Lewis University STEM Undergraduate Research Experience (S.U.R.E.) 2019 Faculty Mentor – Project Application

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Research Project Title: Investigating Cell Toxicity of Amyloid Beta 42 In the Presence of Different Concentrations of Metals and Metal Chelate Complexes Relevant to Alzheimer's

Disease

Alzheimer's Disease (AD) is a neurodegenerative disorder that is currently the sixth leading cause of death in the United States, however, there are no treatment options that effectively cure or prevent progression of the disease. This investigation will focus on study the *in vivo* interactions of the amyloid beta peptide under different conditions in an effort to advance drug development in AD. The amyloid beta peptide (AB) has been discovered in high concentrations in AD patients as aggregated beta sheets or oligomers, and previous investigations suggest this aggregation is promoted by oxidative stress through complex redox reactions involving free metal ions such as Cu(II). Previous work done in my lab has correlated metal ion concentration to hydroxyl radical production (•OH) and Aß peptide aggregation utilizing in vitro studies. This project focuses on investigating similar conditions to study $A\beta_{42}$ aggregation and cell toxicity in vivo using Saccharomyces cerevisiae and Caenorhabditis elegans. C. elegans is a small nematode that can be handled using *in vitro* techniques to provide a full picture of neurotoxicity inside an animal. Saccharomyces cerevisiae (yeast cells) will also be used to investigate Aβ₄₂ aggregation in a single cell system. Both C. elegans and S. cerevisiae have been shown to feature characteristic cell death pathways synonymous with mammals, and are excellent vessels for in vivo investigations of neurodegenerative disorders. The cell toxicity of Aβ₄₂ will be determined using fluorescence techniques and toxicity assays. The results of this study will be used to better understand the role copper and copper chelates in Aß aggregation that leads to cell death.