9.14 class #23: Retinotectal system 3: CNS axonal regeneration.

Readings:

- So, K.-F. & Aguayo, A.J., "Lengthy regrowth of cut axons from ganglion cells after peripheral nerve transplantation into the retina of adult rats", *Brain Research*, 1985, 328: 349-354.
- Chen, D.F., Jhaveri, S. & Schneider, G.E., "Intrinsic changes in developing retinal neurons result in regenerative failure of their axons", *Proc. Natl. Acad. Sci. USA*, 1995, 92: 7287-7291.
- Chen, D.F., Schneider, G.E., Martinou, J.-C. & Tonegawa, S. ,"Bcl-2 promotes regeneration of severed axons in mammalian CNS", *Nature*, 1997, 385: 434-439.
- Barde, Yves-Alain.," Help from within for damaged axons [News and Views]", *Nature*, 1997,385: 391-393. [See also the review by Holm & Isacson in 1999, listed below.]

Also recommended:

- Holm, K. & Isacson, O., "Factors intrinsic to the neuron can induce and maintain its ability to promote axonal outgrowth: a role for BCL2?", *Trends in Neuroscience*, 1999, 22: 269-273.
- David, S. & Aguayo, A.J., "Axonal elongation into peripheral nervous system "bridges" after central nervous system injury in adult rats", *Science*, 1981,214: 931 933.
- So, K.-F., Schneider, G.E. & Ayres, S., "Lesions of the brachium of the superior colliculus in neonate hamsters: correlation of anatomy with behavior", *Experimental Neurology*, 1981,72: 379-400.
- Schnell, L. & Schwab, M.E., " Axonal regeneration in the rat spinal cord produced by an antibody against myelin-associated neurite growth inhibitors", *Nature*, 1990, 343: 269-272.

Cheng, H., Cao, Y. & Olson, L., "Spinal cord repair in adult paraplegic rats: partial restoration of hind limb function", *Science*, 1996, 273: 510 - 513.

Questions:

- 1. Compare and contrast: Schwann cells and oligodendrocytes. What are they, where are they located, and what roles can they play in axonal regeneration?
- 2. After Ramon y Cajal's studies of regeneration and degeneration in the mammalian CNS after injury (book published 1928), it appeared that most CNS axon regeneration was abortive. How did the Aguayo findings lead to a renewal of interest in (and hope for) CNS regeneration?
- 3. Describe the paradigm invented by Kwok-fai So, which has become one of the major experimental paradigms for studying CNS axon regeneration.
- 4. What is the basic finding of Martin Schwab's laboratory concerning oligodendrocytes that appears to fit well with the theories of Aguayo *et al.* ?
- 5. What is the Olsen et al. paradigm, and what have these investigators recently claimed?
- 6. The MIT study by Chen *et al.* (1995) presented a fundamentally different view of CNS axon regeneration. What is it, and how does it explain the Aguayo et al. findings ?
- 7. What is Bcl-2, and how did Chen et al. show its possible importance for axonal growth and regeneration?
- 8. The need for a multifactor approach to the problem of obtaining functionally useful CNS axon regeneration: What is the meaning of each of the "four P's" of regeneration? (Preserve, Permit, Promote, and Plasticity) *In class we will discuss the current status of this field*.