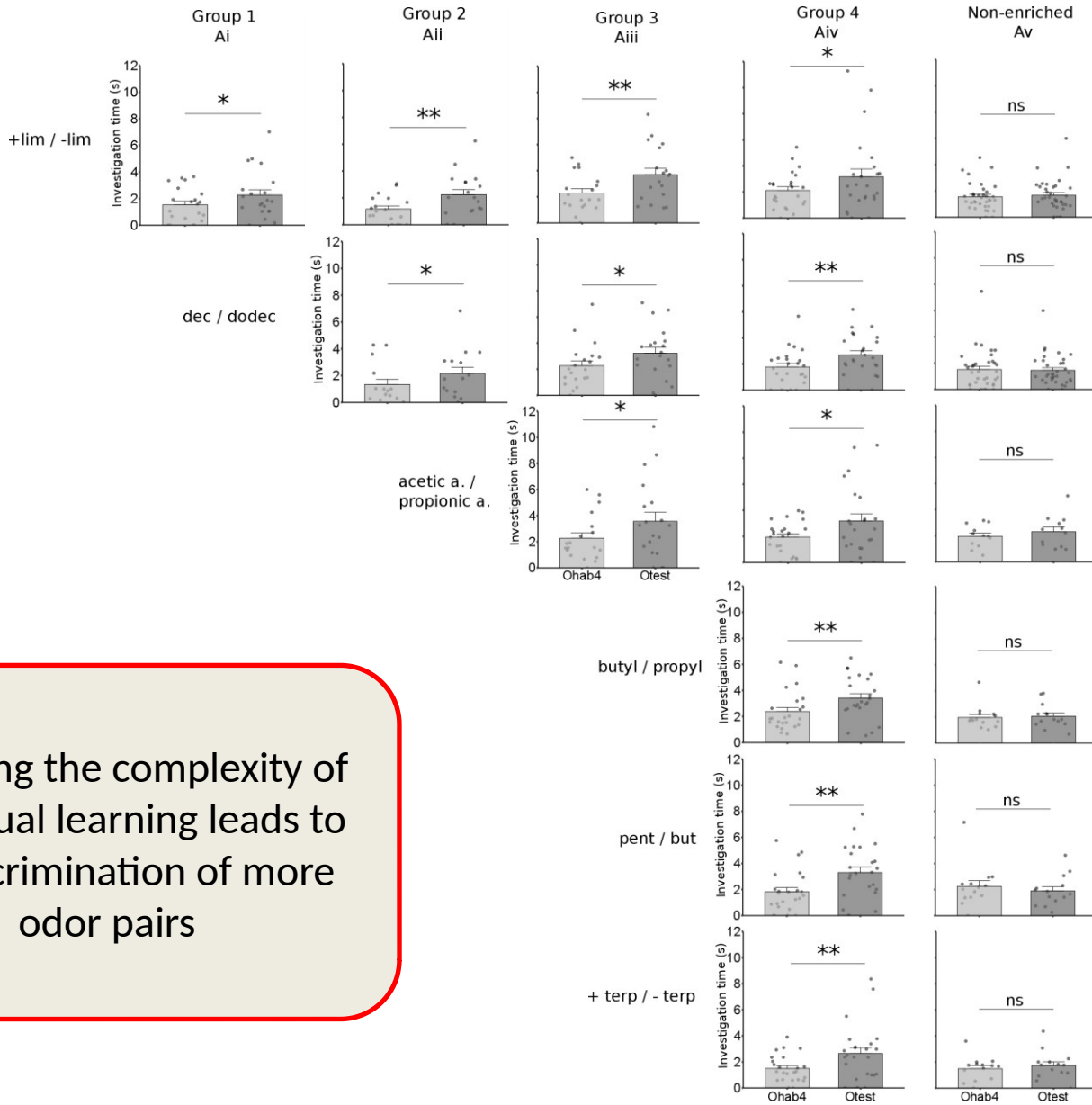
Are adult-born neurons always necessary and/or sufficient?



# Neuronal plasticity in the olfactory bulb during simple and complex learning

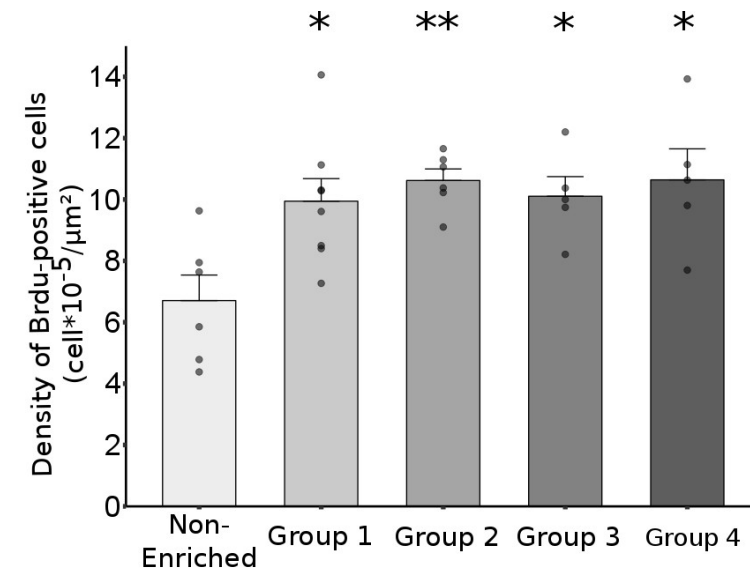
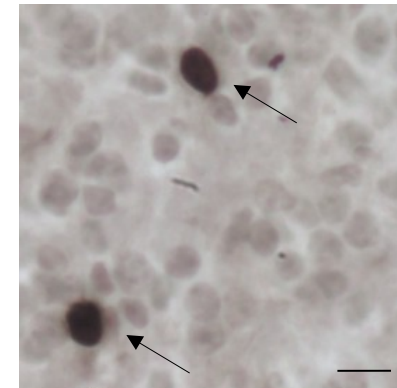
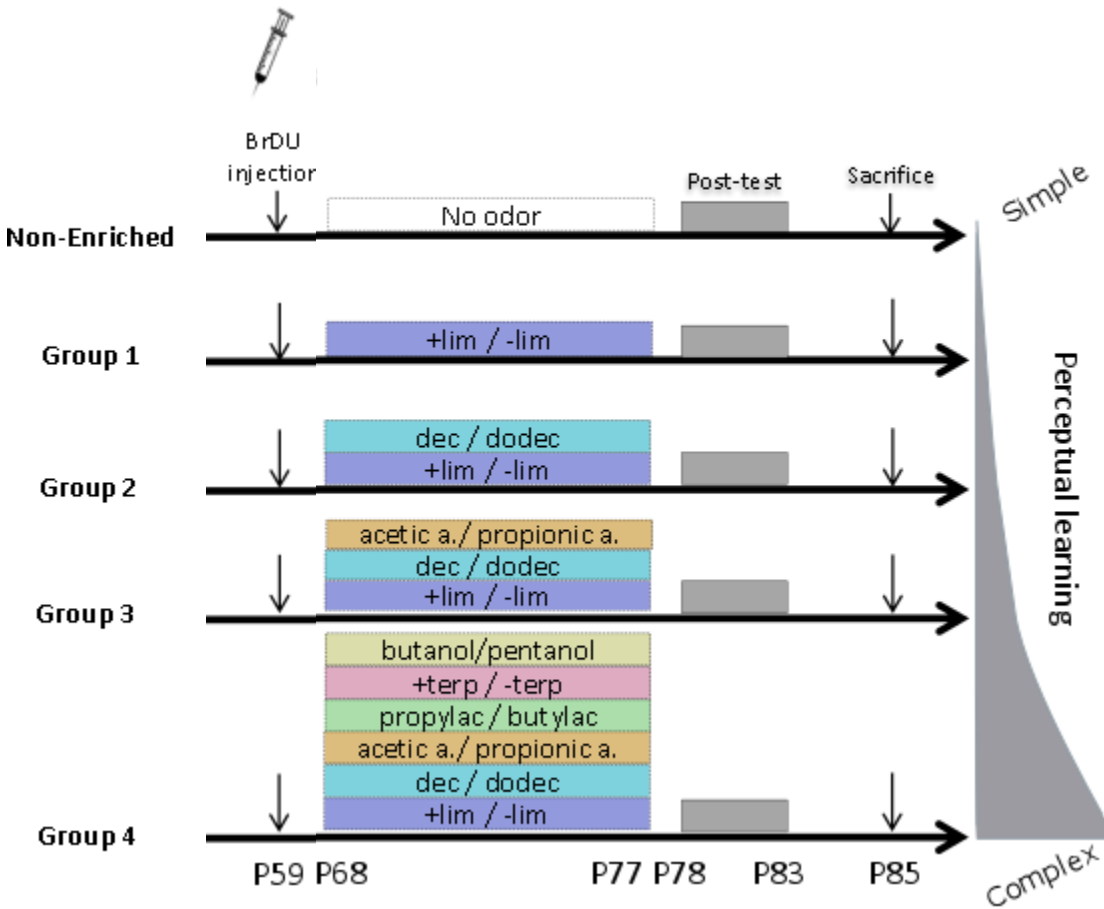


# 1 - Behavior:



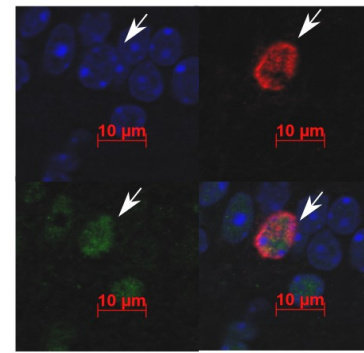
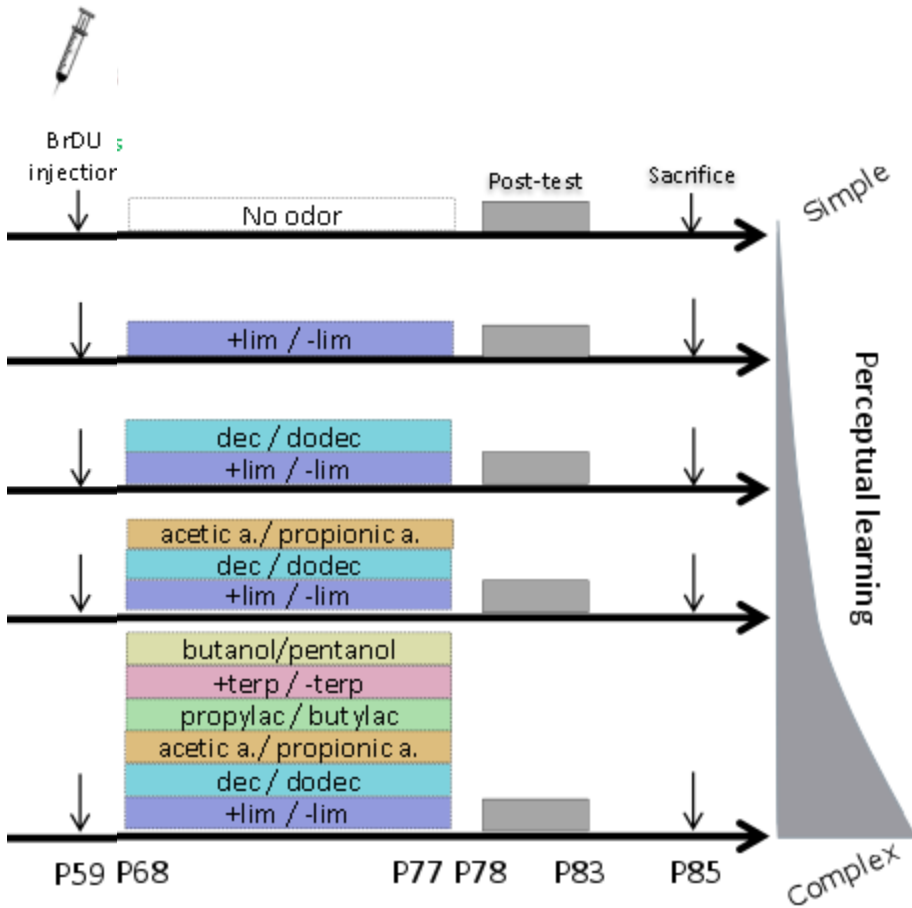
Increasing the complexity of perceptual learning leads to the discrimination of more odor pairs

## 2 - Newborn neurons density:

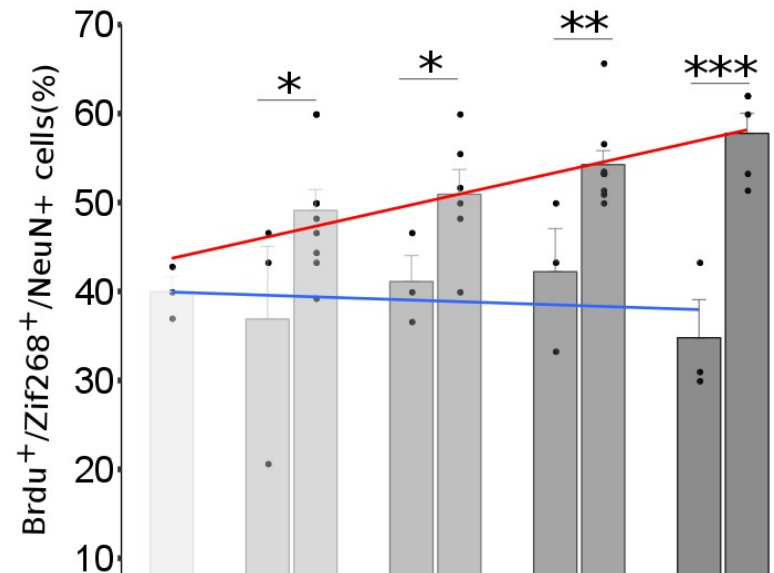


Perceptual learning increased adult-born cell density independently of the enrichment's complexity.

## 2 - Newborn neurons responsiveness:



Adjusted  $R^2=0.35$ ,  $p=0.003$  \*\*  
Adjusted  $R^2=0.20$ ,  $p=0.808$



Increasing the complexity of perceptual learning enhances the recruitment of adult-born neurons in the processing of the learned odorants

Simple

Complex

### 3 - Newborn neurons and preexisting neurons morphology:



Non-Enriched

Group 1

Group 2

Group 3

Group 4

BrDU  
injection

No odor

+lim / -lim

dec / dodec  
+lim / -lim

acetic a. / propionic a.  
dec / dodec  
+lim / -lim

butanol / pentanol  
+terp / -terp  
propylac / butylac  
acetic a. / propionic a.  
dec / dodec  
+lim / -lim

Post-test

Sacrifice

Simple

Perceptual learning

Complex

P59

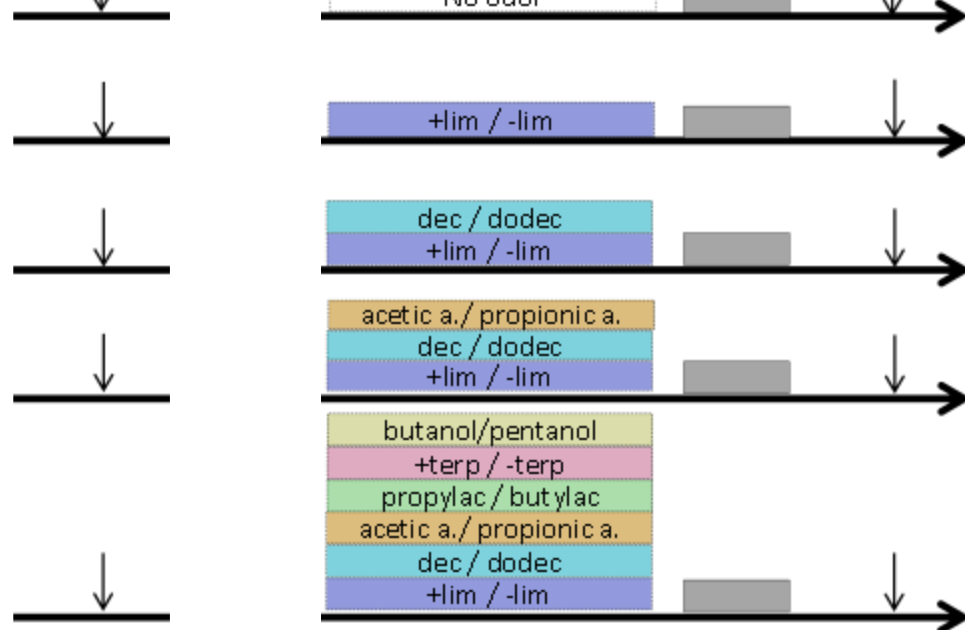
P68

P77

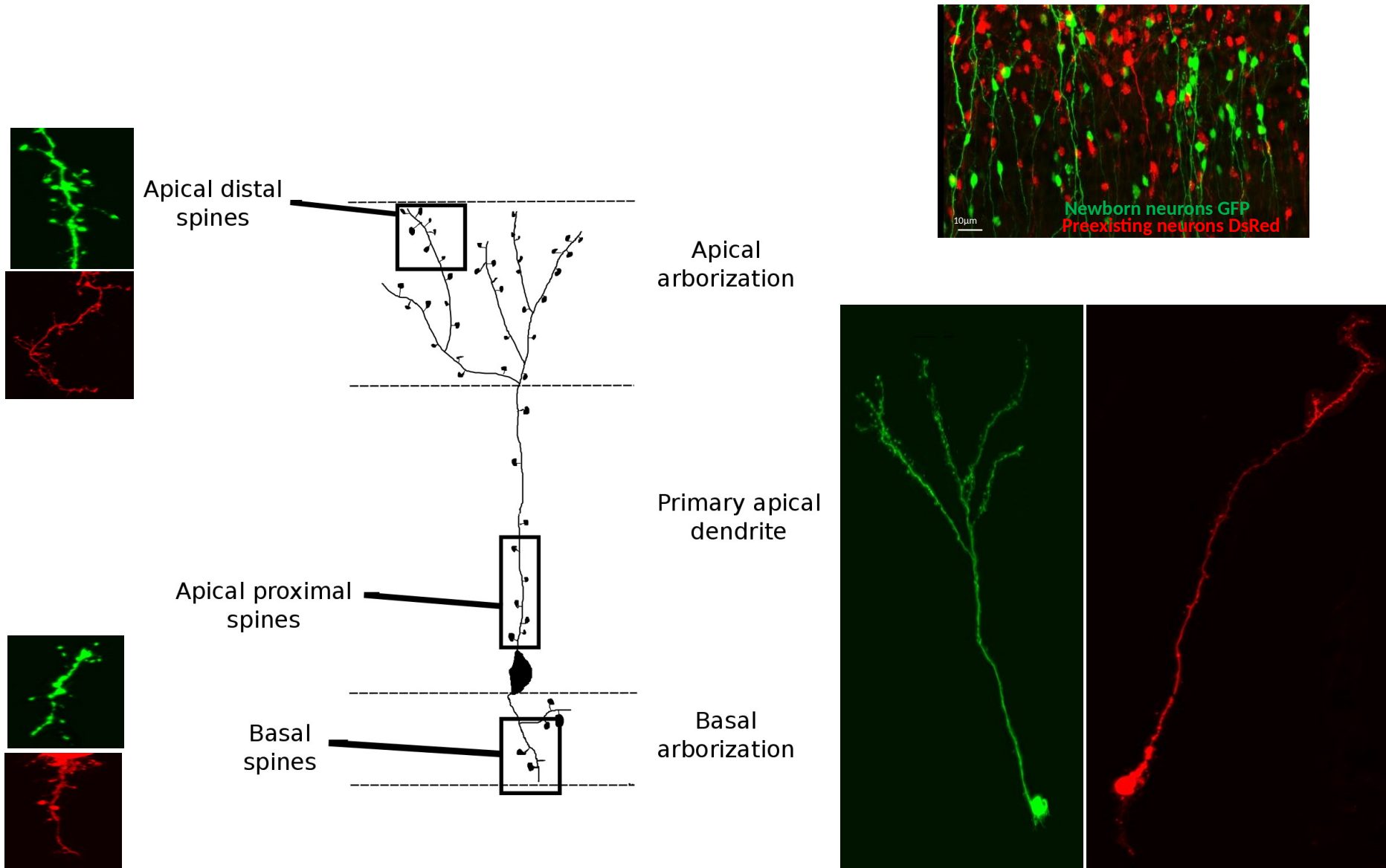
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P83

P85

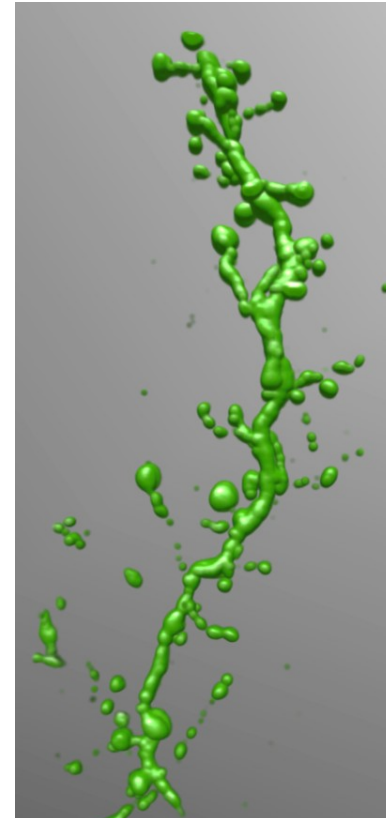
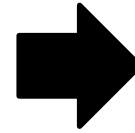
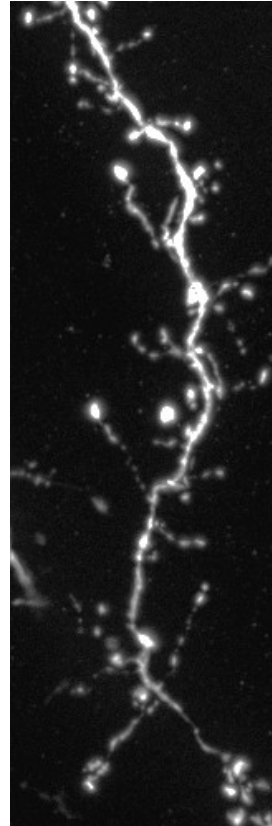
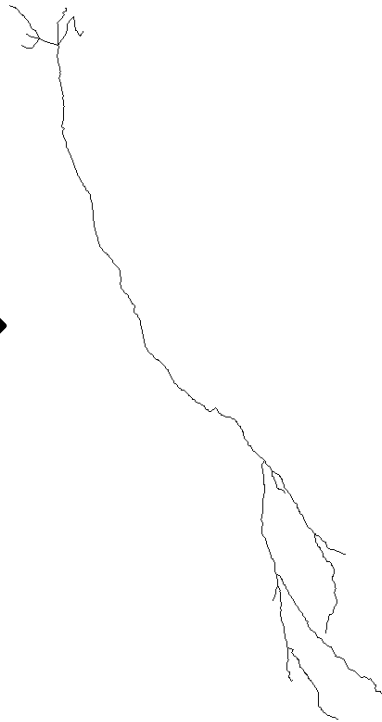
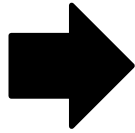
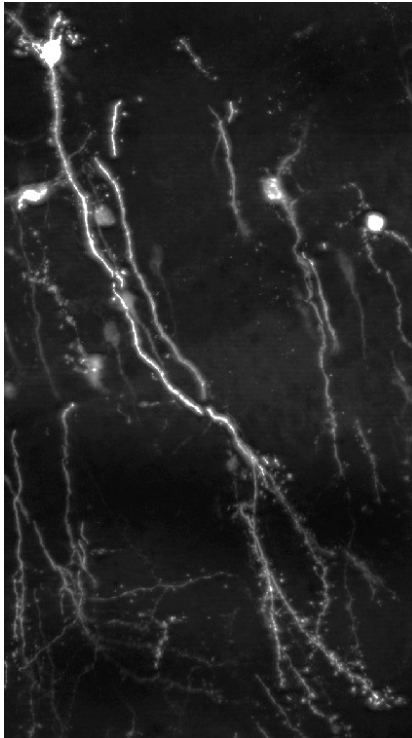


### 3 - Newborn neurons and preexisting neurons morphology:

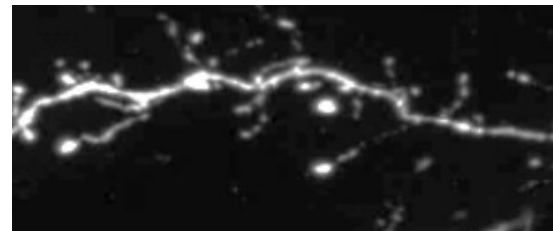
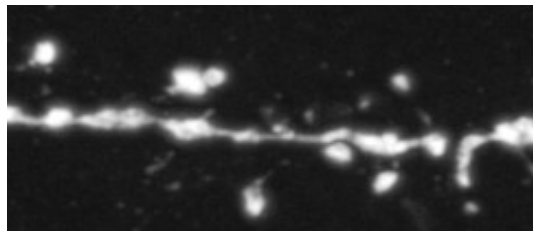
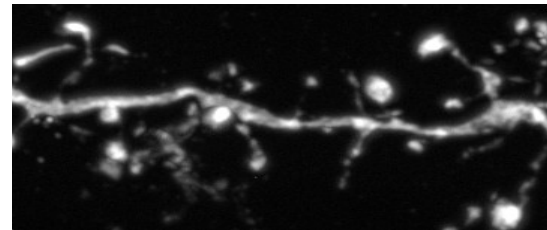
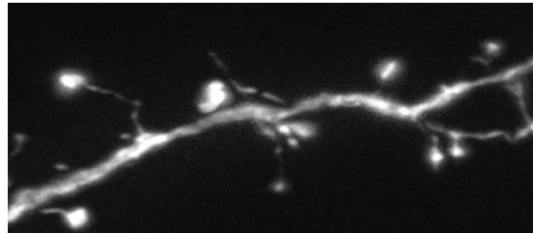
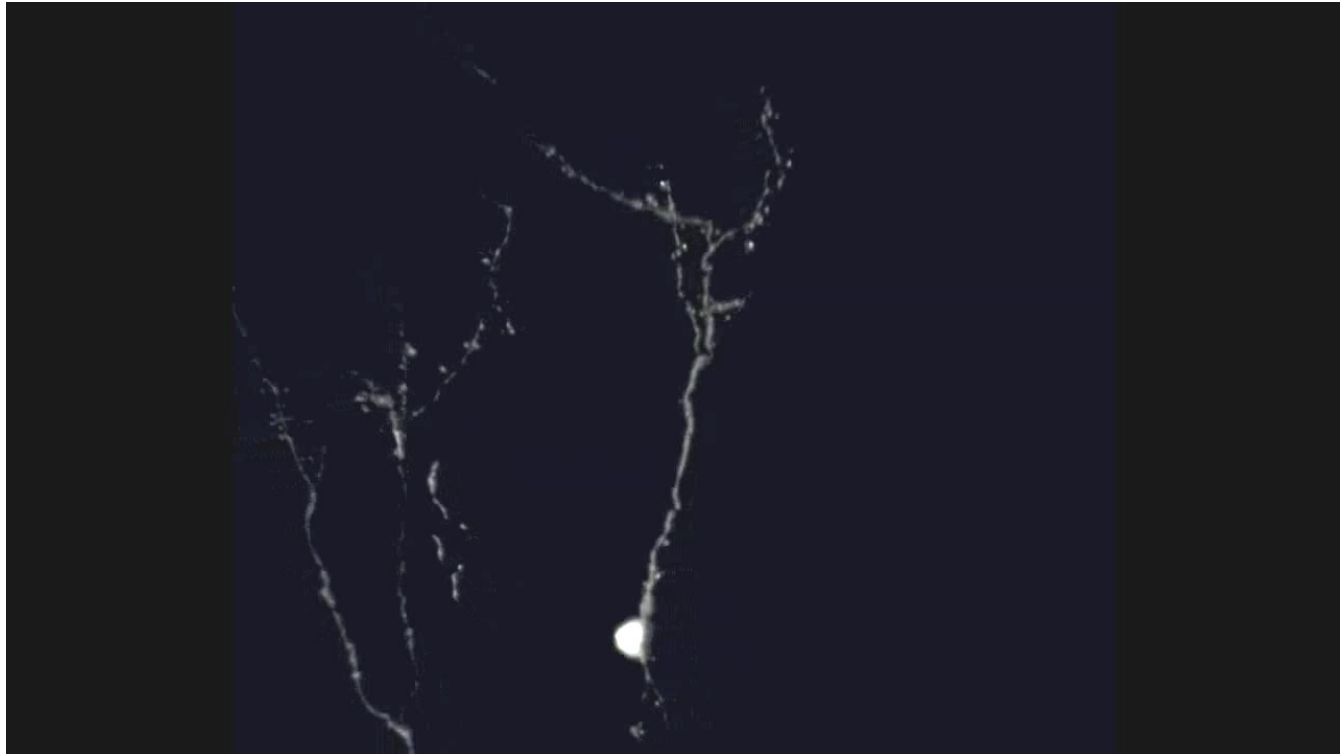


### 3 - Newborn neurons and preexisting neurons morphology:

NeuronStudio

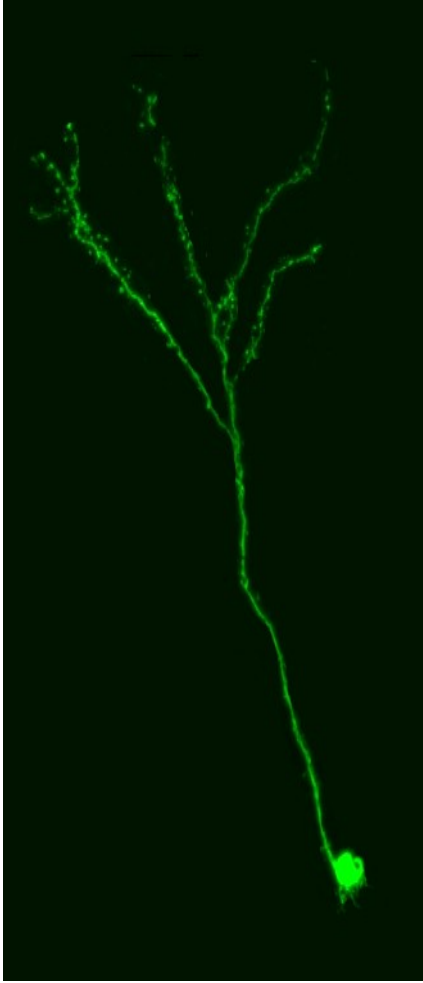


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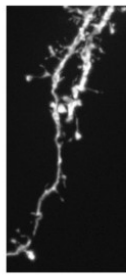
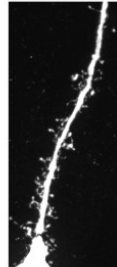
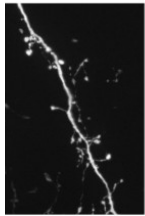
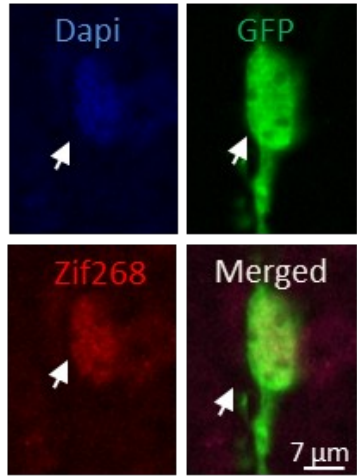




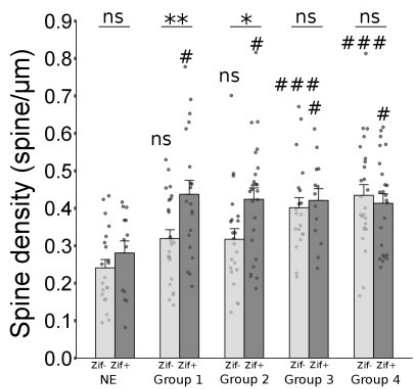
# 3a - Adult-born neurons



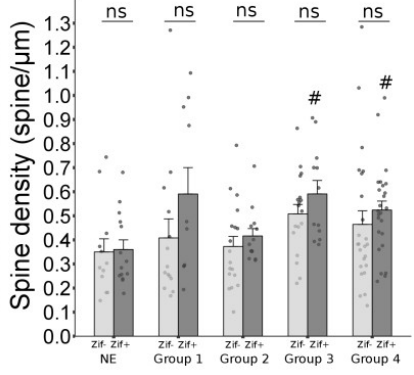
Zif+ neuron



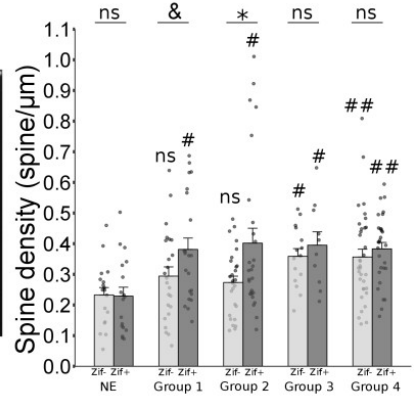
D



E

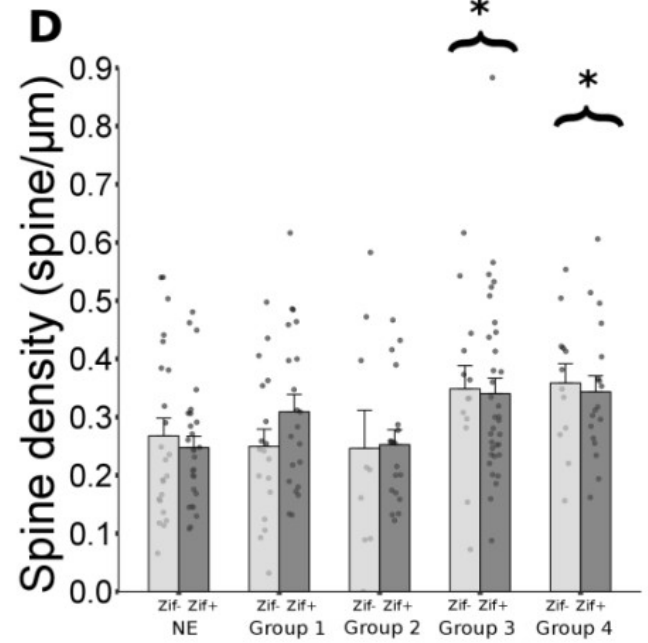
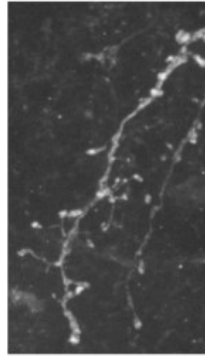
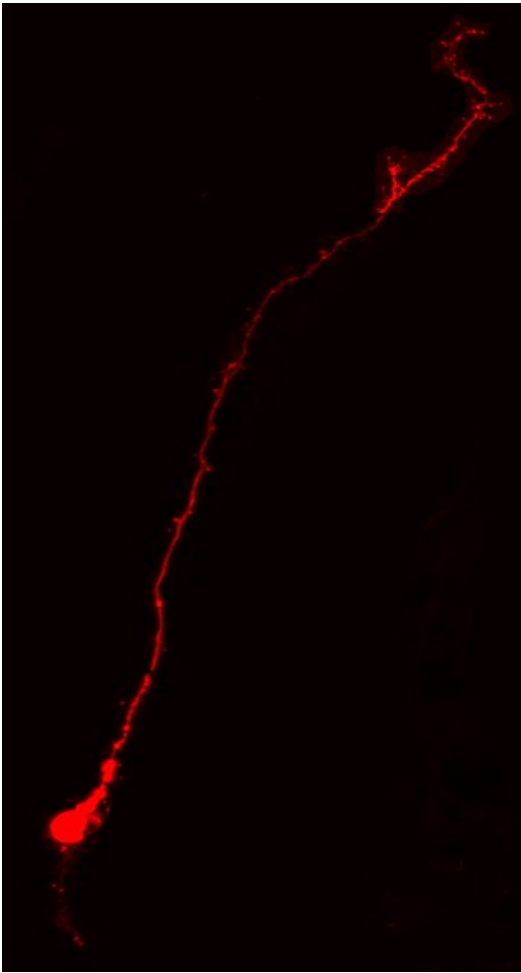


F



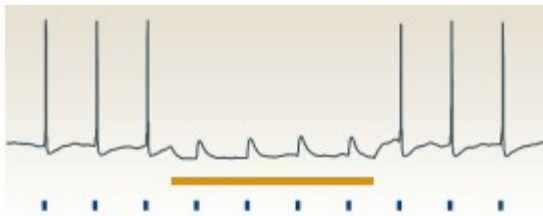
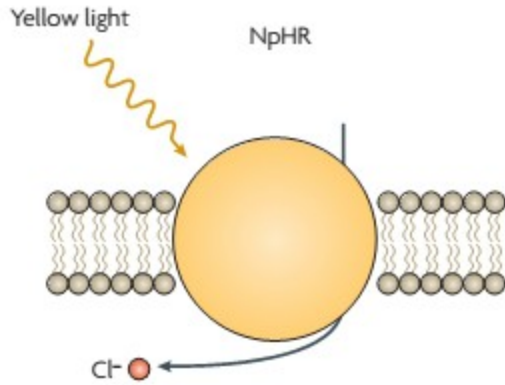
Increasing the complexity of perceptual learning induces a higher structural plasticity of adult-born neurons

### 3b - Preexisting neurons



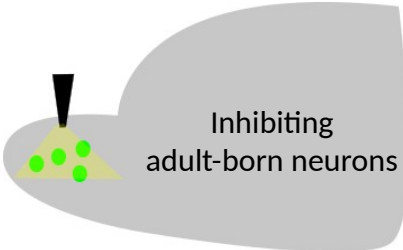
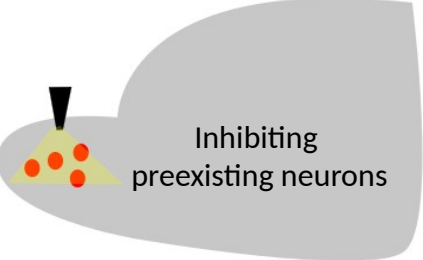


Perceptual learning induces more limited morphological changes in preexisting neurons

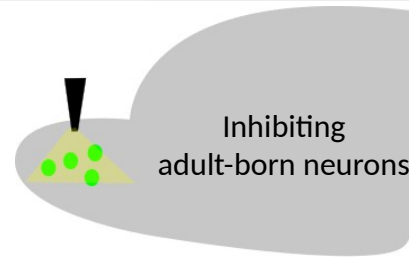
# 4 - Optogenetic



Adapted from Zhang et al. 2007, Nat Reviews Neuroscience

		
 <p>Inhibiting adult-born neurons</p>	<p>NO discrimination ?</p>	<p>NO discrimination ?</p>
 <p>Inhibiting preexisting neurons</p>	<p>Discrimination OK ?</p>	<p>NO discrimination ?</p>

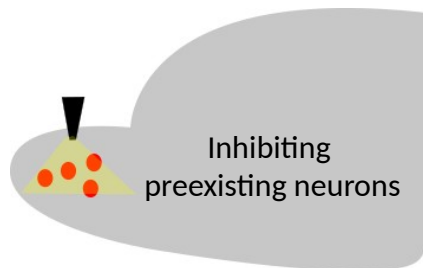
## 4 - Optogenetic



Discrimination



Discrimination



Discrimination



Discrimination



Optogenetically inhibiting either preexisting or adult-born neurons reveals their functionally distinct involvement in simple and complex perceptual learning.

# Conclusion

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- Increased survival of adult-born neurons independently of learning complexity

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- Increased survival of adult-born neurons independently of learning complexity
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- Increased spines density at the apical distal, apical proximal and basal domains of adult-born neurons both in simple and complex learning paradigms

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- Increase spines density at the apical domain of preexisting neurons in complex learning paradigm

And:

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And:

- Adult-born neurons are sufficient and necessary to underlie simple but not complex perceptual learning

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- Increased survival of adult-born neurons independently of learning complexity
- Increased recruitment of adult-born neurons to the processing of the learned odorants with increased complexity
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- Increase spines density at the apical domain of preexisting neurons in complex learning paradigm

And:

- Adult-born neurons are sufficient and necessary to underlie simple but not complex perceptual learning
- Preexisting neurons are necessary for complex but not for simple perceptual learning

## Take home message:

**Adult neurogenesis exhibits limits in its adaptive abilities to answer complex behavioral demands but at the same time can be both necessary and sufficient for simple learnings.**

# AKNOWLEDGEMENTS

## Neuropop Team

