The Effects of a Dopamine D1/D5 Agonist on Local Protein Synthesis in Cultured Hippocampal Neurons

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It is well established that *de-novo* protein synthesis is required for long-term memory storage. Furthermore, the activity-dependent regulation of local protein synthesis in the dendritic compartment is believed to be essential for the persistence of late-phase long term potentiation. Although it has been demonstrated that isolated dendrites of mature hippocampal neurons are able to synthesize proteins (Aakalu and Smith et al., Neuron 2001), the synaptic mechanisms governing this local translation have yet to be elucidated. Previous studies have implicated D1/D5 receptors in a wide range of behavioral and cellular processes: the receptors are critically involved in spatial learning and have also been shown to enhance glutamatergic transmission in the hippocampus. In order to test the hypothesis that D1/D5 receptors regulate dendritic translation, we are examining the effects of the D1/D5-selective agonist (+) SKF-38393 (100 µm) on dendritic GFP synthesis in cultured hippocampal neurons. Between-dish comparisons as well as singlecell time lapse imaging experiments show that bath application of the D1/D5 agonist stimulates GFP synthesis in neurons. Using local perfusion of the agonist to small regions (5-15 microns) of the distal dendritic arbor, we are also investigating the spatial specificity of D1/D5-induced protein synthesis. Our preliminary results indicate that focal application of the agonist results in increased GFP expression that is often restricted to the area of the agonist perfusion.