

Session/Poster#

Presenter

**B11**

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**Structural investigation of sphingomyelin synthase related protein via catalytic domain engineering**

Sphingomyelin (SM), one important lipid on the cell membrane, is synthesized by SM synthases (SMS). The SMS family has three members: SMS1, SMS2, and SMS synthase-related protein (SMSr). Although all three enzymes share four identically conserved regions (D1-D4) with a catalytic domain (Histidine-Histidine-Aspartate) in D3 and D4, SMS1 and SMS2 but not SMSr have SM synthase activity. Thus, there must be certain amino acids surrounding the catalytic domain or within the D1-D4 regions that determine the specificity of the three enzymes. The aim of my work has been to use site-directed mutagenesis to alter amino acids within the SMSr sequence to match those of the SMS1, resulting in an SMSr-mutant isoform with the ability to synthesize SM. Thus far, I have discovered two amino acids within the D3 region that resulted in the partial ability of the SMSr-mutant to produce SM. I have also identified other amino acids within D2-D4 which could be important determinants in enzyme specificity. This work gives clarity into the specific aspects of the enzymes needed to produce SM, which are involved in many metabolic diseases.