



An anatomically accurate circuit for short- and long-term motivational learning in fruit flies

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informatics

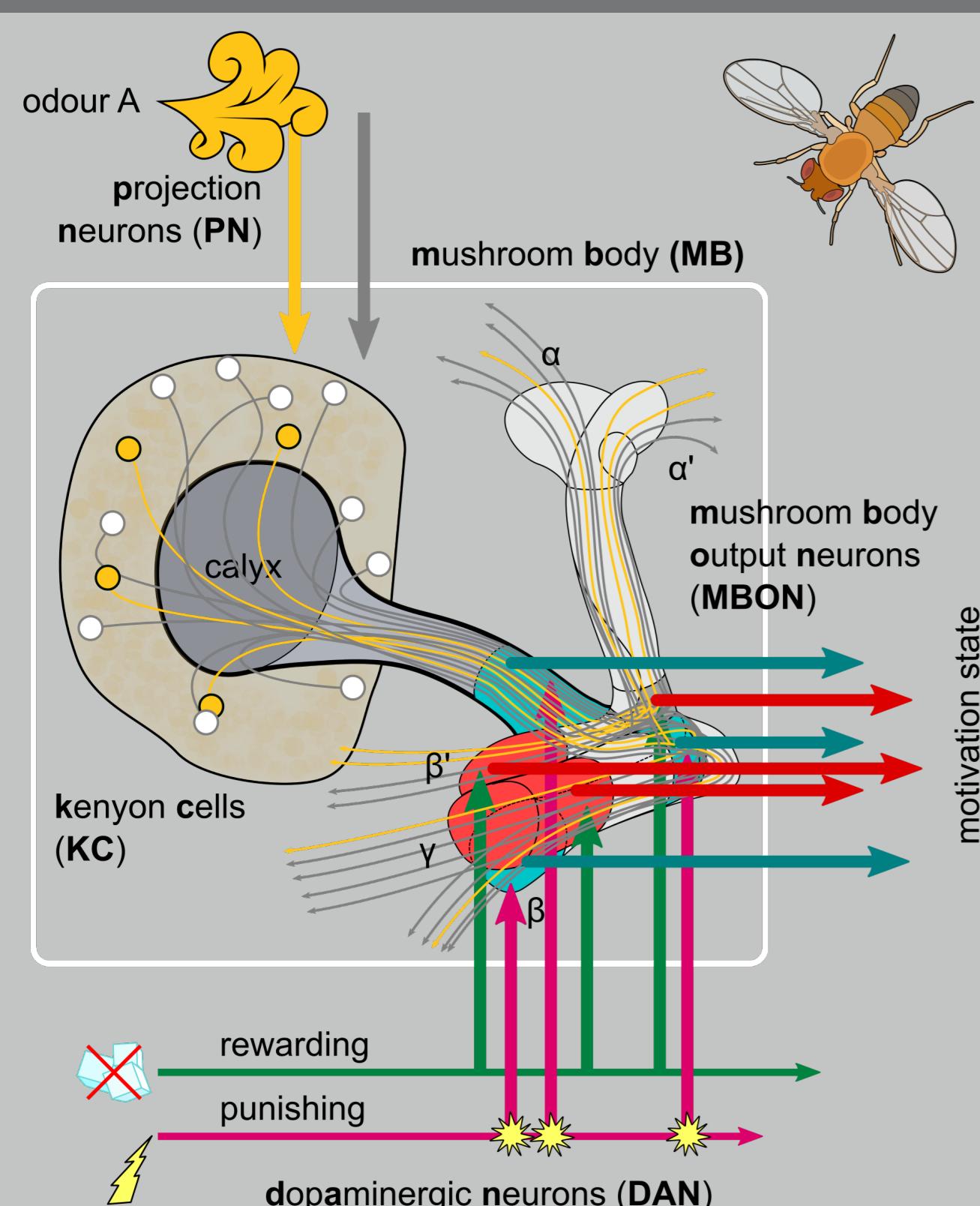


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Abstract

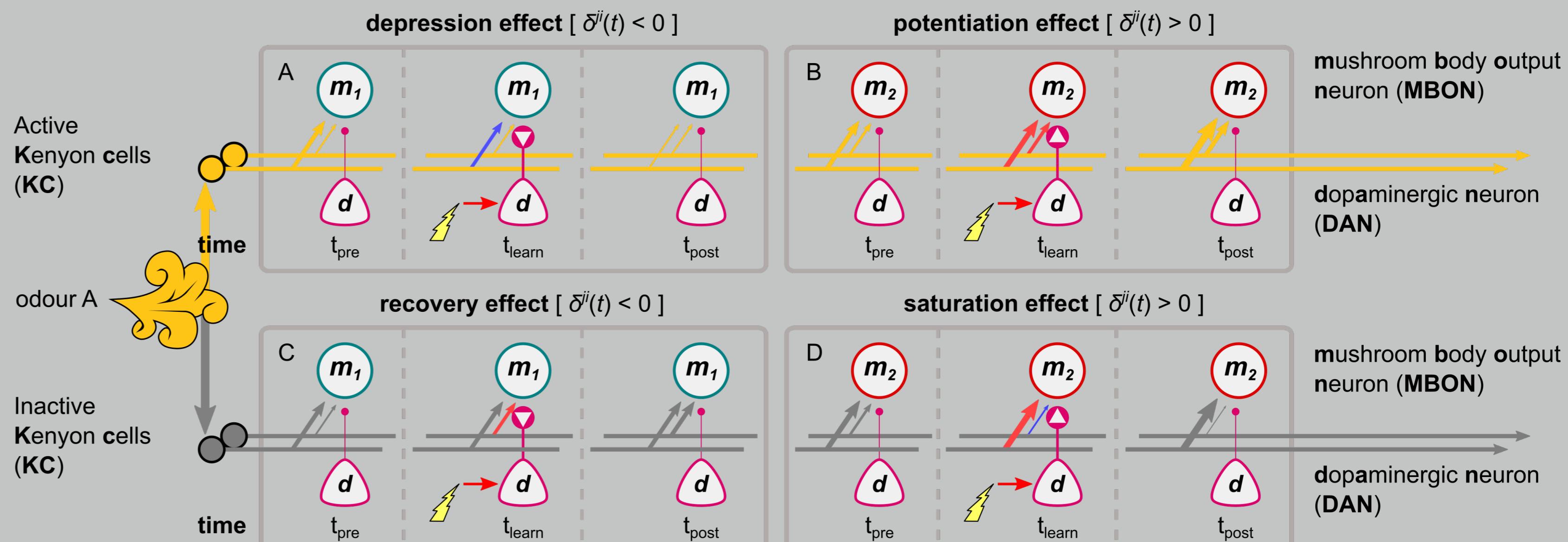
- Key circuits for associative and reinforcement learning have been identified in the mushroom body neuropils of the insect brain [1, 2].
- Detailed imaging, electrophysiological and structural data about the mushroom bodies in *Drosophila melanogaster* has led to the identification of a variety of microcircuits involved in memory.
- In [3], we propose a comprehensive scheme, based on the connectivity and the responses of identified neurons in the mushroom bodies.
 - We link these known microcircuits together as an **incentive circuit** that acquires, forgets and assimilates associative memories over different timescales.
 - We suggest that our novel **dopaminergic plasticity rule** increases the adaptation capabilities of the overall circuit.

About the mushroom bodies



- Sensory input is projected onto the calyxes, from where the numerous kenyon cells (KCs) distribute it to the much fewer output neurons (MBONs).
- Dopaminergic neurons (DANs) deliver multi-dimensional reinforcement signals and modulate the KC-to-MBON synaptic weights.

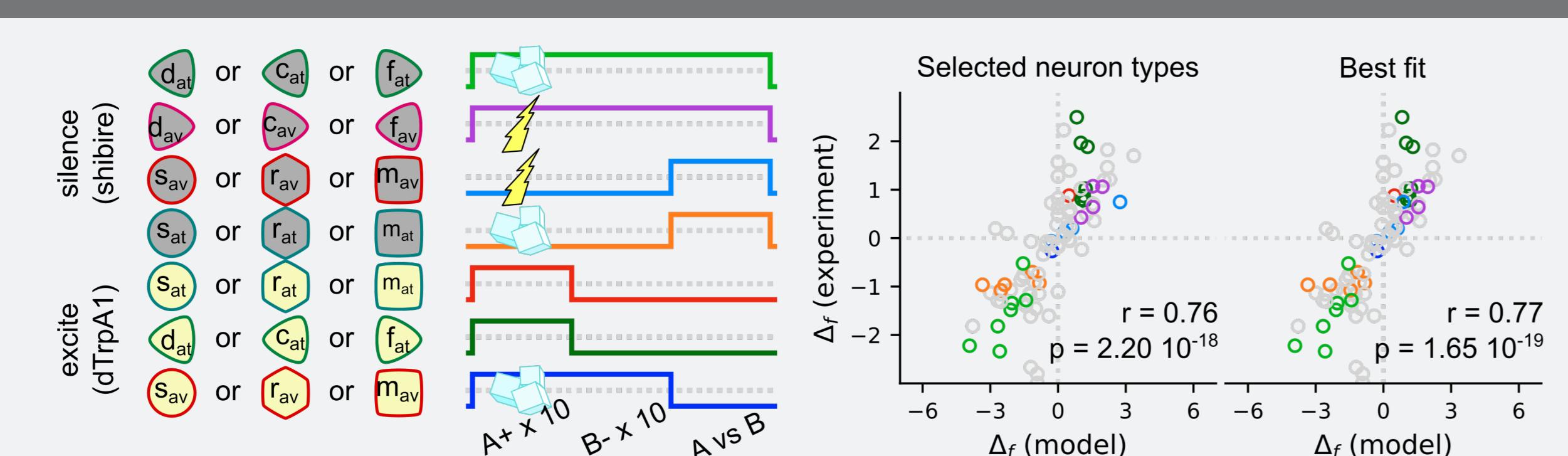
Model: the dopaminergic plasticity rule



- Update of the KC-to-MBON synaptic weights:

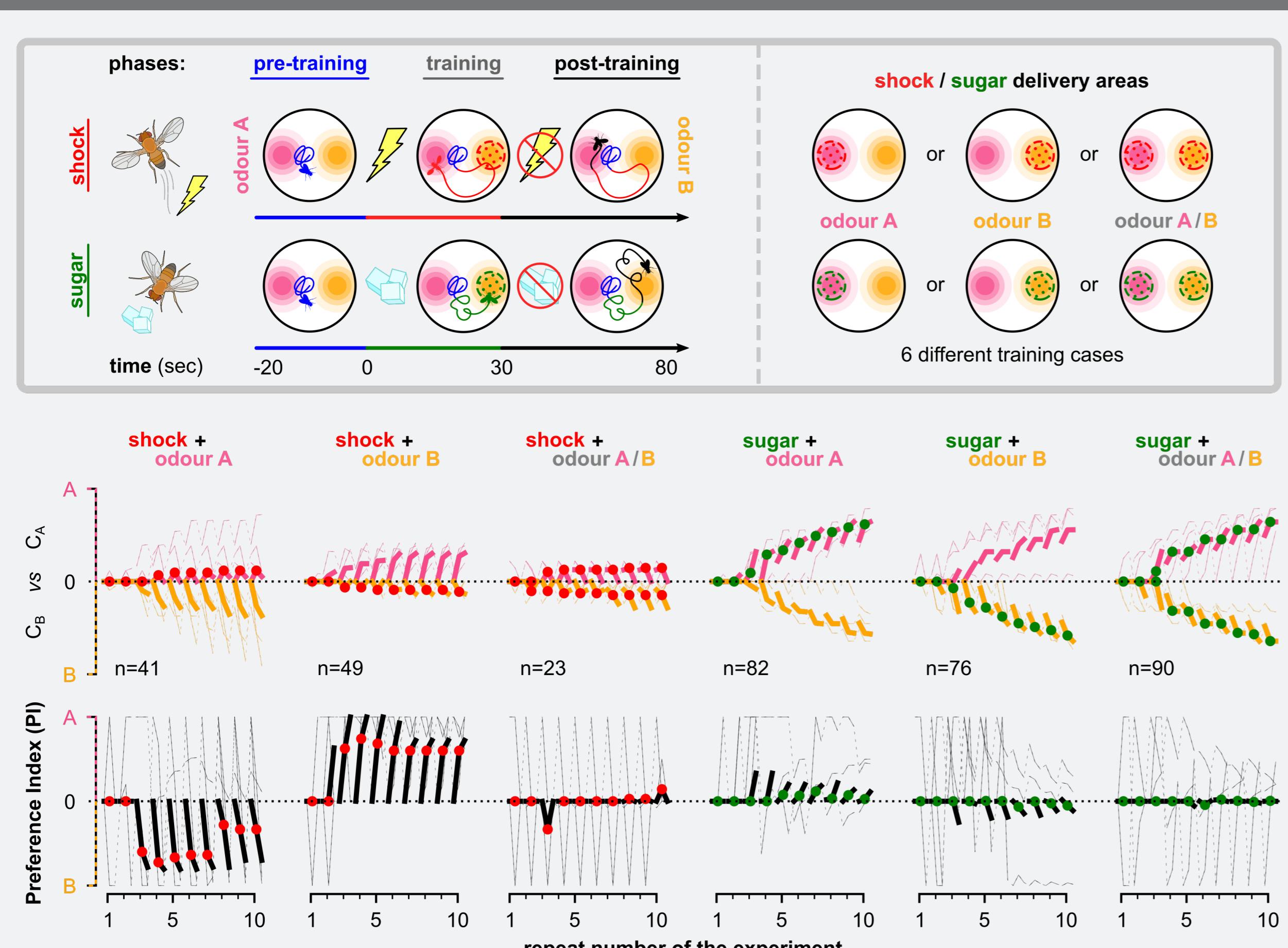
$$\Delta W_{k2m}^{ij}(t) = \delta^{ij}(t)[k^i(t) + W_{k2m}^{ij}(t) - w_{rest}]$$
- $\delta^{ij}(t) = D_{\Delta}^{ij}(t) - D_{\nabla}^{ij}(t)$ controls the learning at the KCⁱ-to-MBON^j synapse.
- $k^i(t)$ is the KC activity.
- $w_{rest} = 1$ is the resting value.
- $D_{\Delta}^{ij}(t)$ and $D_{\nabla}^{ij}(t)$ are components of the dopaminergic signal, with a short and a long time-constant respectively, that are key for explaining backward learning [5].
- Depending on conditions, this synaptic modulation causes weights to stabilise or to increase with positive feedback
- We exploit these properties to enable short- and long-term memories, respectively, to be formed and forgotten in parallel for different contexts.

Results: classical olfactory conditioning



- Our predictions correlate with 92 intervention experiments from 14 studies (data and method from [4]).

Results: modelling the behaviour

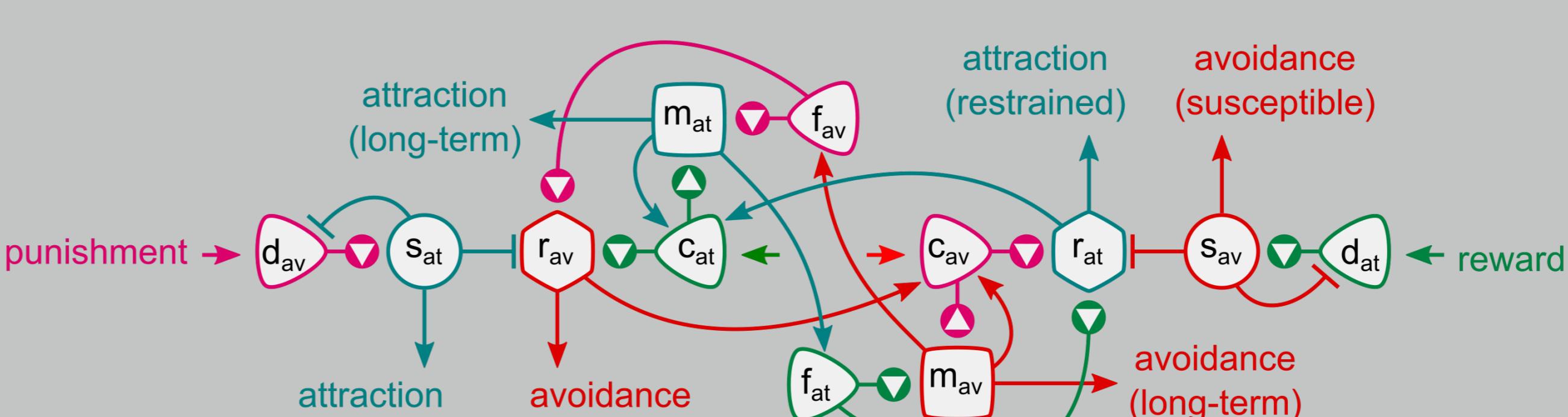


- We further verify the function of the incentive circuit by demonstrating how the reproduced responses of the output neurons could drive the behaviour of a virtual fruit fly, creating similar odour preferences to the real flies.

References

- [1] Y. Aso, D. Hattori, Y. Yu, et al., "The neuronal architecture of the mushroom body provides a logic for associative learning," *eLife*, vol. 3, p. e04577, 2014.
- [2] F. Li, J. W. Lindsey, E. C. Marin, et al., "The connectome of the adult *Drosophila* mushroom body provides insights into function," *eLife*, vol. 9, p. e62576, 2020.
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- [4] J. E. M. Bennett, A. Philippides, and T. Nowotny, "Learning with reinforcement prediction errors in a model of the *Drosophila* mushroom body," *Nature Communications*, vol. 12, no. 1, p. 2569, 2021.
- [5] A. Handler, T. G. Graham, R. Cohn, et al., "Distinct Dopamine Receptor Pathways Underlie the Temporal Sensitivity of Associative Learning," *Cell*, vol. 178, no. 1, pp. 60–75.e19, 2019.

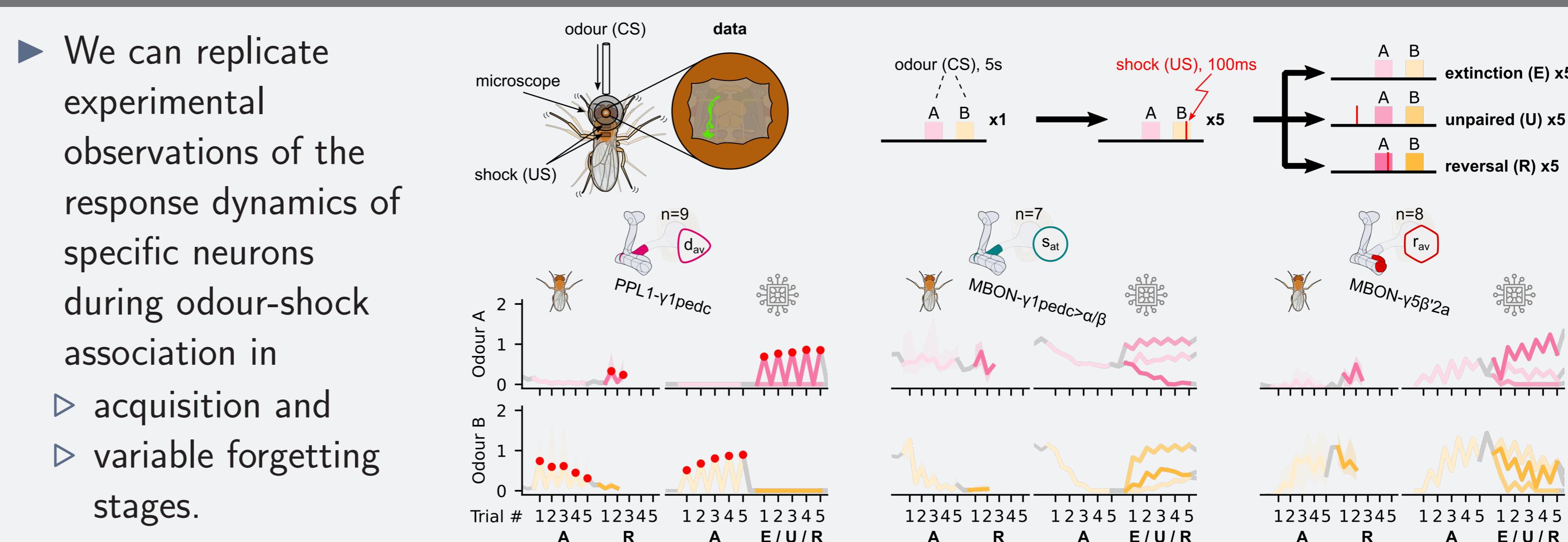
Model: the incentive circuit



- Susceptible (s) MBONs underlie primitive memories.
- Restrained (r) MBONs underlie flexible memories.
- Long-term memory (m) MBONs.
- Anatomically validated: each connection exists in the fly brain.
- Discharging (d) DANs.
- Charging (c) DANs.
- Forgetting (f) DANs.
- Note: KCs are not shown.

Results: modelling the neural responses

- We can replicate experimental observations of the response dynamics of specific neurons during odour-shock association in
 - acquisition and
 - variable forgetting stages.



Conclusion

- The dopaminergic plasticity rule is an alternative to prediction error plasticity rules, and within the incentive circuit can support acquisition, forgetting and assimilation of memories.
- Different MBONs hold primitive, flexible or long-term memories, supporting flexible exploration/exploitation trade-off in an olfactory conditioning task.

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