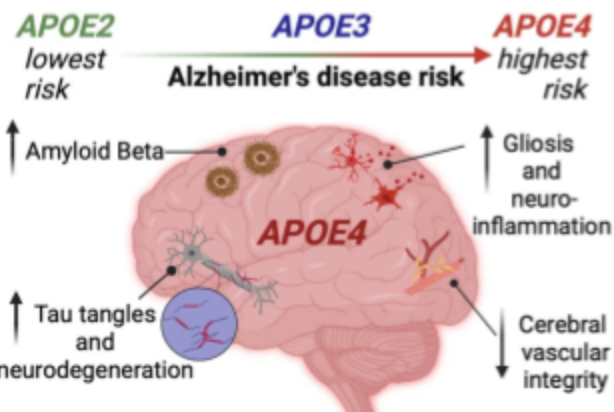
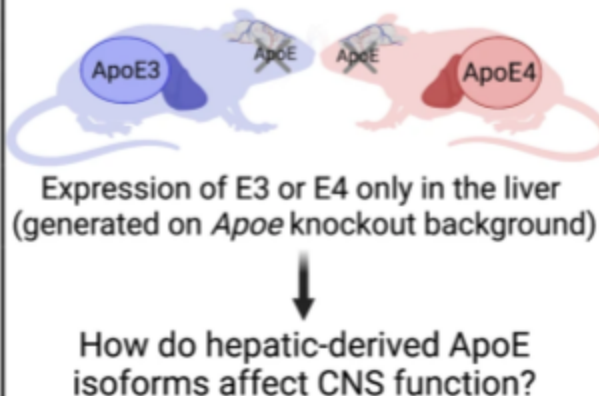


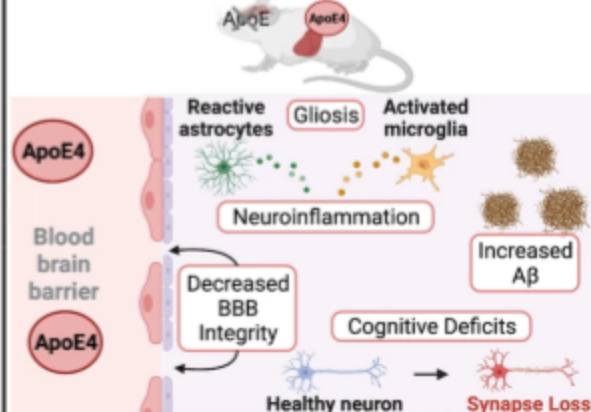
### 1. ApoE4 increases AD risk



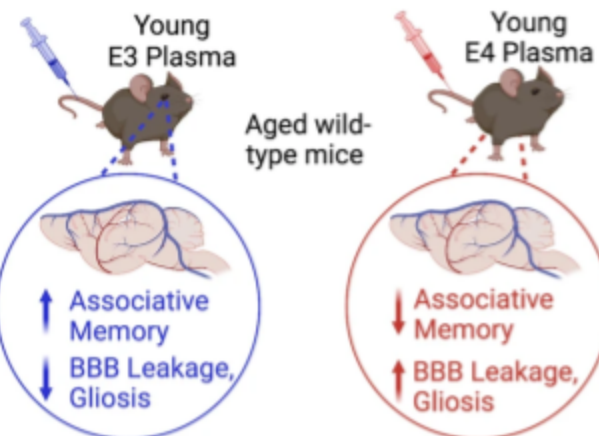
### 2. What is the role of peripheral ApoE4?



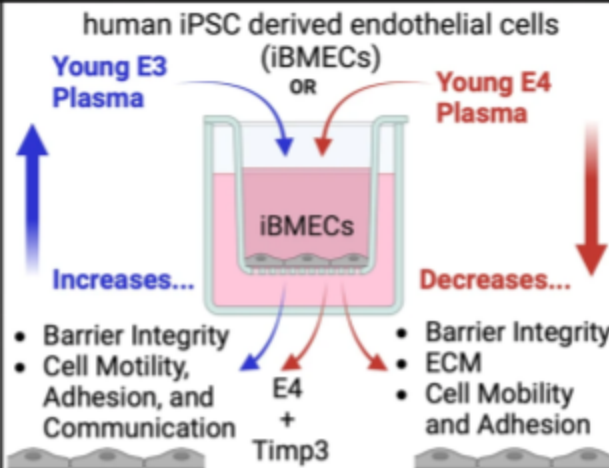
### 3. Peripheral ApoE4 has pathogenic effects



### 4. Young E3, but not E4, Plasma Rescues Age-Related Deficits



### 5. BBB Integrity Impaired by Endothelial Cell Dysfunction



### Remaining Questions

1. Hepatic E2 vs. E3 vs. E4? Other *APOE* variants?
2. Is hepatic E4 still harmful in the presence of brain E2 or E3?
3. Are peripheral E4's gain-of-toxic-function effects dose dependent? How much E4 is too much?