Microtubule force generation in axon growth cones

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Background	Model	Results
	Model, modifications to the base model	Results, what did this tell us.

Background starting with neurons -> axons -> growth cones -> base model Much of this comes first from BPS poster for the neuron to axon (condensed) Then from proposal

Future Work

This model is constructed on a population based steady state model with temporal attributes included as an after-thought. A more rigorous approach would likely include an agent based model with mt motions, protein binding events, and actin network activity taking place with defined rates. An agent based simulation constructed in this way could explore more precise relationships between actin treadmilling, adhesion, and mt force generation by sliding and polymerization. Constructed properly, this agent based simulation could explore the two-dimensional landscape of the growth cone and investigate the role of mt force generation in growth cone guidance or lack thereof. Furthermore, such a model would have the advantage of being double validated by the original population model in the growth cone and agent based simulations in the axon.

References

- [1] Erin M. Craig, Jonathan Stricker, Margaret Gardel, and Alex Mogilner. Model for adhesion clutch explains biphasic relationship between actin flow and traction at the cell leading edge. Physical Biology, 12(3):035002, May 2015.
- [2] Erin M. Craig, David Van Goor, Paul Forscher, and Alex Mogilner. Membrane Tension, Myosin Force, and Actin Turnover Maintain Actin Treadmill in the Nerve Growth Cone. Biophysical Journal, 102(7):1503–1513, April 2012.
- [3] Sara de Vincentiis, Alessandro Falconieri, Marco Mainardi, Valentina Cappello, Vincenzo Scribano, Ranieri Bizzarri, Barbara Storti, Luciana Dente, Mario Costa, and Vittoria Raffa. Extremely Low Forces Induce Extreme Axon Growth.
- Journal of Neuroscience, 40(26):4997–5007, June 2020. [4] Katherine Kalil and Erik W. Dent.
- Touch and go: Guidance cues signal to the growth cone cytoskeleton. Current Opinion in Neurobiology, 15(5):521-526, October 2005.
- [5] F. J. Nédélec, T. Surrey, A. C. Maggs, and S. Leibler. Self-organization of microtubules and motors. Nature 1997 389:6648, 389(6648):305-308, 1997.
- [6] Vittoria Raffa.
- Force: A messenger of axon outgrowth. Seminars in Cell & Developmental Biology, 140:3–12, May 2023.
- [7] Carlos Sánchez-Huertas and Eloísa Herrera. With the Permission of Microtubules: An Updated Overview on Microtubule Function During Axon Pathfinding. Frontiers in Molecular Neuroscience, 14, December 2021.

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