

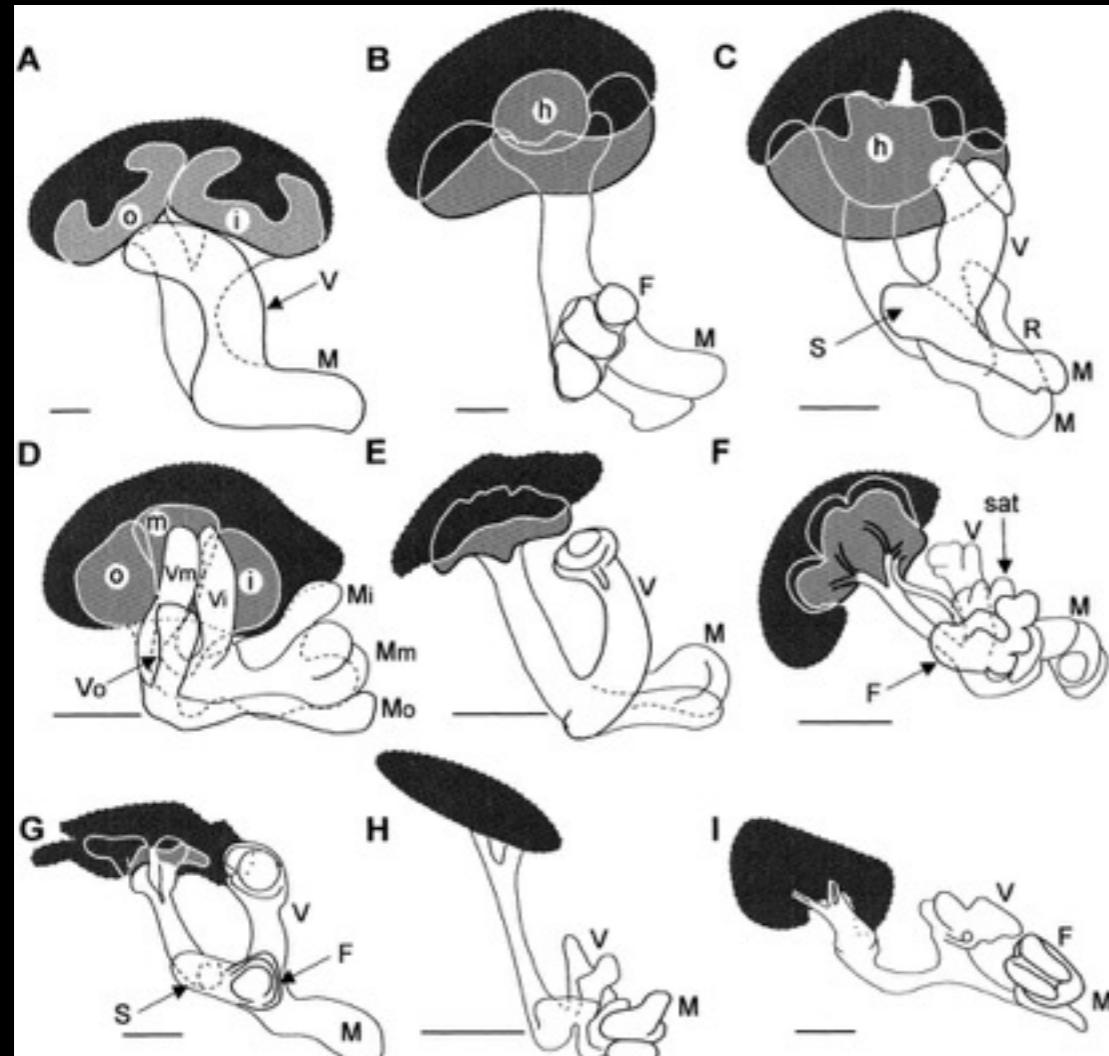


Jürgen Berger

Courtesy of Jurgen Berger. Used with permission.

# Mushroom body endows organisms with a degree of free will or intelligent control over instinctive actions

- Dujardin, 1850



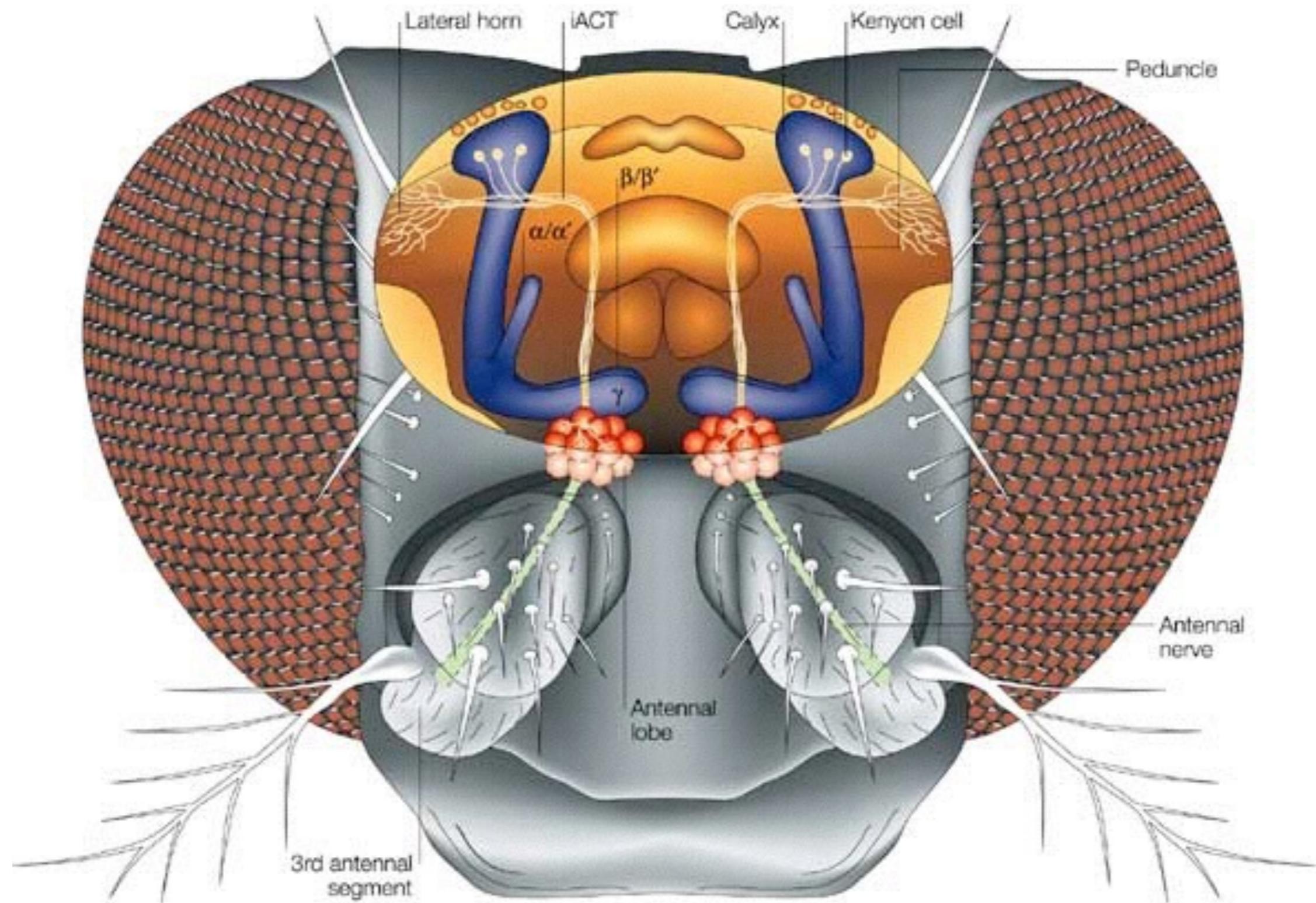
(Strausfeld et al., 1998)

Courtesy of Cold Spring Harbor Laboratory Press. Used with permission.

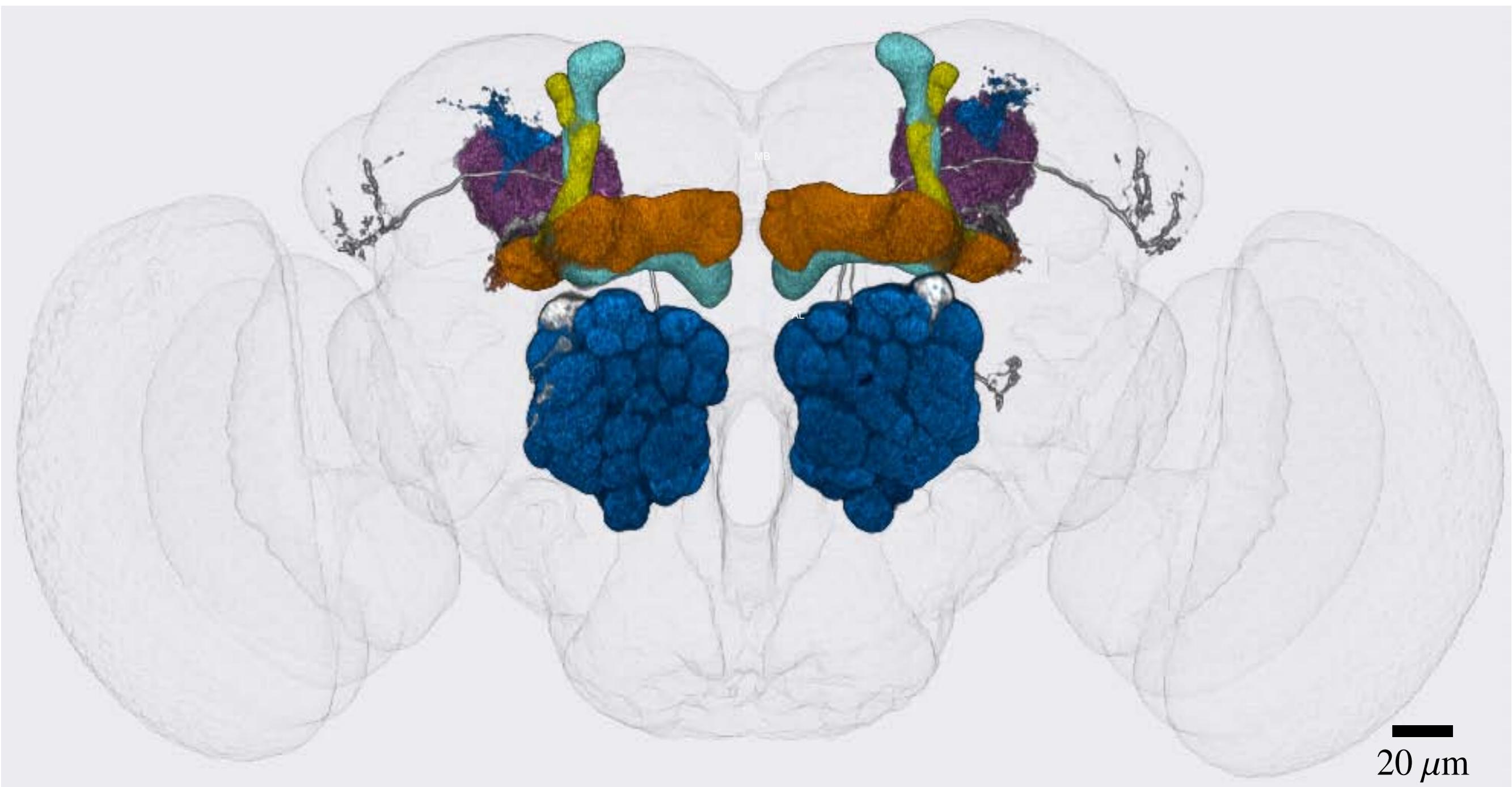


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Photos removed due to copyright restrictions. Please see the video.

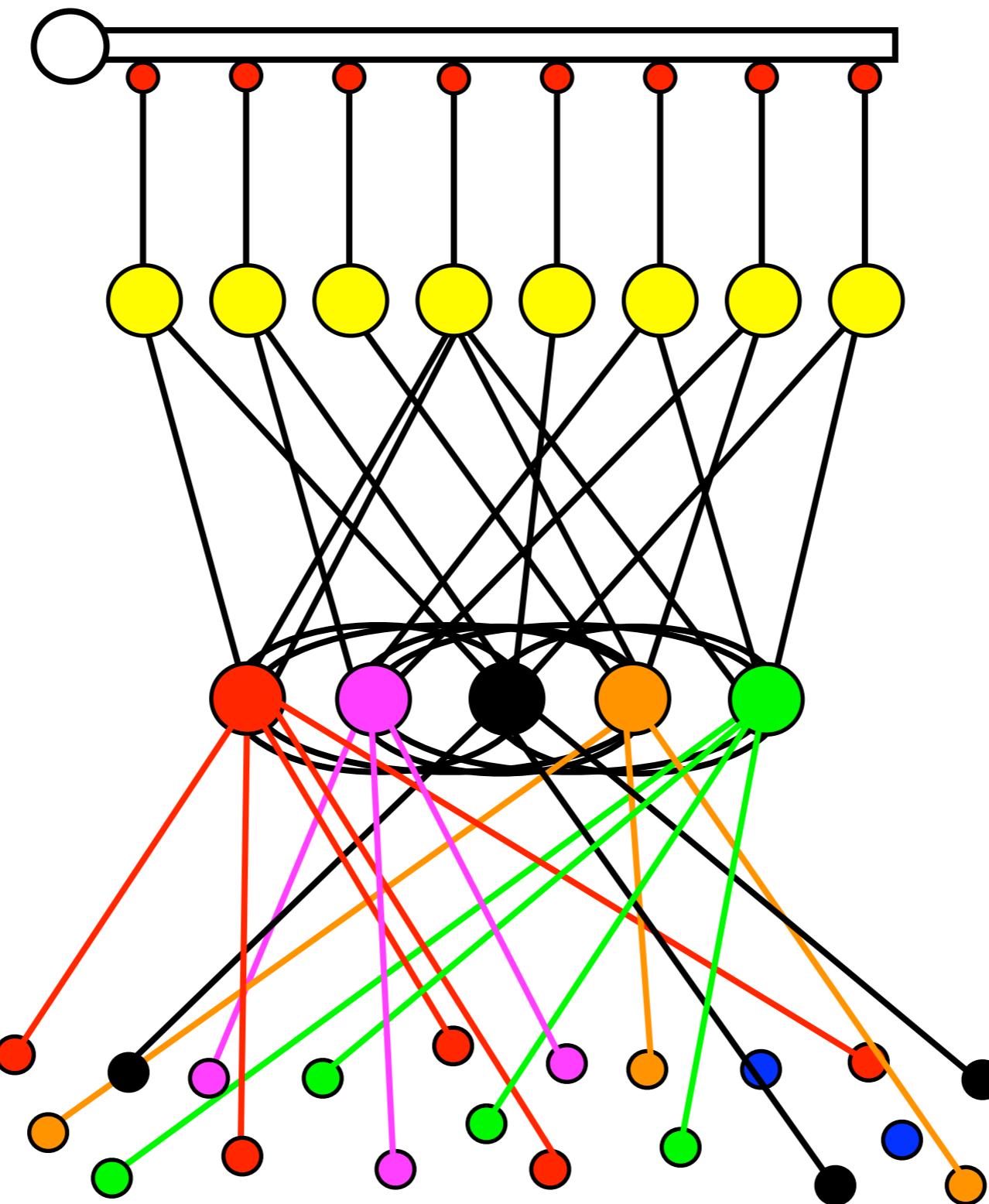


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Source: Heisenberg, Martin. "Mushroom body memoir: From maps to models."  
Nature Reviews Neuroscience 4, no. 4 (2003): 266-275. ©2003.



20  $\mu$ m

Courtesy of eLife. Used with permission.



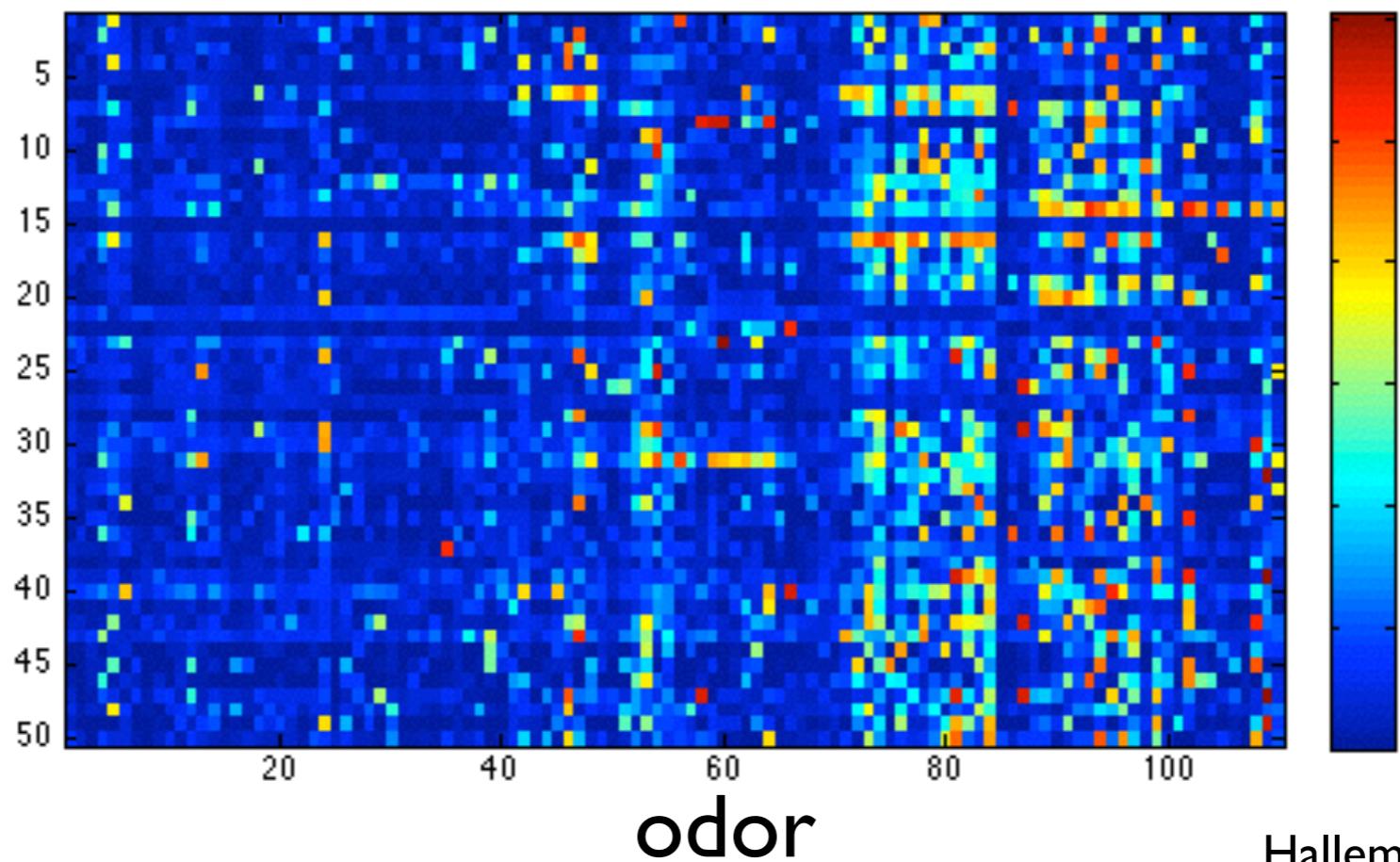
PN  
 $N \sim 200/50$   
 $D \sim 35$

ORN  
 $N \sim 1000/50$   
 $D \sim 30$

pooling  
noise  
reduction  
normalization  
input

**D = 30**

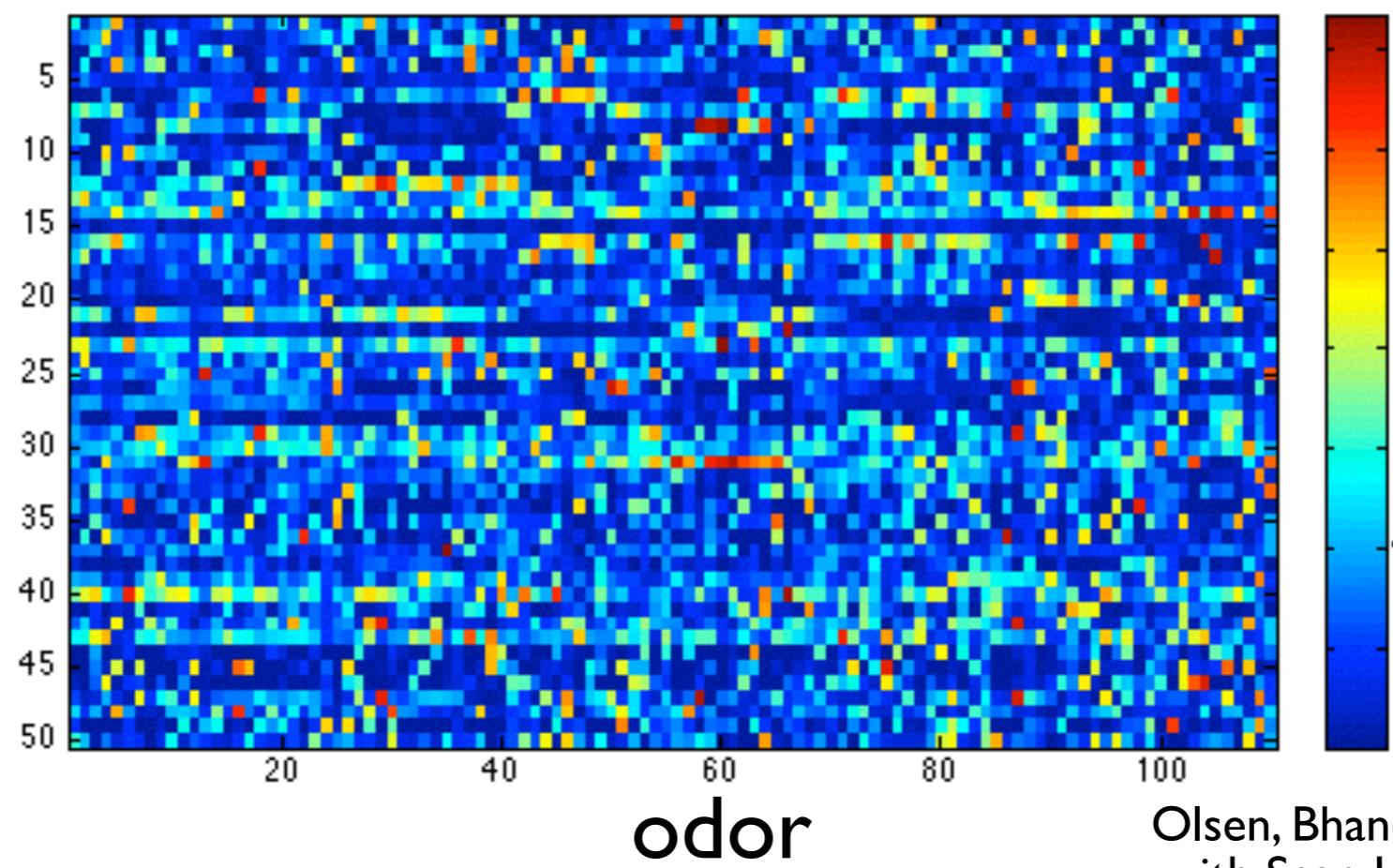
**ORN**



Hallem & Carlson 2006

**D = 35**

**PN**



Olsen, Bhandawat, Wilson 2010  
with Sean Luo, Richard Axel

Courtesy of PNAS. Used with permission.

**MBON**

$N \sim 34/21$

$D \sim 20$

**KC**

$N \sim 2000/7$

$D \sim 1000$

**PN**

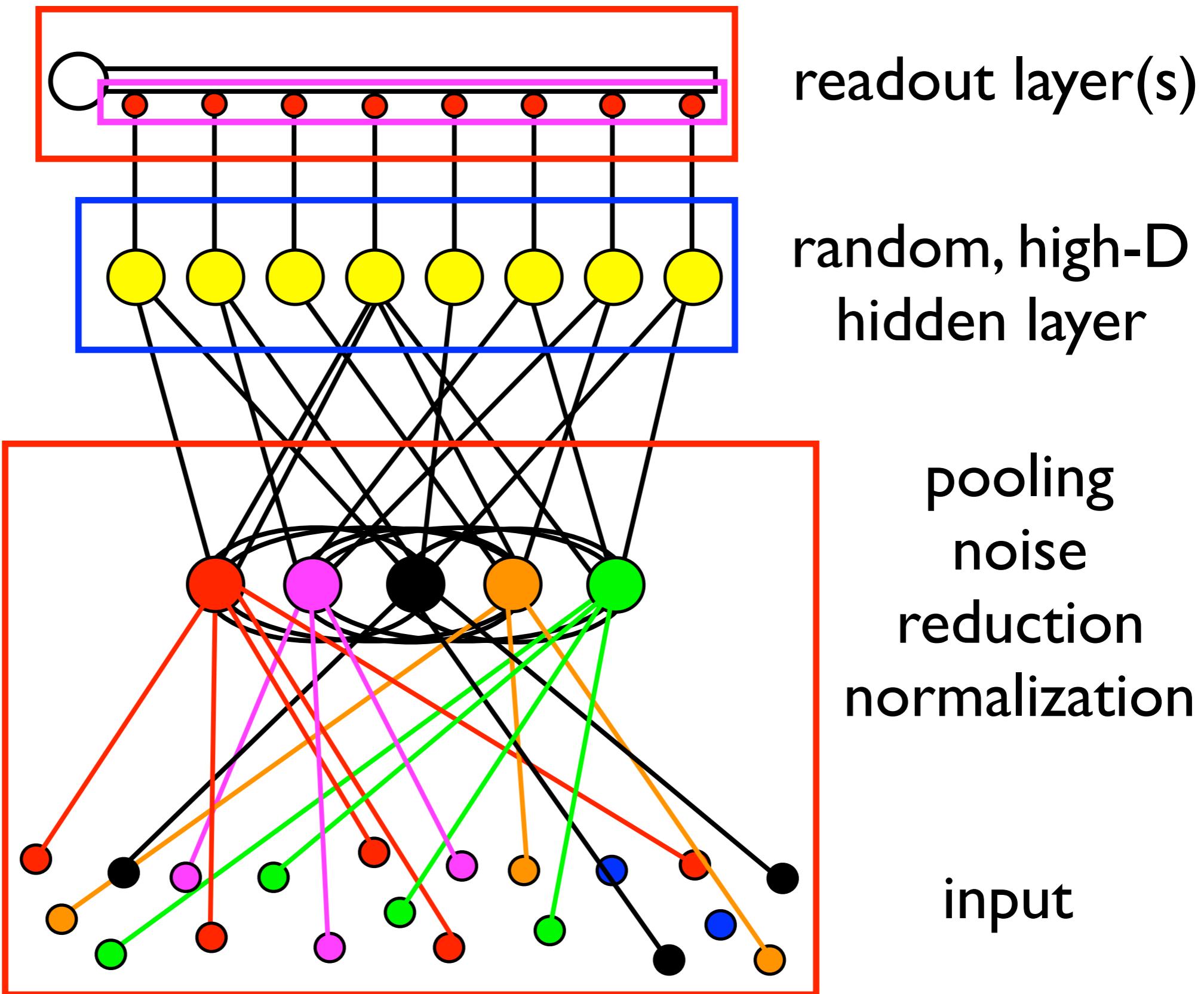
$N \sim 200/50$

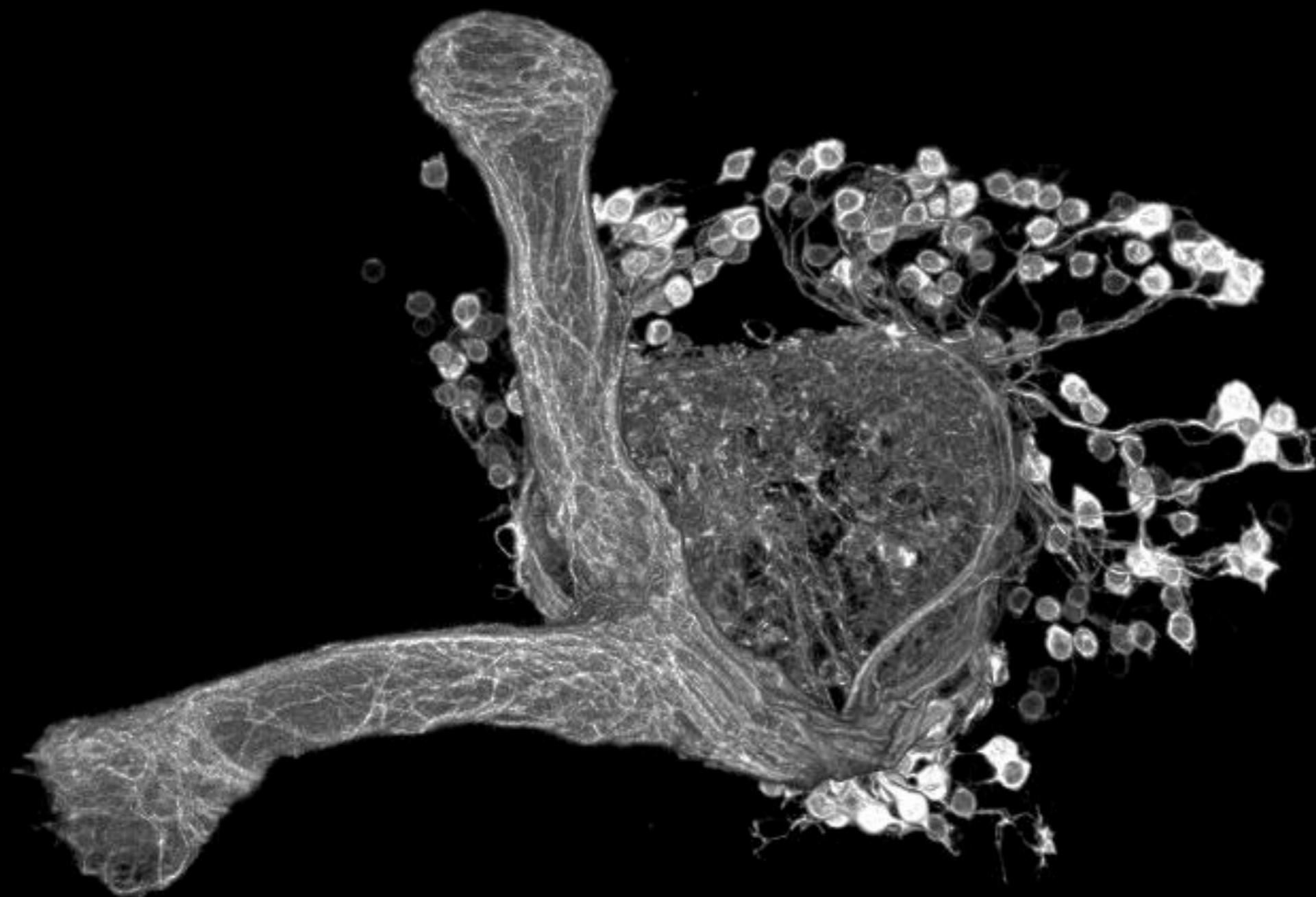
$D \sim 35$

**ORN**

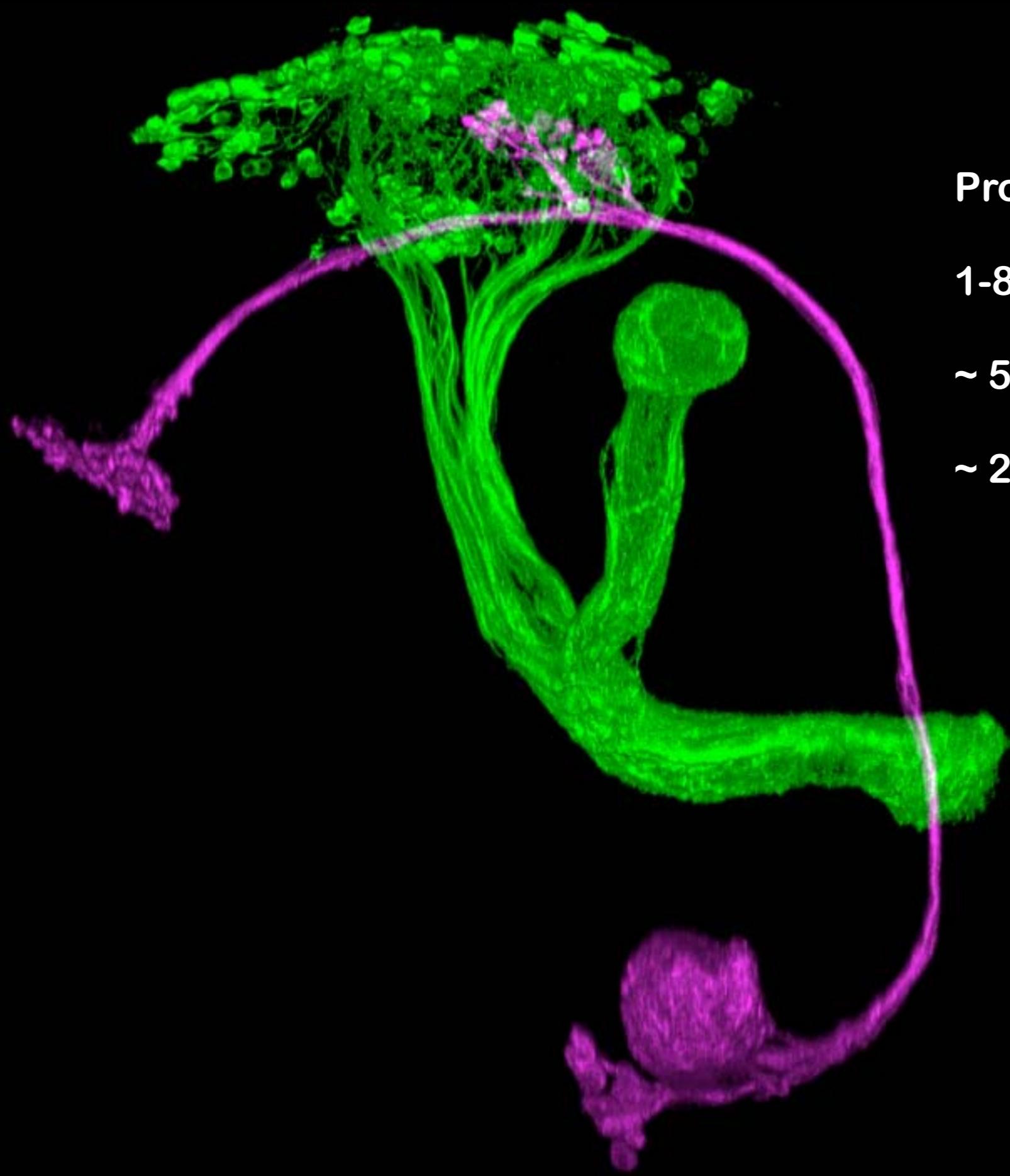
$N \sim 1000/50$

$D \sim 30$





Courtesy of Yoshi Aso. Used with permission.



**Projection neurons**

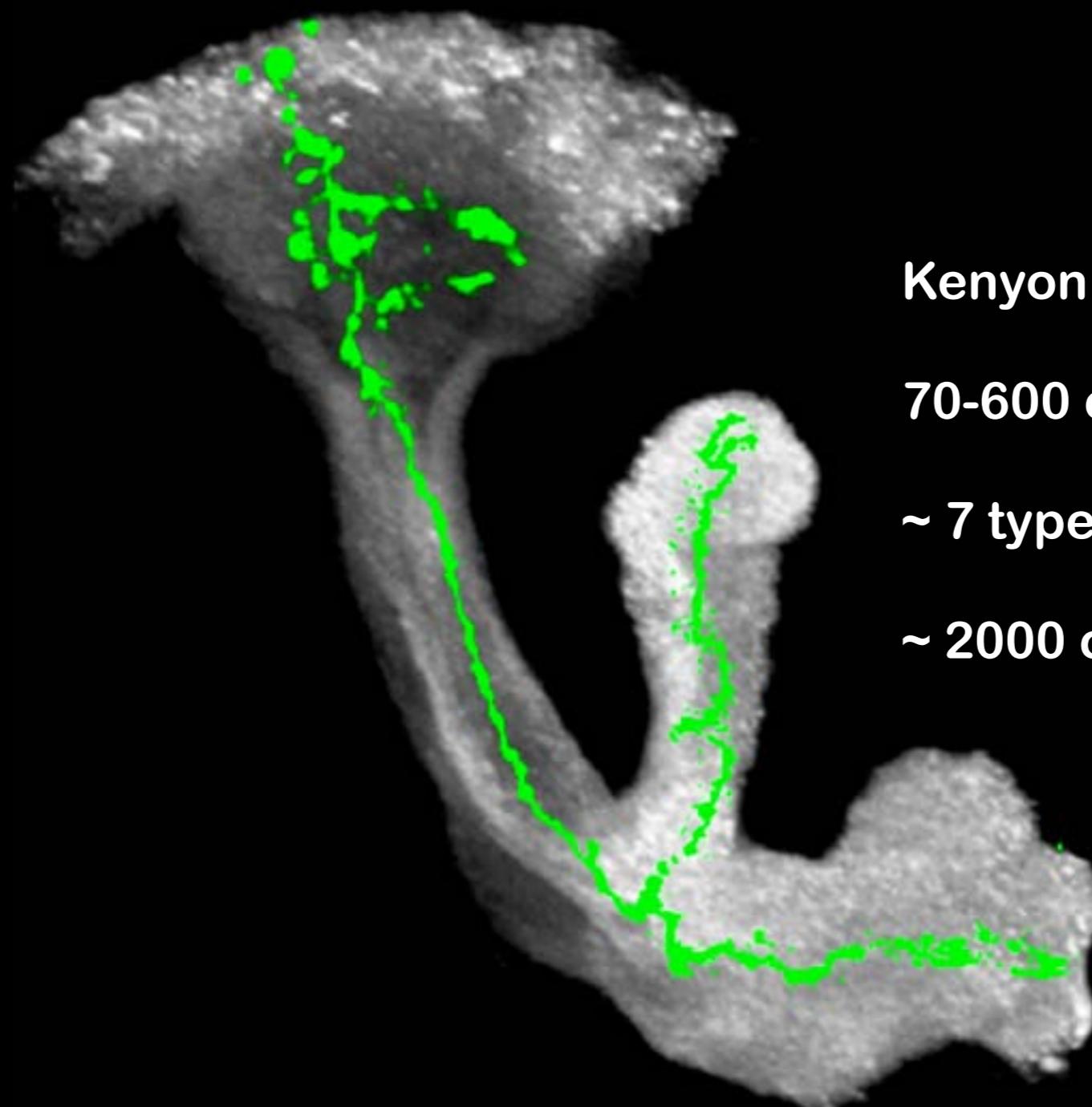
**1-8 cells/type**

**~ 50 types**

**~ 200 cells**

**Yoshi Aso, Daisuke Hattori**

Courtesy of Yoshi Aso. Used with permission.



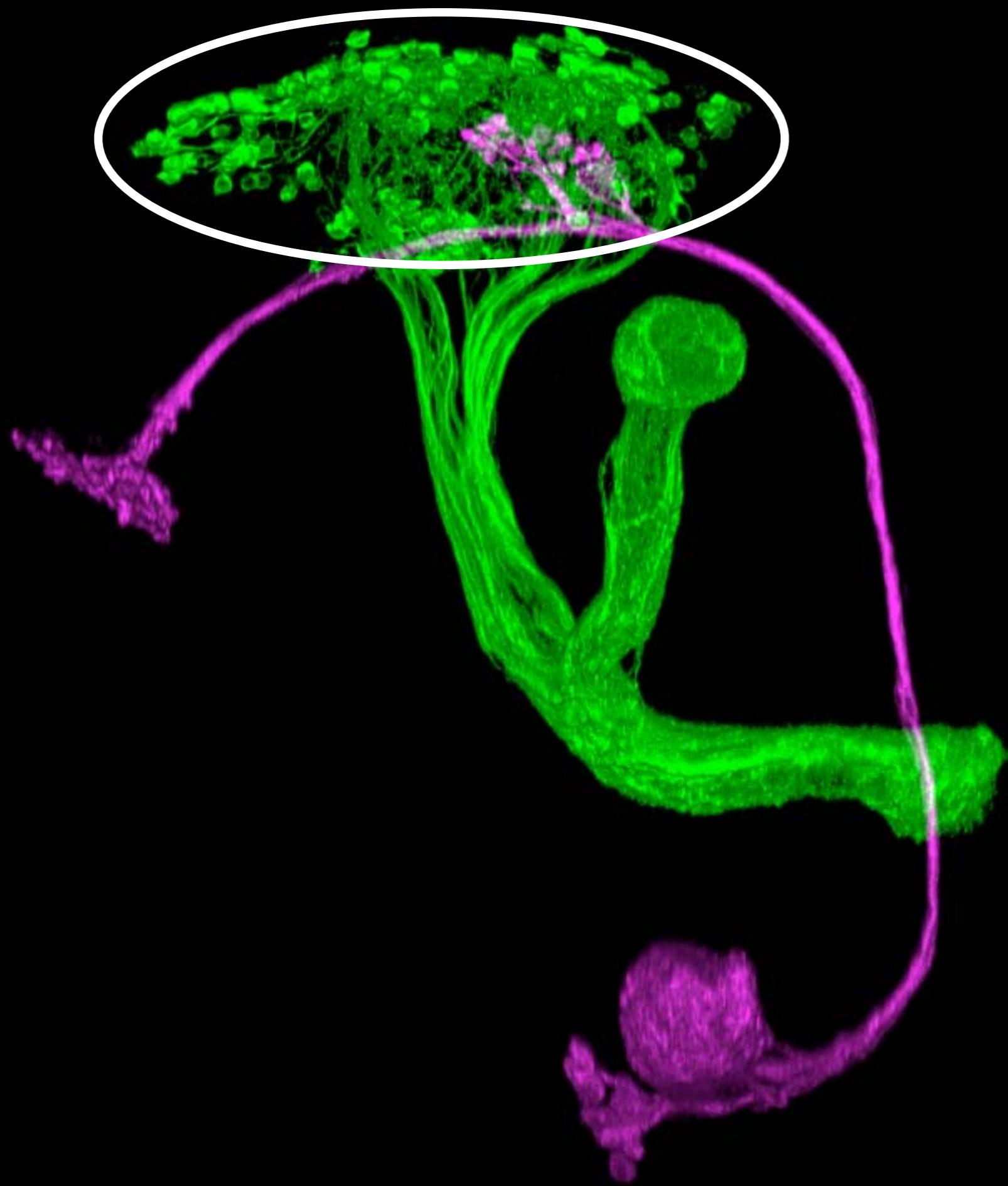
**Kenyon cells**

**70-600 cells/type**

**~ 7 types**

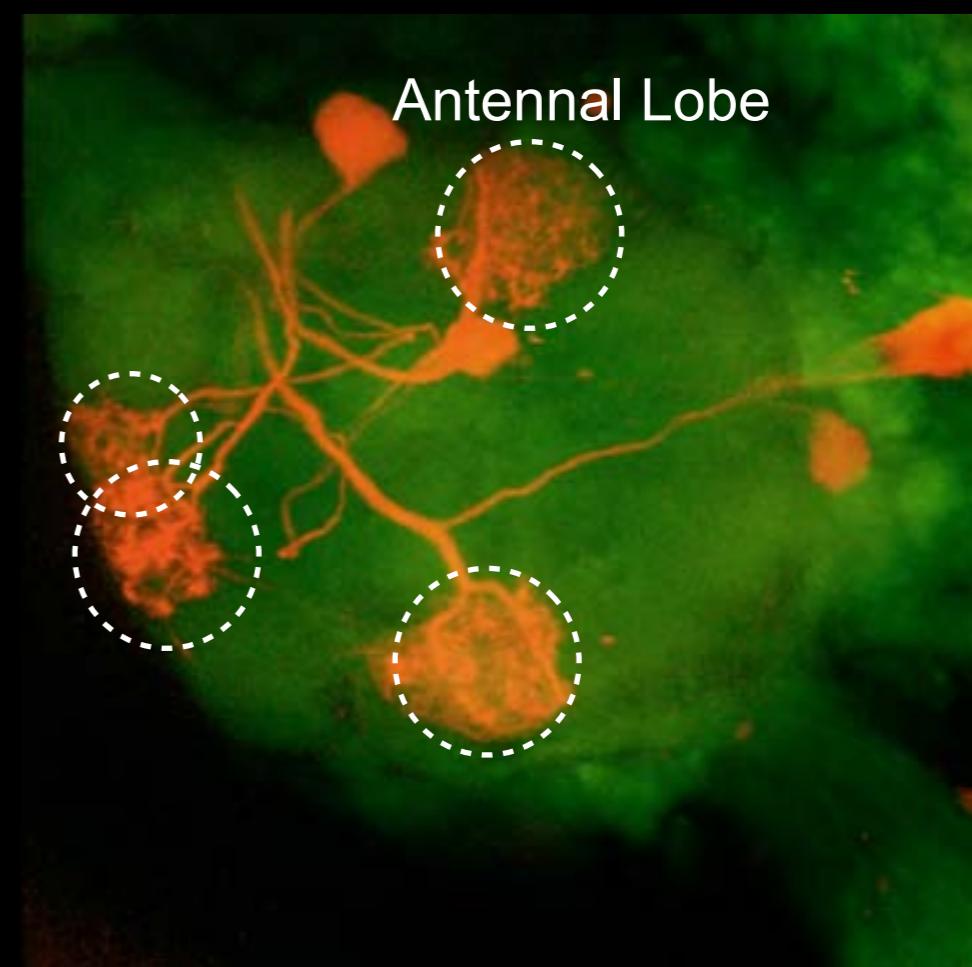
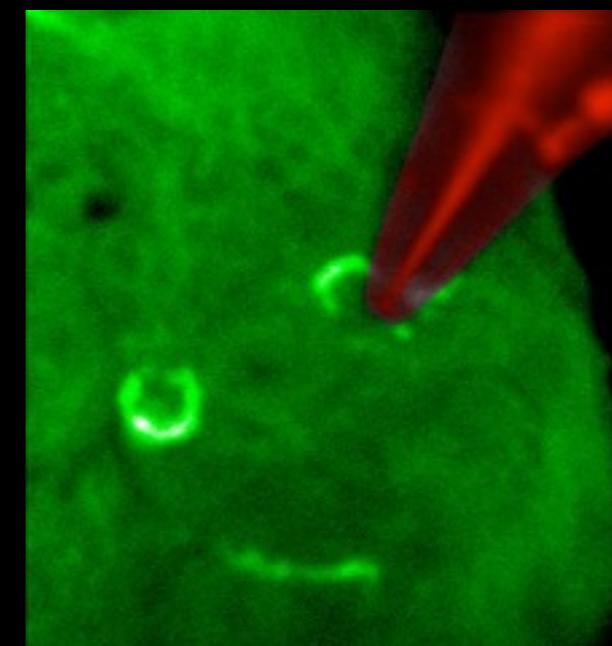
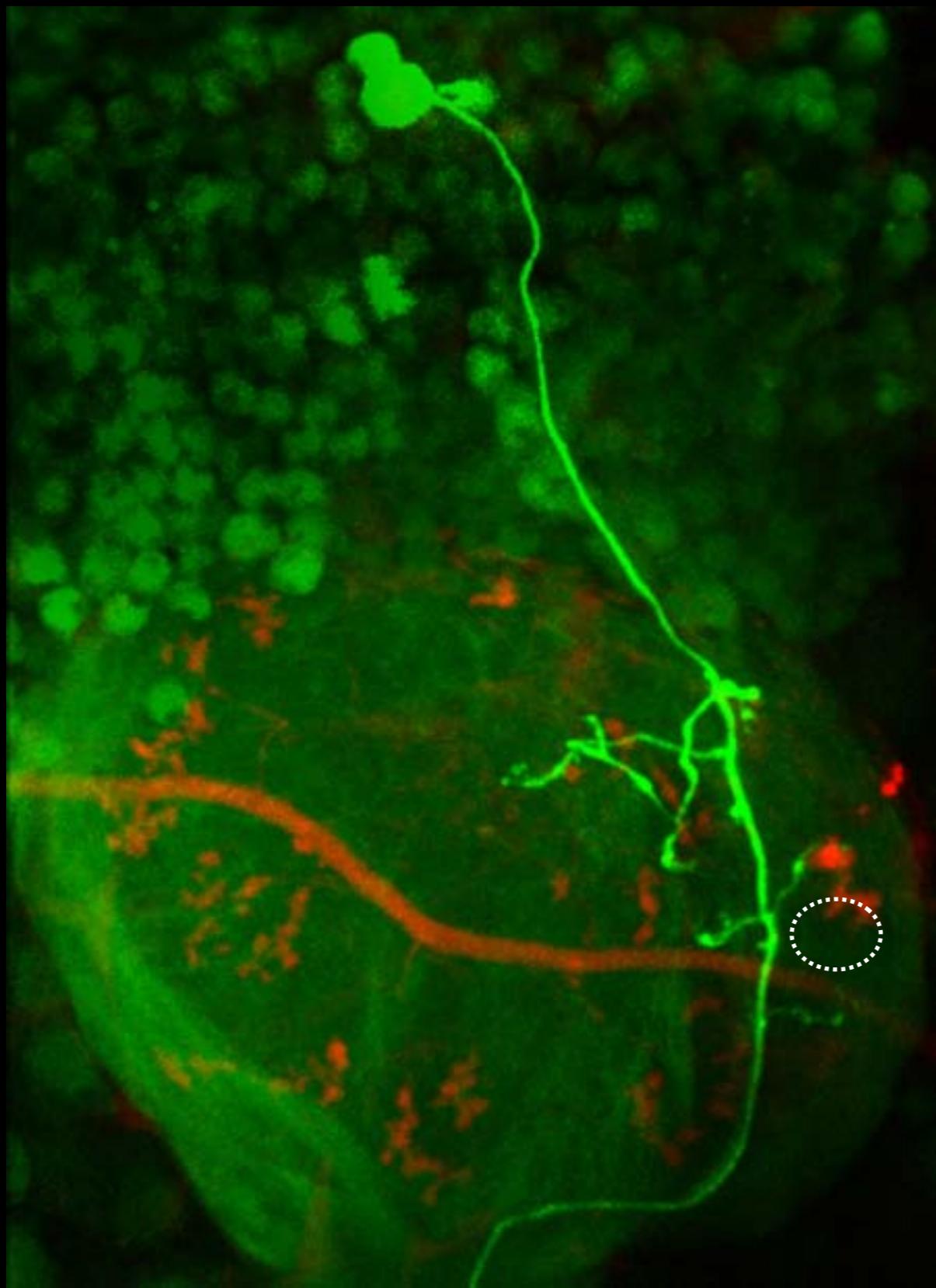
**~ 2000 cells**

Courtesy of Yoshi Aso. Used with permission.



Yoshi Aso, Daisuke Hattori

Courtesy of Yoshi Aso. Used with permission.

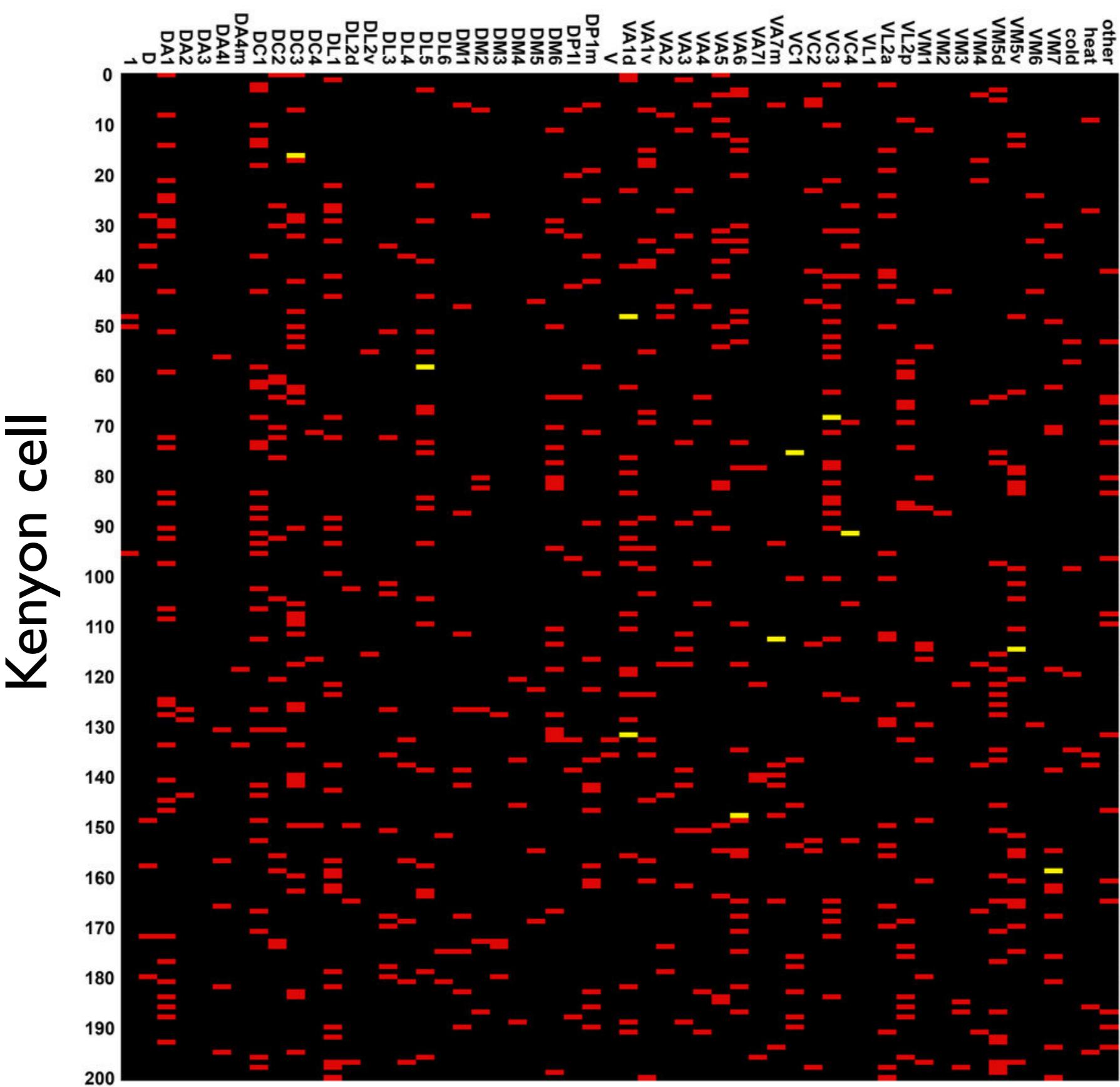


Reprinted by permission from Macmillan Publishers Ltd: Nature.

Source: Caron, Sophie JC, Vanessa Ruta, L. F. Abbott, and Richard Axel. "Random convergence of olfactory inputs in the Drosophila mushroom body." *Nature* 497, no. 7447 (2013): 113-117. © 2013.

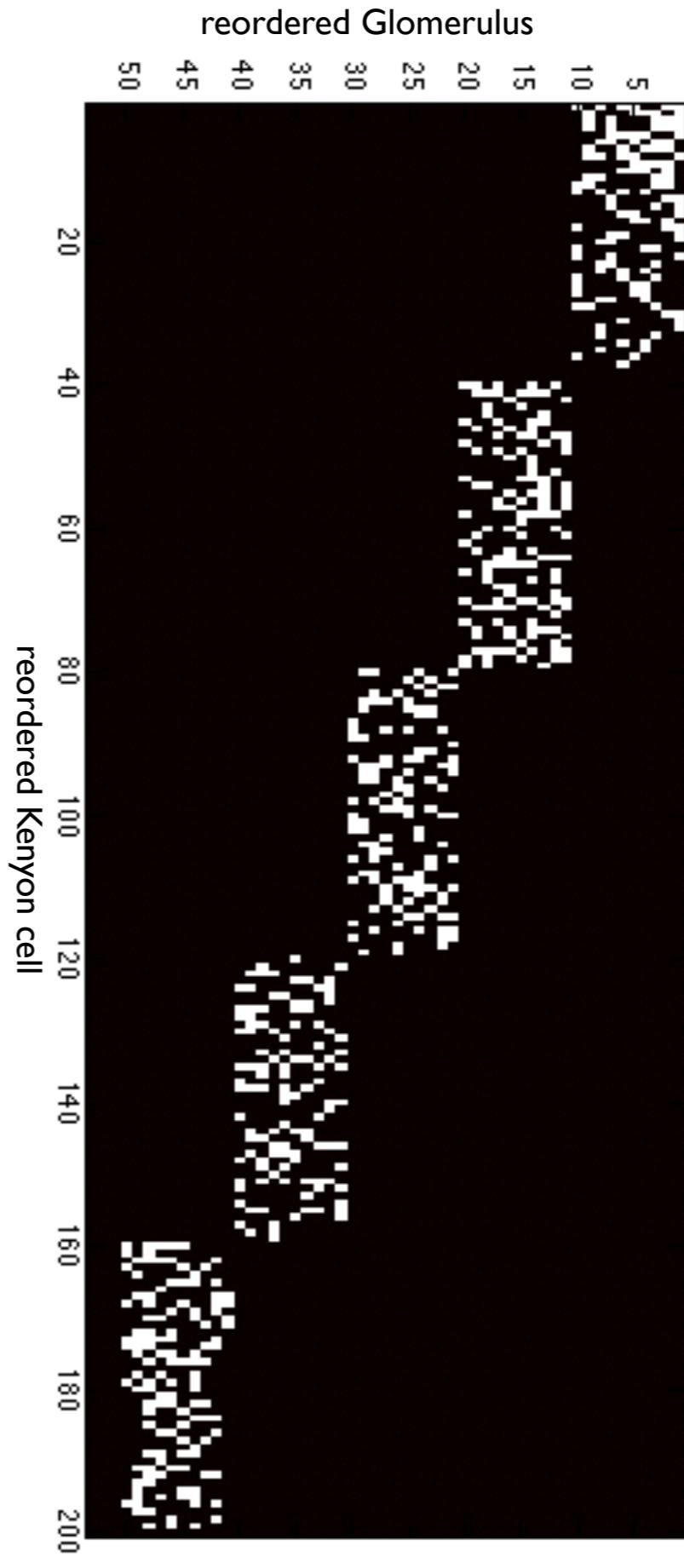
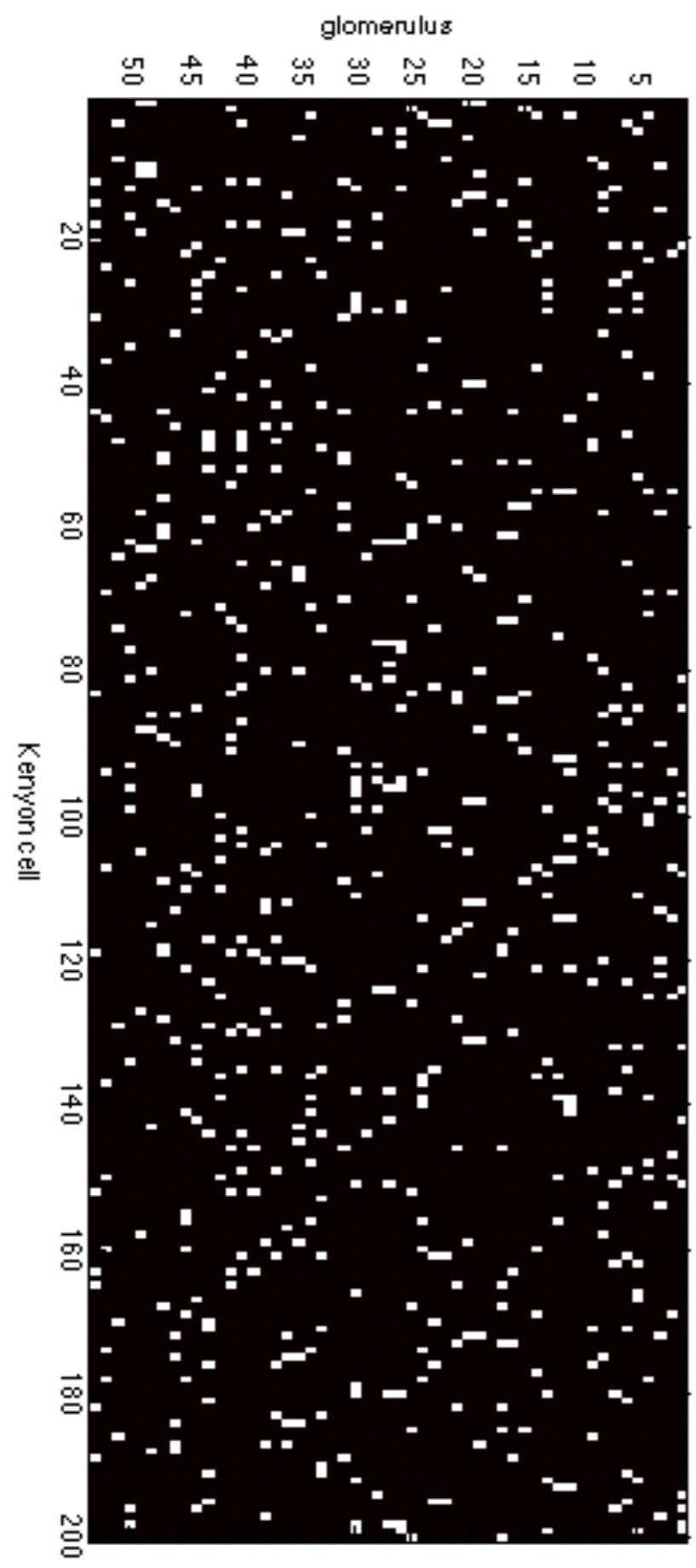
Caron, Ruta, Abbott, Richard Axel, 2013

# glomerulus

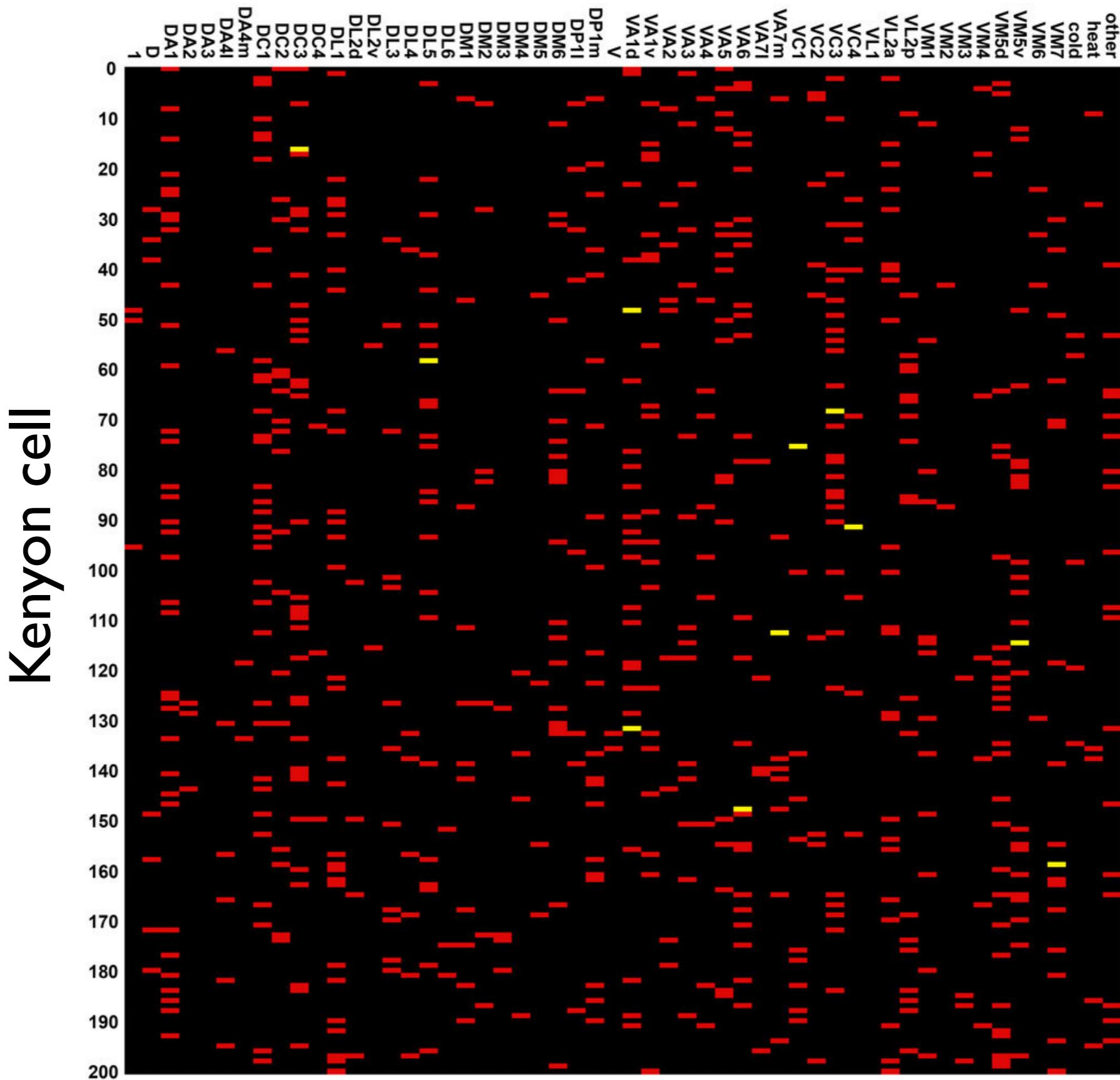


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Source: Caron, Sophie JC, Vanessa Ruta, L. F. Abbott, and Richard Axel. "Random convergence of olfactory inputs in the Drosophila mushroom body." Nature 497, no. 7447 (2013): 113-117. © 2013.



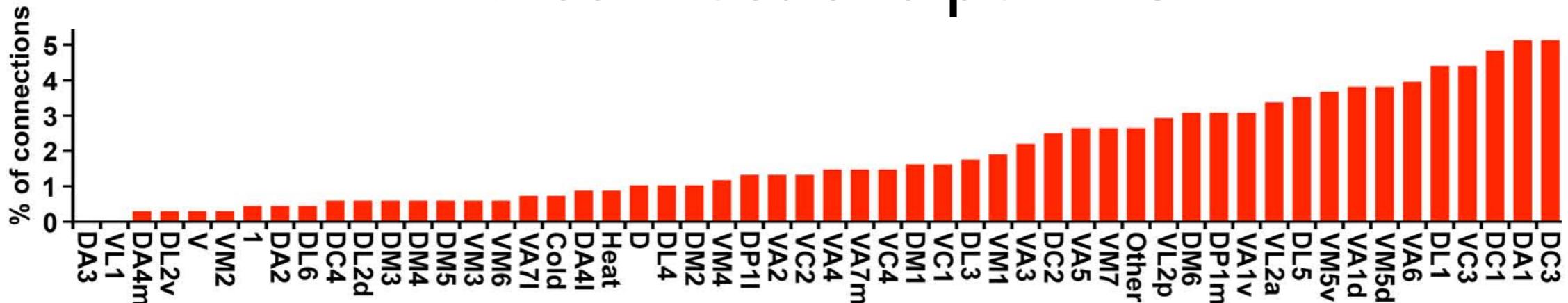
# glomerulus



Reprinted by permission from Macmillan Publishers Ltd: Nature.

Source: Caron, Sophie JC, Vanessa Ruta, L. F. Abbott, and Richard Axel. "Random convergence of olfactory inputs in the Drosophila mushroom body." *Nature* 497, no. 7447 (2013): 113-117. © 2013.

~7 connections per KC



Further structure?

No

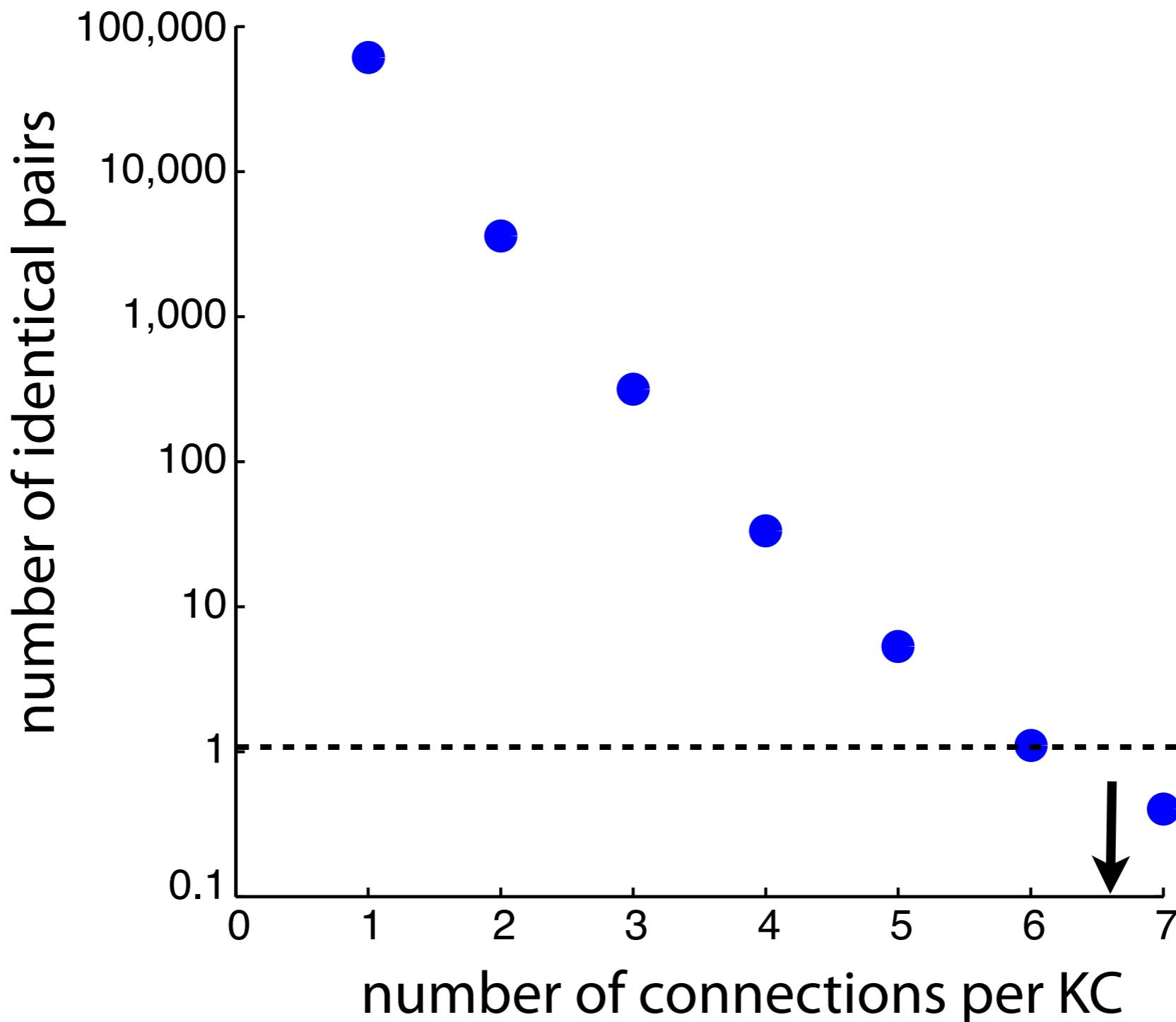
all tests consistent with random draws  
from this glomerular distribution

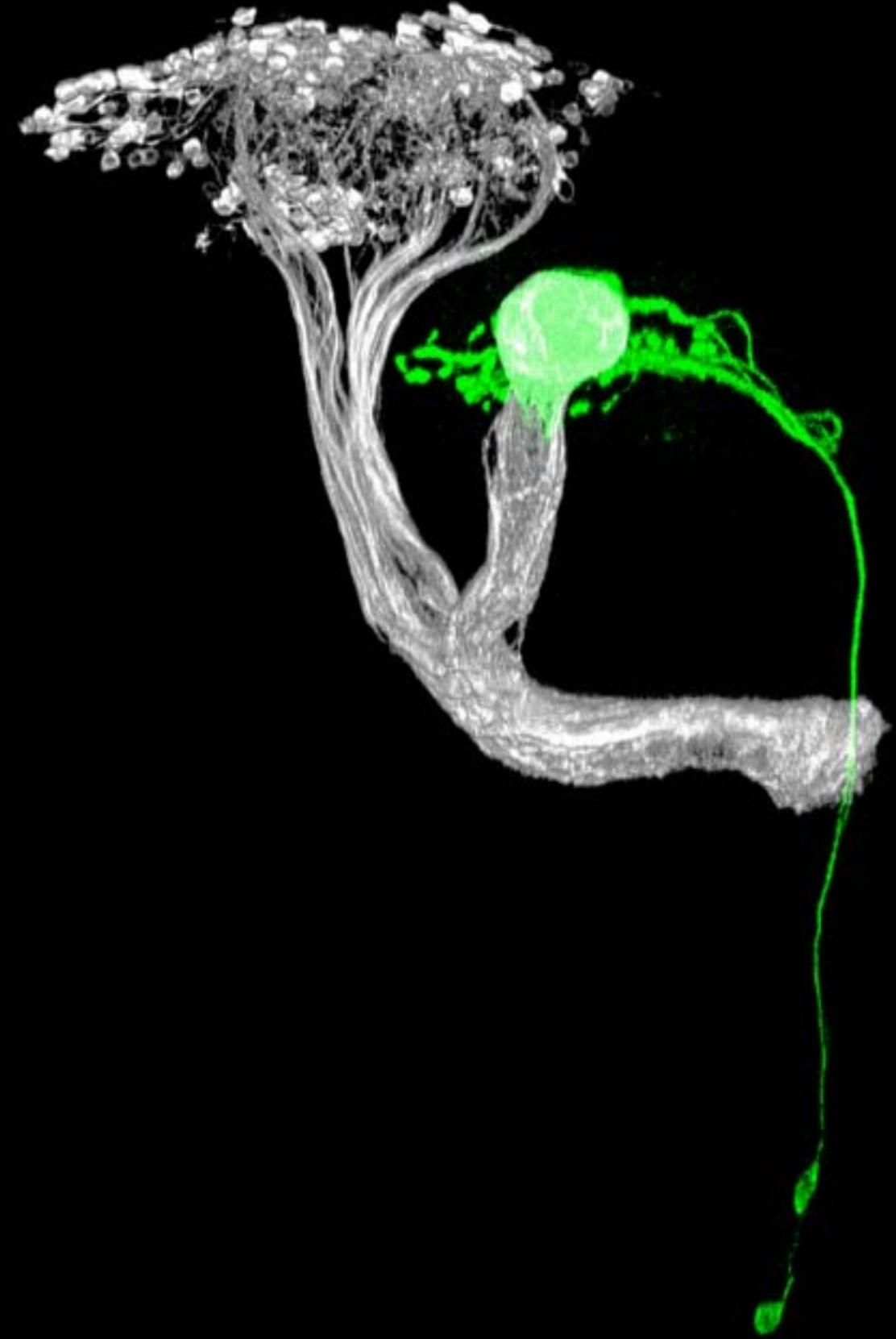
Murthy, Fiete, Laurent 2008

Caron, Ruta, Abbott, Richard Axel 2013

Gruntman, Turner 2013

# Why 7 connections?





**MB-Output neurons**

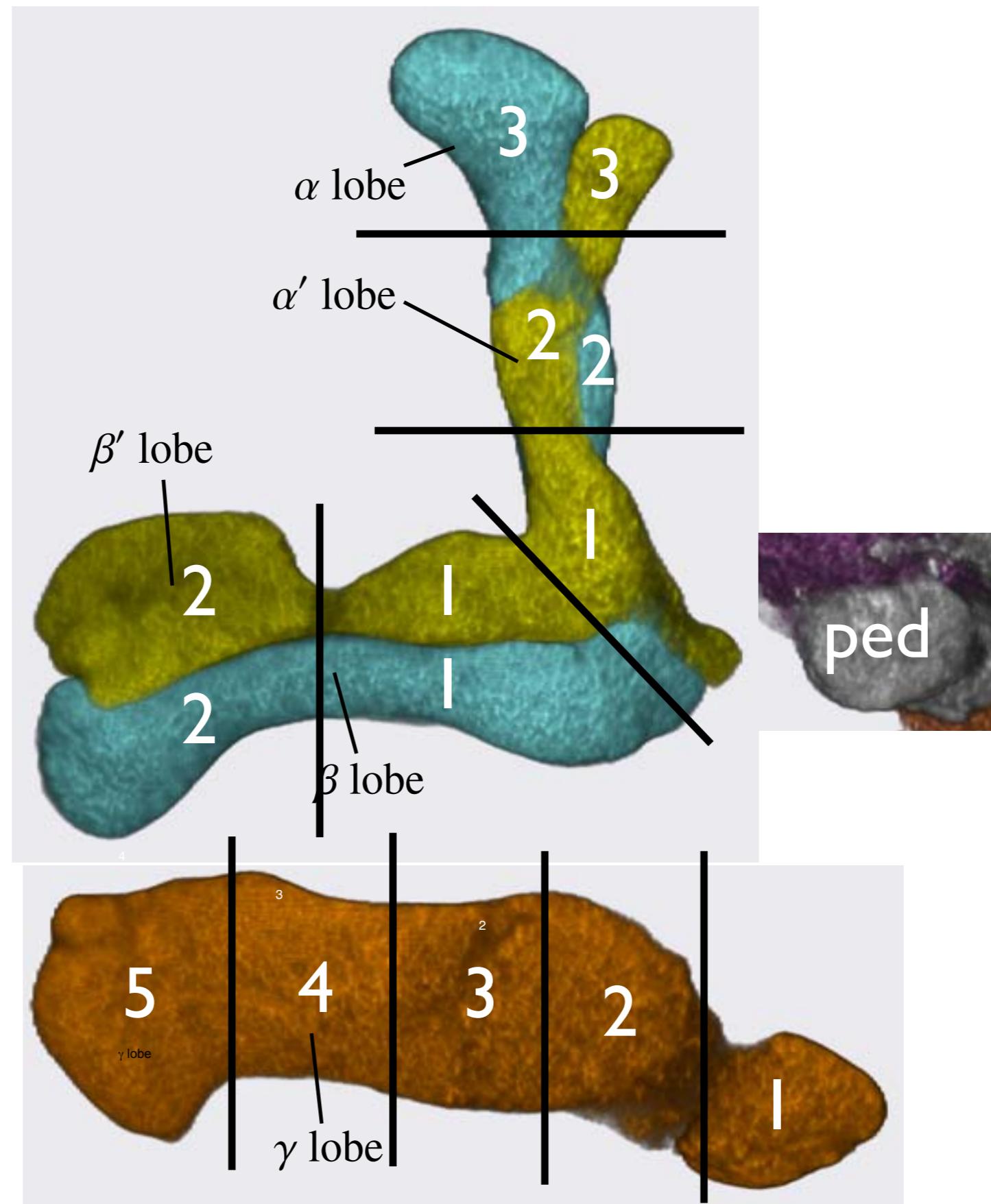
**1-7 cells/type**

**21 types**

**34 cells**

Courtesy of Yoshi Aso. Used with permission.

Yoshi Aso, Daisuke Hattori

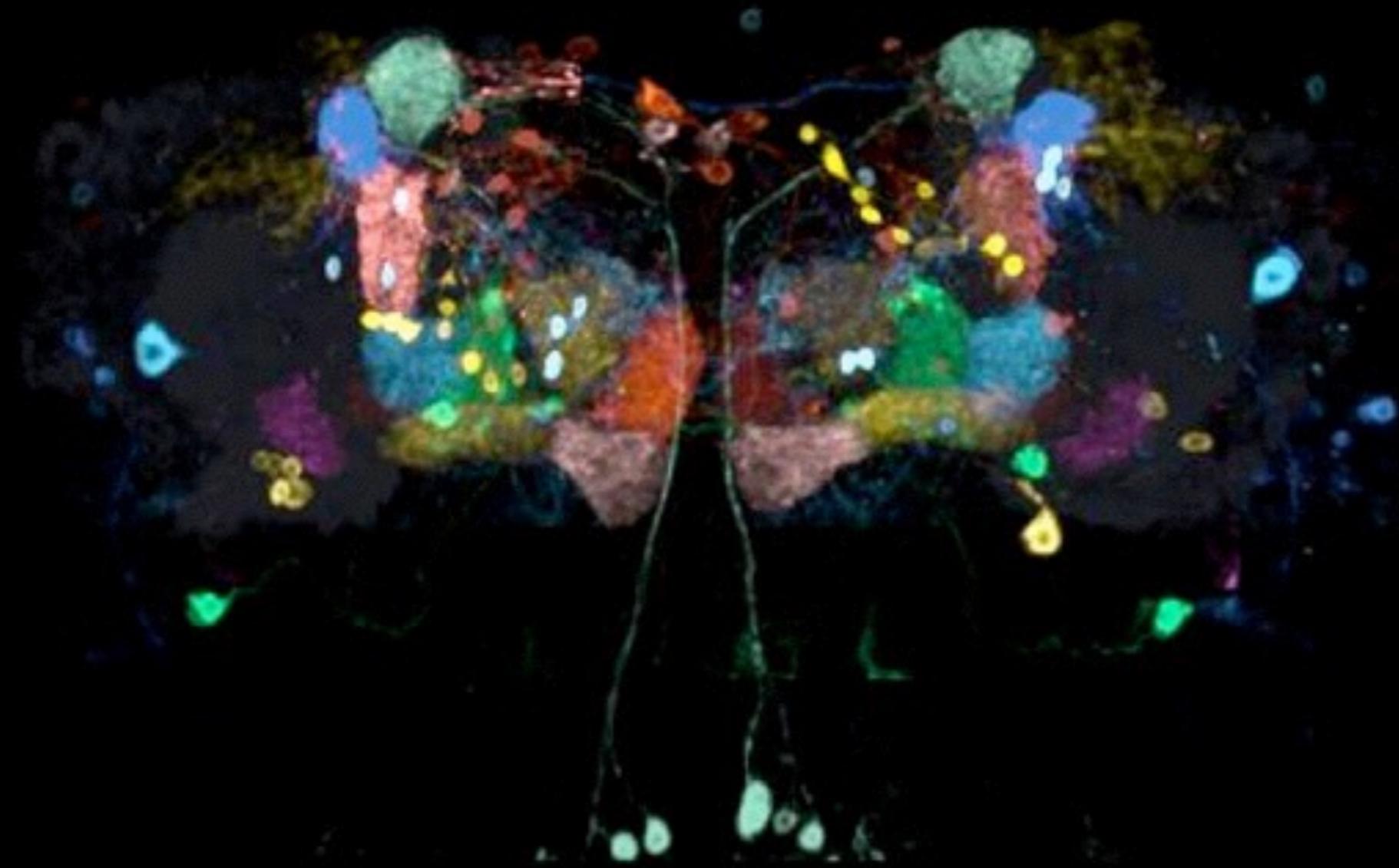


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Tanaka, Tanimoto, Ito 2008

# Mushroom Body Extrinsic Neurons

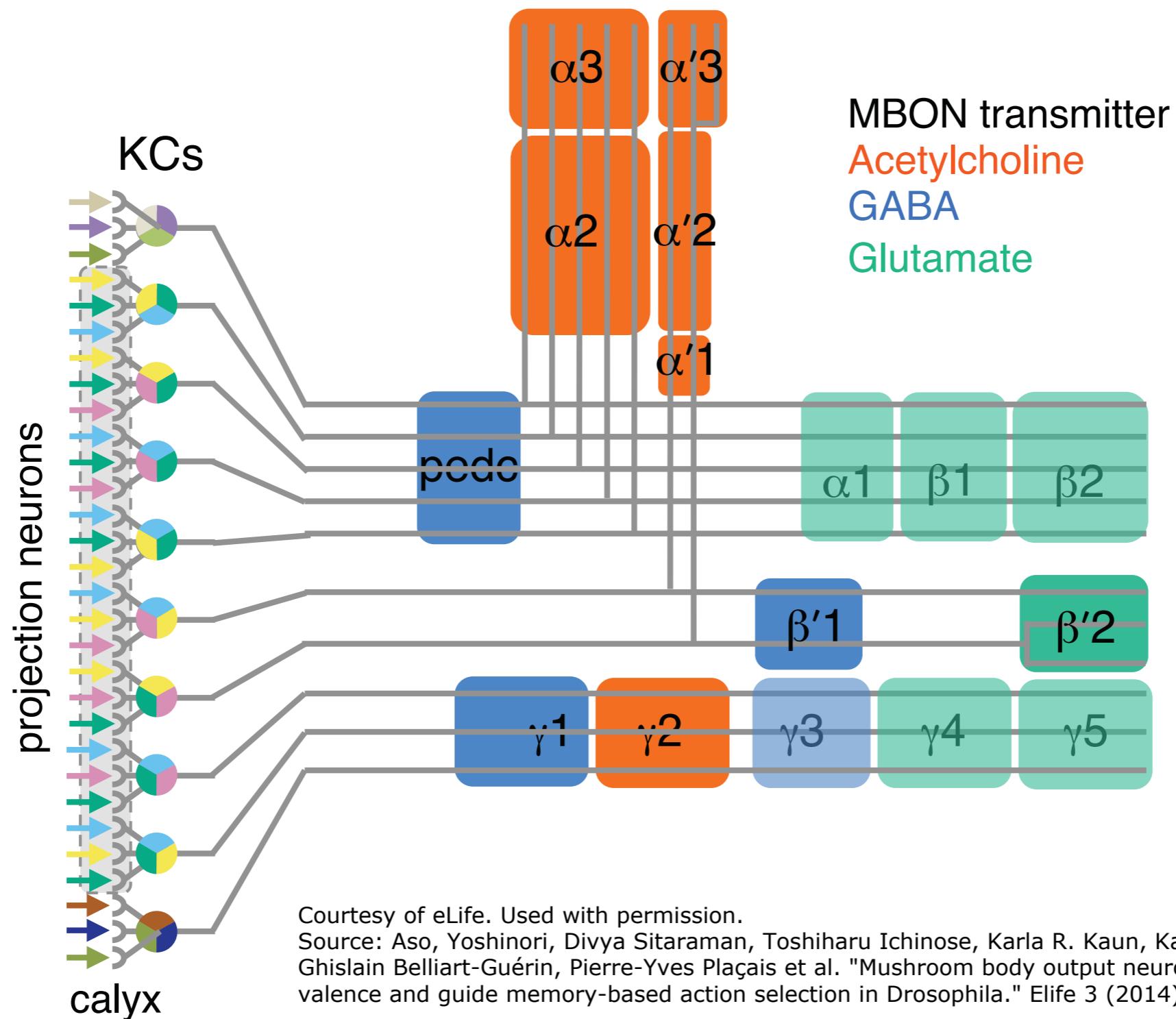
Registration of 13 Gal4 lines



Courtesy of eLife. Used with permission.

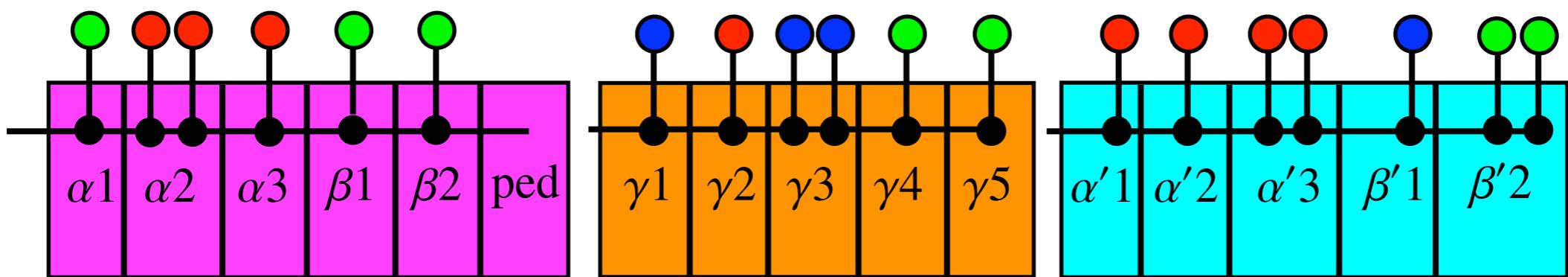
Source: Aso, Yoshinori, Divya Sitaraman, Toshiharu Ichinose, Karla R. Kaun, Katrin Vogt, Ghislain Belliart-Guérin, Pierre-Yves Plaçais et al. "Mushroom body output neurons encode valence and guide memory-based action selection in Drosophila." *Elife* 3 (2014): e04580.

Yoshinori Aso, Daisuke Hattori, Yang Yu, Rebecca M Johnston, Nirmala A Iyer, Teri-TB Ngo, Heather Dionne, LF Abbott, Richard Axel, Hiromu Tanimoto, Gerald M Rubin, 2014



● cholinergic ● glutamatergic ● GABAergic

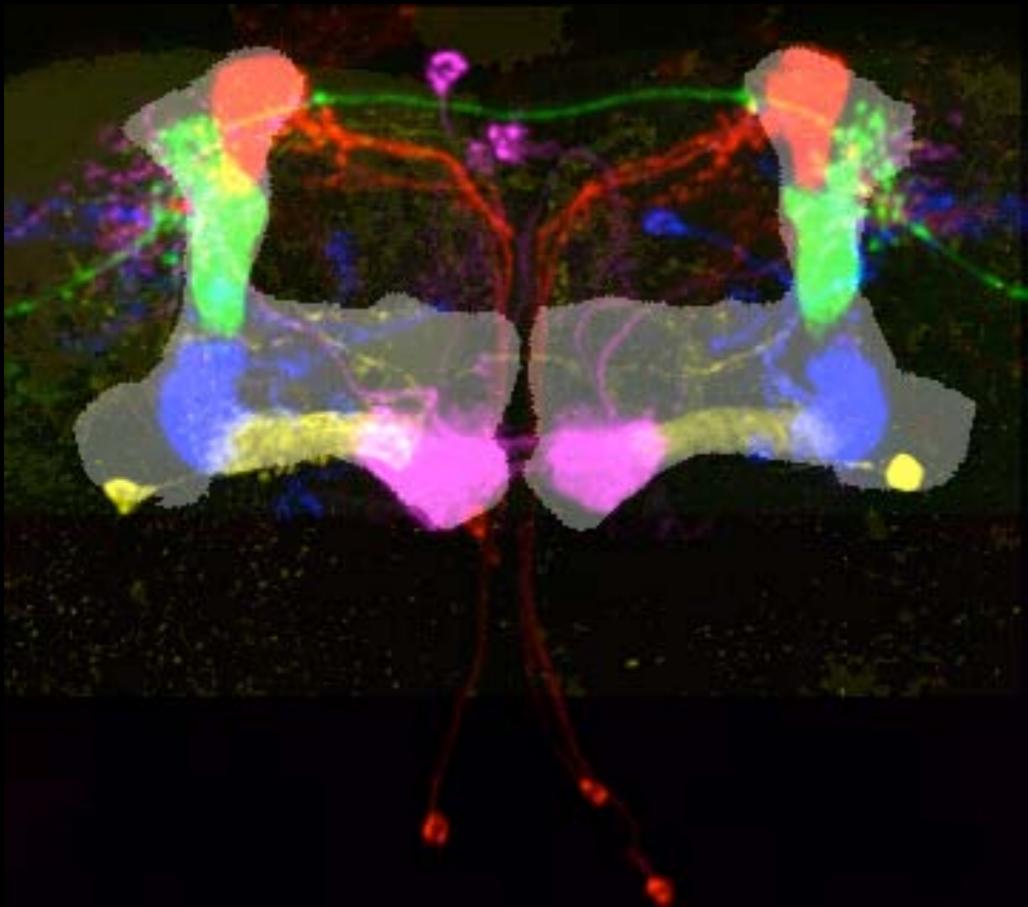
output neurons



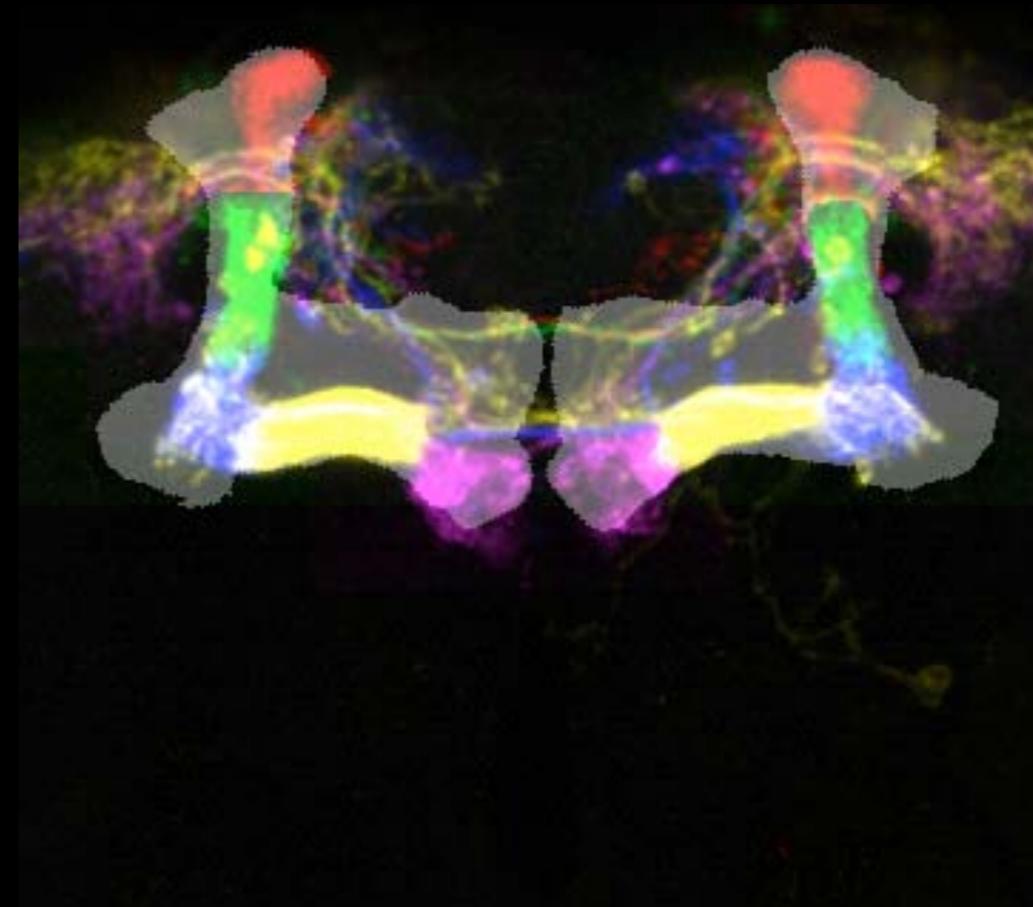
# Mushroom body lobe synaptic units

Neurons innervating  $\alpha/\beta$  lobes

Output neurons



Dopaminergic neurons



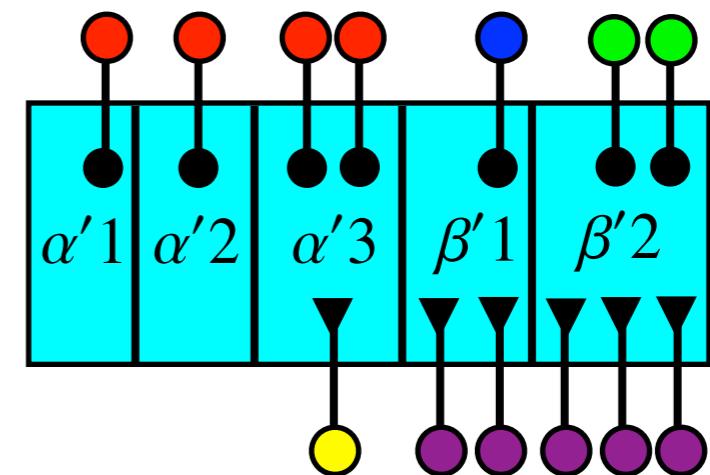
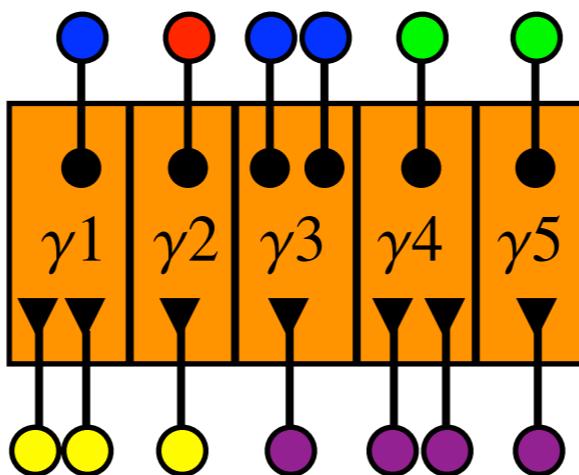
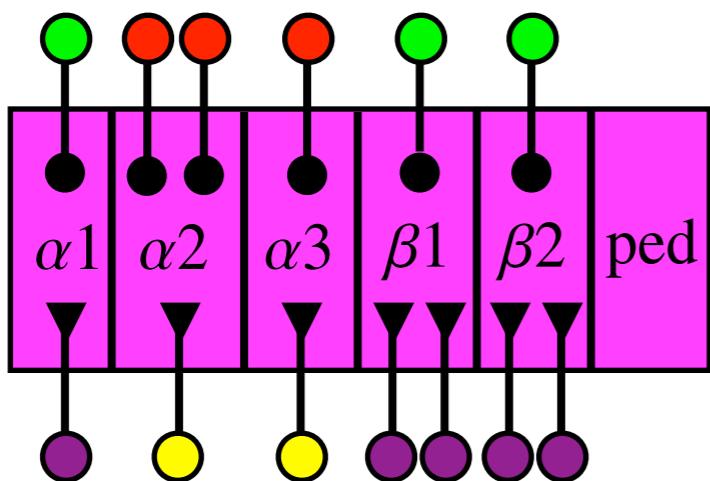
Courtesy of eLife. Used with permission.

Source: Aso, Yoshinori, Divya Sitaraman, Toshiharu Ichinose, Karla R. Kaun, Katrin Vogt, Ghislain Belliard-Guerin, Pierre-Yves Plaçais et al. "Mushroom body output neurons encode valence and guide memory-based action selection in Drosophila." *Elife* 3 (2014): e04580.

Yoshinori Aso, Daisuke Hattori, Yang Yu, Rebecca M Johnston, Nirmala A Iyer, Teri-TB Ngo, Heather Dionne, LF Abbott, Richard Axel, Hiromu Tanimoto, Gerald M Rubin, 2014

● cholinergic ● glutamatergic ● GABAergic

output neurons

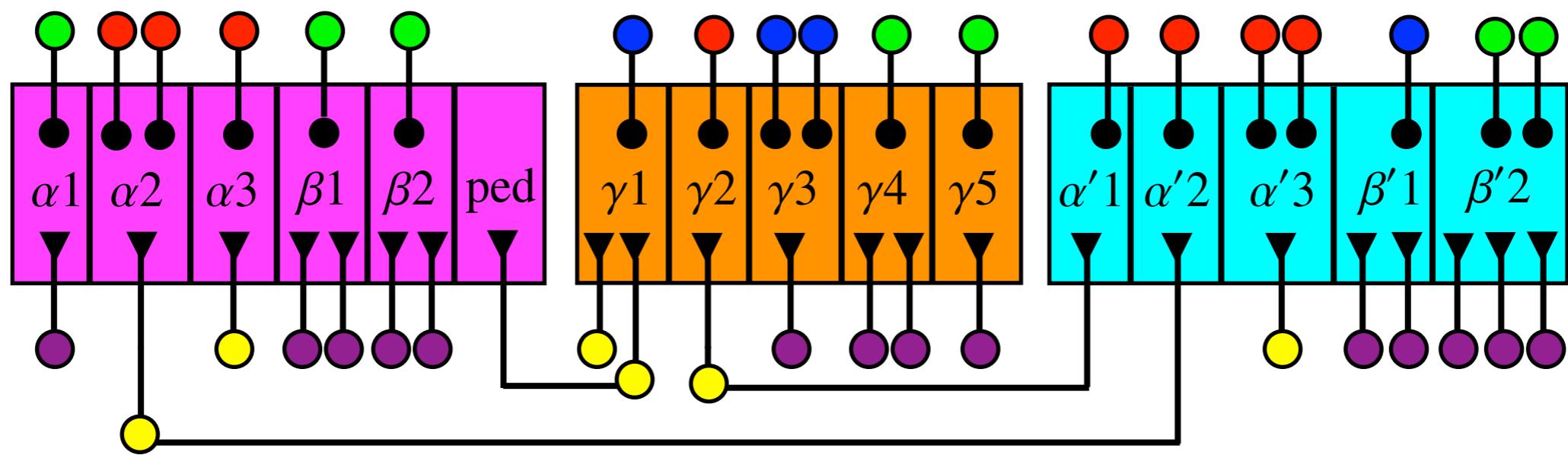


● PAM ● PPLI

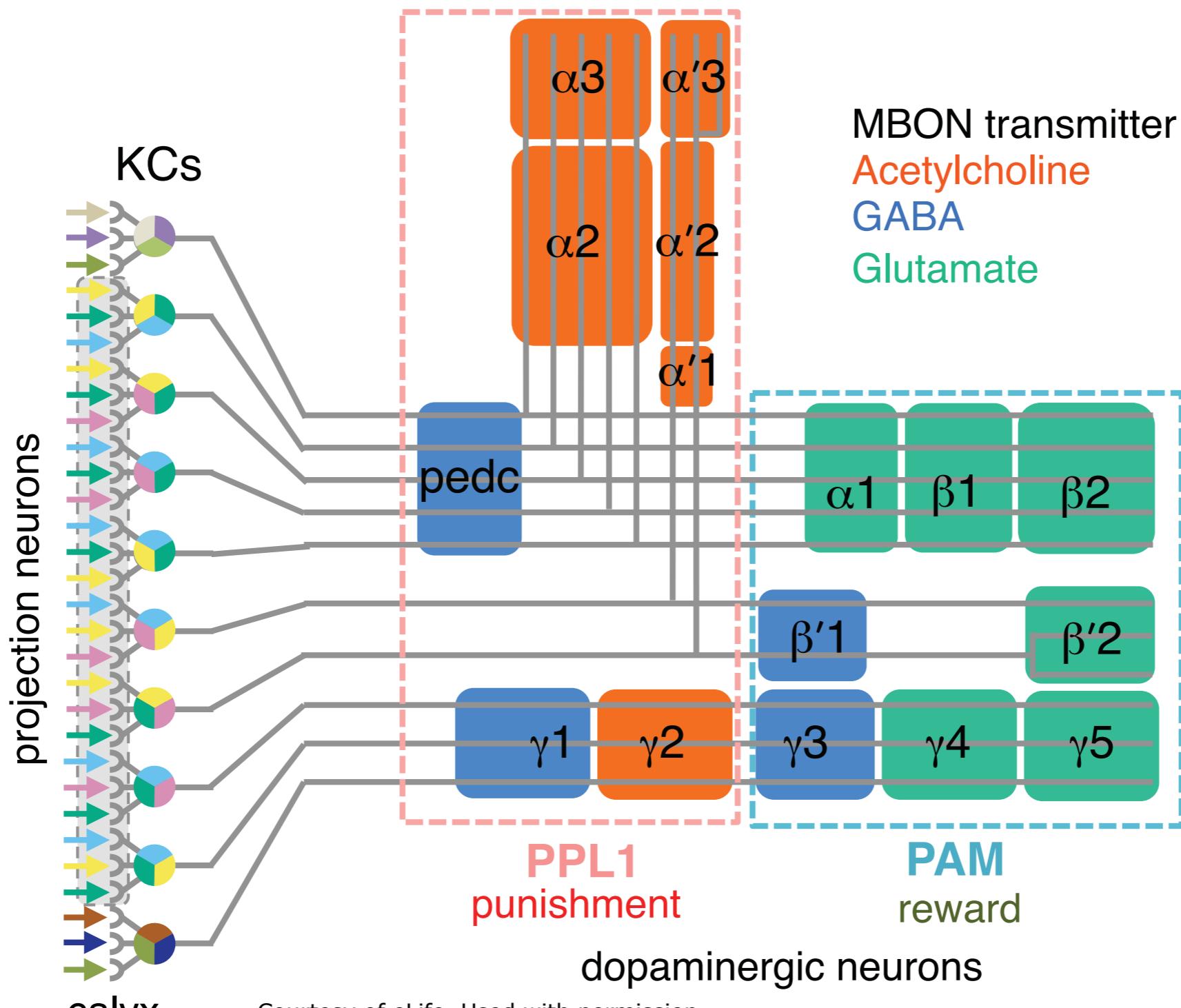
dopamine neurons

● cholinergic ● glutamatergic ● GABAergic

output neurons



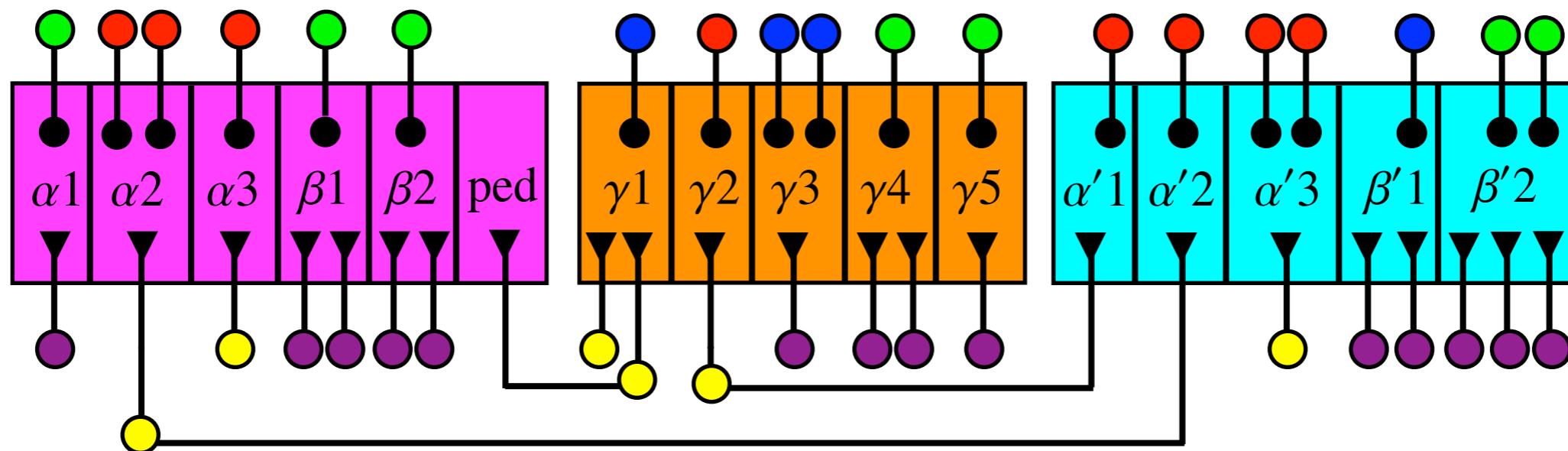
dopamine neurons

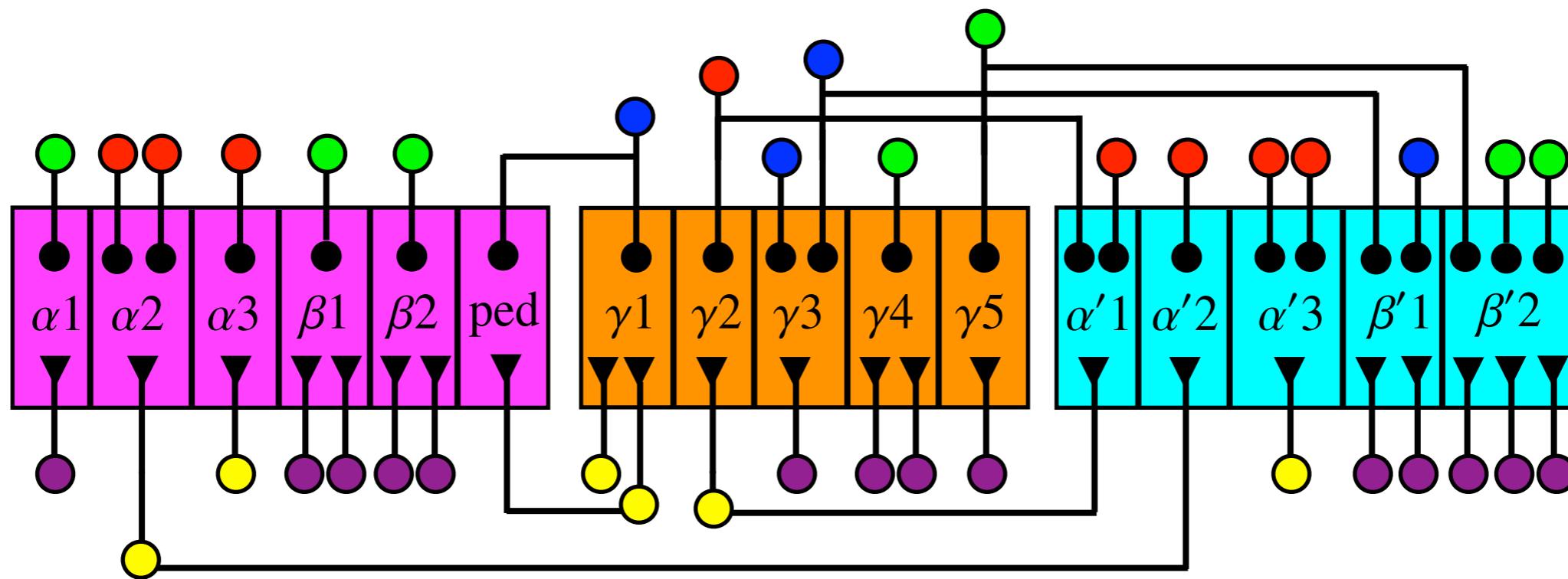


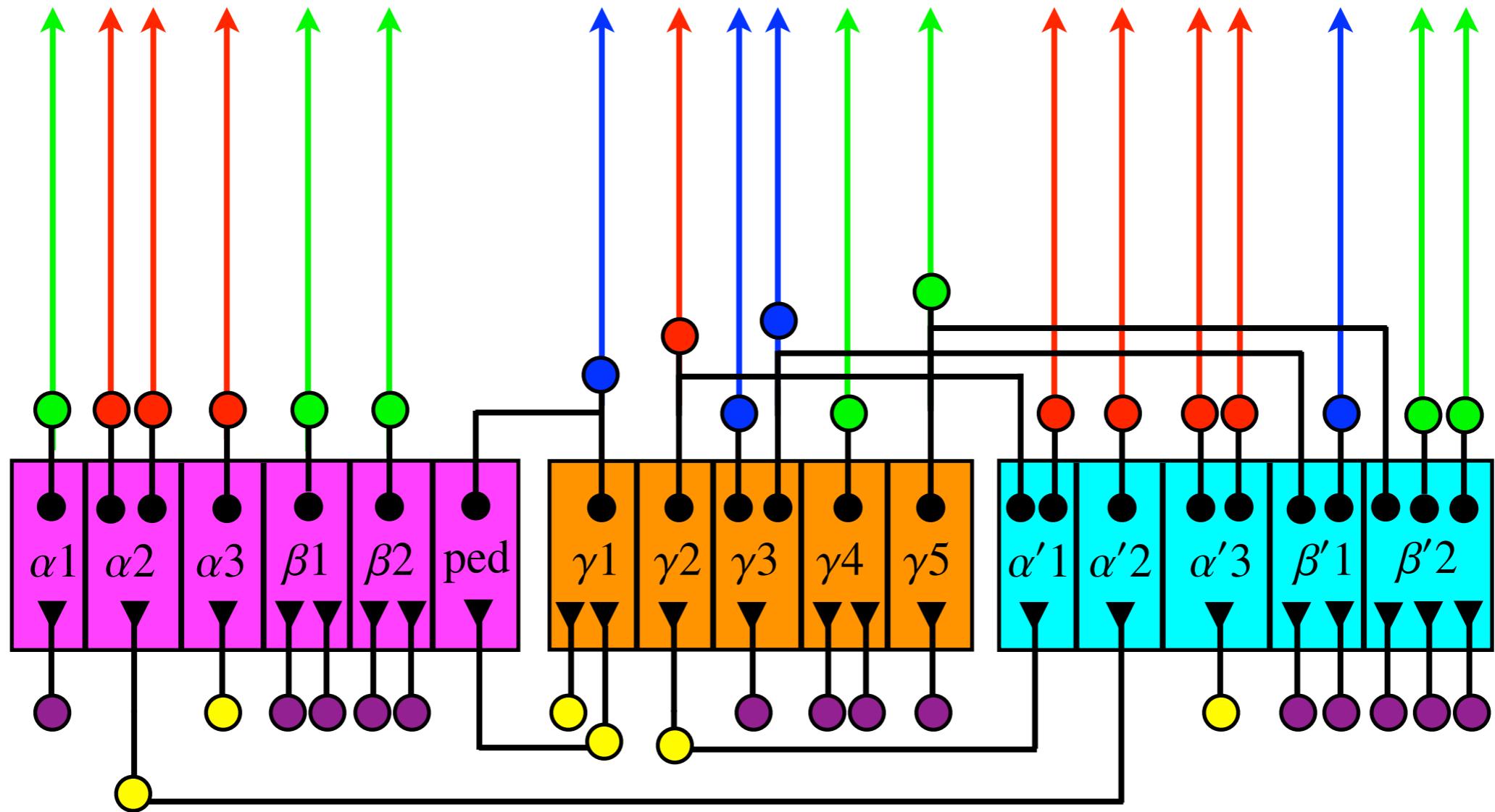
Courtesy of eLife. Used with permission.

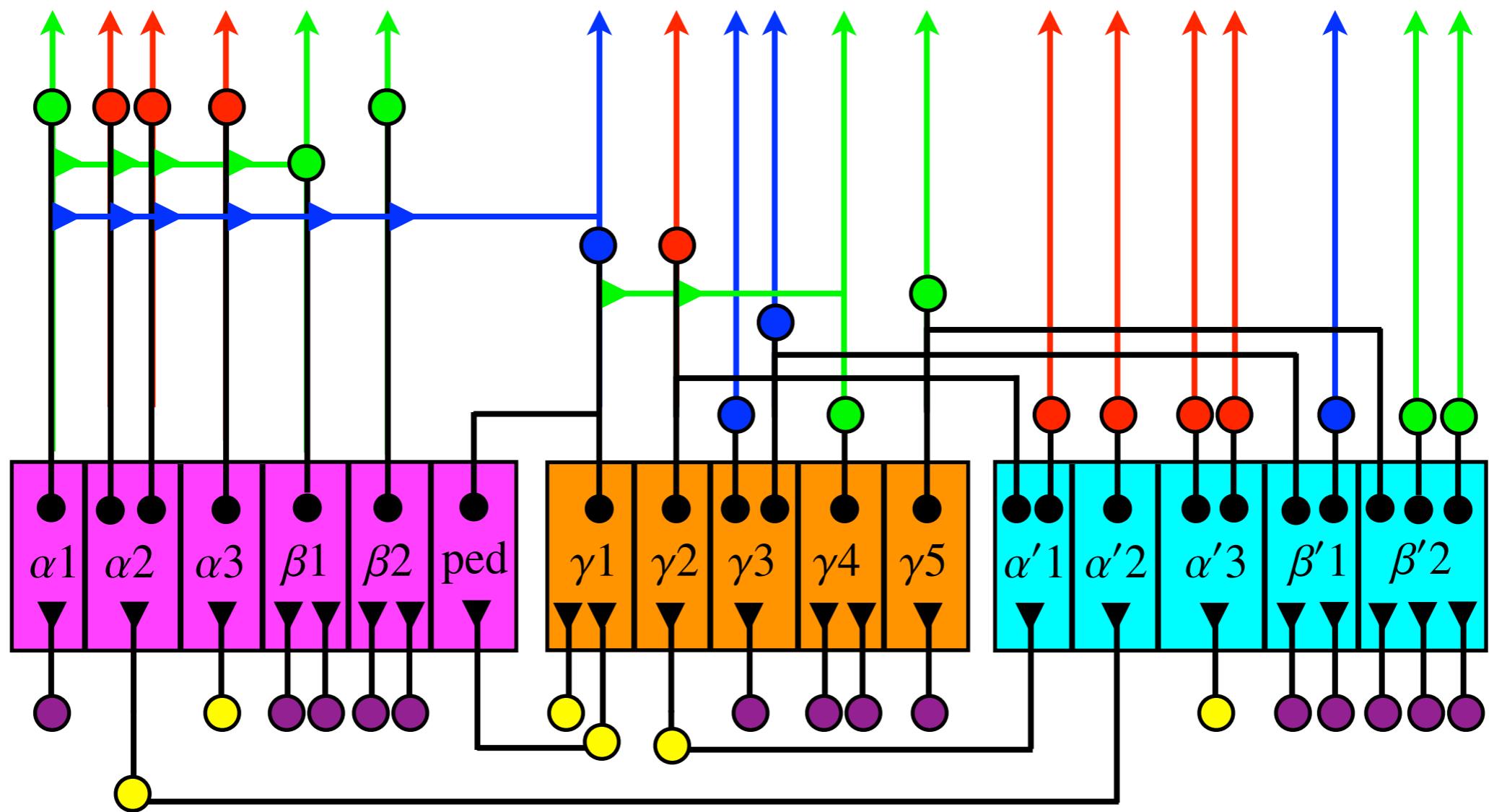
Source: Aso, Yoshinori, Divya Sitaraman, Toshiharu Ichinose, Karla R. Kaun, Katrin Vogt, Ghislain Belliard-Guérit, Pierre-Yves Plaçais et al. "Mushroom body output neurons encode valence and guide memory-based action selection in *Drosophila*." *Elife* 3 (2014): e04580.

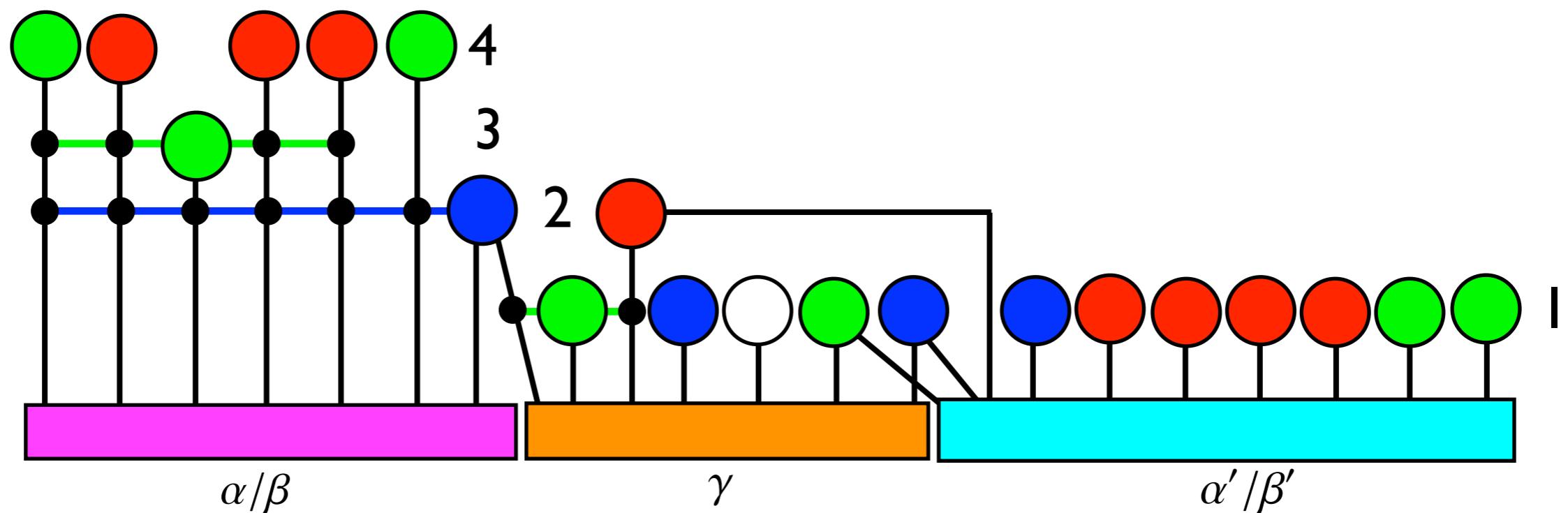
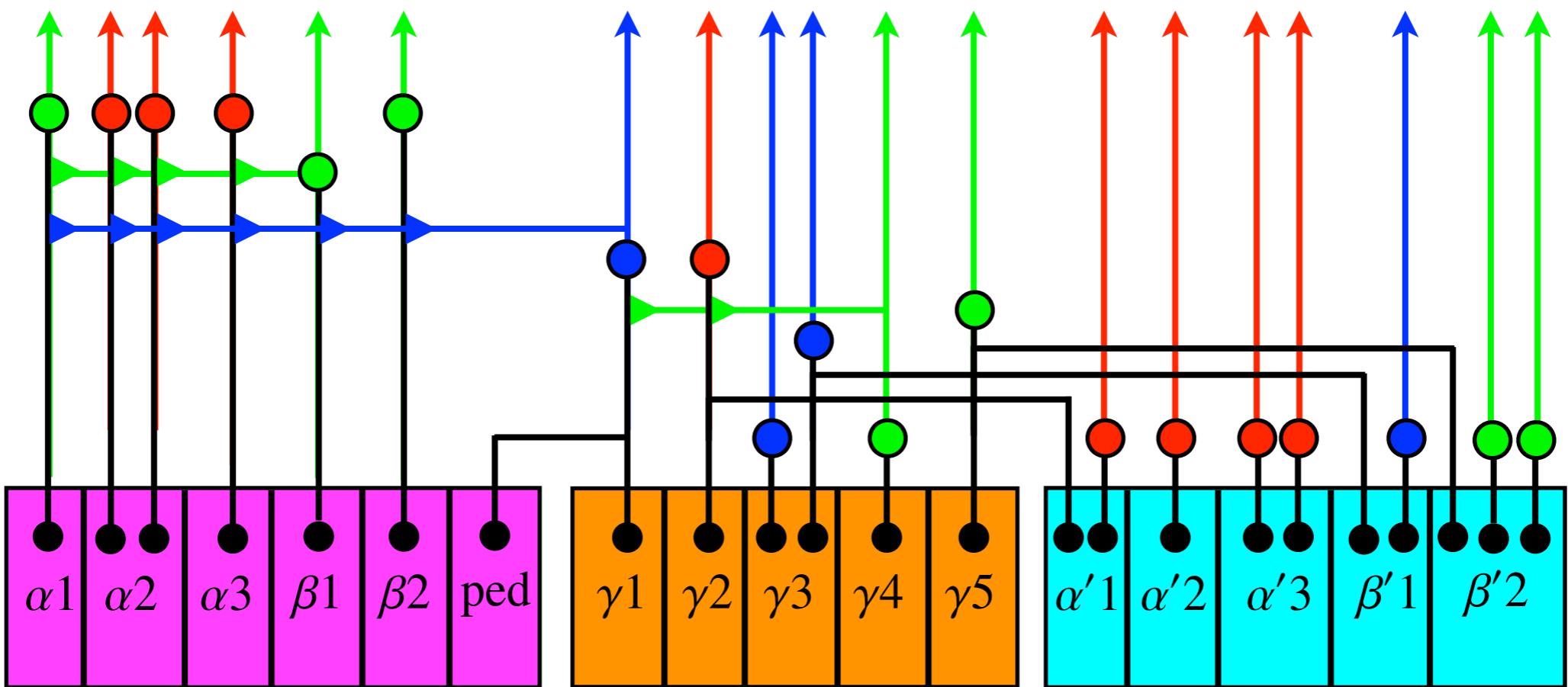
● cholinergic ● glutamatergic ● GABAergic



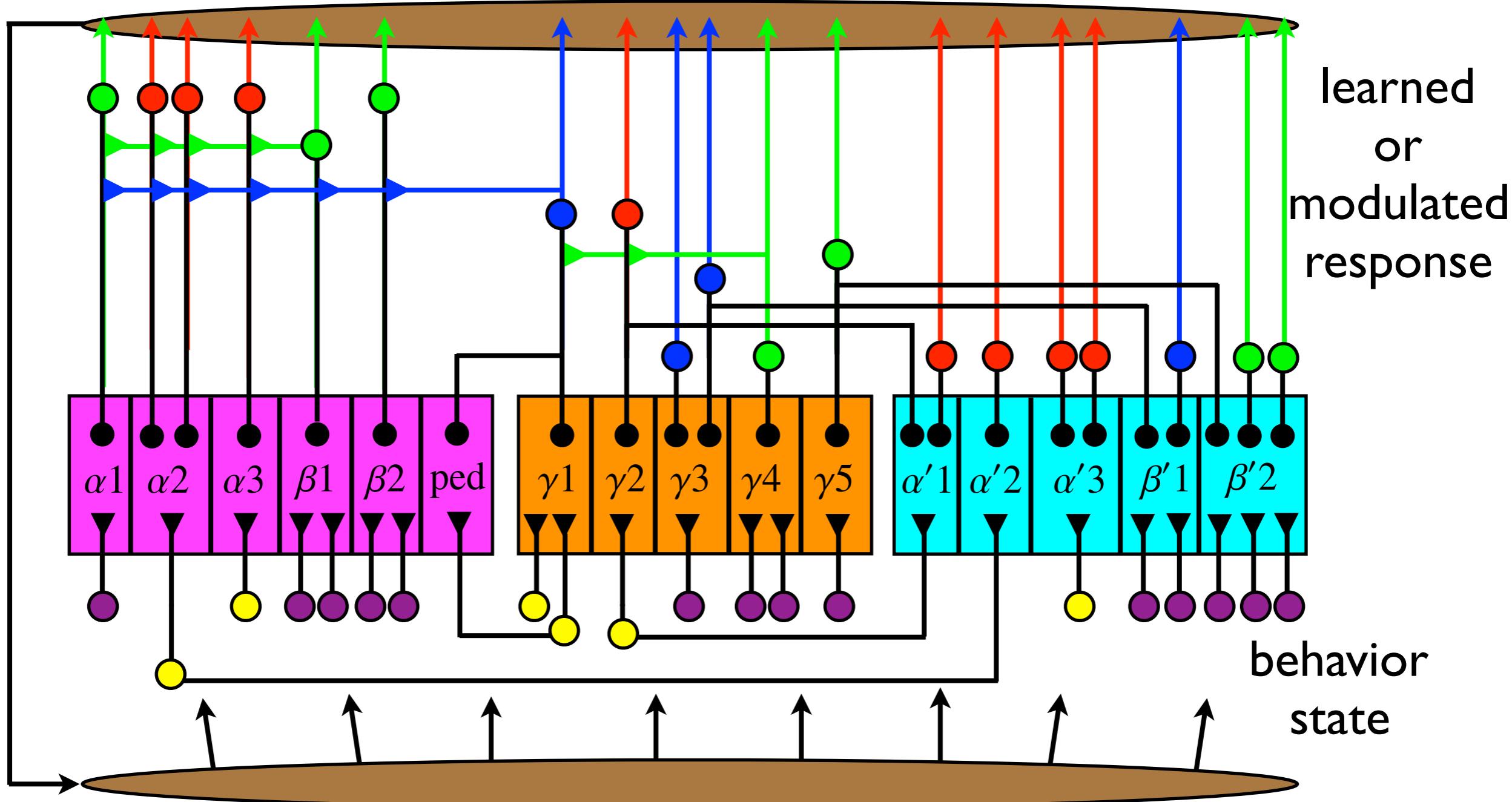




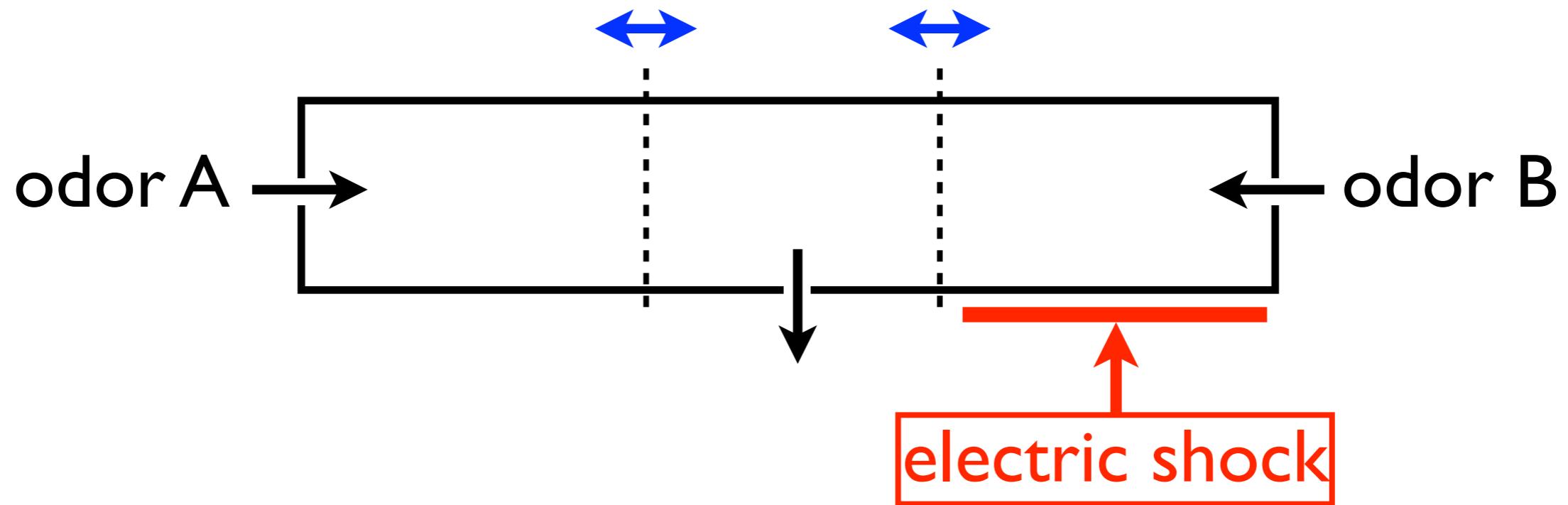




crepine, superior medial, intermediate and lateral protocerebrum, lateral horn

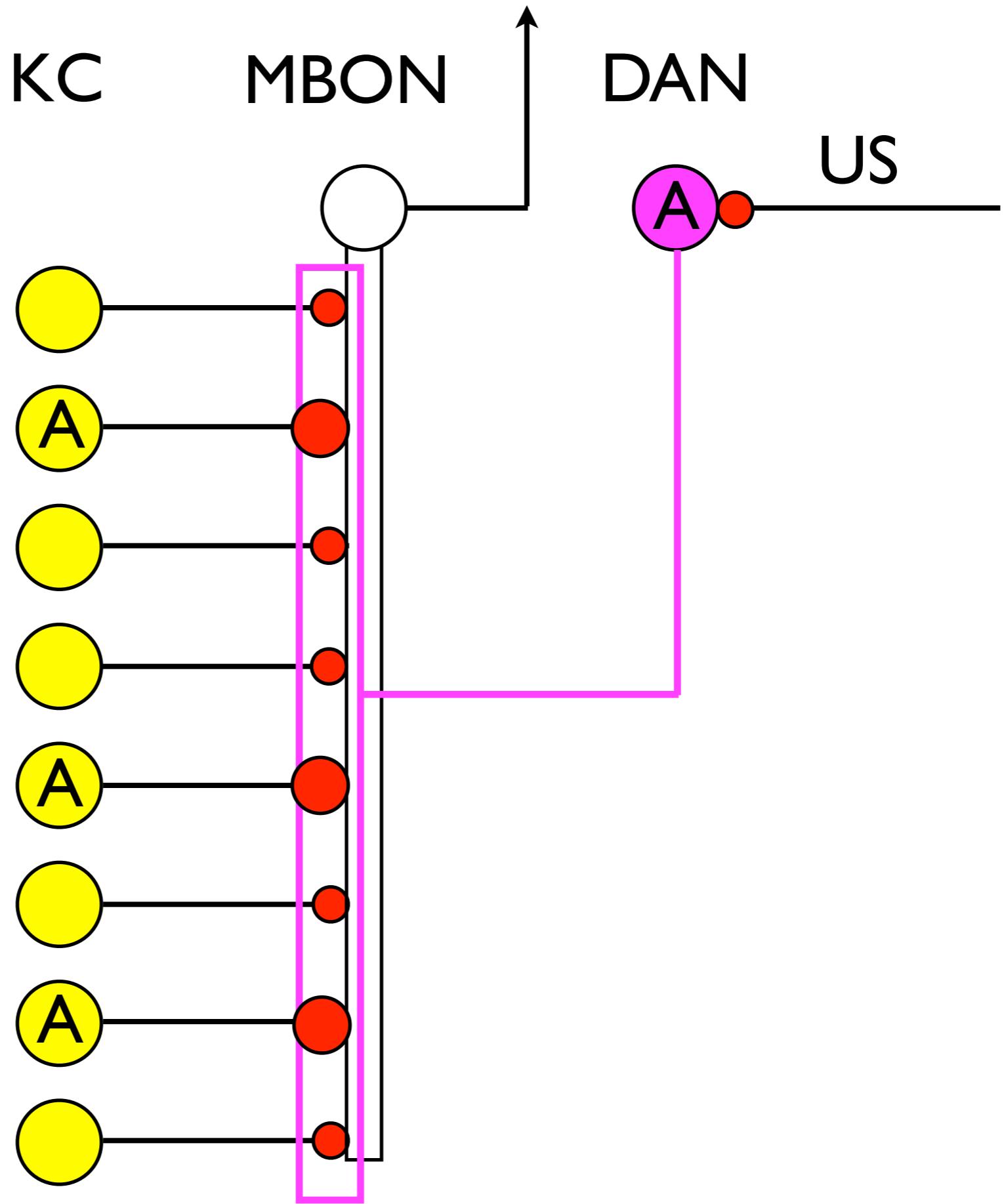


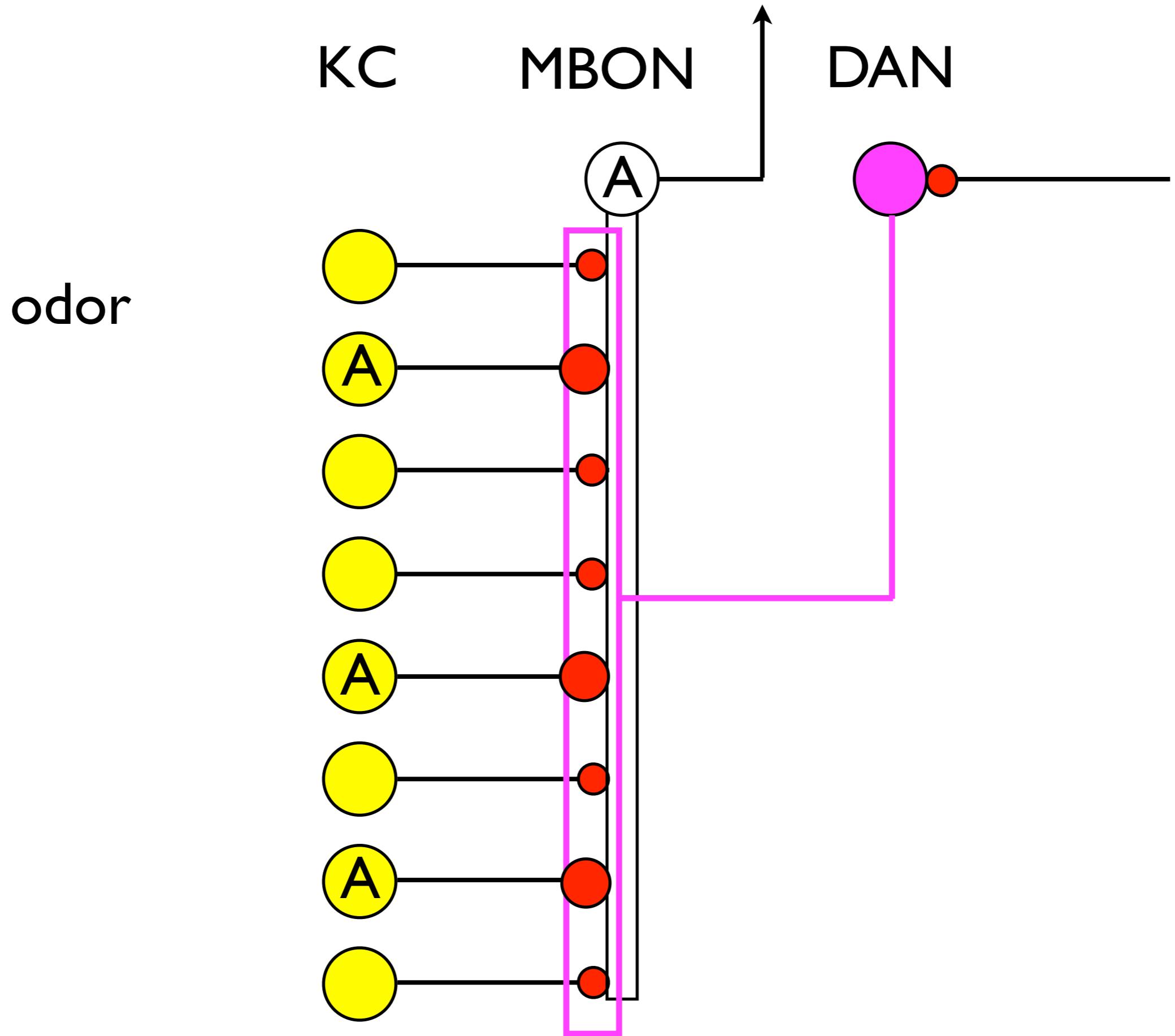
# classical conditioning

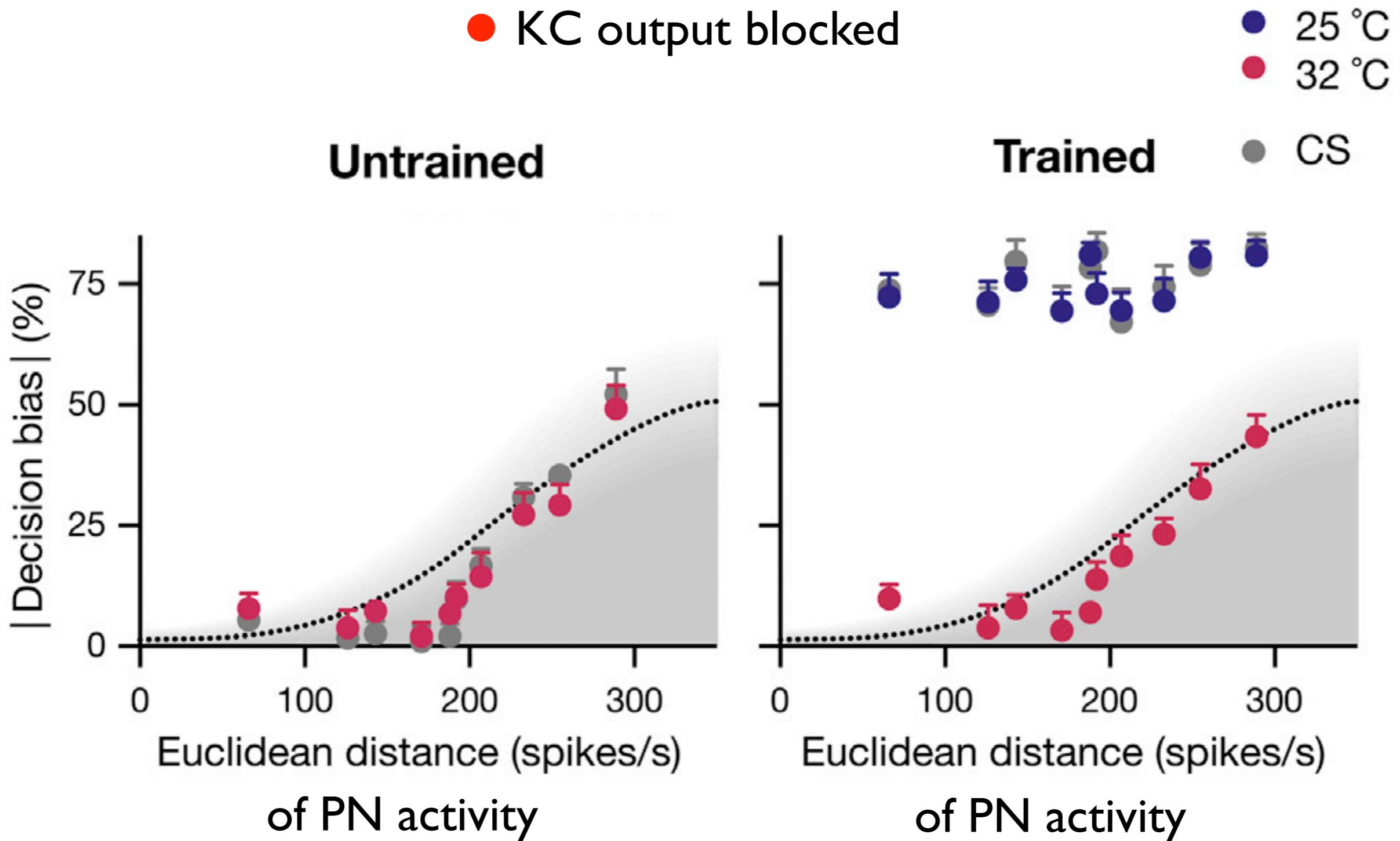


Moshe Parnas, Andrew C. Lin, Wolf Huettneroth, Gero Miesenbock 2013

odor - CS



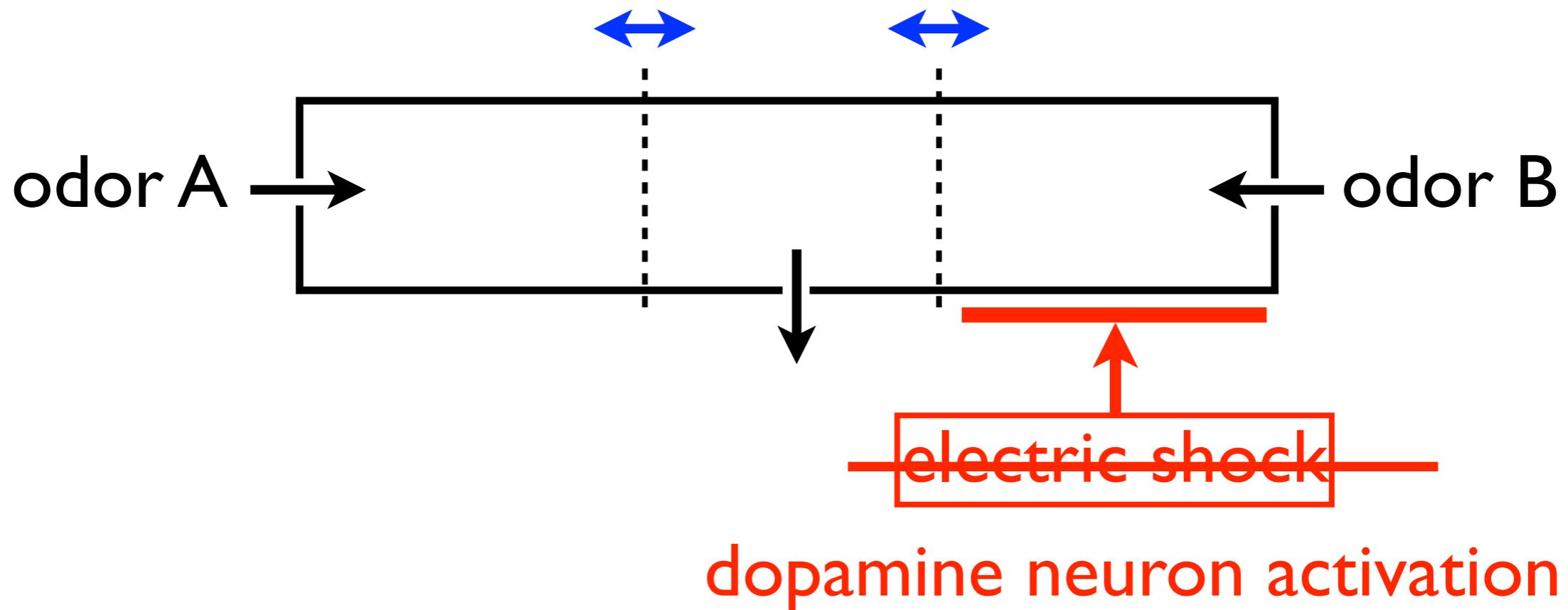




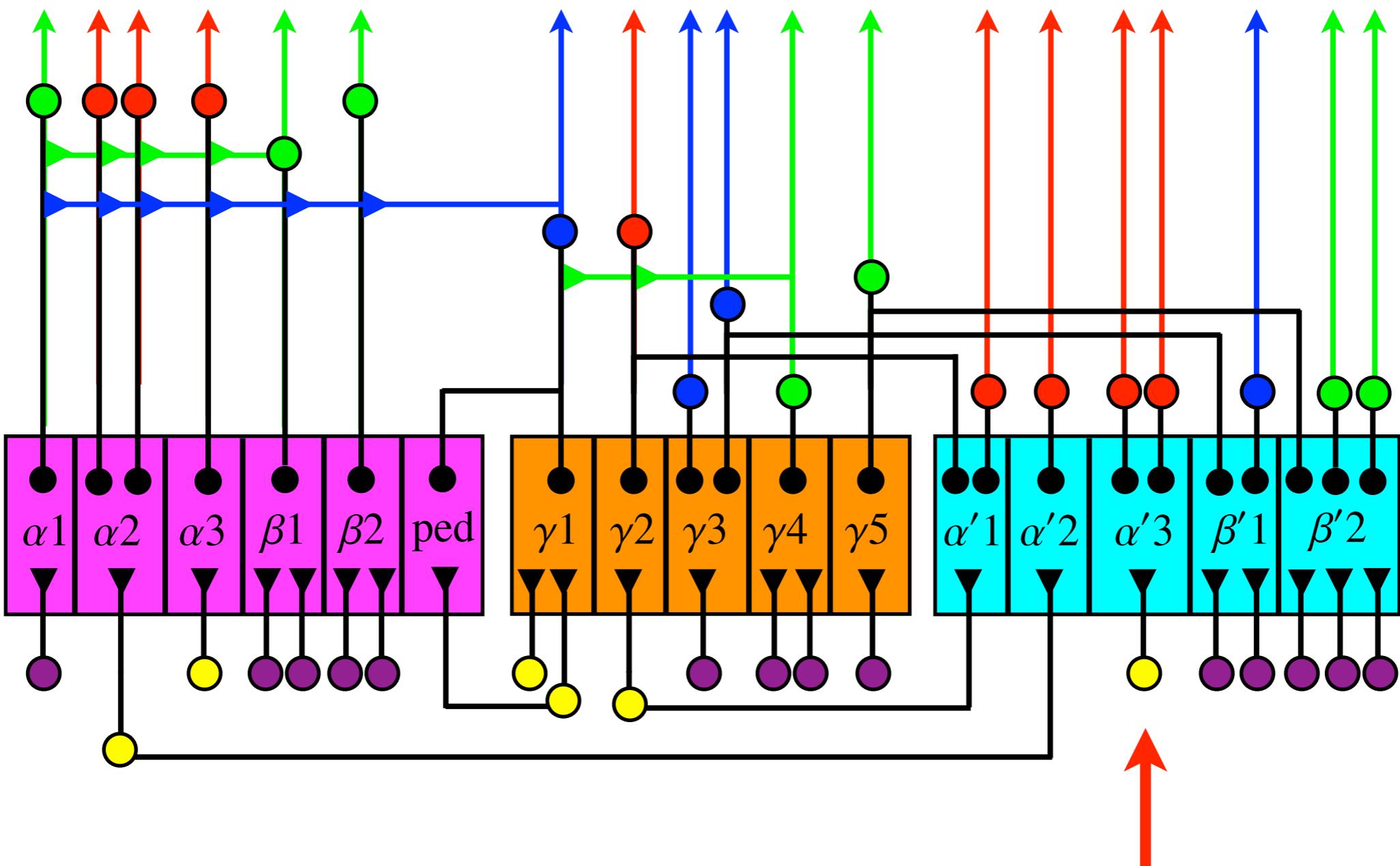
Courtesy of Elsevier, Inc., <http://www.sciencedirect.com>. Used with permission.

Source: Parnas, Moshe, Andrew C. Lin, Wolf Huettneroth, and Gero Miesenböck. "Odor discrimination in Drosophila: From neural population codes to behavior." *Neuron* 79, no. 5 (2013): 932-944.

# classical conditioning

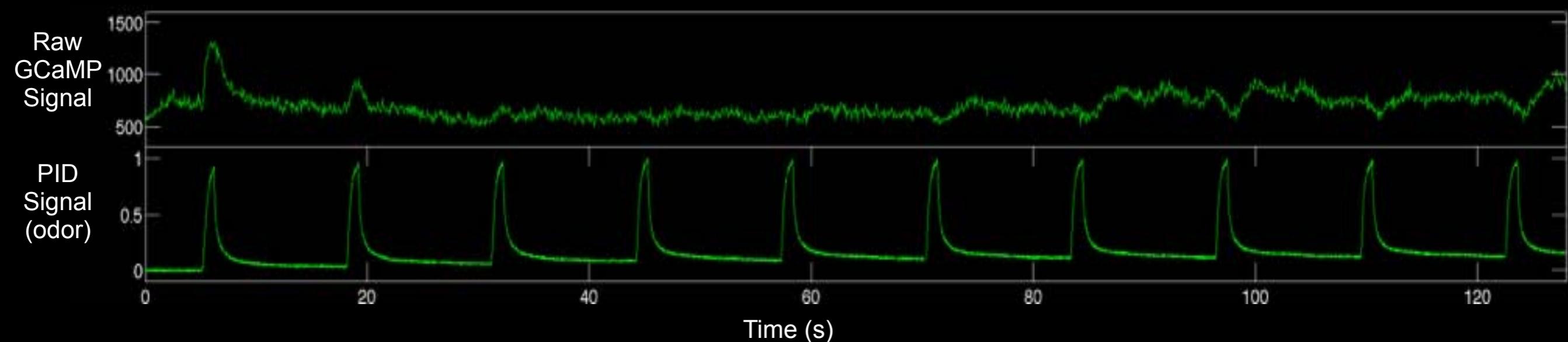


# recognition memory

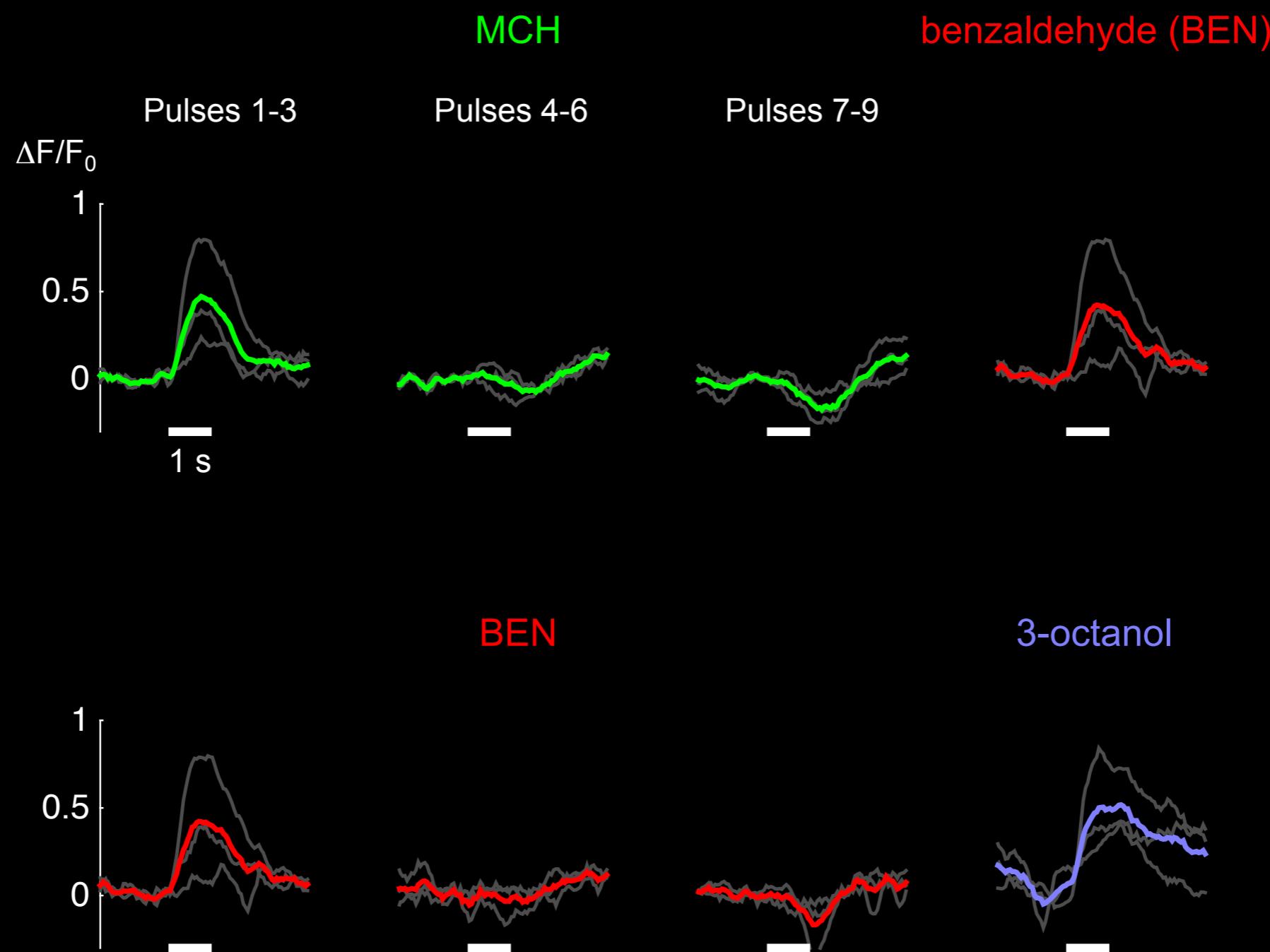


# MBON- $\alpha'3$ activity depresses upon repetitive odor stimulation

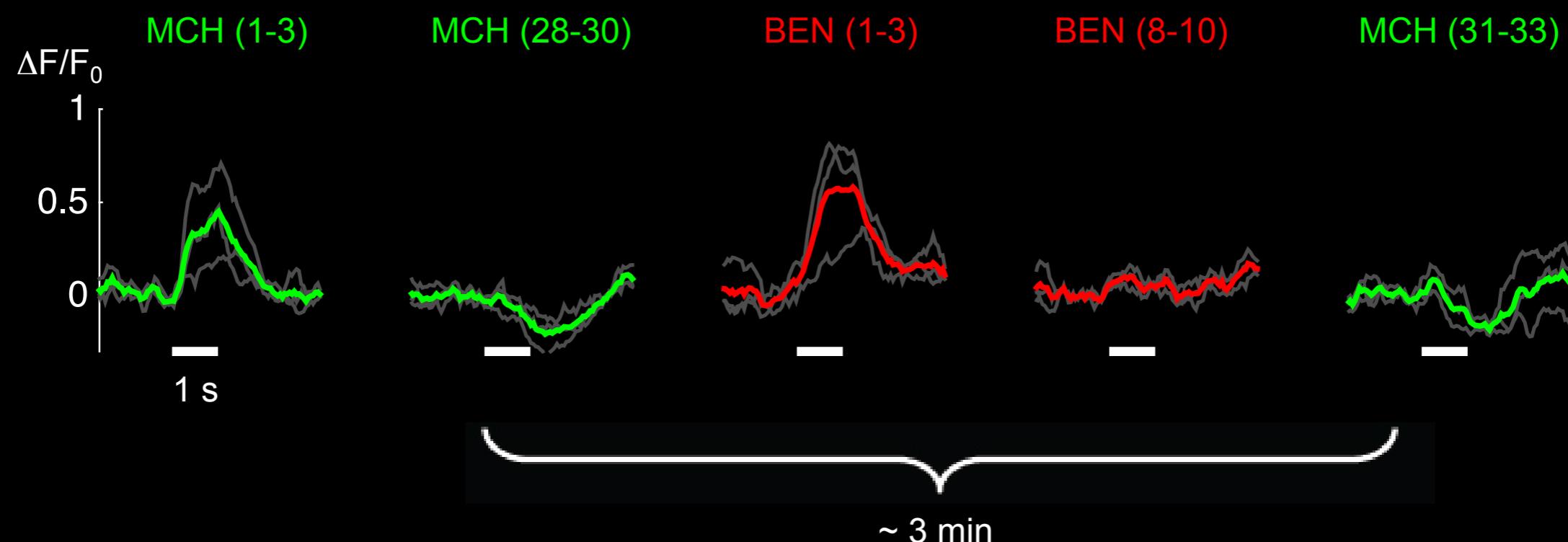
4-methylcyclohexanol (MCH)



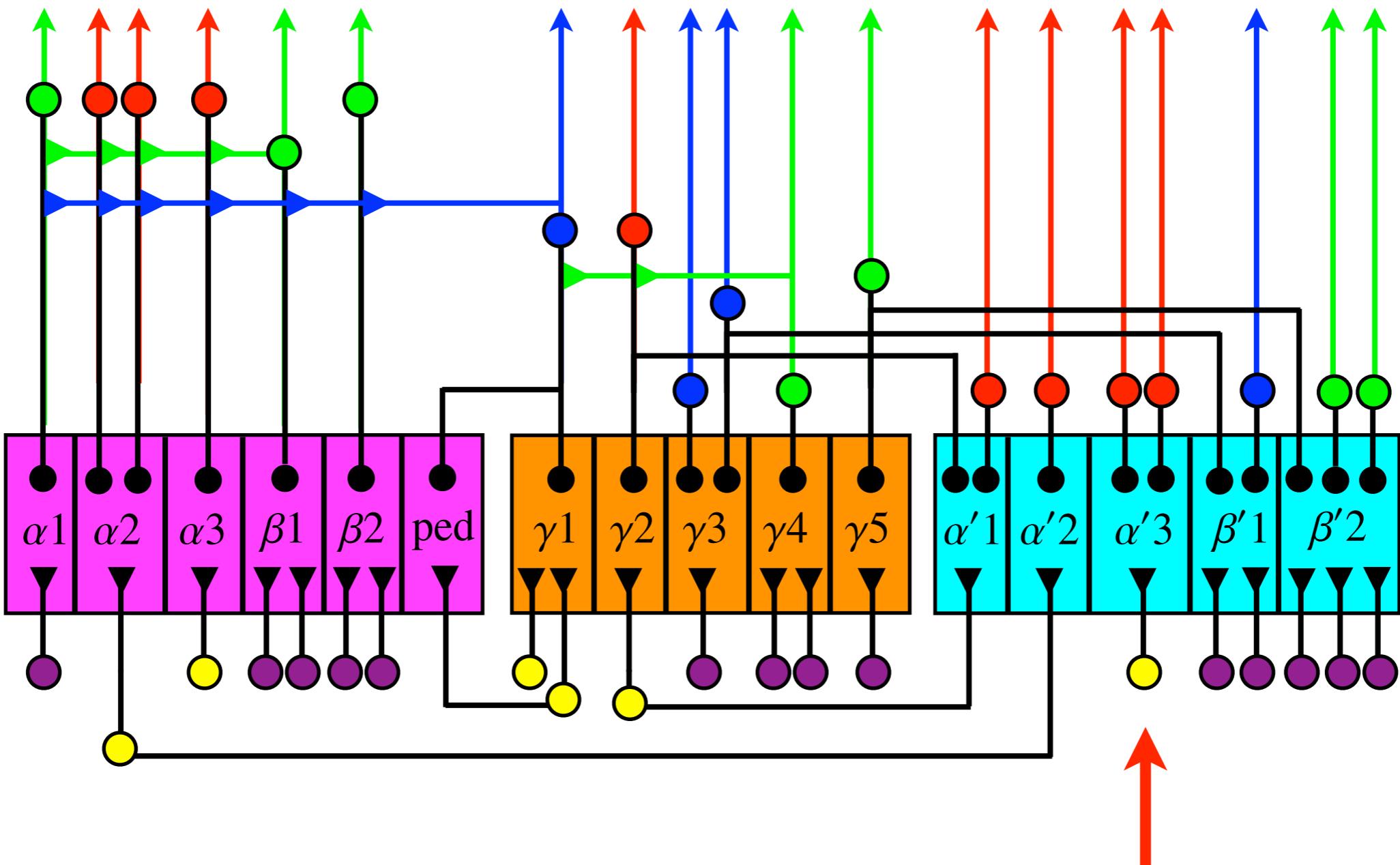
# MBON- $\alpha'3$ depression is specific to repeated odor



# MBON- $\alpha'3$ depression is persistent

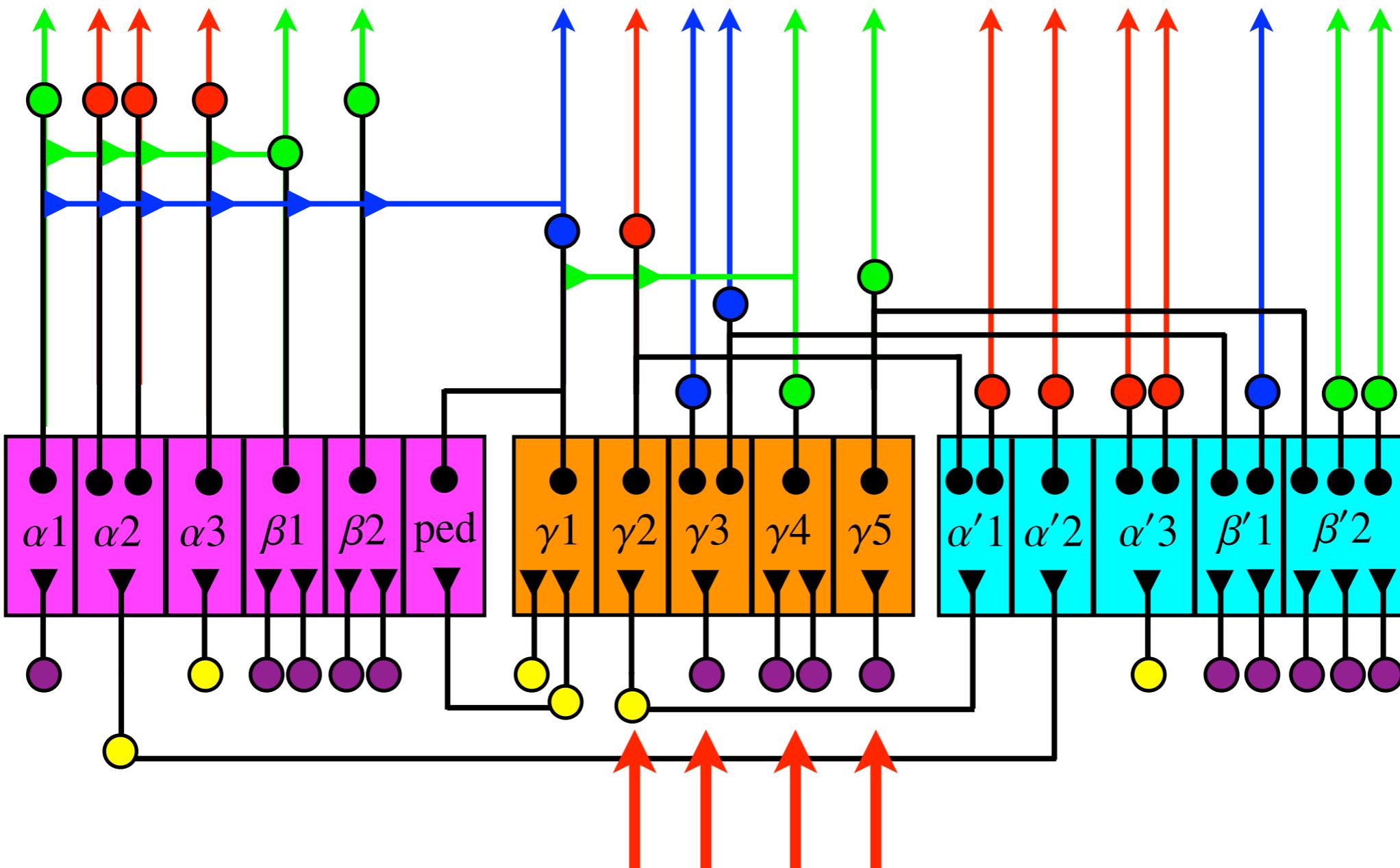


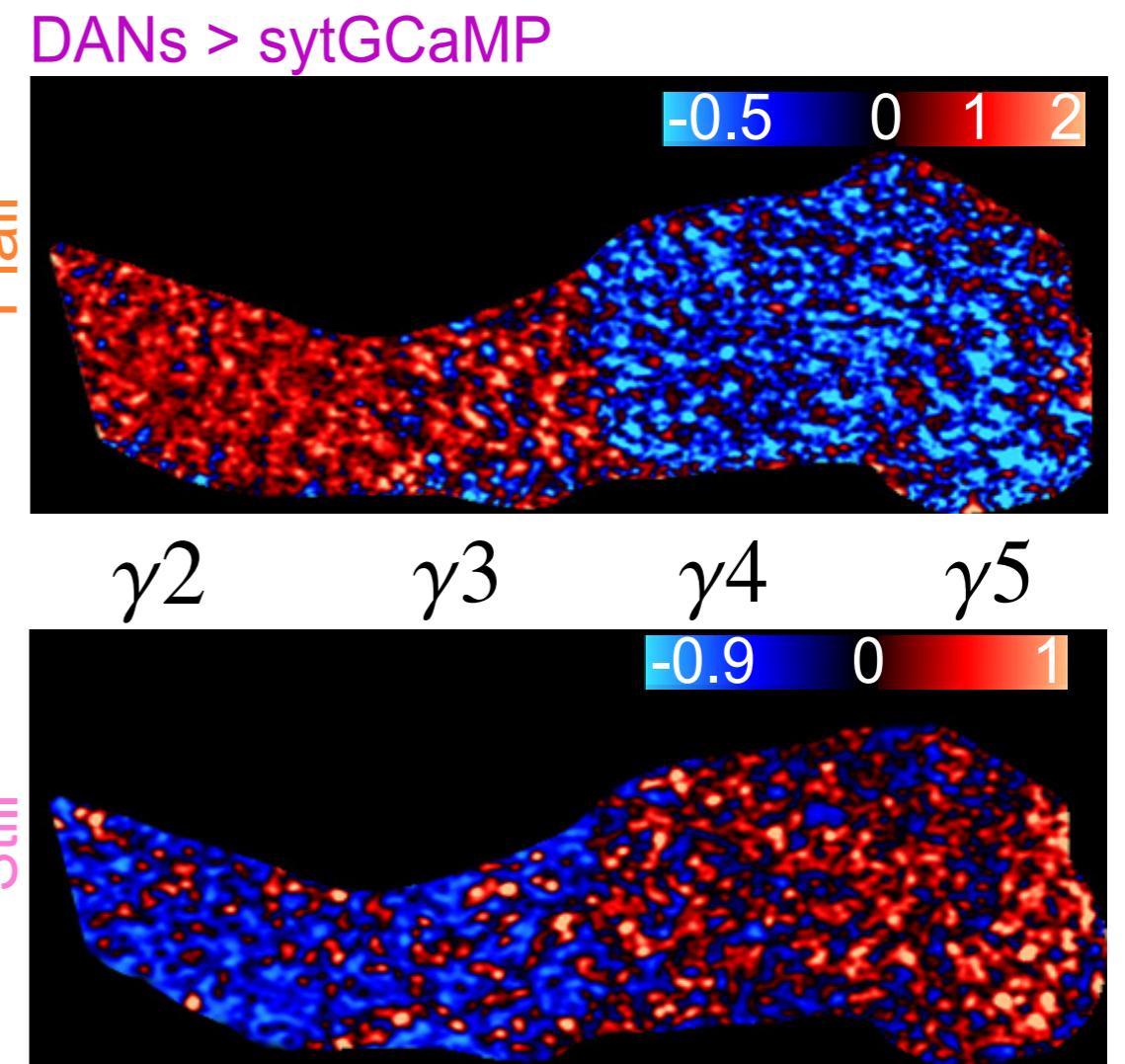
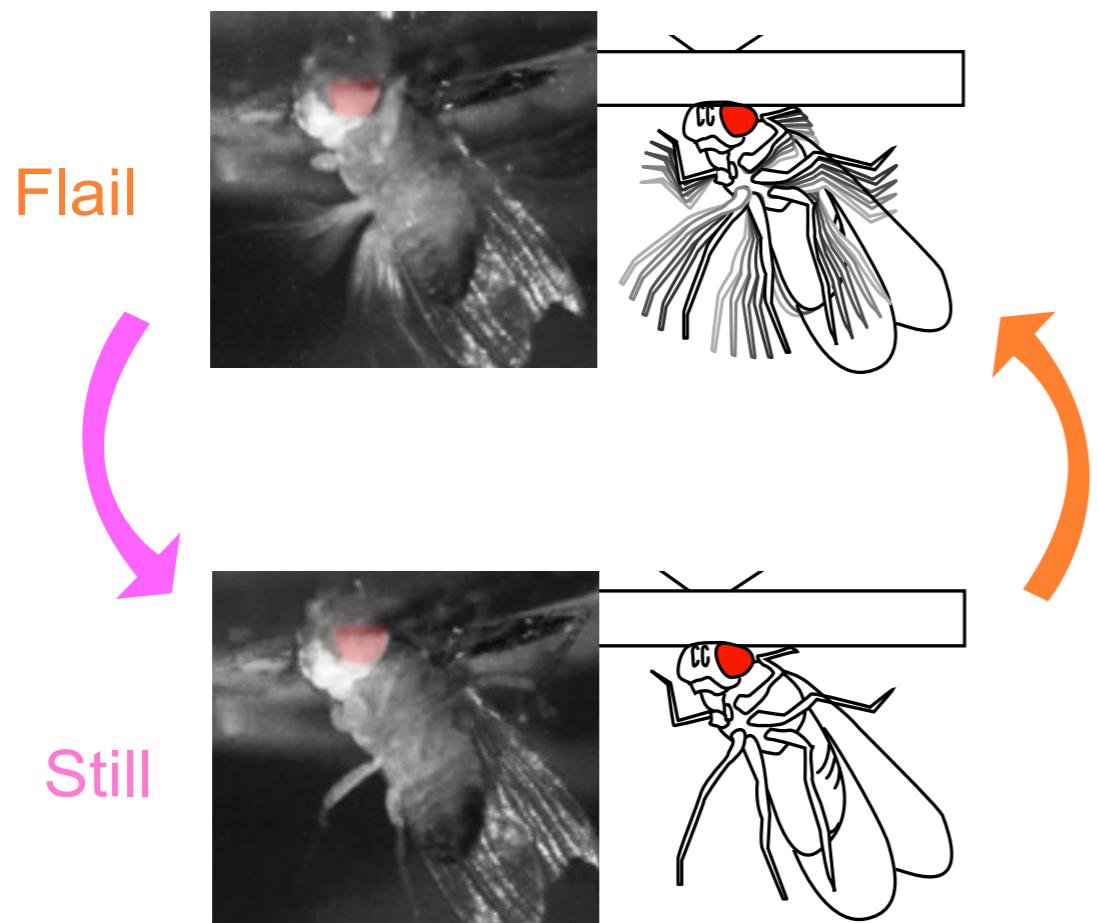
# recognition memory



Daisuke Hattori, Richard Axel

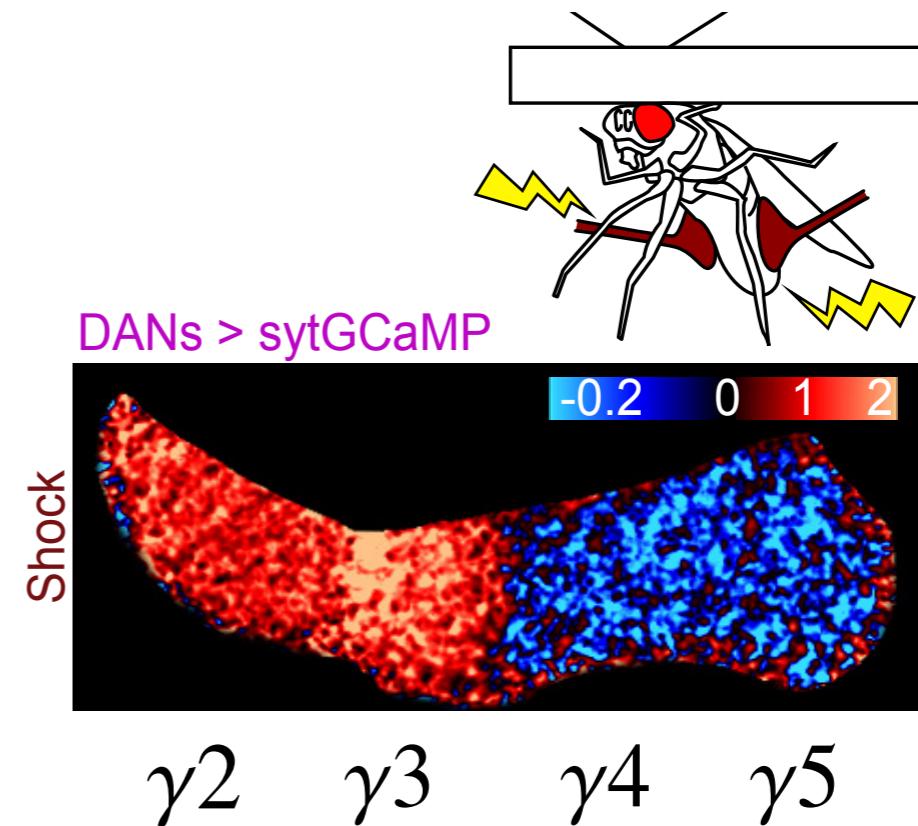
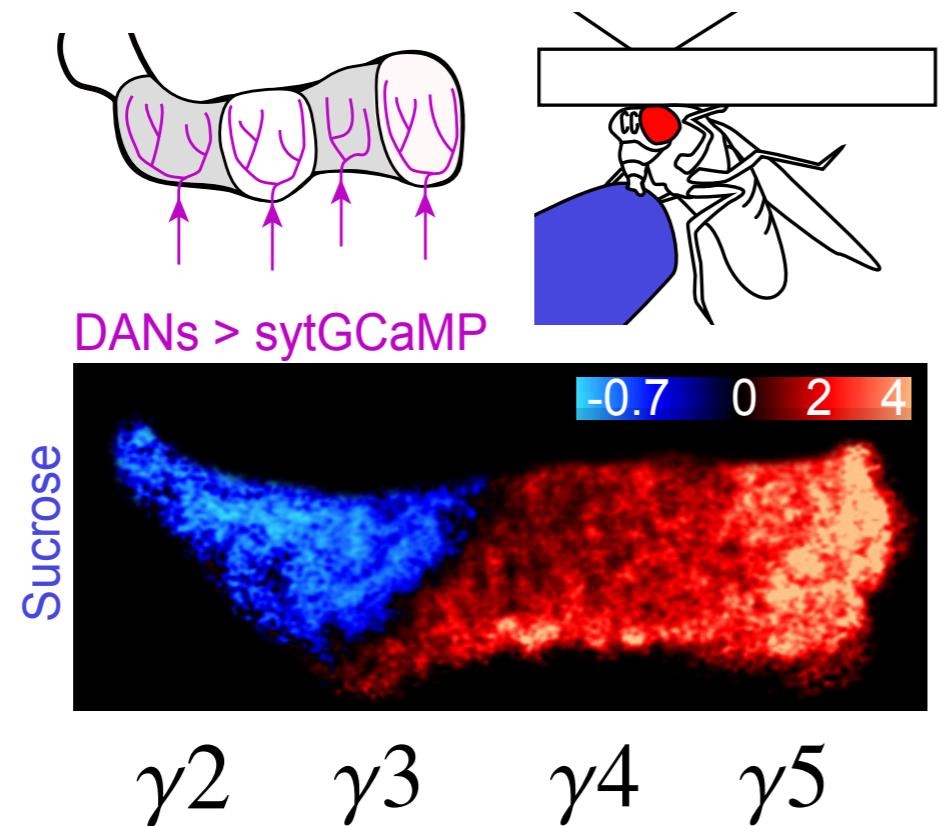
# internal state





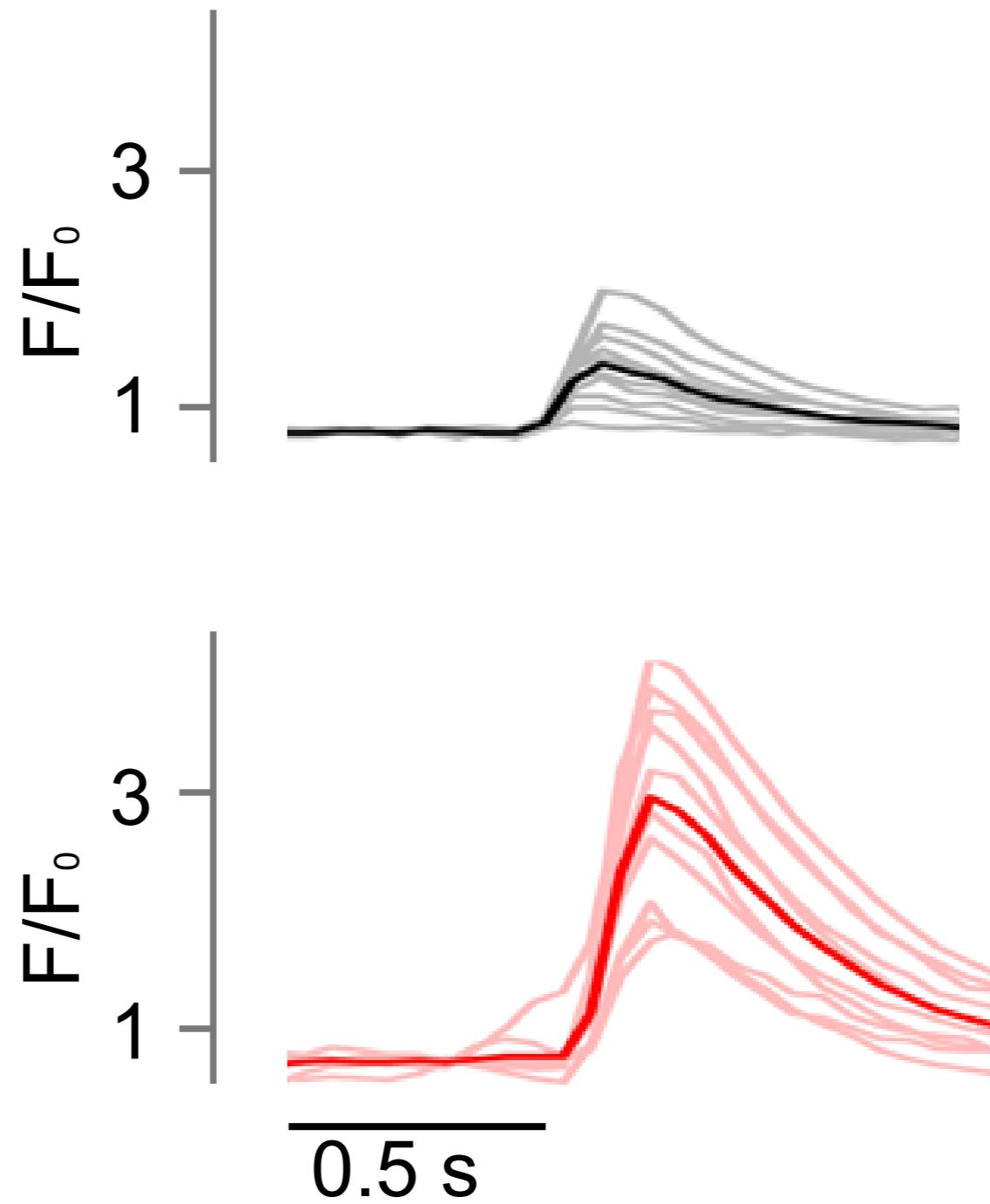
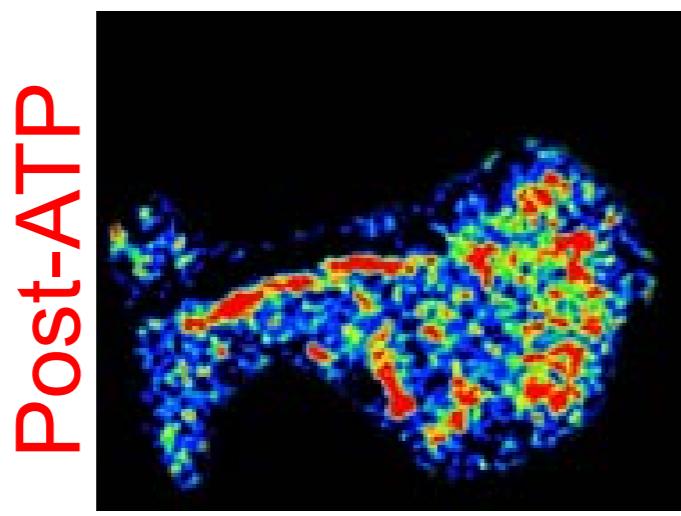
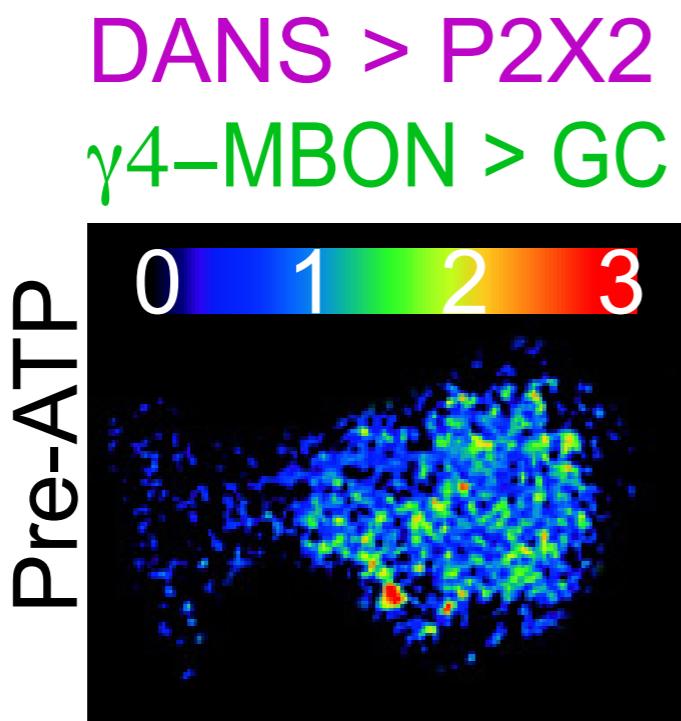
Courtesy of Elsevier, Inc., <http://www.sciencedirect.com>. Used with permission.

Source: Cohn, Raphael, Ianessa Morante, and Vanessa Ruta. "Coordinated and compartmentalized neuromodulation shapes sensory processing in Drosophila." *Cell* 163, no. 7 (2015): 1742-1755.



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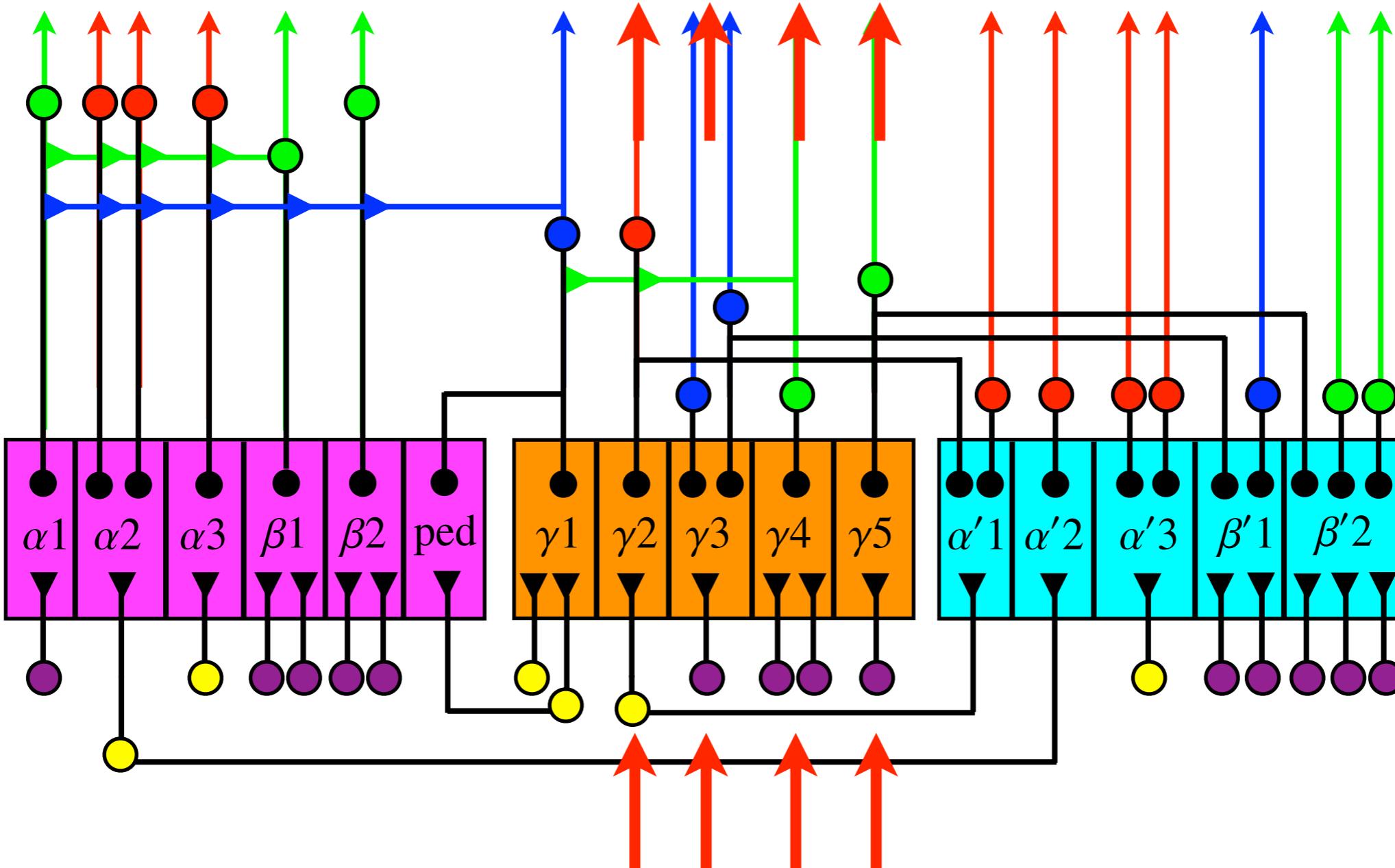
Source: Cohn, Raphael, Ianessa Morante, and Vanessa Ruta. "Coordinated and compartmentalized neuromodulation shapes sensory processing in Drosophila." *Cell* 163, no. 7 (2015): 1742-1755.



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Source: Cohn, Raphael, Ianessa Morantte, and Vanessa Ruta. "Coordinated and compartmentalized neuromodulation shapes sensory processing in Drosophila." Cell 163, no. 7 (2015): 1742-1755.

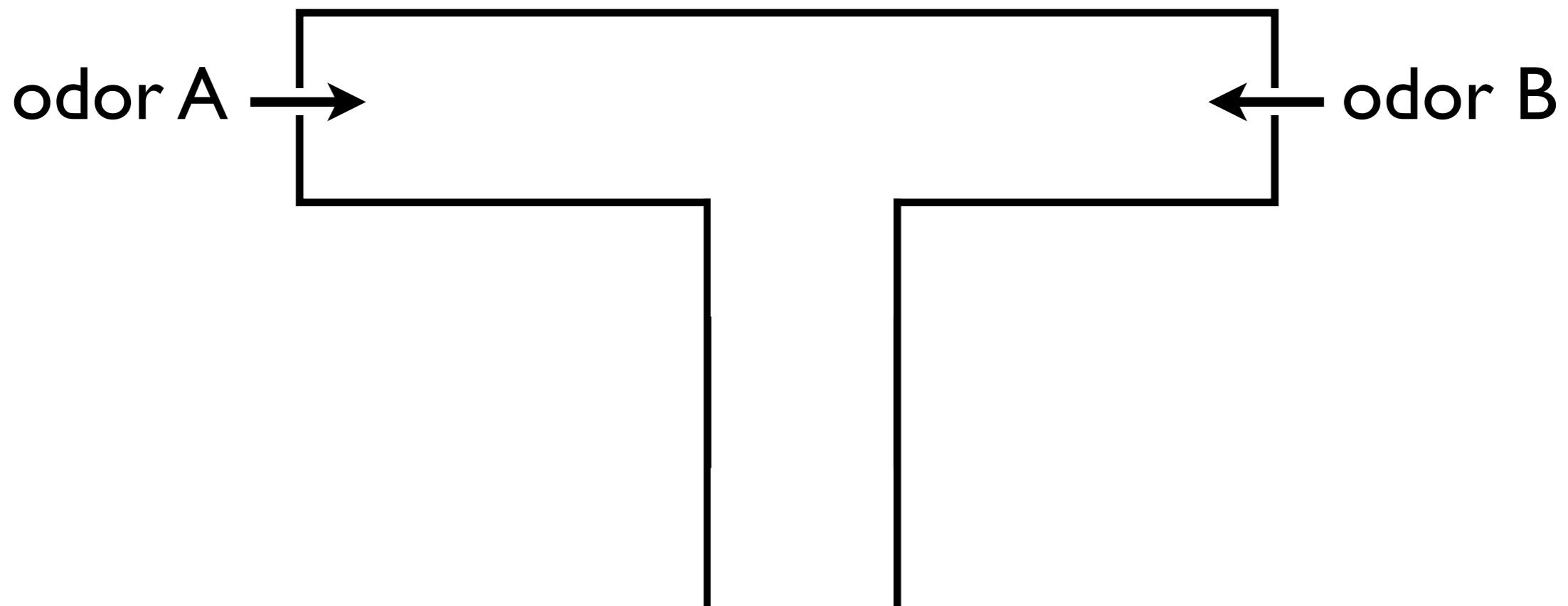
# internal state affects routing

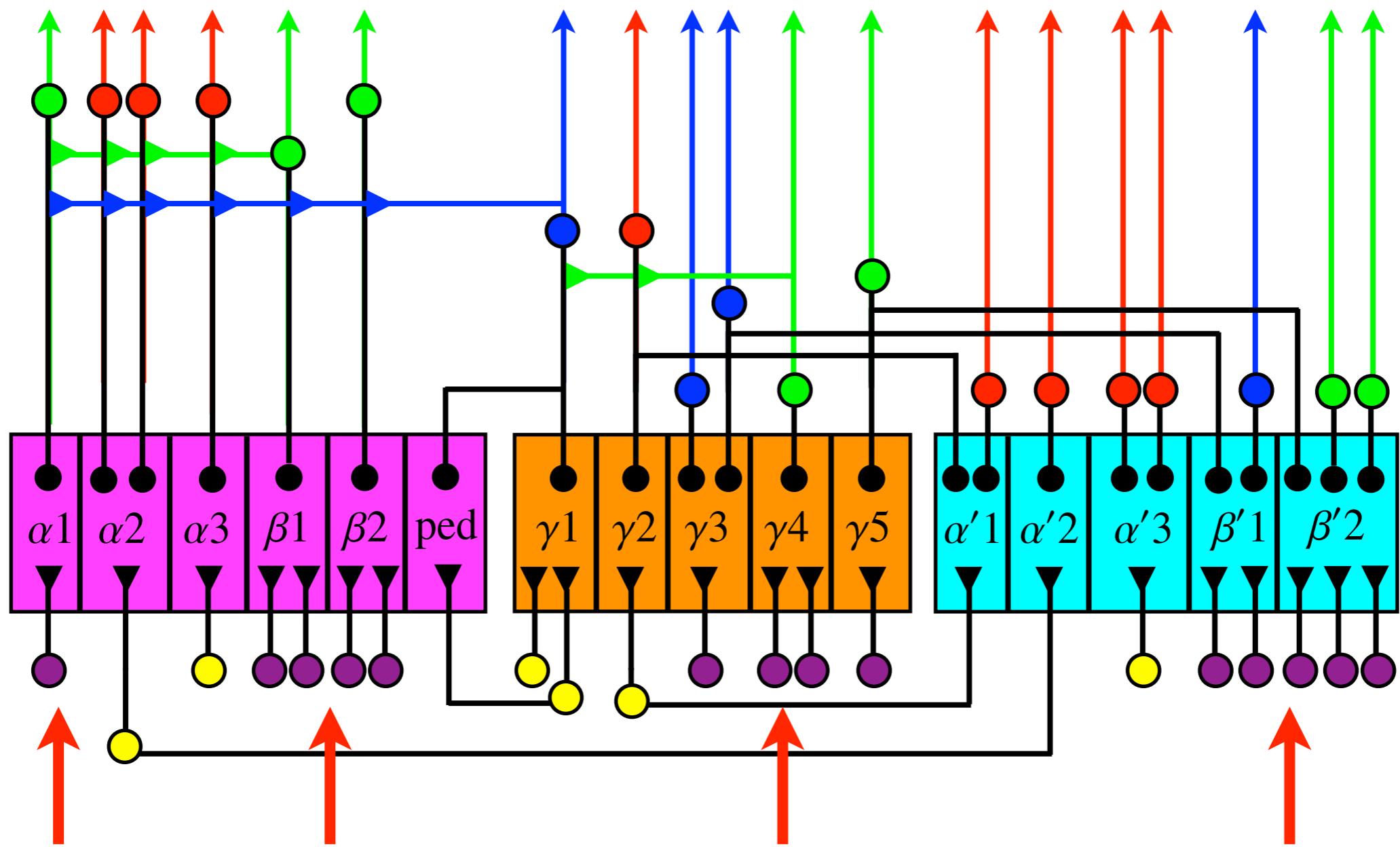


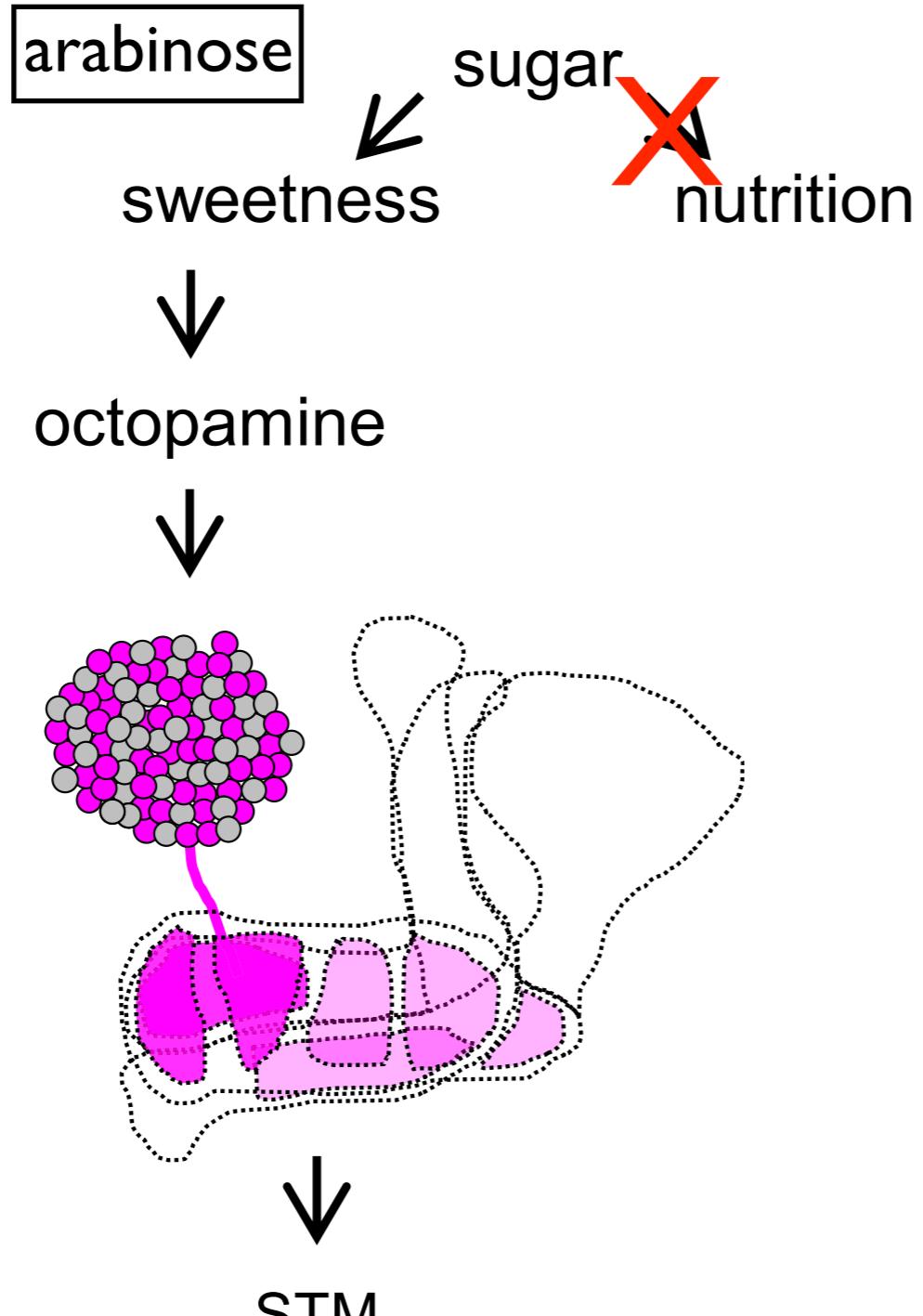
# internal state affects memory

odor A + “sweet”

hungry

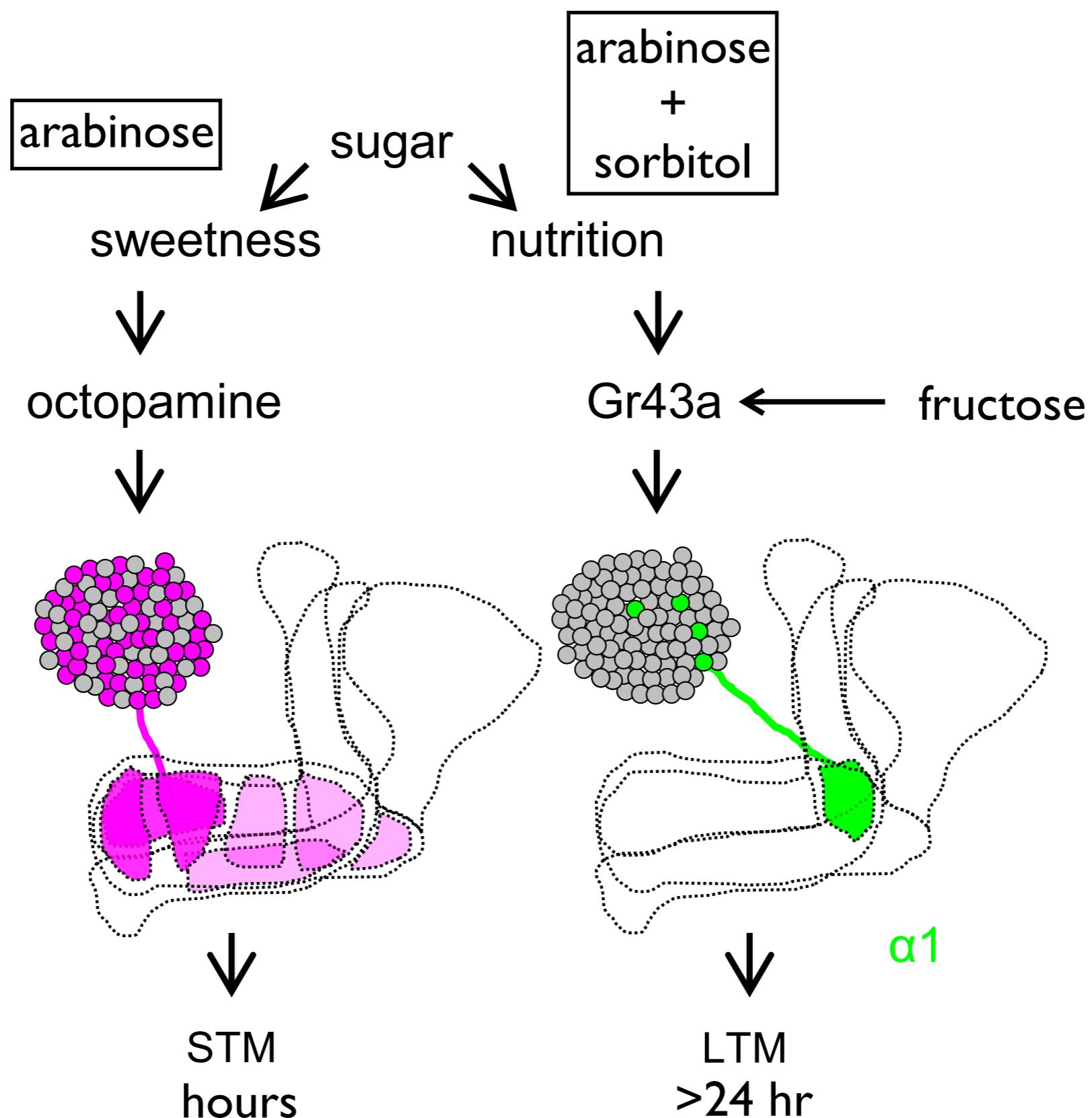






Courtesy of Proceedings of the National Academy of Science. Used with permission  
 Source: Yamagata, Nobuhiro, Toshiharu Ichinose, Yoshinori Aso, Pierre-Yves Plaçais, Anja B. Friedrich, Richard J. Sima, Thomas Preat, Gerald M. Rubin, and Hiromu Tanimoto. "Distinct dopamine neurons mediate reward signals for short-and long-term memories." *Proceedings of the National Academy of Sciences* 112, no. 2 (2015): 578-583.

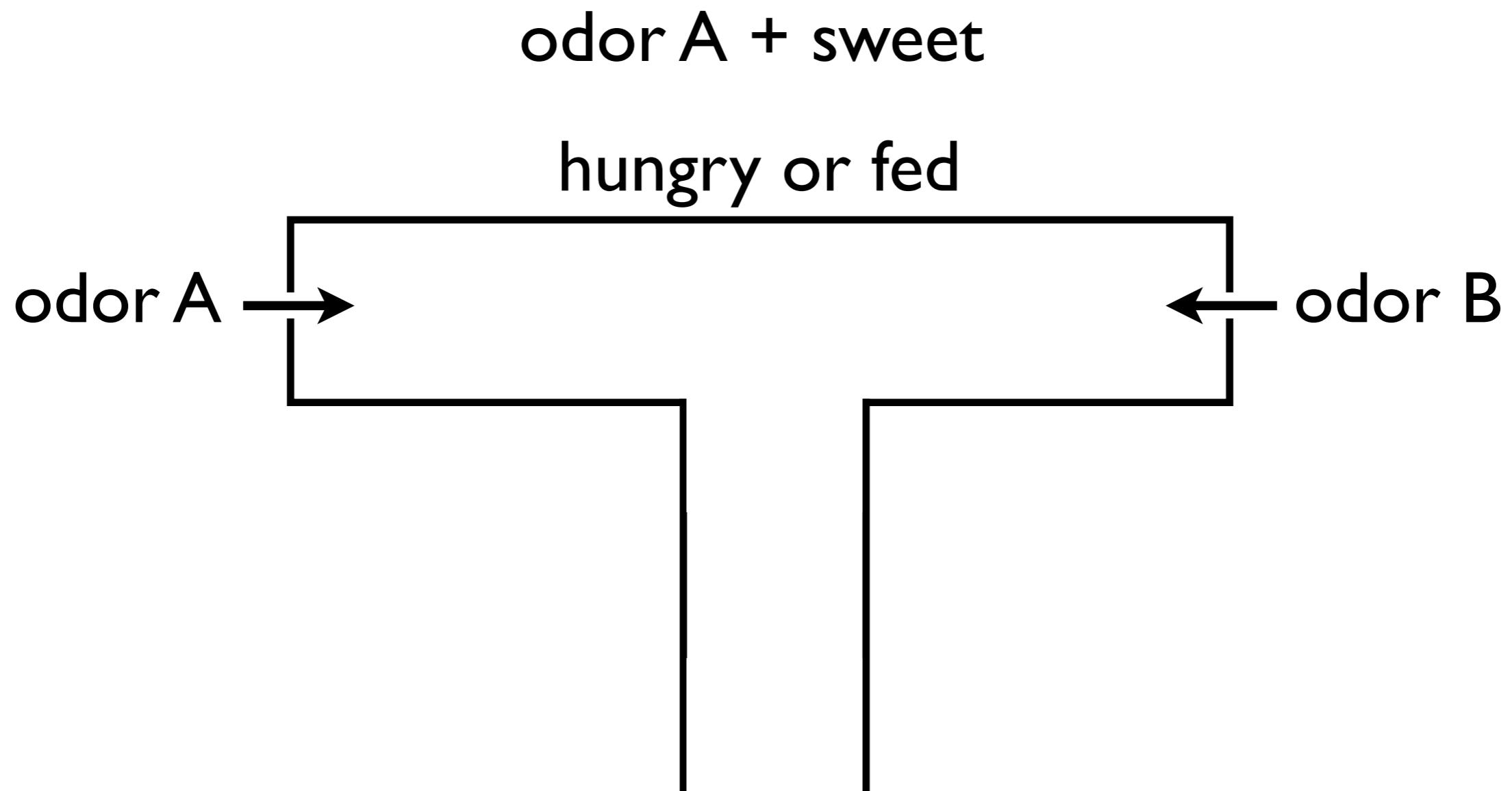
Nobuhiro Yamagata, Toshiharu Ichinose, Yoshinori Aso, Pierre-Yves Plaçais, Anja B. Friedrich, Richard J. Sima, Thomas Preat, Gerald M. Rubin, Hiromu Tanimoto, 2014



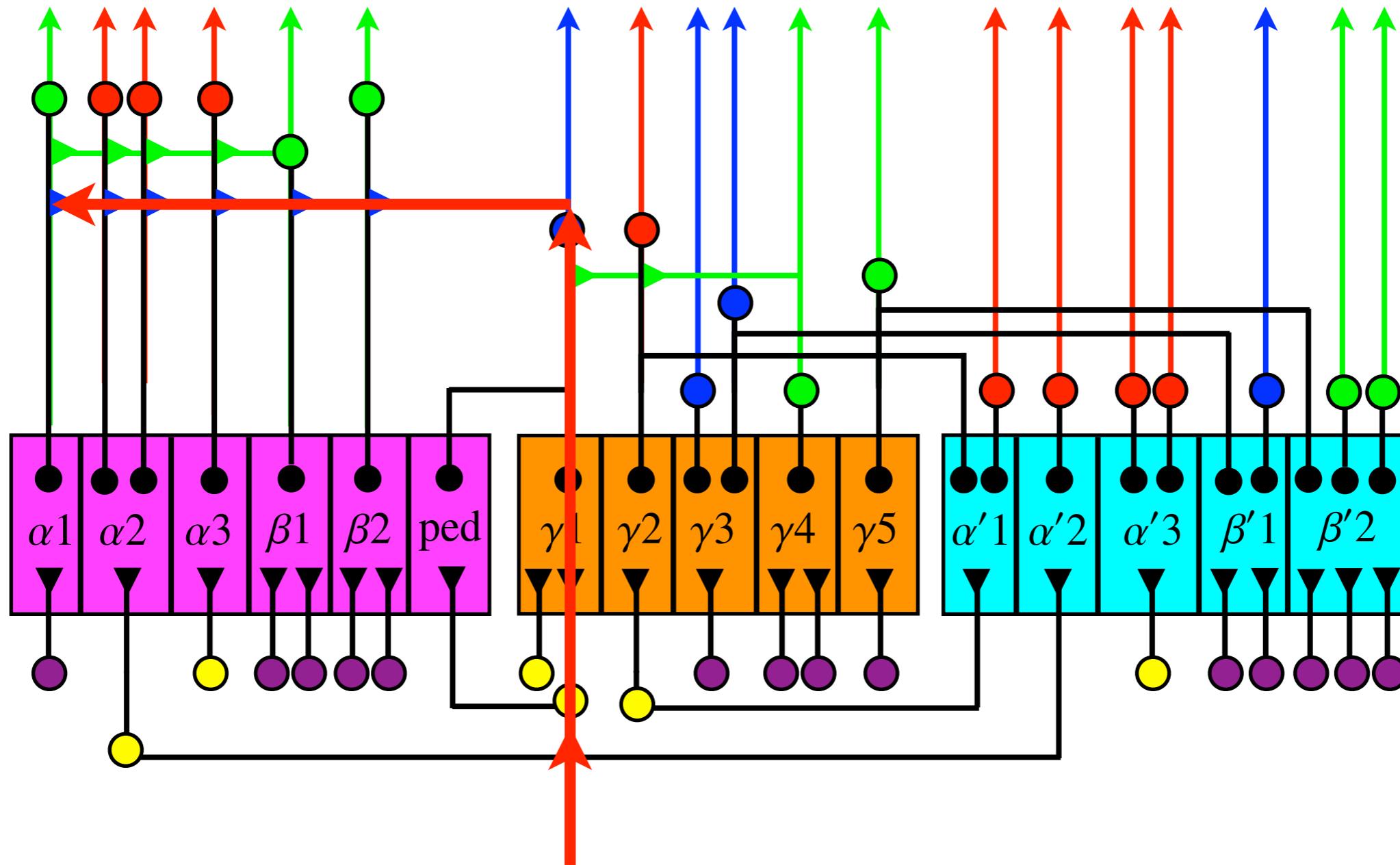
Courtesy of Proceedings of the National Academy of Science. Used with permission  
 Source: Yamagata, Nobuhiro, Toshiharu Ichinose, Yoshinori Aso, Pierre-Yves Plaçais, Anja B. Friedrich, Richard J. Sima, Thomas Preat, Gerald M. Rubin, and Hiromu Tanimoto. "Distinct dopamine neurons mediate reward signals for short-and long-term memories." *Proceedings of the National Academy of Sciences* 112, no. 2 (2015): 578-583.

Nobuhiro Yamagata, Toshiharu Ichinose, Yoshinori Aso, Pierre-Yves Plaçais, Anja B. Friedrich, Richard J. Sima, Thomas Preat, Gerald M. Rubin, Hiromu Tanimoto, 2014

# internal state and gating



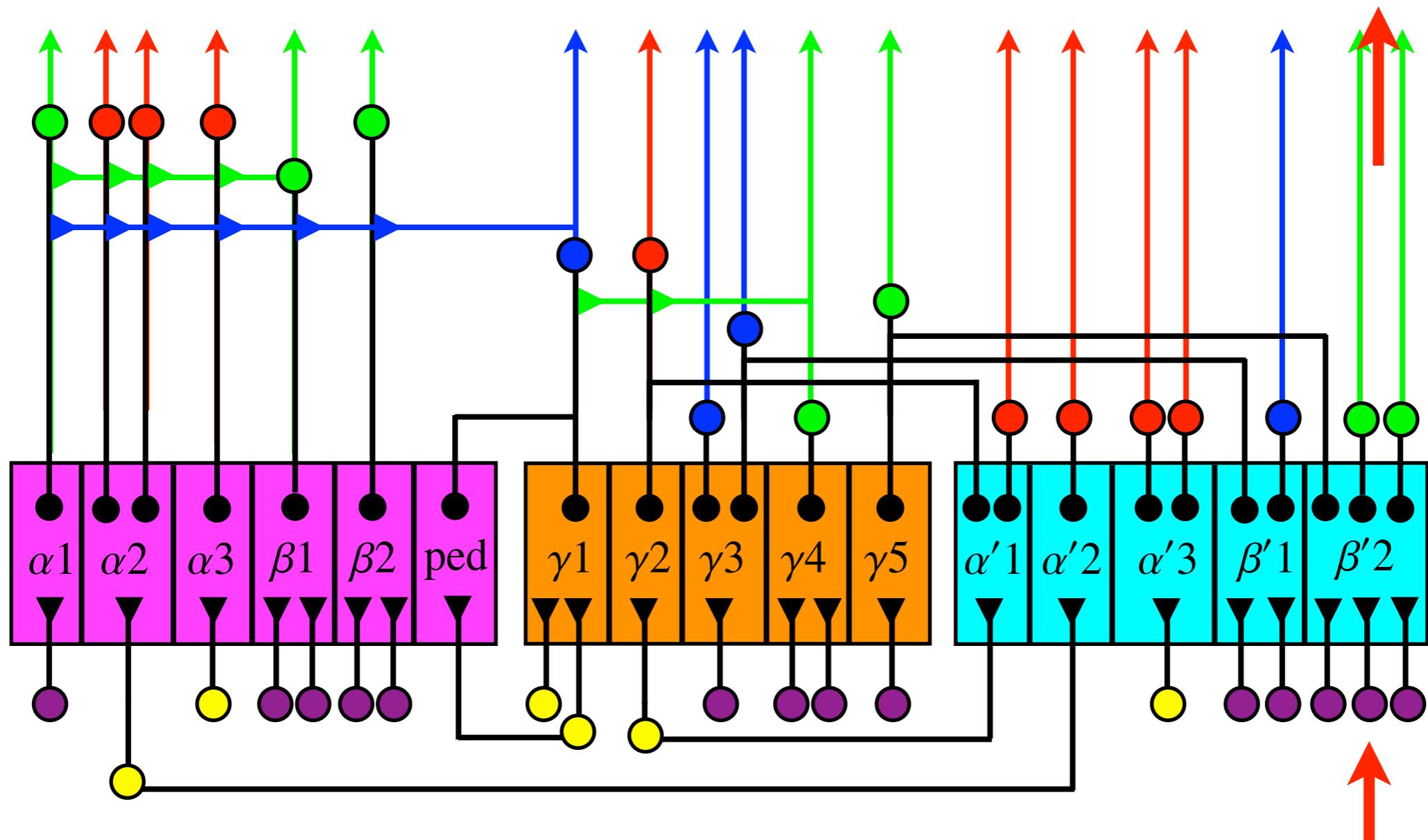
Michael J. Krashes, Shamik DasGupta, Andrew Vreede,  
Benjamin White, J. Douglas Armstrong, Scott Waddell, 2009



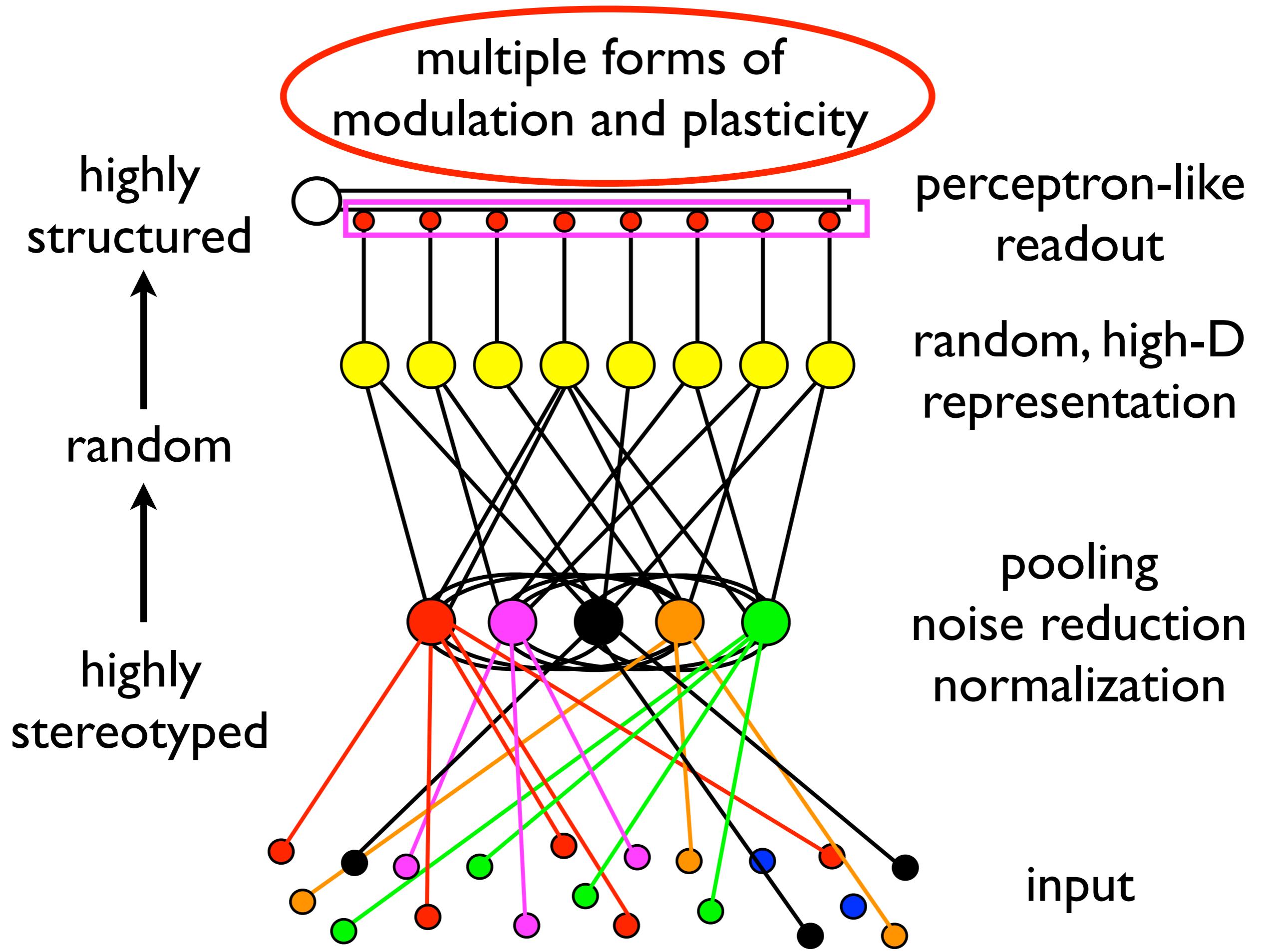
activated in fed state  
silenced in hungry state

Michael J. Krashes, Shamik DasGupta, Andrew Vreede,  
Benjamin White, J. Douglas Armstrong, Scott Waddell, 2009

# CO<sub>2</sub> avoidance



modulation  
by food & hunger



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