

Chem 412: BMB Seminar

May 7, 2001

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Glaucoma: "Sneak Thief of Sight"

Glaucoma is the third leading cause of preventable blindness. Roughly 5.2 million people have the disease and 2 million of these people don't know they have it. Glaucoma is characterized by high intraocular pressure eventually causing irreversible optic nerve damage. The high intraocular pressure arises from unequal flow of aqueous humor into the anterior chamber and out through the trabecular meshwork. The trabecular meshwork is located at the angle where the cornea and iris meet. A high intraocular pressure will develop if the angle is too narrow or if it's completely blocked. The molecular mechanisms of the blockage is unknown, but current research is targeting cells and proteins associated in the trabecular meshwork. One gene has been cloned and closely linked to glaucoma, GLCA1, also known as MYOC and TIGR. The protein associated with the gene is called TIGR, Trabecular Meshwork Inducible Glucocorticoid Response. The function of the protein is unknown, but some structural properties have been discovered and speculations of what its function could be. Glaucoma can be treated with eye drops if caught early enough. Researchers theorize glaucoma is caused by an interaction of many proteins. Future goals are aimed at discovering these proteins and the genes associated with these proteins. Once various proteins are discovered the mechanism will no longer be a mystery.

- Fautsch, MP., et. al. Recombinant TIGR/MYOC Increases Outflow Resistance in the Human Anterior Segment. *Invest. Ophthalmology Visual Sciences*. 41(13): 4163-8 (Dec, 2000).
- Hogg, P., et. al. Aqueous Humor Stimulates the Migration of Human Trabecular Meshwork Cells In Vitro. *Invest. Ophthalmology*. 41(1); 1091-8 (Apr. 2001).
- Huang, W. et.al. Expression of the TIGR gene in the Iris, Ciliary body, and Trabecular Meshwork of the Human Eye. *Ophthalmic Genetics*. 21(3): 155-69 (Sep. 2000).
- Jacobson N., Andrews, et.al. Non-secretion of Mutant Proteins of the Glaucoma Gene Myocilin in Cultured Trabecular Meshwork Cells and in Aqueous Humor. *Human Molecular Genetics*. 10(2): 117-25 (Jan 15, 2001).
- Neufeld, Arthur H. et.al. Inhibition of Nitric-Oxide Synthase 2 by Aminoguanidine provides Neuroprotection of Retinal Ganglion Cells in a Rat Model of Chronic Glaucoma. *Pharmacology*. 96(17): 9944-9948 (Aug. 17, 1999).
- Nguyen, Thai D., et. al. Gene Structure and Properties of TIGR, an Olfactomedin-related Glycoprotein Cloned from Glucocorticoid-induced Trabecular Meshwork Cells. *Journal of Biological Chemistry*. 273(11); 6341-6350 (March 13, 1998).
- Polansky, J.R. et. al. Regulation of TIGR/MYOC Gene Expression in Human Trabecular Meshwork Cells. *Eye*. Jun 14, 2000: 503-514.
- Stone, Edwin M. et. al. Identification of a Gene that Causes Primary Open Angle Glaucoma. *Science*. 275(31): 668-670 (Jan. 1997).