

Exploring novel multi-modal photonic techniques for live imaging of synaptic activity

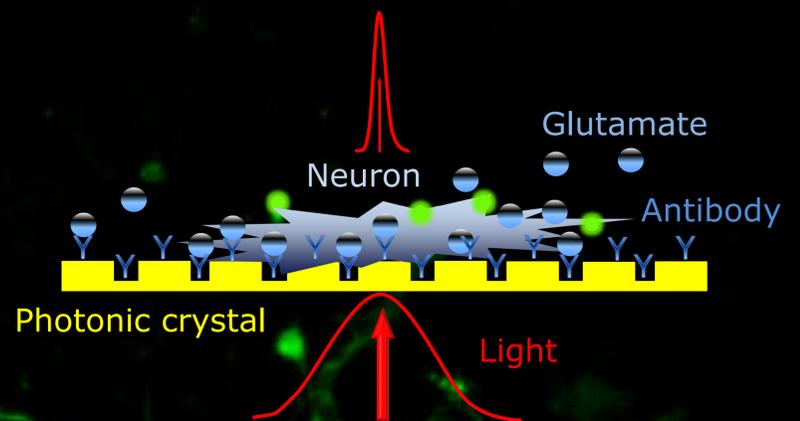
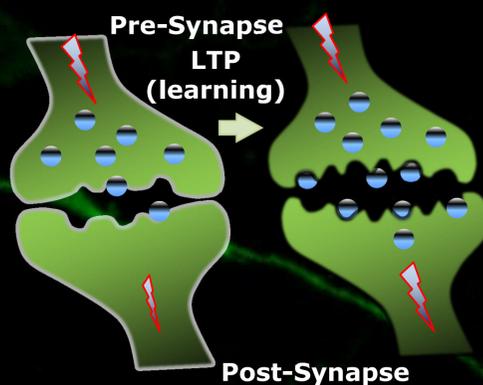
Graham J. Triggs^a, Matthias Fischer^a, Thomas F. Krauss^a, and Gareth J.O. Evans^b

^aDepartment of Physics; ^bDepartment of Biology; University of York, YO10 5DD

Objective: Endogenous observation of a chemical neurotransmitter released by active synapses during the course of learning by using a silicon-based photonic crystal biosensor

Background:

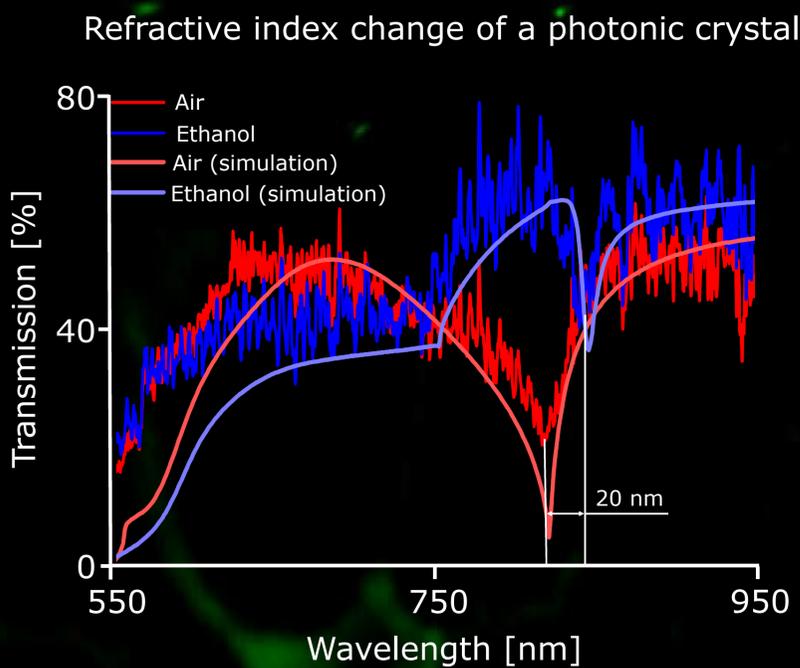
The fundamental cellular mechanisms of learning in the brain cause strengthening of synaptic connections between neurons following sensory stimuli. Learning-induced increases in synaptic activity are due to changes on both sides of the synapse: (I) enhanced neurotransmitter (glutamate) release from the pre-synaptic cell and (II) enhanced detection of glutamate in the post-synaptic cell.



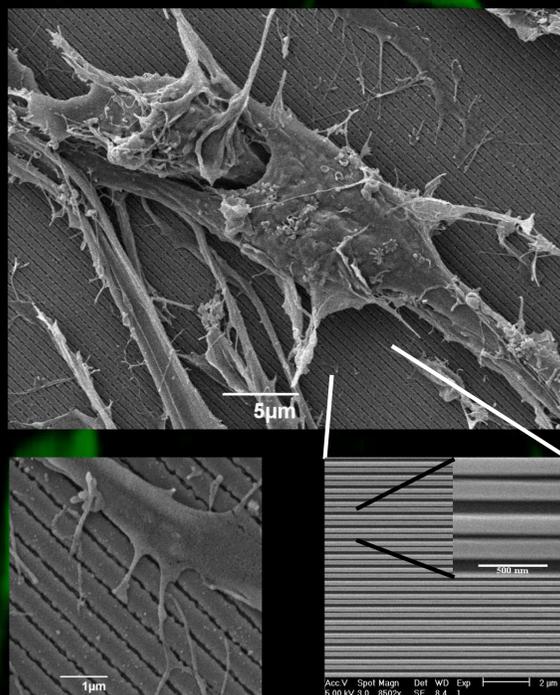
For detection of synaptic glutamate release, we utilise a photonic crystal surface that has been functionalised with anti-glutamate antibodies. The photonic crystal exhibits a resonance shift in wavelength in response to glutamate binding on the surface.

In combination with real-time imaging and expression of a fluorescent synaptic marker to visualise the location of all synapses, a map of the activity of individual synapses over time will be generated, allowing changes in synaptic activity to be monitored.

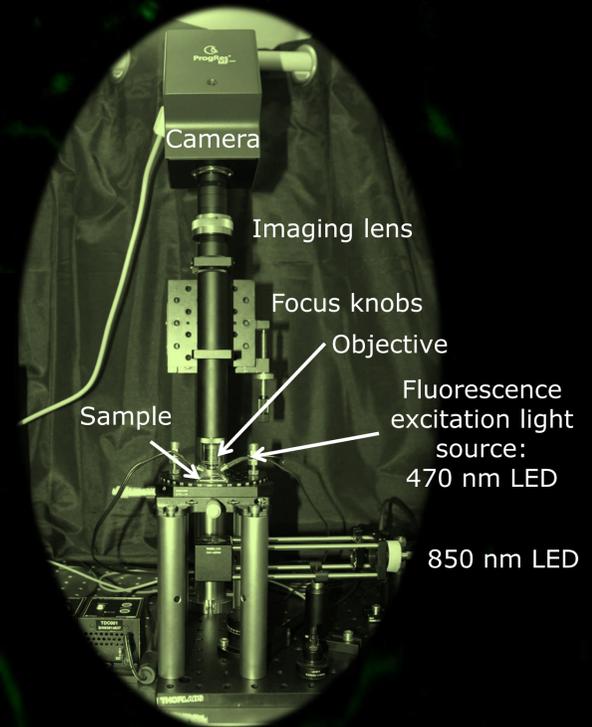
Results:



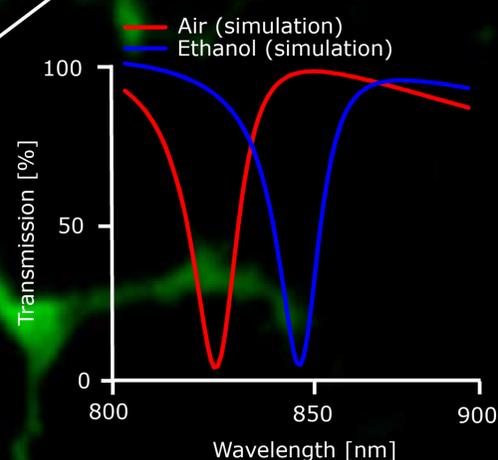
SEM image of neurons cultured on a photonic crystal surface



Microscope



Fluorescence image of a neuron



Conclusion:

- Resonant photonic crystal biosensor for label-free monitoring of neuronal activity being developed
- Combination of fluorescence microscopy with resonant detection demonstrated
- Neuron growth on sensor surface demonstrated
- Shift in resonance observed

Outlook:

- Fabrication of an optimised photonic crystal
- Optimisation of fluorescence excitation
- Real-time and simultaneous measurement of glutamate release and activity of individual synapses by electrical field stimulation at ~ 0.1 Hz that mimics learning
- Provide a model for the future study of synaptic strengthening during learning

References: E.A. Lidstone et al., (2011) Analyst. 136(18); M.A. Cousin and G.J.O. Evans (2011) J. Physiol. 589