

Investigating the functional roles of δ -catenin

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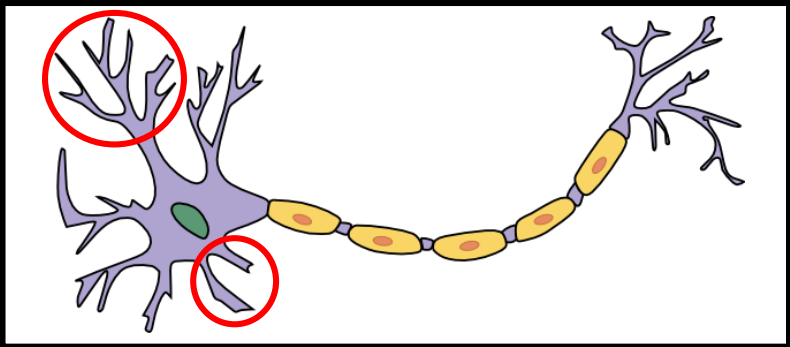
Molecular, Cellular, and Developmental Biology

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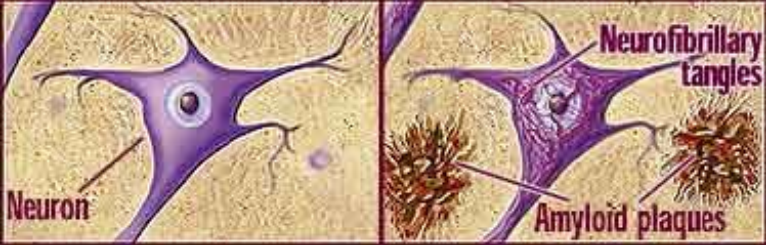
Mutations lead to neurological disorders

Too few or too many synapses



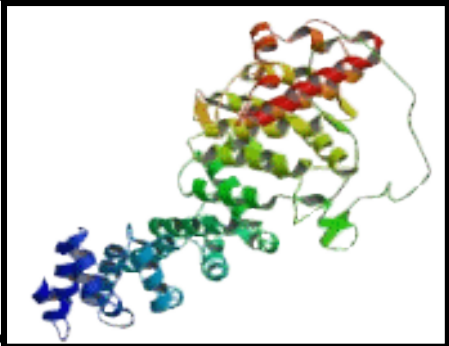
Autism

Plaques and tangles



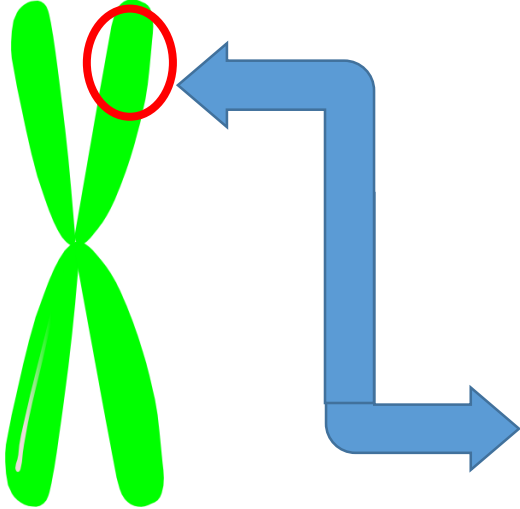
Alzheimer's disease

Delta-catenin 2

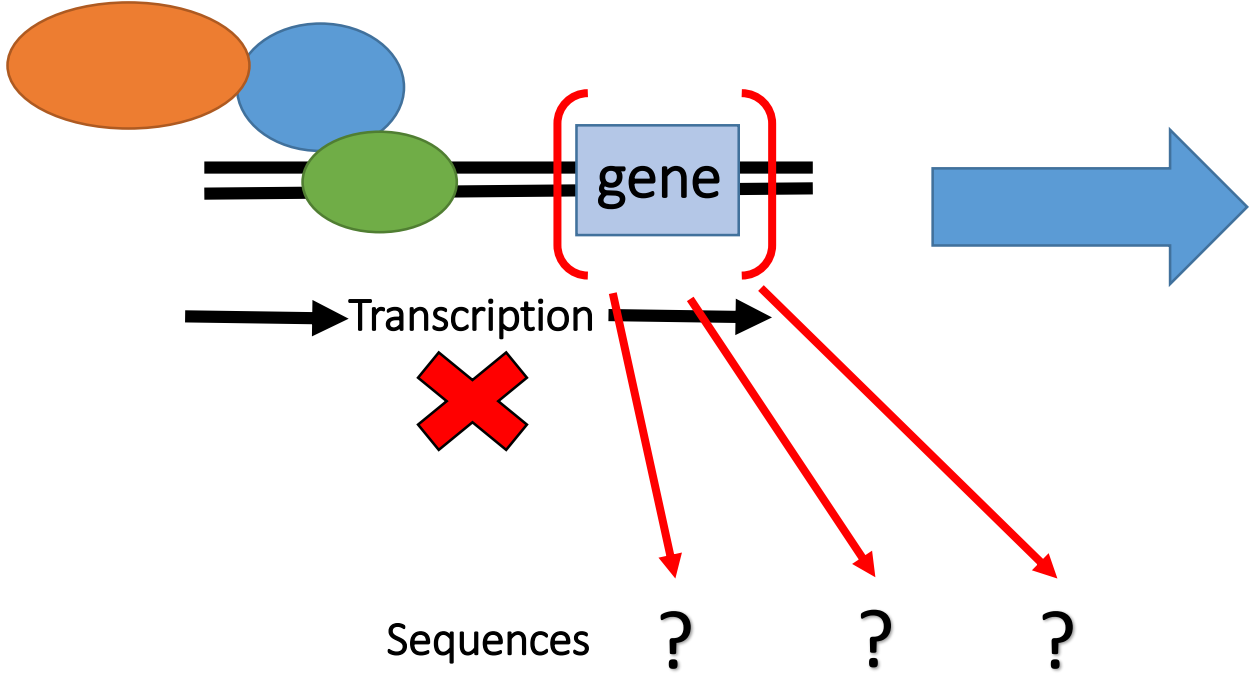


Studying affect of δ -catenin by knocking down

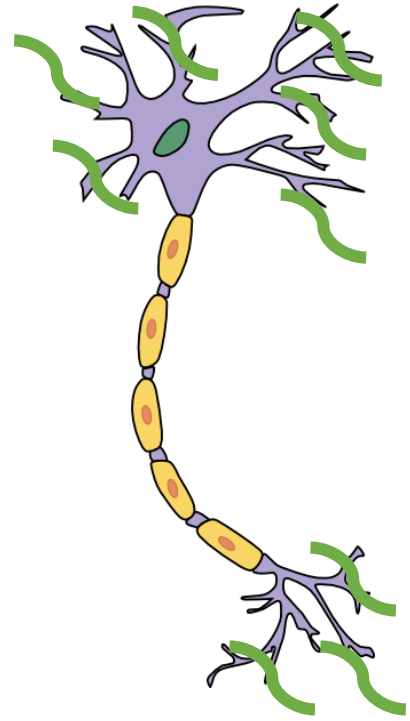
Chromosome 5p



δ -catenin gene

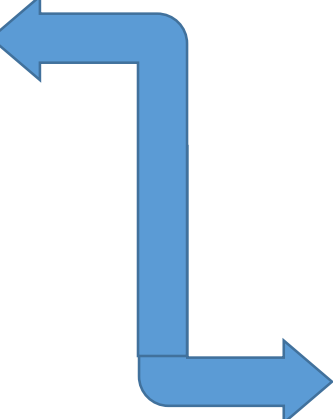
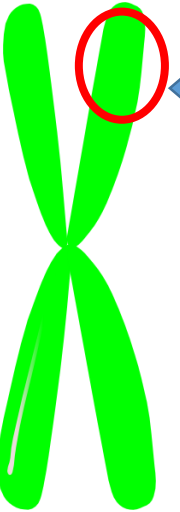


δ -catenin protein

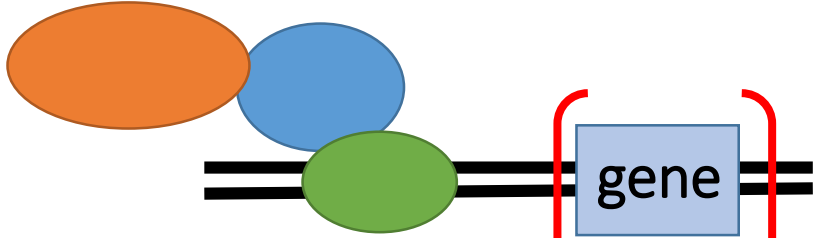


Studying affect of δ -catenin by knocking down

Chromosome 5p



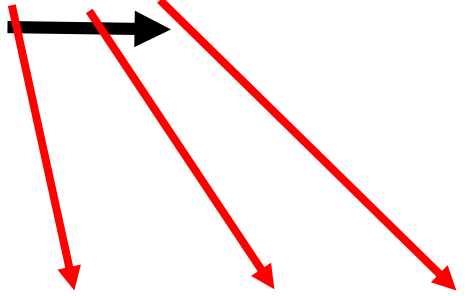
δ -catenin gene



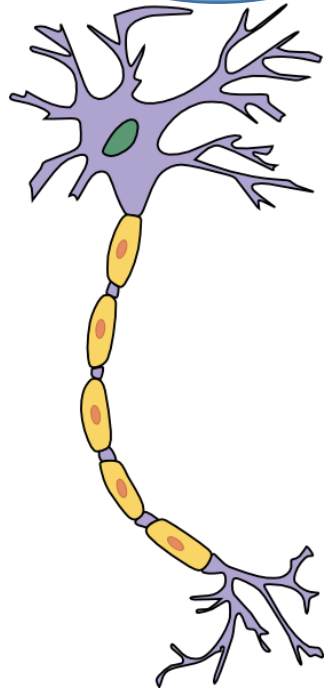
Transcription



Sequences ? ? ?

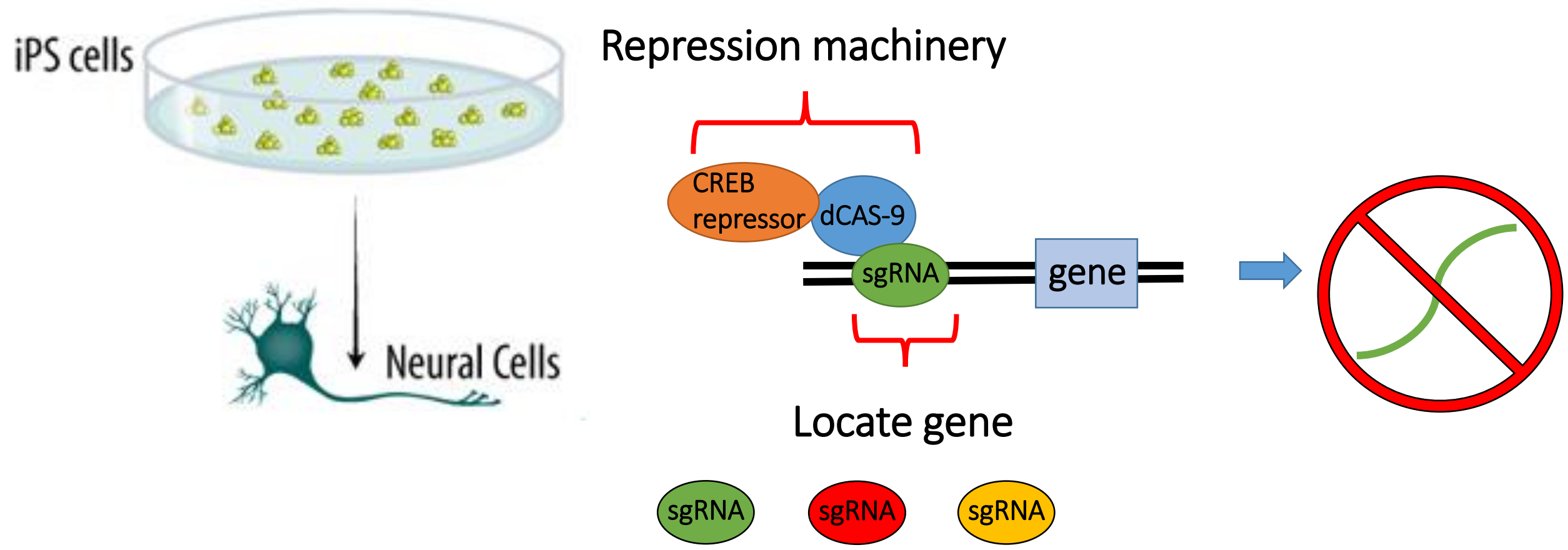


~~δ -catenin protein~~



Step 1

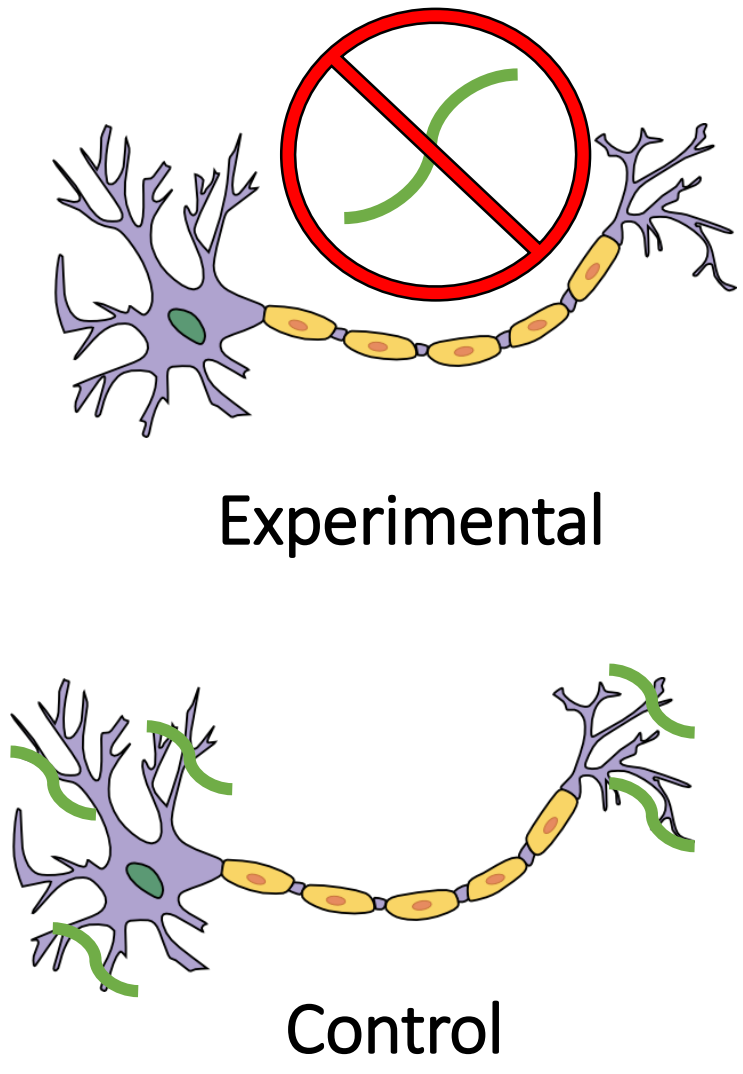
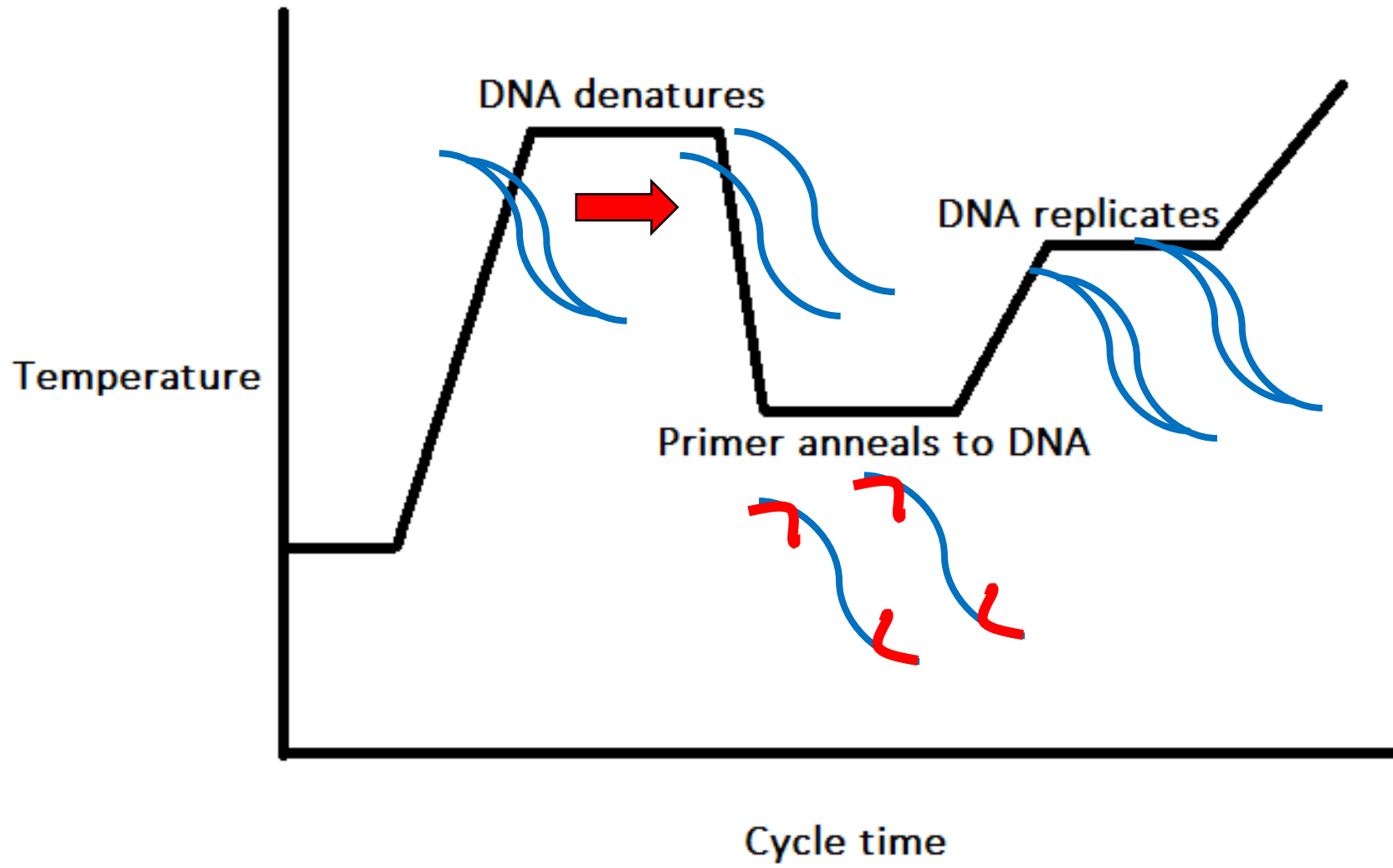
Repressing δ -catenin gene in stem cells with CRISPRi



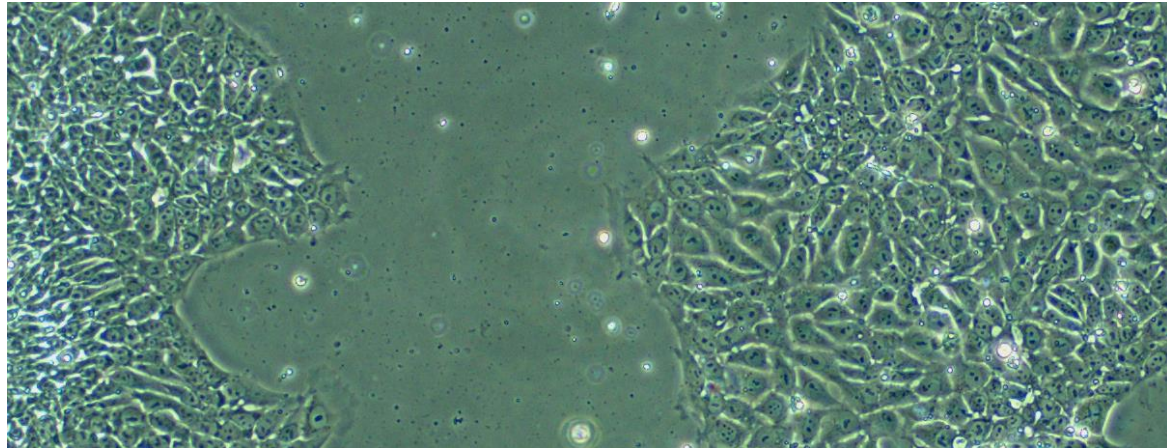
"dead" Cas9 system = binding, no cutting

Step 2

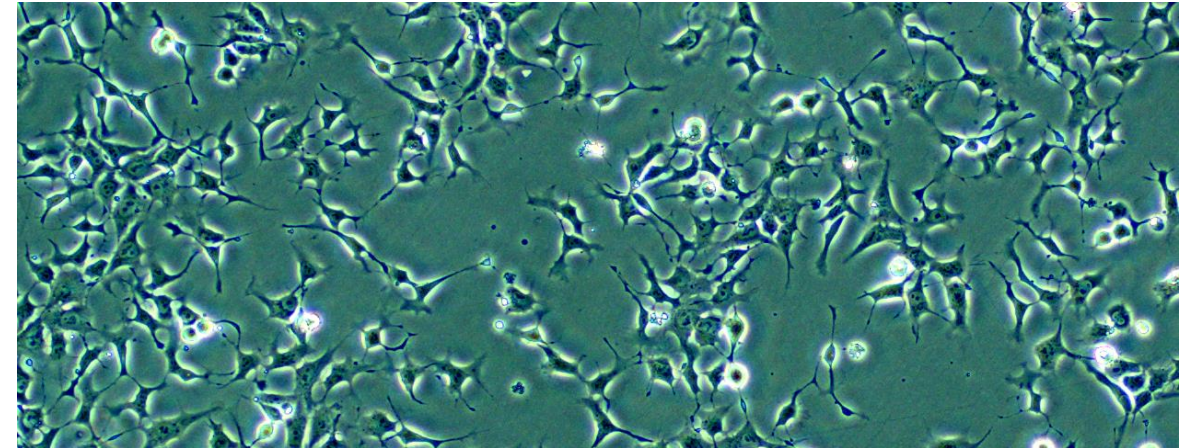
Amplifying δ -catenin mRNA with qPCR



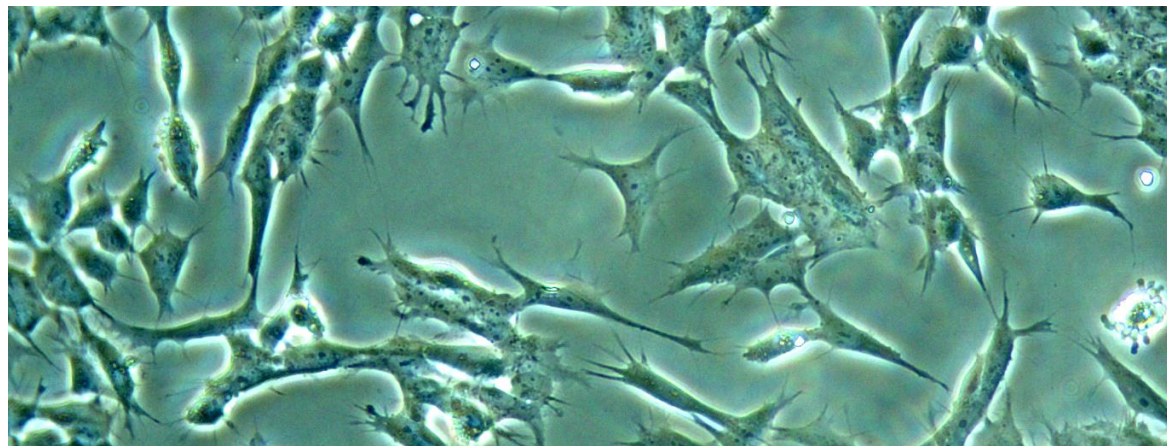
Stem cell differentiation into human neurons



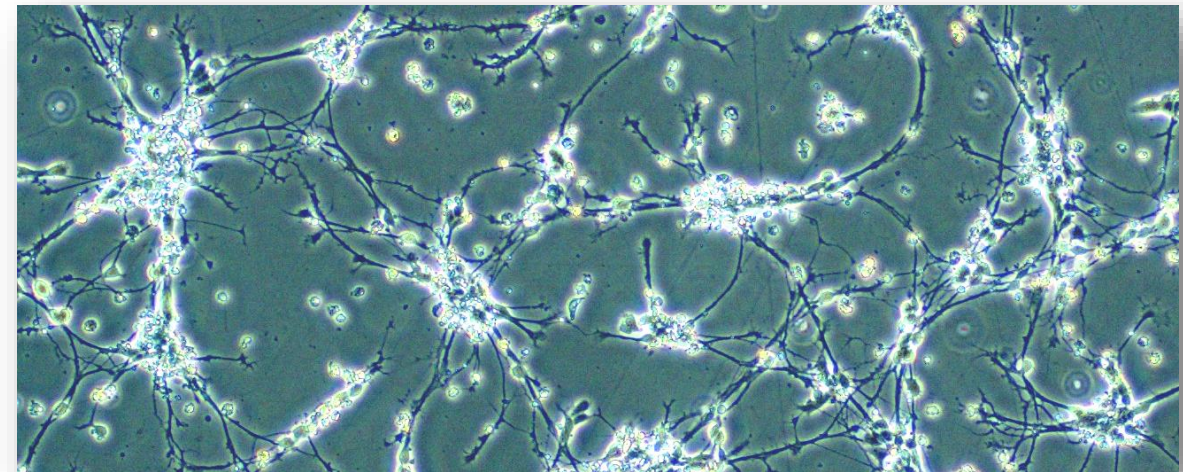
Before differentiation (10x)



1st day after differentiation (10x)

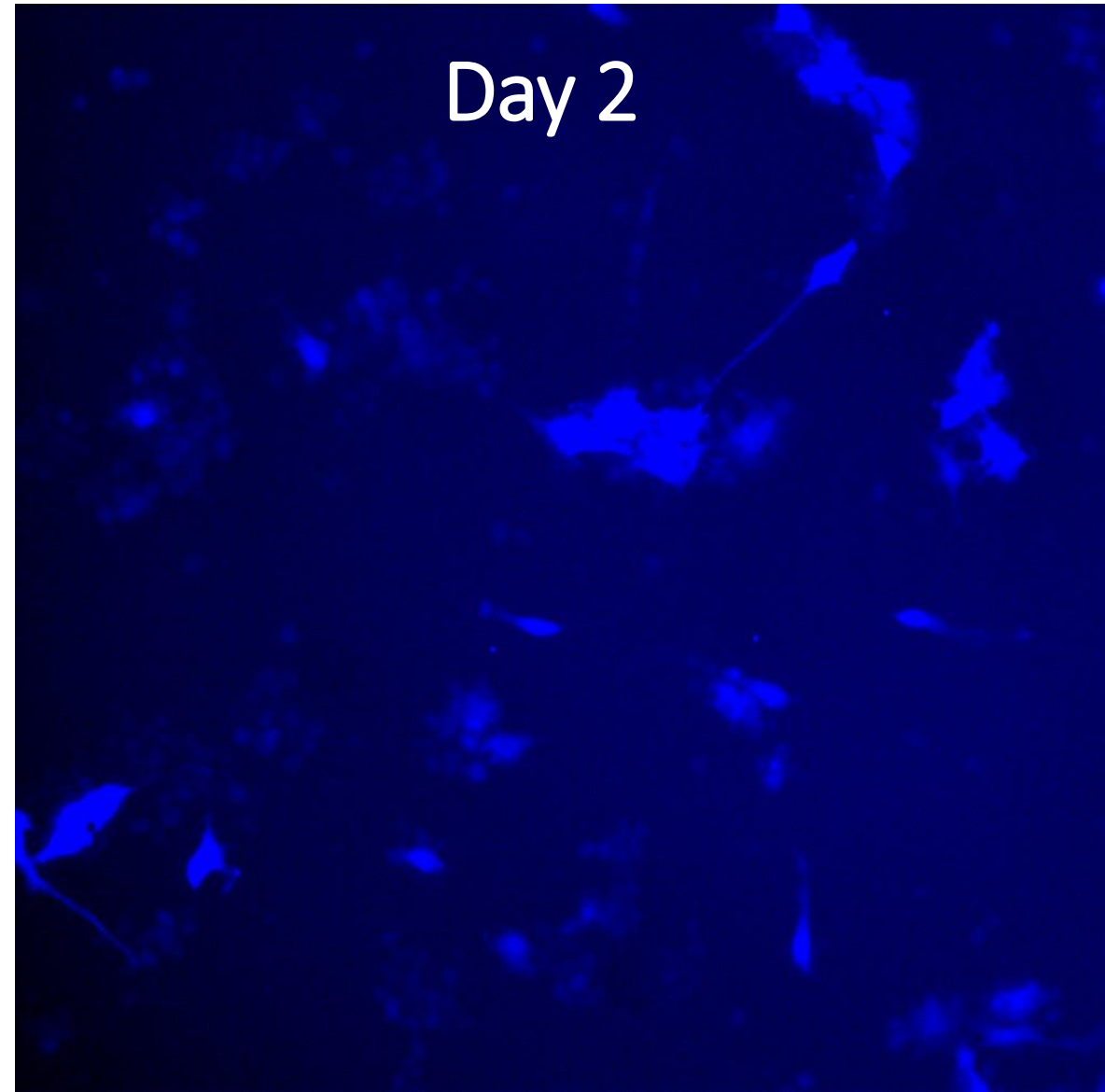
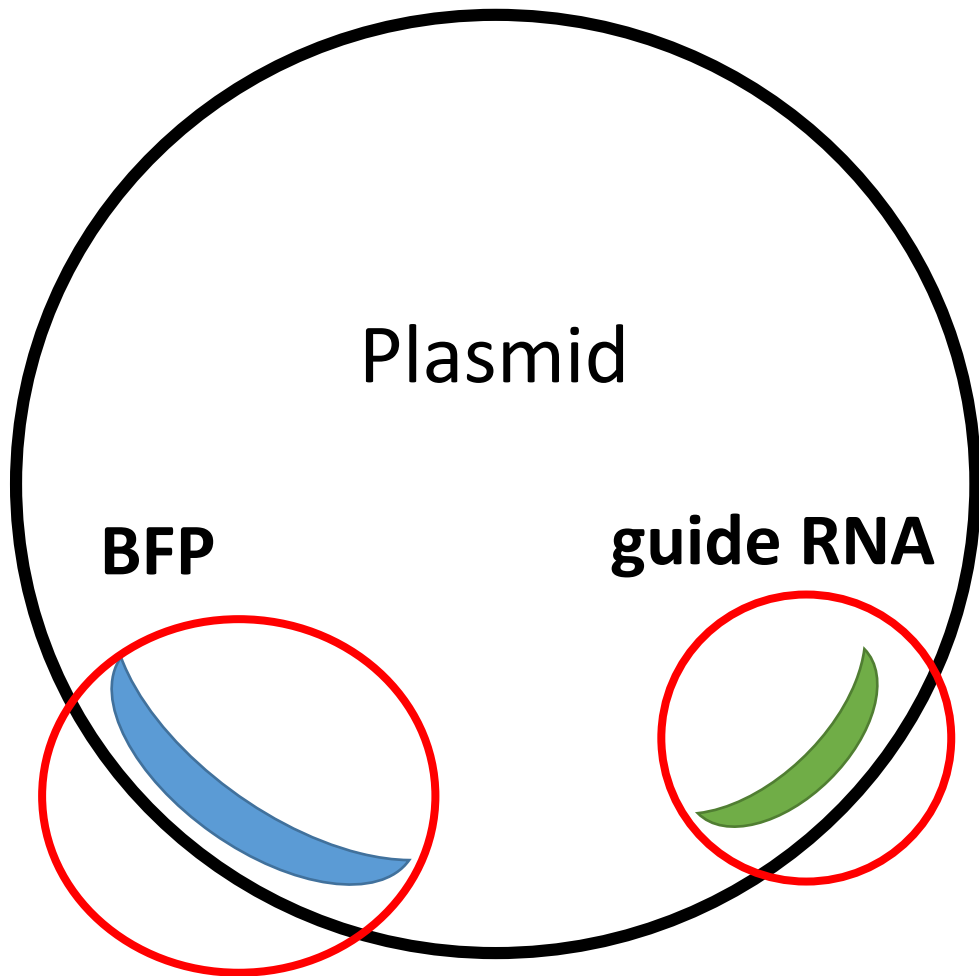


2nd day after differentiation (20x)



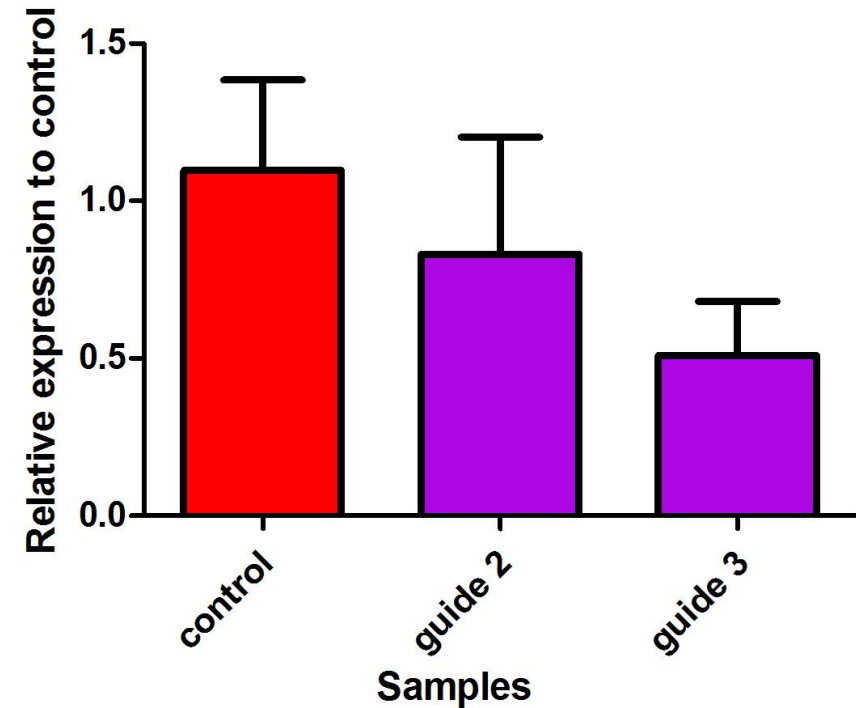
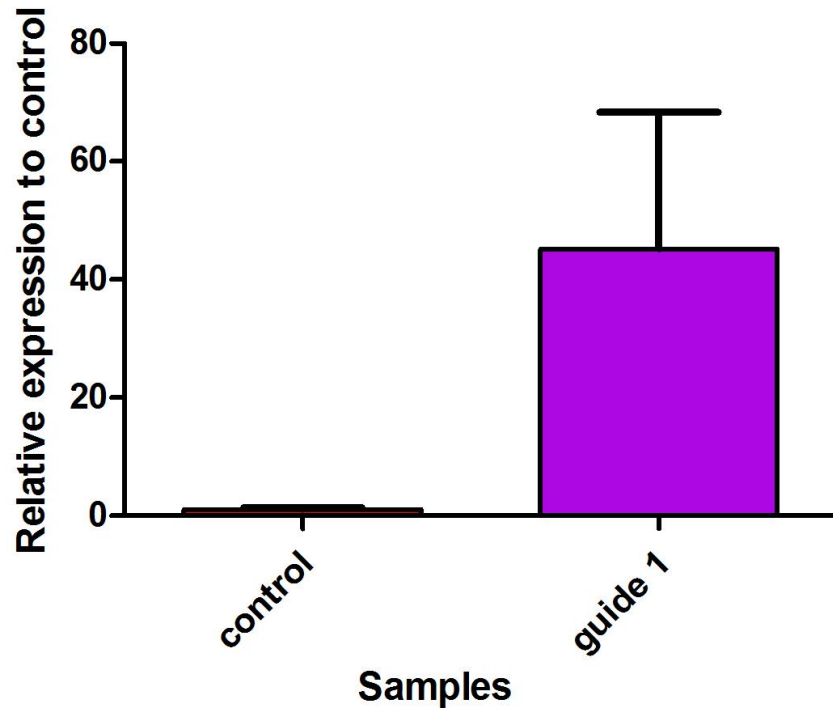
7th day after differentiation (20x)

Confirming presence of guide RNA in H4 cells

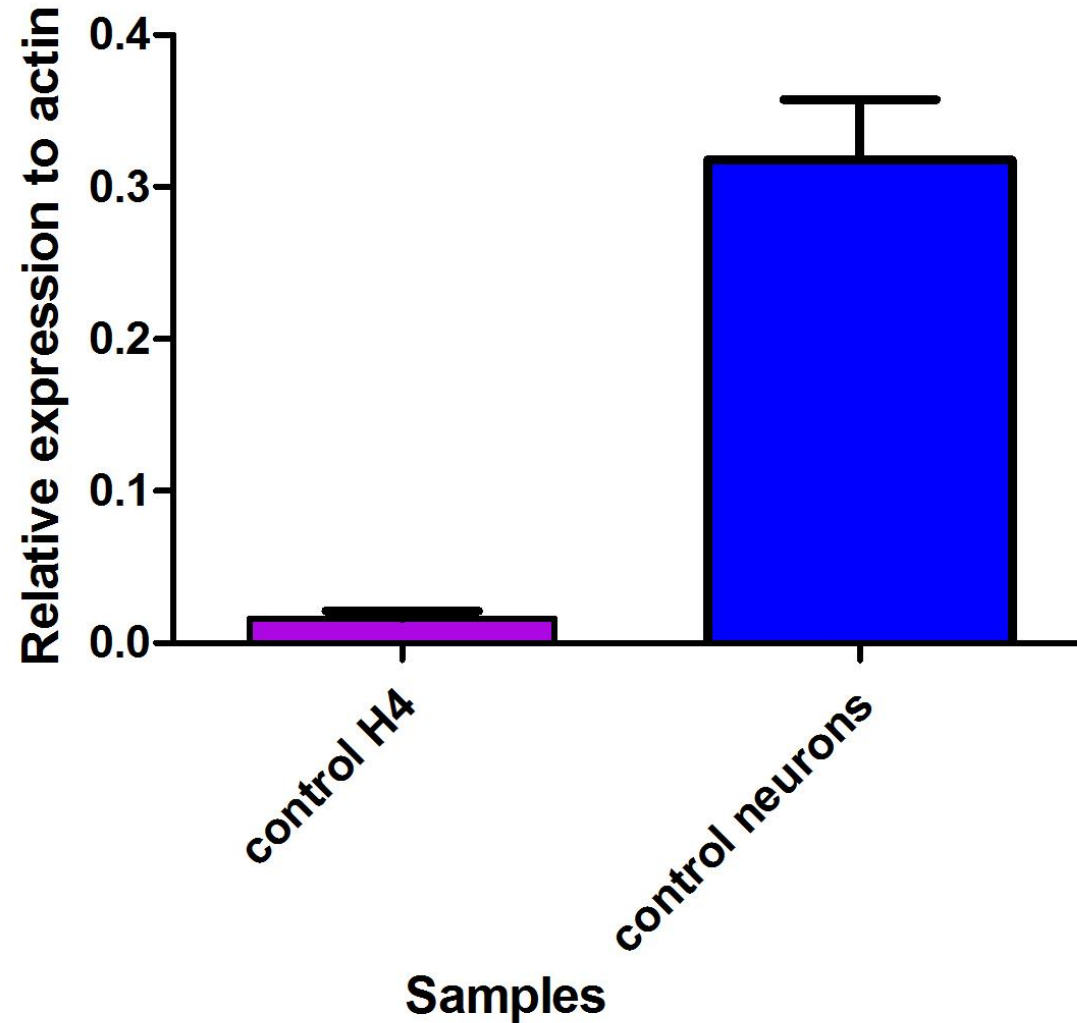


δ -catenin knockdown in H4 cells inconclusive

One-way ANOVA test shows no significant differences

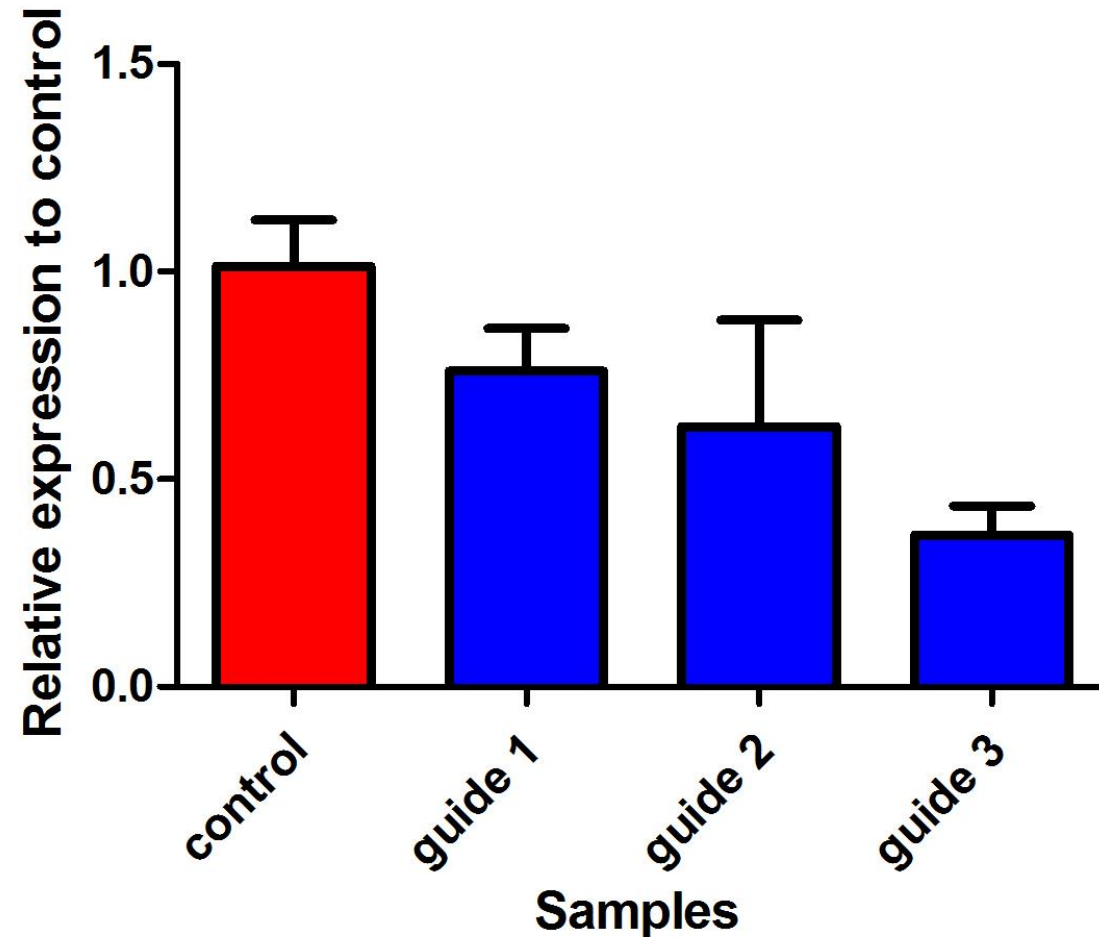


δ -catenin more highly expressed in neurons



Knockdown of δ -catenin confirmed in neurons

T-test shows significant difference between guide 3 and control

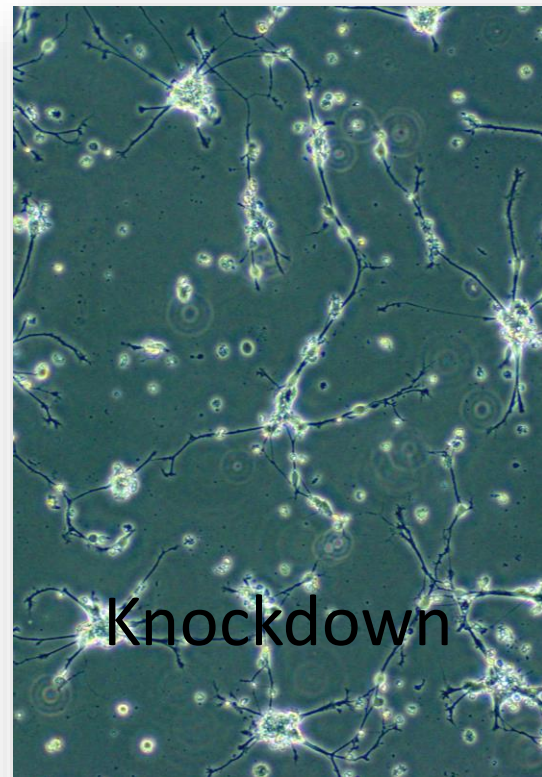
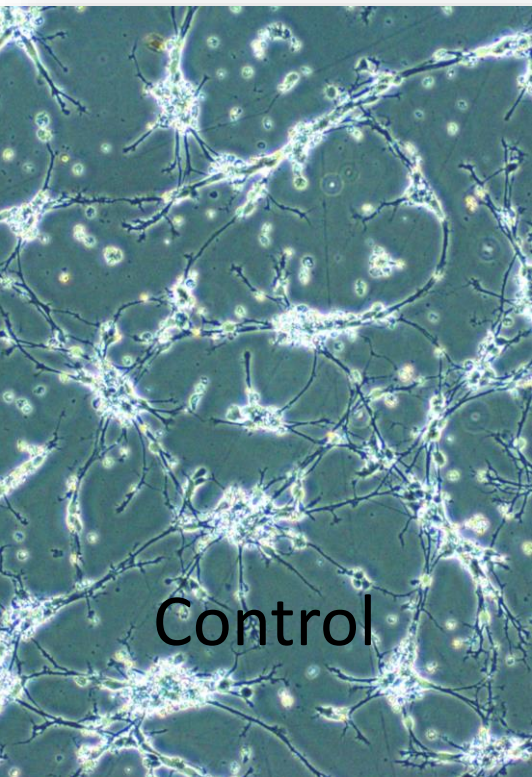


Guide RNA 3 most effective at repression

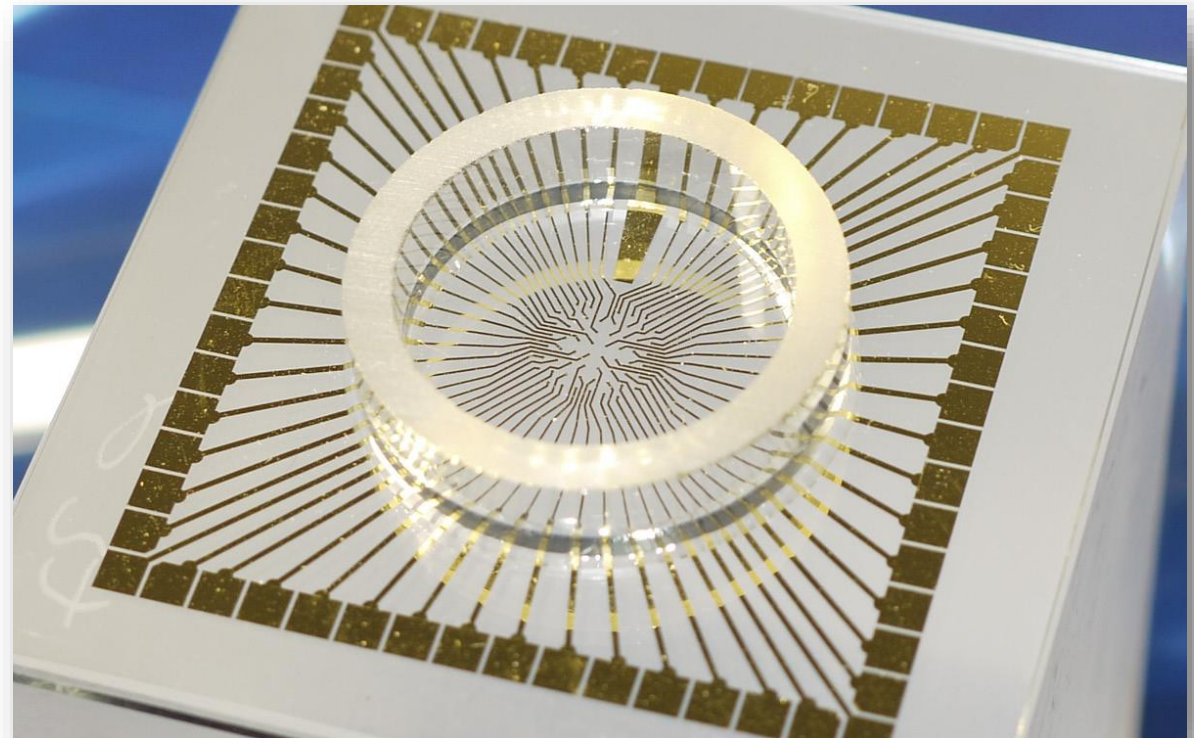
- Confirmation of most effective δ -catenin knockdown

Future plans:

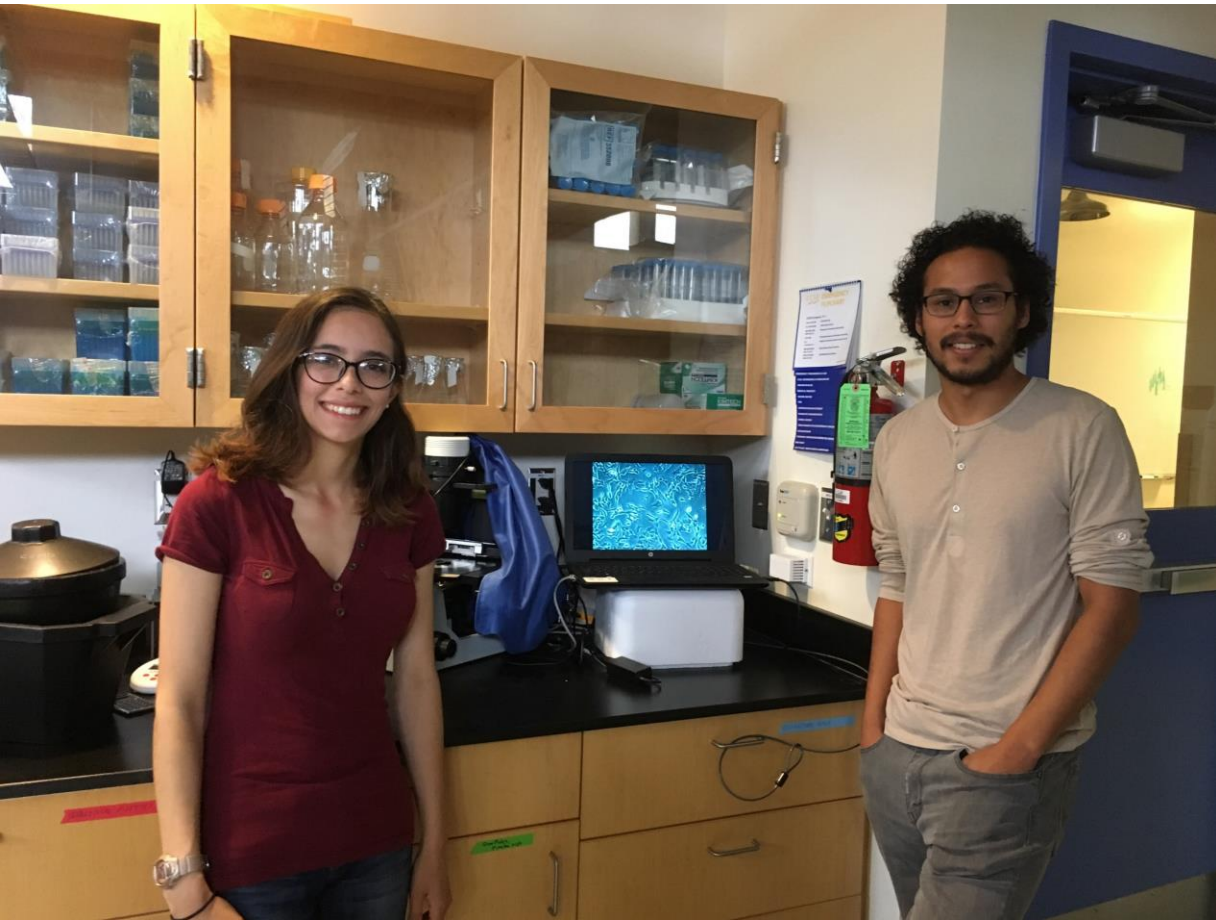
Morphology



Electrophysiological properties



Acknowledgements



- Elmer Guzman, Mentor
- Dr. Ken Kosik, Principal Investigator
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- My program peers

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