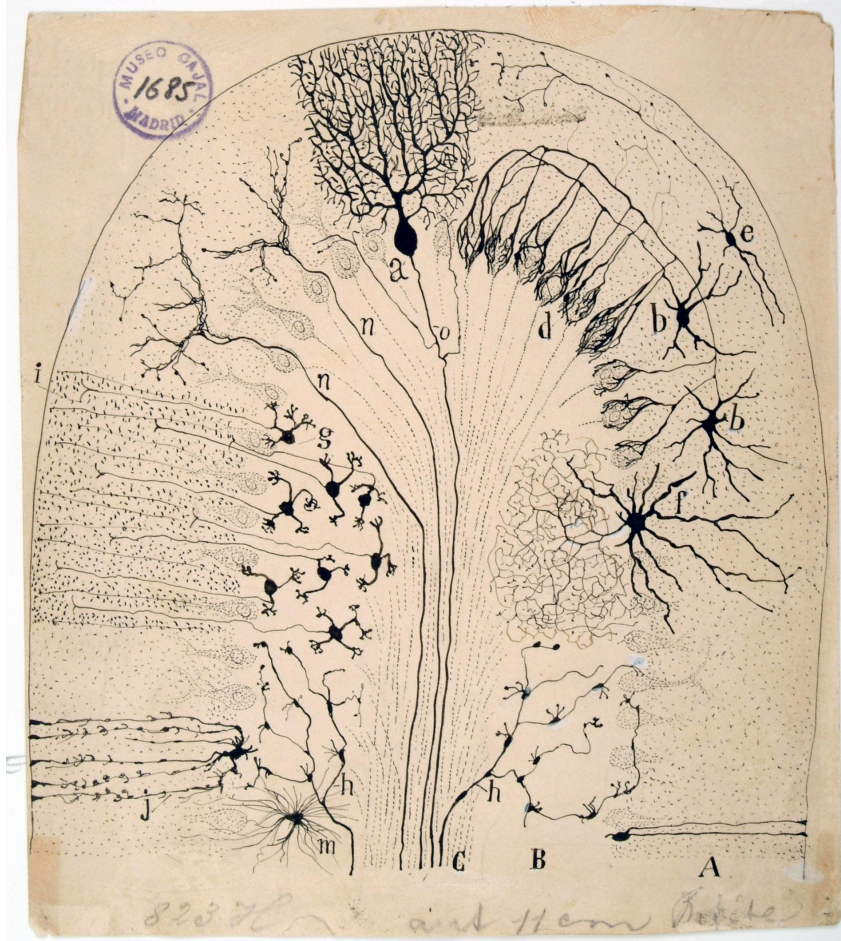


Circuits and Mechanistic Neuroscience

Adam Marblestone

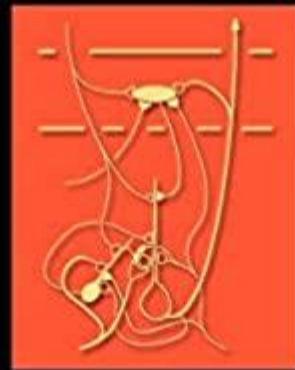
Submit questions here:

<https://app.sli.do/event/92gy6nuo>



The Synaptic Organization of the Brain

FIFTH EDITION



EDITED BY

Gordon M. Shepherd

Do we know *in circuit-level detail* how brain implements at least one ML algorithm?

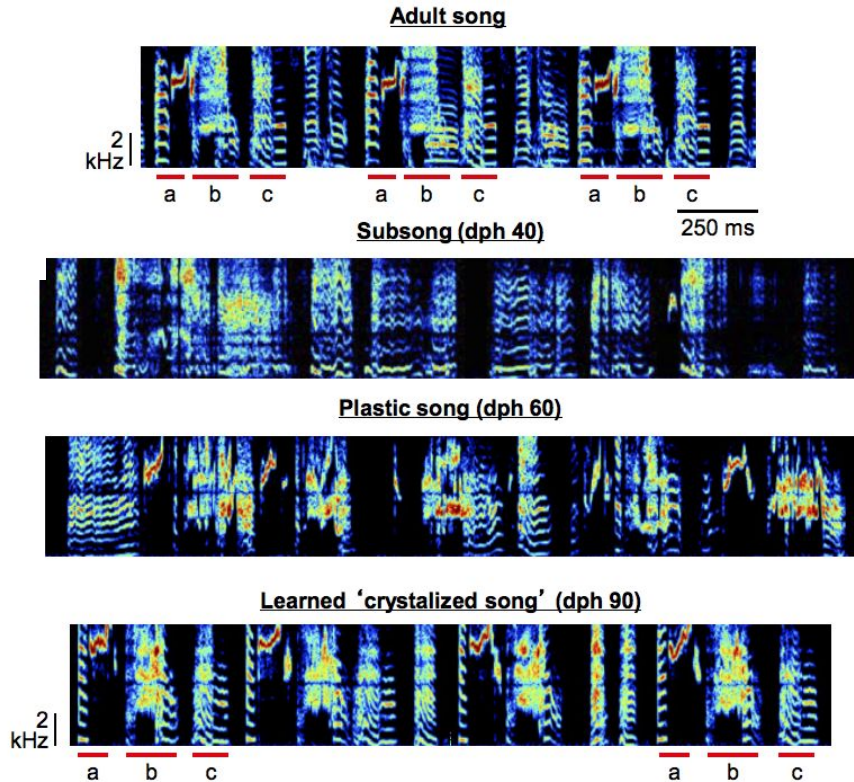
Do we know *in circuit-level detail* how brain implements at least one ML algorithm?

Yes: RL

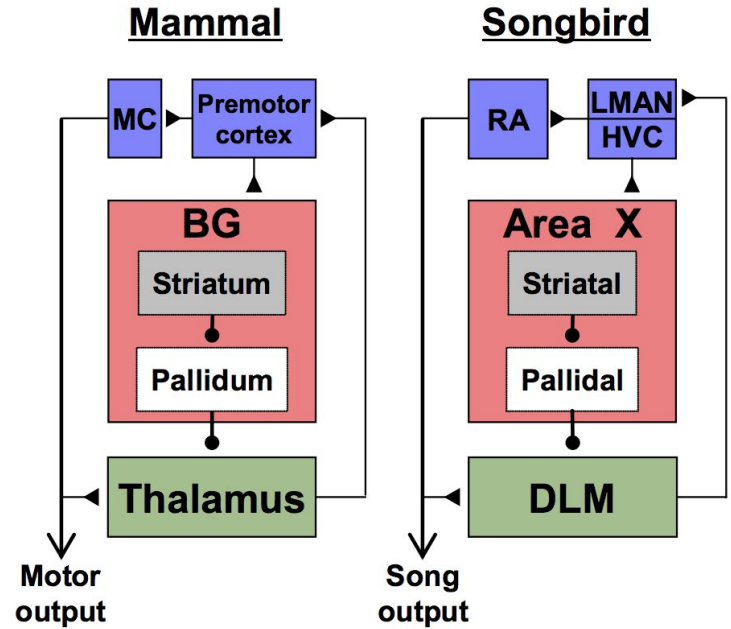
Do we know *in circuit-level detail* how brain implements at least one ML algorithm?

Yes: RL (or at least a simple instance)

Circuit-level understand of RL

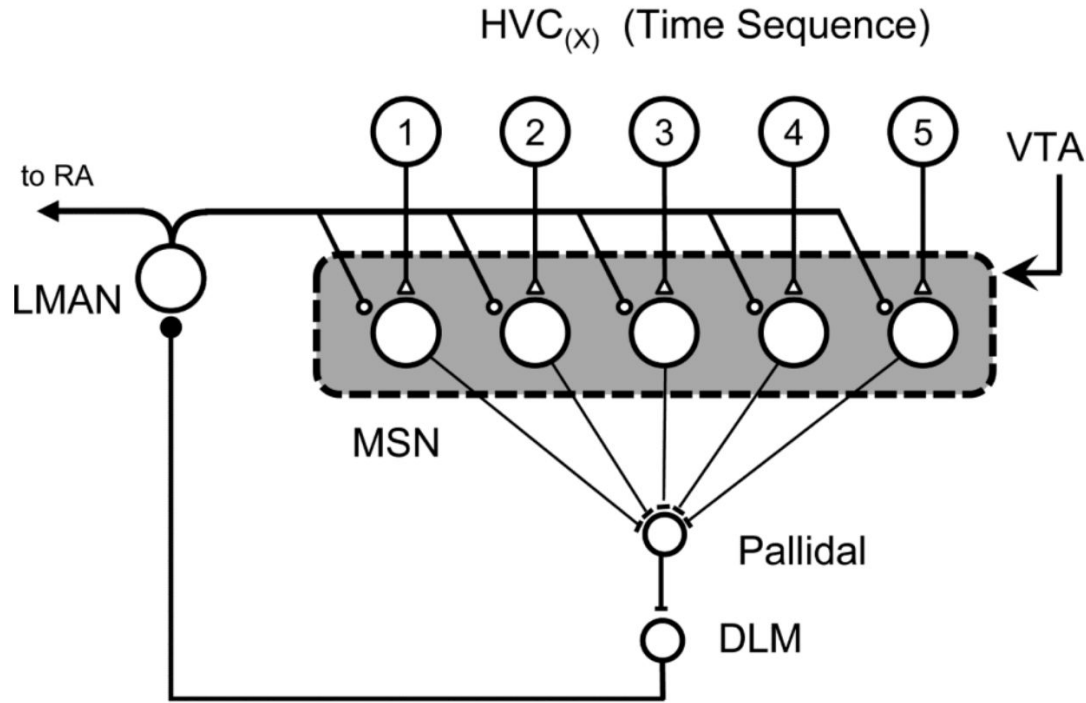


Variability
 Similarity to tutor



A hypothesis for basal ganglia-dependent reinforcement learning in the songbird

Circuit-level understand of RL

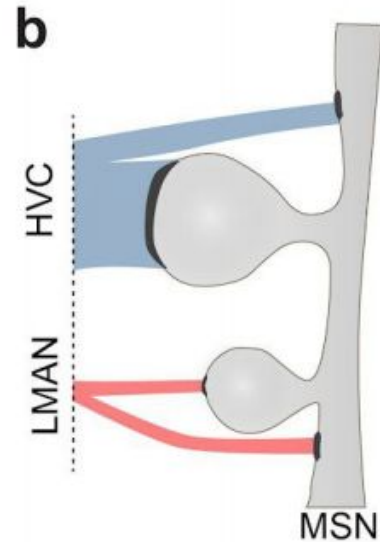
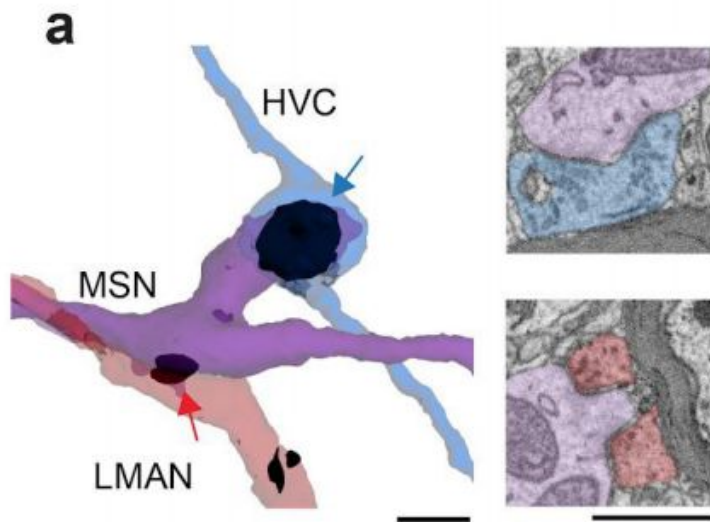


“Node perturbation” in each of many parallel “channels” through striatum

Ultimately “consolidated” into cortico-cortical connections

A hypothesis for basal ganglia-dependent reinforcement learning in the songbird

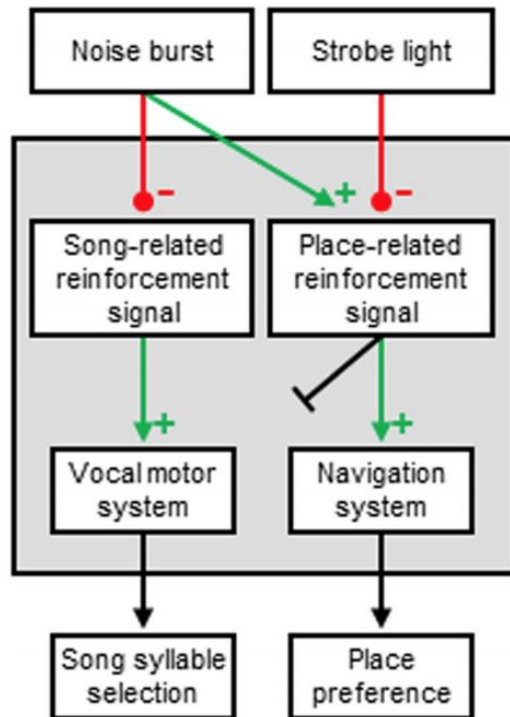
Circuit-level understand of RL



Deep synapse-level circuit reconstructions validate predictions of this theory

An anatomical substrate of credit assignment in reinforcement learning

RL in the (songbird) brain does not use one monolithic reward



Spatially segregated circuits see distinct reinforcers

Reinforcers are internally-generated
(e.g., based on comparison to stored tutor song template)

Place preference and vocal learning rely on distinct reinforcers in songbirds

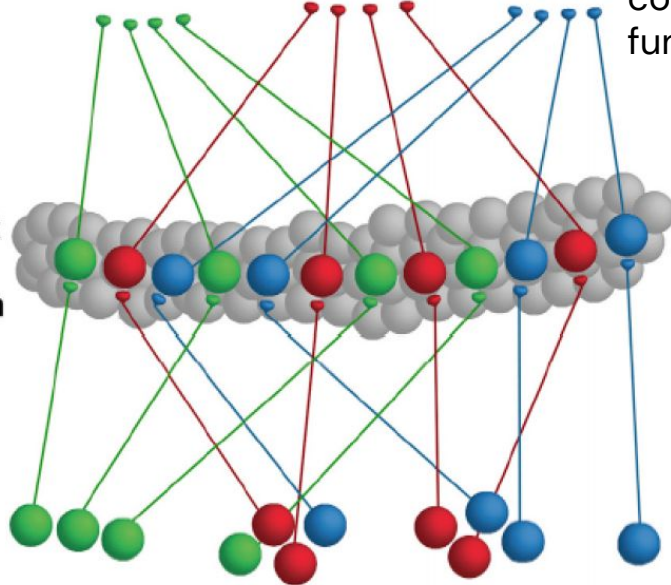
What about in mammals?

EnvA- Δ G-rabies infection

prefrontal cortex basolateral amygdala motor cortex

Various neural pathways such as the acetylcholine system could support something like multiple cost or reward functions applied to different sub-circuits.

cre-dependent viral spread in basal forebrain



A Cholinergic Mechanism for Reward Timing within Primary Visual Cortex

Alexander A. Chubykin³, Emma B. Roach³, Mark F. Bear^{1,2}, Marshall G. Hussain Shuler^{1,2}
³ These authors contributed equally to this work

Nucleus basalis-enabled stimulus-specific plasticity in the visual cortex is mediated by astrocytes (i.e., glia not neurons)

Naiyan Chen^{a,1}, Hiroki Sugihara^{a,1}, Jitendra Sharma^{a,b}, Gertrudis Perea^a, Jeremy Petracvic^a, Chuong Le^a, and Mriganka Sur^{a,2}

afferent labeling

lateral septum

central amygdala

caudate putamen

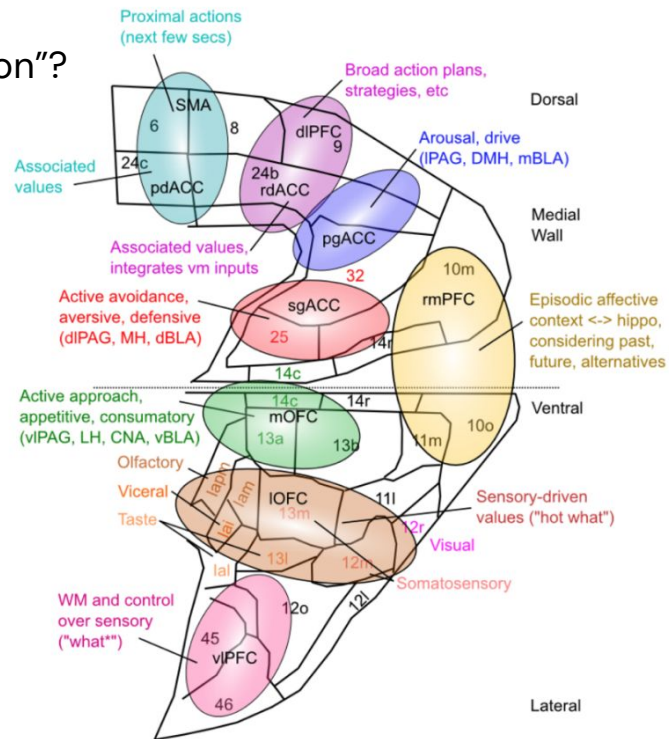
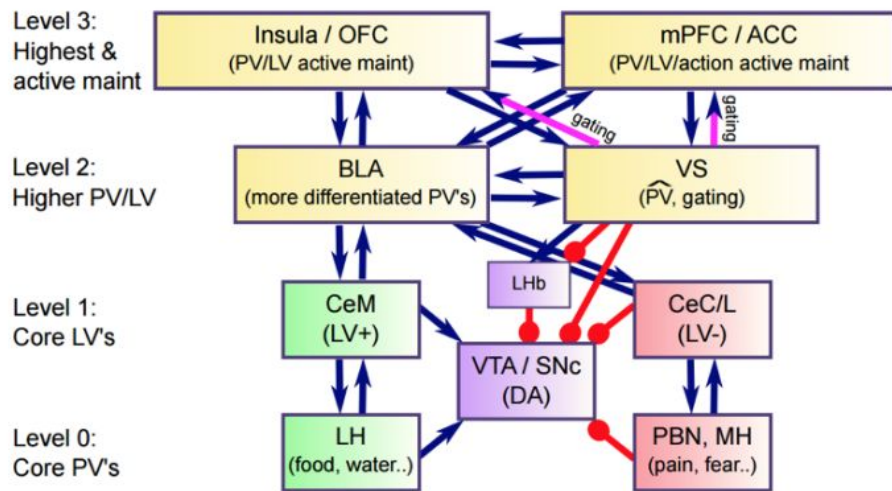
somato-sensory cortex

The Input-Output Relationship of the Cholinergic Basal Forebrain

Matthew R. Gielow and Laszlo Zaborszky

What about in mammals?

Complex cortical networks feeding into something like a “value function”?



Do we know *in circuit-level detail* how brain implements a multi-layer credit assignment mechanism of similar power to backprop?

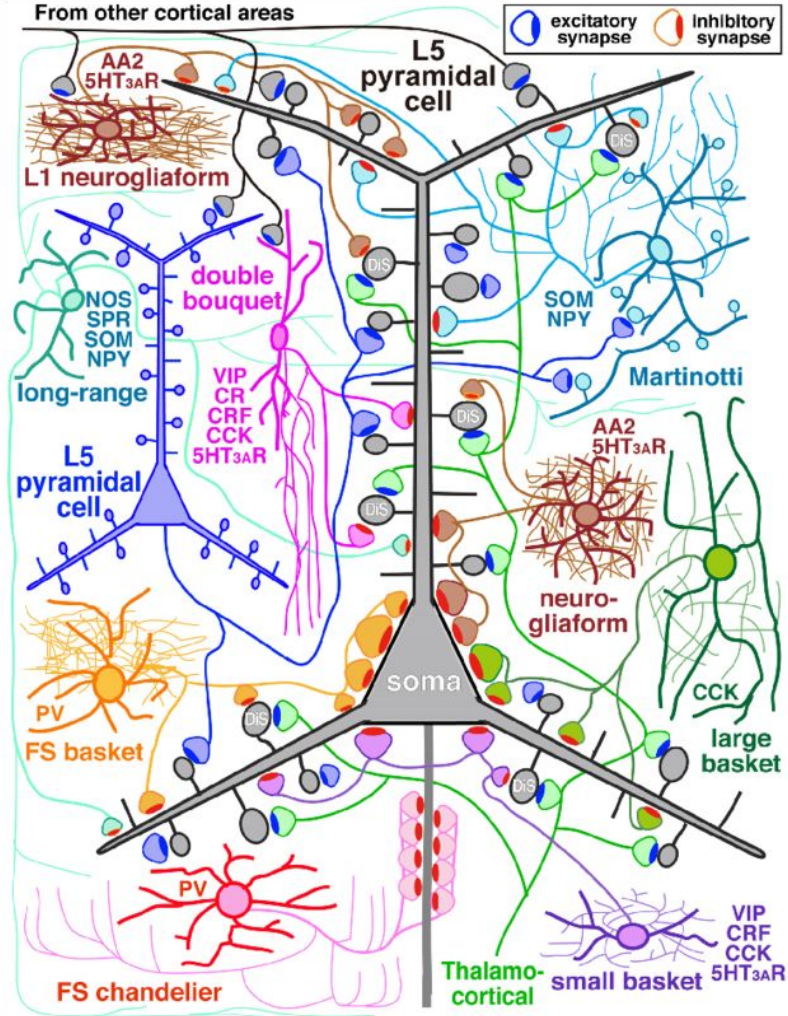
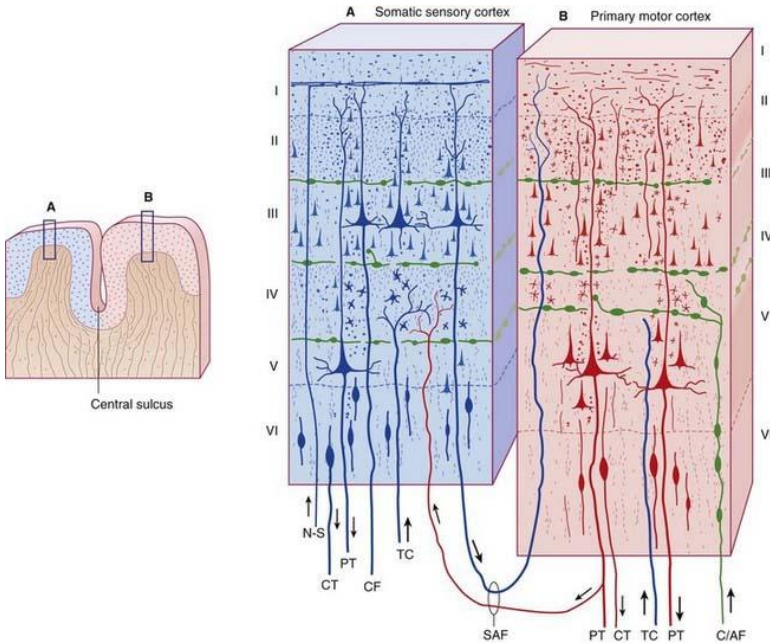
Do we know *in circuit-level detail* how brain implements a multi-layer credit assignment mechanism of similar power to backprop?

Trick question: not sure it does at all

Do we know *in circuit-level detail* how brain implements a multi-layer credit assignment mechanism of similar power to backprop?

Trick question: but there are ideas...

Does the need to implement a powerful learning algorithm underly the cellular complexity of cortical micro-circuits?



Does the need to implement a powerful learning algorithm underly the cellular complexity of cortical micro-circuits?

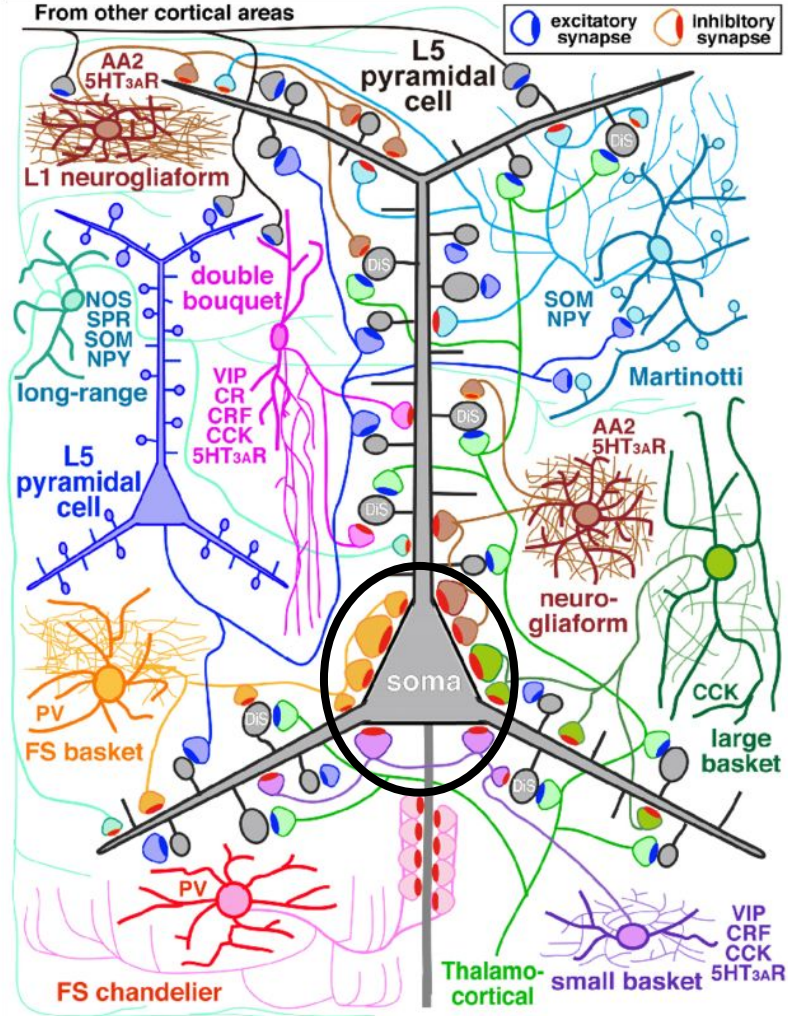
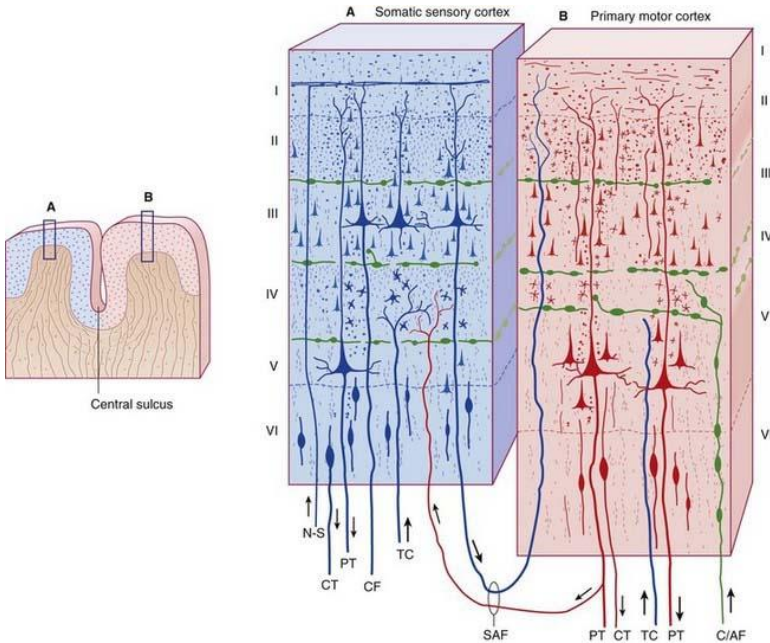


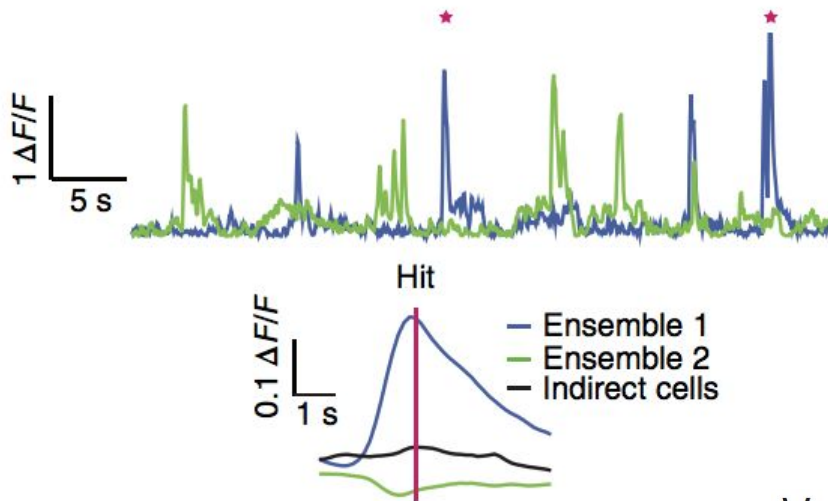
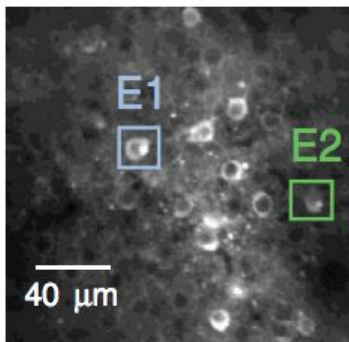
Table 1. Comparison of Models

		Temporal-error model		Explicit-error model	
		Contrastive learning	Continuous update	Predictive coding	Dendritic error
Properties ^a	Control signal	Required	Required	Not required	Not required
	Connectivity	Unconstrained	Unconstrained	Constrained	Constrained
	Propagation time	L-1	L-1	2L-1	L-1
	Pre-training	Not required	Not required	Not required	Required
Error encoded in		Difference in activity between separate phases	Rate of change of activity	Activity of specialised neurons	Apical dendrites of pyramidal neurons
Data accounted for		Neural responses and behaviour in a variety of tasks	Typical spike-time-dependent plasticity	Increased neural activity to unpredicted stimuli	Properties of pyramidal neurons
MNIST performance ^b		~2-3	-	~1.7	~1.96

Review

Theories of Error Back-Propagation in the Brain

Can such ideas be validated?



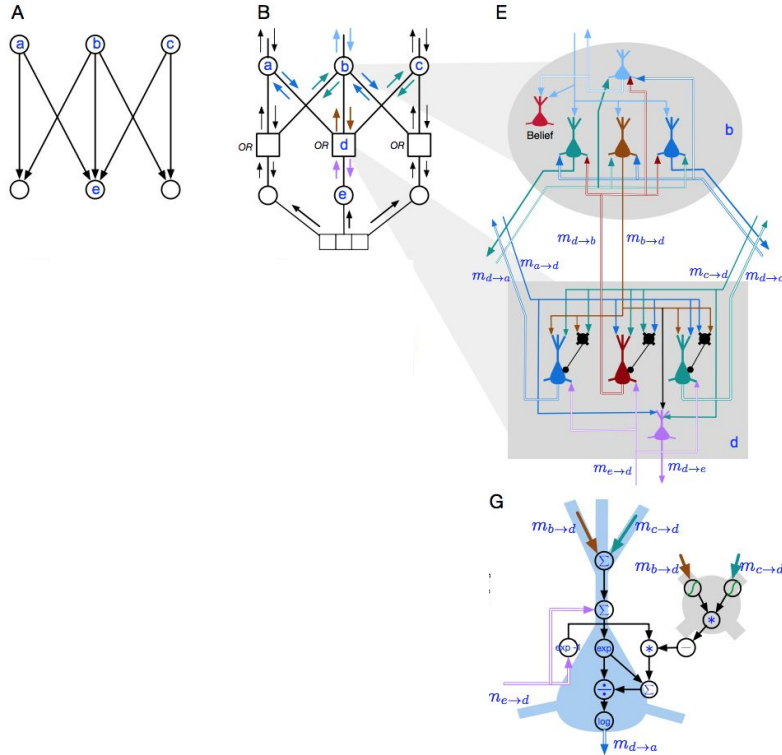
Clever experimental design with “brain computer interface” experiment in animals might allow us to tease out credit assignment mechanisms from *movies of neurons firing (calcium imaging)*

Rewards are made contingent on the firing of particular neurons on the circuit

Volitional modulation of optically recorded calcium signals during neuroprosthetic learning

Kelly B Clancy^{1,7}, Aaron C Koralek^{2,7}, Rui M Costa³,
Daniel E Feldman^{2,4} & Jose M Carmena^{2,5,6}

Other interpretations of cortical circuit



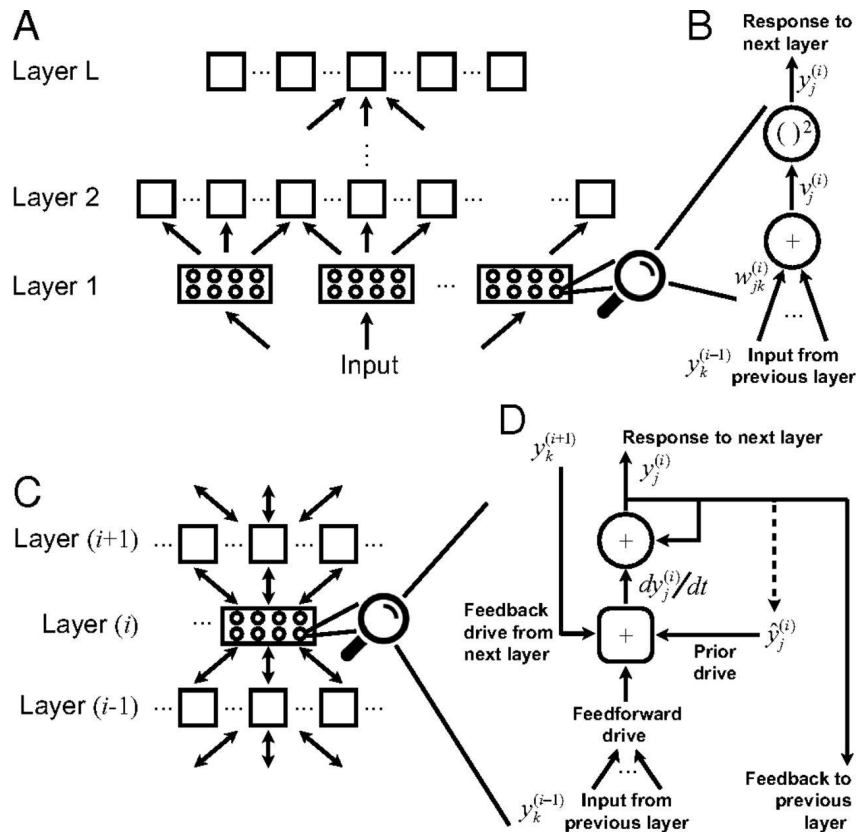
In other interpretations, cortical circuit architecture is determined more by the need for complex inference (e.g., as in message passing algorithms for graphical models of particular structures)

Rather than by the need for weight transport for deep learning-like multilayer credit assignment

A detailed mathematical theory of thalamic and cortical microcircuits based on inference in a generative vision model

 Dileep George,  Miguel Lázaro-Gredilla, Wolfgang Leirach, Antoine Dedieu, Guangyao Zhou

Other interpretations of cortical circuit

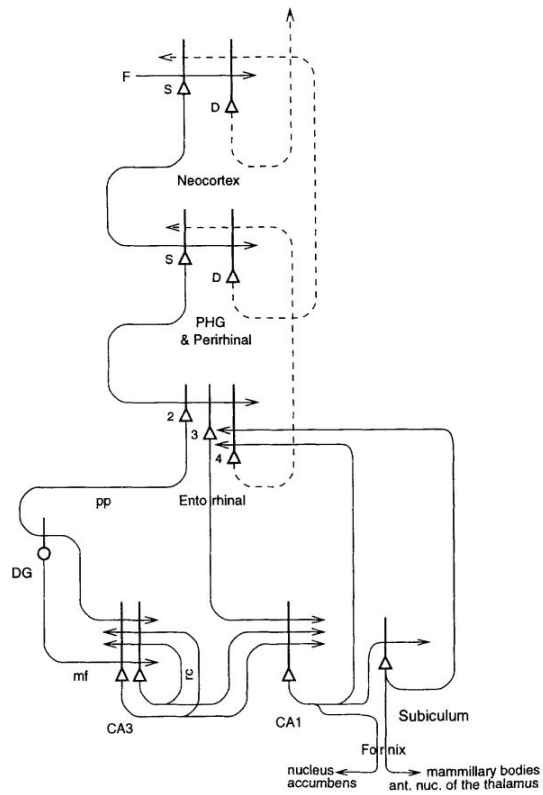
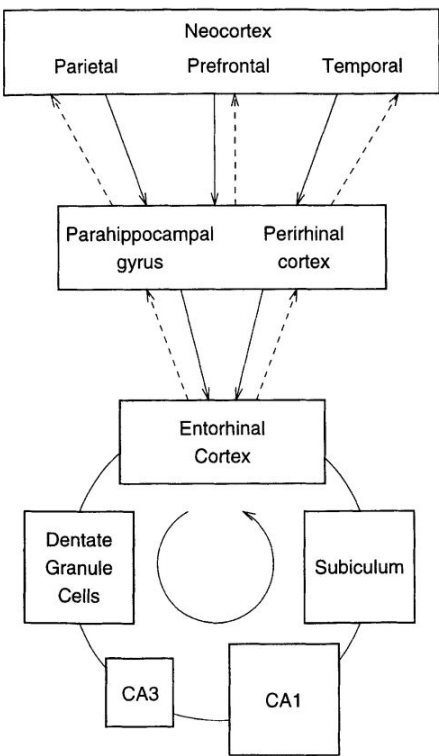


In yet other interpretations, cortical circuits reflect energy-based models that incorporate both top-down prior information and bottom-up sensory information into the energy function

Such models are meant to explain perceptual phenomena in a "perception as inference" framework

Theory of cortical function

Content-addressable memory in brain



The hippocampus looks like a content-addressable memory system taking input from many higher cortical areas

Pattern separation (dentate gyrus), *pattern completion* (CA3), and *reconstruction* (CA1)

Not just for spatial navigation...

A Theory of Hippocampal Function in Memory

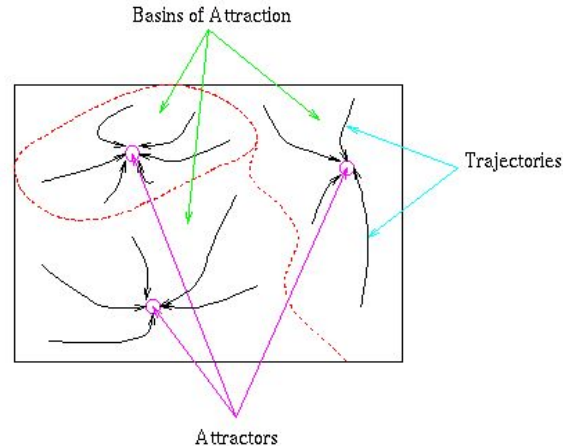
Edmund T. Rolls

Content-addressable memory in brain

Abstract

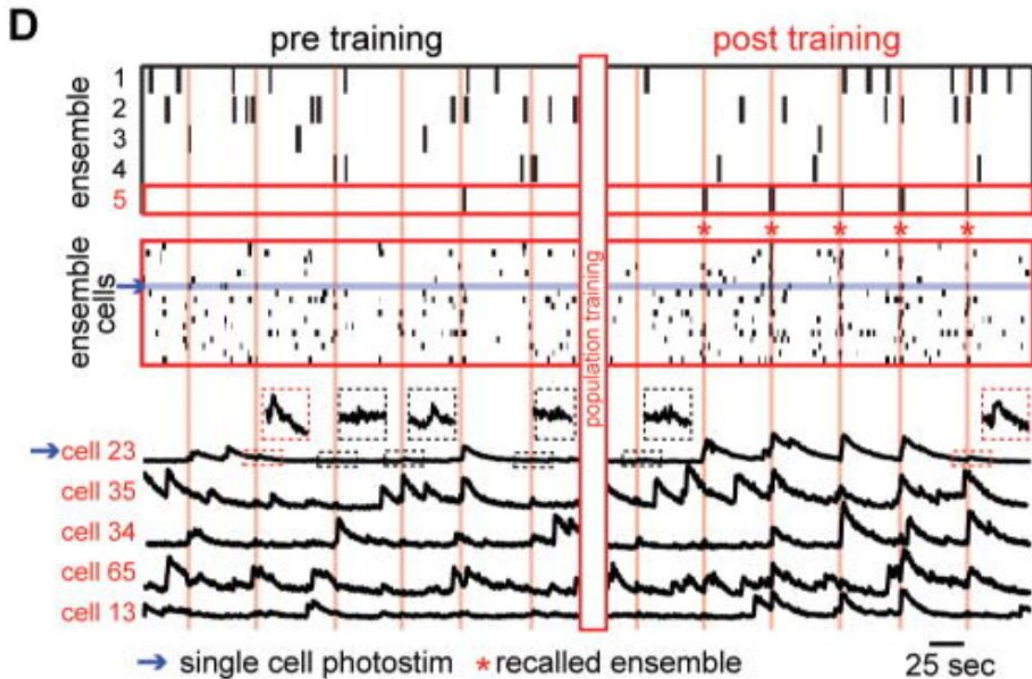
Neuronal circuits produce self-sustaining sequences of activity patterns, but the precise mechanisms remain unknown. Here we provide evidence for autoassociative dynamics in sequence generation. During sharp-wave ripple (SWR) events, hippocampal neurons express sequenced reactivations, which we show are composed of discrete attractors. Each attractor corresponds to a single location, the representation of which sharpens over the course of several milliseconds, as the reactivation focuses at that location. Subsequently, the reactivation transitions rapidly to a spatially discontinuous location. This alternation between sharpening and transition occurs repeatedly within individual SWRs and is locked to the slow-gamma (25 to 50 hertz) rhythm. These findings support theoretical notions of neural network function and reveal a fundamental discretization in the retrieval of memory in the hippocampus, together with a function for gamma oscillations in the control of attractor dynamics.

To a degree one can even see “pattern completion” happening in activity recordings...



Autoassociative dynamics in the generation of sequences of hippocampal place cells

Content-addressable memory in brain



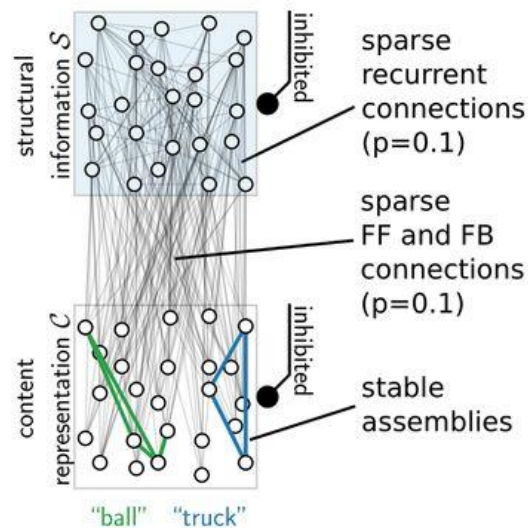
To a degree one can even see “pattern completion” happening in activity recordings...

Imprinting and Recalling Cortical Ensembles

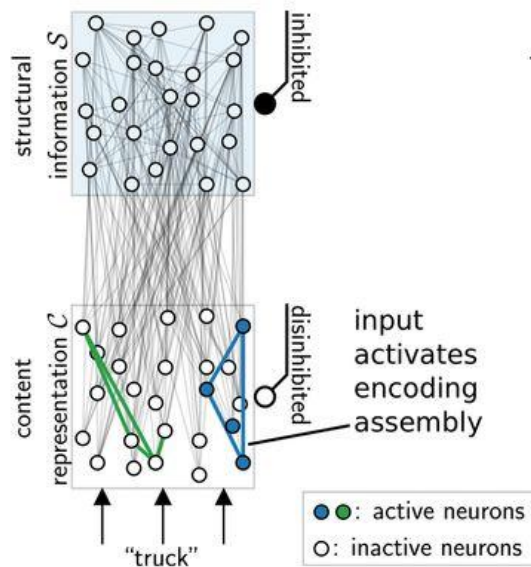
[Luis Carrillo-Reid](#), [Weijian Yang](#), [Yuki Bando](#), [Darcy S. Peterka](#), and [Rafael Yuste](#)

Using such mechanisms for symbolic processing?

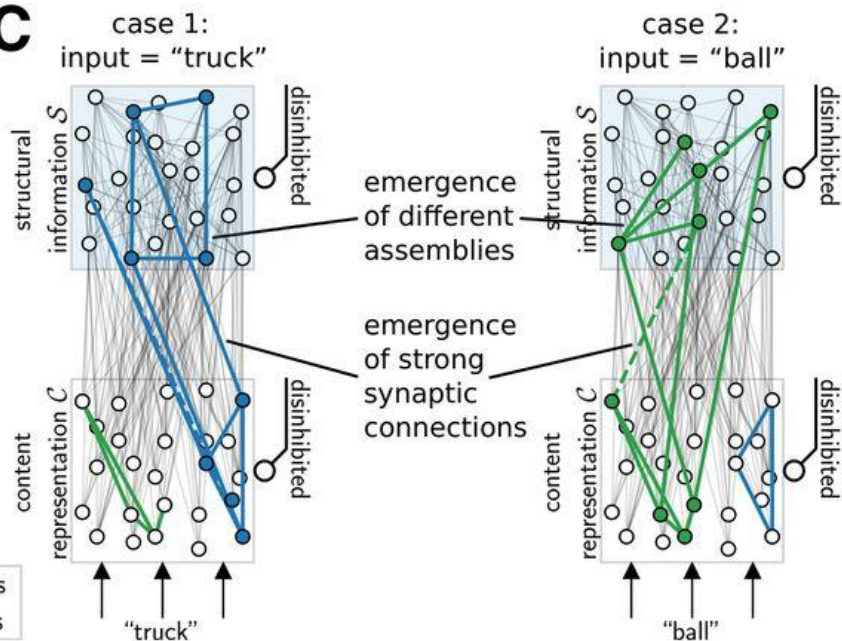
A



B



C

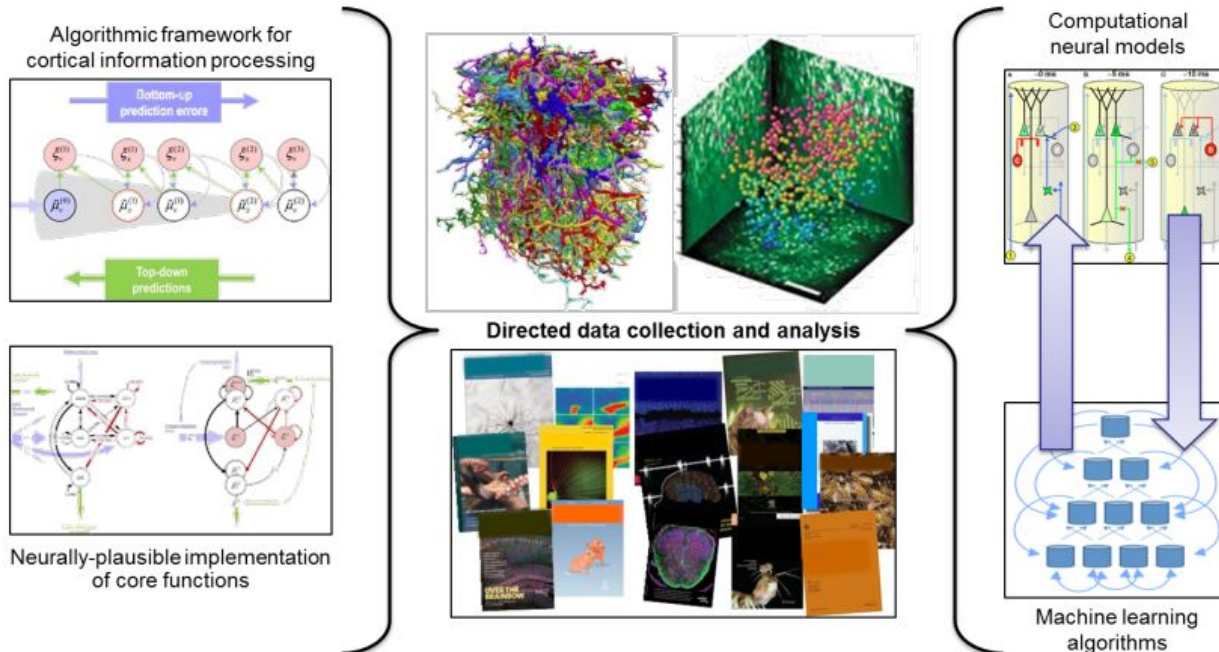


A Model for Structured Information
Representation in Neural Networks of the Brain

Looking to the future: better circuit level data

IARPA MICRONS (2015–2020) 1mm³ scale circuits (mouse brain ~500 mm³, human ~500,000 mm³)

MICrONS will combine neuroscience and data science to advance machine learning by uncovering how the cortex performs computations at the mesoscale



Looking to the future: better circuit level data

Visualizing differences in brain volume scale (if 1,000 cubic microns is proportional to 1cm)

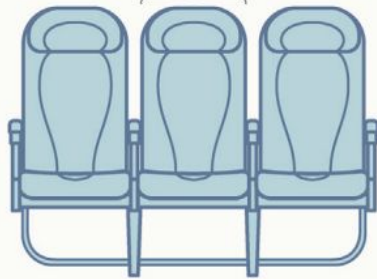
A

Caenorhabditis elegans



Brain volume:
 5×10^4 cubic microns
Equivalent length:
50 cm

Average airline seat width ~50 cm



B

Drosophila melanogaster



Brain volume:
 5×10^7 cubic microns
Equivalent length:
500 m

6.5 Boeing 747 airliners ~500 m



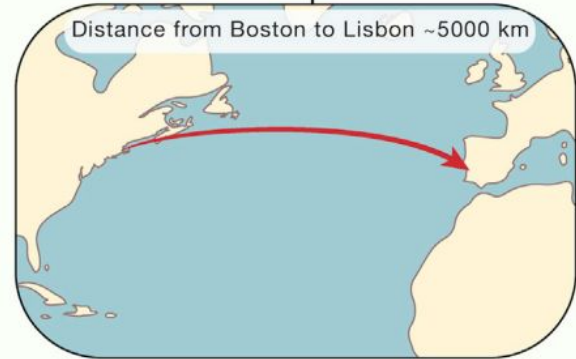
C

Mus musculus



Brain volume:
 5×10^{11} cubic microns
Equivalent length:
5000 km

Distance from Boston to Lisbon ~5000 km



1980s
Connectome
~2020
Full activity map

~2020
Connectome

The Mind of a Mouse ~2025-2030?

Larry F. Abbott¹, Davi D. Bock², Edward M. Callaway³, Winfried Denk^{4, 25}, Catherine Dulac⁵, Adrienne L. Fairhall⁶, Ila Fiete⁷, Kristen M. Harris⁸, Moritz Helmstaedter⁹, Viren Jain¹⁰,
²⁵ ㉔, Narayanan Kasthurj¹¹, Yann LeCun¹², Jeff W. Lichtman^{13, 25} ㉔, Peter B.
Littlewood¹⁴, Liqun Luo¹⁵, John H.R. Maunsell¹⁶, R. Clay Reid^{17, 25}, Bruce R. Rosen¹⁸ ...
David C. Van Essen²⁴

Take-home messages: circuits part

Some algorithms being used in ML (and we've only covered a subset here) have biologically plausible hypothesized mappings onto neural circuits, but much remains to be understood and validated

For some ML concepts, like reinforcement learning or pattern completion, there are detailed mapping to particular types and circuits of cells in the brains of birds or mammals

There are many basic things we still don't know, like how uniform the cortex is, whether some approximation of backprop is used by the brain, how the brain represents probabilities or does something like Bayesian inference, whether it uses symbolic mechanisms, and so on

Technologies have scaled activity recording from a handful of cells to thousands and can start to be more useful to address these issues, while connectomics has scaled to the level of a fly brain and there is a major push to scale it to a mammal brain

Still early days, but we see hints that circuit neuroscience can inform AI more deeply in next 10-15 years...