

Neural Circuit Development Notes

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1 Thoughts

brain part info from:

1. <http://www.news-medical.net/health/What-does-the-Thalamus-do.aspx>
2. <http://neuroscience.uth.tmc.edu/s4/chapter06.html> – talks about fear response and amygdala

Parts of the brain:

1. cerebral cortex (cerebrum)
 - frontal lobe (top front) – reasoning, planning, parts of speech, movement, emotions, problem solving
 - parietal lobe (top middle) – movement, orientation, recognition, perception of stimuli
 - occipital lobe – visual processing
 - temporal lobe – perception and recognition of auditory stimuli, memory, and speech
2. cerebellum (little brain)
 - associated with regulation and coordination of movement, posture, and balance
 - evolutionarily really old; reptiles have this as more or less their full brain
3. limbic system (emotional brain) – buried within cerebrum, like cerebellum, fairly old
 - Thalamus - relays sensory impulses from receptors in various parts of the body to the cerebral cortex. Experts think of it as a gate. 98% of sensory input is relayed by it (not olfaction? – maybe olfaction is a more primitive sense that routes to cerebellum, and is similar to chemosensors in *c. elegans*?).
 - Hypothalamus – controls release of 8 major hormones, involved in temperature regulation, control of food and water intake, sexual behavior, daily cycles in physiological state and behavior, and mediation of emotional responses
 - Amygdala – integrative center for emotions, emotional behavior, and motivation. where memory and emotions are “combined”. combines many different sensory inputs.
 - Amygdalofugal Pathway (link whereby motivation and drives can influence responses, and where responses are learned, rewards and punishments), stria terminalis (similar to fornix) – both important, come back to this.
 - Hippocampus – associated primarily with memory. looks like a seahorse.

4. Brain Stem – underneath limbic system, responsible for basic vital life functions such as breathing, heartbeat, and blood pressure.

Midbrain – (tectum, the 'roof', and tegmentum, in front of the tectum). Tectum responsible for visual reflexes. Tegmentum coordinates sensorimotor information.

Pons – connects the spinal cord to higher brain levels, and transfers info from cerebrum to cerebellum, some of which are part of the reticular formation, which regulates alertness, sleep, and wakefulness.

Medulla – transmits signals between the spinal cord and higher parts of the brain, controls autonomic functions like heartbeat and respiration. Also holds part of reticular formation.

So, basically, input goes into thalamus, and is then relayed, in this way, associations can be built. Thalamus has three groups of cells:

1. Sensory relay nuclei – These include the ventral posterior nucleus and lateral and medial geniculate body. Relay primary sensations by passing specific sensory information to the corresponding cortical area.
2. Association nuclei – receive input from specific areas of the cortex, which is projected back to the cortex to “somewhat” generalized association areas, where they regulate activity.
3. non-specific nuclei (intralaminar and midline thalamic), which receive input from cerebral cortex and project information diffusely through it. Most of these interconnect brain activity between different areas of the brain and play a role in general functions such as alerting.

Ortus basic premise: entire system works similarly to the CO₂ and O₂ regulation mechanism. Aim is to keep a balance. E.g., if IFEAR goes up, this should inherently be bad. Could be that the reason for this is that it is tied to a very basic system, like breathing. So, if system is wired such that as INO₂ increases, IFEAR increases, and an increase in INO₂ causes an intake of O₂, which decreases INO₂, the system *inherently* wants to minimize INO₂ and IFEAR. Everything should build off of and/or expand this basic idea/structure.

Take C. elegans, for example. It only has 302 neurons, and is a relatively simple organism, with its connectome nearly entirely known. Despite its relative simplicity, it is capable of avoiding toxins (cite toxin avoidance), and withdrawing from a touch to the head. Both of these actions show a tendency to minimize certain conditions. In the context of an organism as simple as C. elegans, it becomes clear that this tendency arises from a circuit configuration that causes certain “pre-wired” responses to be preside over others. **This another premise of Ortus:** The idea of “emotions”, as we know them, are simply the rise and fall in activation of different groups of neurons, tied to very fundamental behaviors. The concepts of “good” and “bad” sensations or emotions only carry meaning to us because of their associations to circuits that are either desirable or undesirable from a longevity perspective.

1. perhaps use a “chemical” to signal that a synapse may be created
2. may also need to factor location in... that would be a pain, because each synapse would need a 3D coordinate.
3. classical conditioning – two stimuli paired, instrumental conditioning – stimulus -> response -> reward

4. as things get repeated, the pathway between input and output shortens (creates a “reflexive reaction”, though not the same as a real reflex, like a knee jerk)

Ortus premise: Essentially, complex behaviors are more nuanced reflexes. A reflex goes from a sensory neuron to the spinal cord where interneurons redirect the signal to a motoneuron. Complex behaviors originate from some combination of existing neural activity and sensory input, which combine and, after being passed through a number of different interneurons, end up as signals to motor neurons, or loop back around to continue the “thought” process.

5. Should have a loop that re-energizes (in a decaying way) neural pathways/circuits that were recently used. In this way, perhaps we can implement instrumental learning, and time-based/sequence-based knowledge.
6. grey matter:

2 Neural reuse: a fundamental organizational principle of the brain. [1]

Test...

3 Functional roles of short-term synaptic plasticity with an emphasis on inhibition [2]

Test...

4 Emerging roles of astrocytes in neural circuit development. [3]

Test...

5 Synapse Formation in Developing Neural Circuits [4]

Test...

6 Astrocytes: Orchestrating synaptic plasticity? [5]

Test...

7 Two matrix metalloproteinase classes reciprocally regulate synaptogenesis [6]

Test...

8 Neural circuits on a chip [7]

Test...

9 Understanding synaptogenesis and functional connectome in C. elegans by imaging technology [8]

Test...

10 Single-Cell Memory Regulates a Neural Circuit for Sensory Behavior [9]

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11 Neural circuit rewiring: insights from DD synapse remodeling. [10]

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12 Rules for shaping neural connections in the developing brain [11]

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13 Correlated Synaptic Inputs Drive Dendritic Calcium Amplification and Cooperative Plasticity during Clustered Synapse Development [12]

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14 Neuronal development: Signalling synaptogenesis [13]

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15 Circuit Mechanisms of Sensorimotor Learning [14]

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16 Towards reverse engineering the brain: Modeling abstractions and simulation frameworks [15]

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17 Biologically based neural circuit modelling for the study of fear learning and extinction [16]

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18 A feedback neural circuit for calibrating aversive memory strength [17]

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19 Neurotrophin regulation of neural circuit development and function [18]

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20 Neural plasticity across the lifespan [19]

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21 The Purkinje cell as a model of synaptogenesis and synaptic specificity [20]

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22 Synapse biology in the 'circuit-age' paths toward molecular connectomics [21]

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23 Activity-Dependent Inhibitory Synaptogenesis in Cerebellar Cultures [22]

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24 Theoretical Models of Neural Circuit Development [23]

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25 The development of cortical circuits for motion discrimination. [24]

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26 The interplay between neurons and glia in synapse development and plasticity [25]

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27 Timing Rules for Synaptic Plasticity Matched to Behavioral Function [26]

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28 Neural plasticity and behavior ??? sixty years of conceptual advances [27]

Test...

29 Homeostatic Plasticity of Subcellular Neuronal Structures: From Inputs to Outputs [28]

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30 Mechanisms of Neural Circuit Formation [29]

(Note: this is a book comprised of research articles, title of relevant articles as subsections)

30.1 Introduction to mechanisms of neural circuit formation

Topics in book:

1. cell adhesion molecules (and downstream roles in cell identity, recognition, and synaptic specificity)
2. axon guidance, formation of terminals, and dendritic arborization
3. formation of synaptic structures themselves (remains subject to remodeling and plasticity throughout development and even in adult animals)

30.2 Wired for Behaviors: from development to function of innate limbic system circuitry

31 Synaptogenesis: A synaptic bridge [30]

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References

- [1] M. L. Anderson, “Neural reuse: a fundamental organizational principle of the brain.,” *The Behavioral and brain sciences*, vol. 33, no. 4, pp. 245–266; discussion 266–313, 2010.
- [2] H. Anwar, X. Li, D. Bucher, and F. Nadim, “Functional roles of short-term synaptic plasticity with an emphasis on inhibition,” *Current Opinion in Neurobiology*, vol. 43, pp. 71–78, 2017.
- [3] L. E. Clarke and B. a. Barres, “Emerging roles of astrocytes in neural circuit development.,” *Nature reviews. Neuroscience*, vol. 14, no. 5, pp. 311–21, 2013.

- [4] D. A. Colón-Ramos, “Synapse Formation in Developing Neural Circuits,” *Current Topics in Developmental Biology*, vol. 87, no. 09, pp. 53–79, 2009.
- [5] M. De Pittà, N. Brunel, and A. Volterra, “Astrocytes: Orchestrating synaptic plasticity?,” *Neuroscience*, vol. 323, pp. 43–61, 2016.
- [6] M. L. Dear, N. Dani, W. Parkinson, S. Zhou, and K. Broadie, “Two matrix metalloproteinase classes reciprocally regulate synaptogenesis,” *Development*, pp. 75–87, 2015.
- [7] M. F. Hasan and Y. Berdichevsky, “Neural circuits on a chip,” *Micromachines*, vol. 7, no. 9, pp. 1–15, 2016.
- [8] J. H. Hong and M. Park, “Understanding synaptogenesis and functional connectome in *C. elegans* by imaging technology,” *Frontiers in Synaptic Neuroscience*, vol. 8, no. JUN, pp. 1–10, 2016.
- [9] K. Kobayashi, S. Nakano, M. Amano, D. Tsuboi, T. Nishioka, S. Ikeda, G. Yokoyama, K. Kaibuchi, and I. Mori, “Single-Cell Memory Regulates a Neural Circuit for Sensory Behavior,” *Cell Reports*, vol. 14, no. 1, pp. 11–21, 2016.
- [10] N. Kurup and Y. Jin, “Neural circuit rewiring: insights from DD synapse remodeling,” *Worm*, vol. 5, no. 1, p. e1129486, 2016.
- [11] E. Kutsarova, M. Munz, and E. S. Ruthazer, “Rules for shaping neural connections in the developing brain,” *Frontiers in Neural Circuits*, vol. 10, no. January, p. 111, 2016.
- [12] K. F. H. Lee, C. Soares, J. P. Thivierge, and J. C. Béique, “Correlated Synaptic Inputs Drive Dendritic Calcium Amplification and Cooperative Plasticity during Clustered Synapse Development,” *Neuron*, vol. 89, no. 4, pp. 784–799, 2016.
- [13] S. Lewis, “Neuronal development: Signalling synaptogenesis,” *Nature Reviews Neuroscience*, p. 2016, 2016.
- [14] H. Makino, E. J. Hwang, N. G. Hedrick, and T. Komiyama, “Circuit Mechanisms of Sensorimotor Learning,” *Neuron*, vol. 92, no. 4, pp. 705–721, 2016.
- [15] J. M. Nageswaran, M. Richert, N. Dutt, and J. L. Krichmar, “Towards reverse engineering the brain: Modeling abstractions and simulation frameworks,” in *Proceedings of the 2010 18th IEEE/IFIP International Conference on VLSI and System-on-Chip, VLSI-SoC 2010*, pp. 1–6, 2010.
- [16] S. S. Nair, D. Paré, and A. Vicentic, “Biologically based neural circuit modelling for the study of fear learning and extinction,” *Nature Publishing Group*, vol. 1, no. July, pp. 1–7, 2016.
- [17] T. Ozawa, E. A. Ycu, A. Kumar, L.-F. Yeh, T. Ahmed, J. Koivumaa, and J. P. Johansen, “A feedback neural circuit for calibrating aversive memory strength,” *Nature Neuroscience*, vol. 20, no. November, pp. 1–11, 2016.
- [18] H. Park and M. M. Poo, “Neurotrophin regulation of neural circuit development and function,” *Nat Rev Neurosci*, vol. 14, no. 1, pp. 7–23, 2013.
- [19] J. D. Power and B. L. Schlaggar, “Neural plasticity across the lifespan,” *Wiley Interdisciplinary Reviews: Developmental Biology*, vol. 6, no. February, pp. 1–9, 2016.

- [20] M. Sasso-Pognetto and A. Patrizi, “The Purkinje cell as a model of synaptogenesis and synaptic specificity,” *Brain Research Bulletin*, 2016.
- [21] D. Schreiner, J. N. Savas, E. Herzog, N. Brose, and J. de Wit, “Synapse biology in the circuit-age paths toward molecular connectomics,” *Current Opinion in Neurobiology*, vol. 42, pp. 102–110, 2017.
- [22] F. J. Seil, “Activity-Dependent Inhibitory Synaptogenesis in Cerebellar Cultures,” *Brain Plasticity*, vol. 1, no. 2, pp. 207–214, 2016.
- [23] H. D. Simpson, D. Mortimer, and G. J. Goodhill, *Chapter 1 Theoretical Models of Neural Circuit Development*, vol. Volume 87. Elsevier Inc., 1 ed., 2009.
- [24] G. B. Smith, A. Sederberg, Y. M. Elyada, S. D. Van Hooser, M. Kaschube, and D. Fitzpatrick, “The development of cortical circuits for motion discrimination,” *Nature neuroscience*, vol. 18, no. 2, pp. 252–61, 2015.
- [25] J. A. Stogsdill and C. Eroglu, “The interplay between neurons and glia in synapse development and plasticity,” *Current Opinion in Neurobiology*, vol. 42, pp. 1–8, 2017.
- [26] A. Suvrathan, H. Payne, and J. Raymond, “Timing Rules for Synaptic Plasticity Matched to Behavioral Function,” *Neuron*, vol. 92, no. 5, pp. 959–967, 2016.
- [27] J. D. Sweatt, “Neural plasticity and behavior ??? sixty years of conceptual advances,” *Journal of Neurochemistry*, vol. 139, pp. 179–199, 2016.
- [28] W. Wefelmeyer, C. J. Puhl, and J. Burrone, “Homeostatic Plasticity of Subcellular Neuronal Structures: From Inputs to Outputs,” *Trends in Neurosciences*, vol. 39, no. 10, pp. 656–667, 2016.
- [29] J. A. Weiner, R. W. Burgess, and J. Jontes, *Mechanisms of Neural Circuit Formation*. 2015.
- [30] D. Yates, “Synaptogenesis: A synaptic bridge,” *Nat Rev Neurosci*, vol. 17, no. 3, p. 135, 2016.