Propositions accompanying the dissertation:

Unraveling Molecular Mechanisms Underlying Alzheimer's disease and its Related Endophenotypes

- 1. The combined effects of common variants associated with Alzheimer's disease modify the high risk of Apolipoprotein E ϵ_4 homozygote carriers. (This thesis)
- 2. Protein truncating mutations in the Sortilin Related Receptor 1 (SORL1) gene play a crucial role in the pathogenesis of Alzheimer's disease. (This thesis)
- 3. The mutation p.P522R in the Phospholipase C Gamma 2 (*PLCG*2) gene protects against Alzheimer's disease and implicates innate immunity in the pathophysiology of the disease. (This thesis)
- 4. Between 30 and 40% of the heritability of brain lobar volumes can be explained by common genetic variation. (This thesis)
- 5. Docosahexaenoic acid (DHA) and free cholesterols in high density cholesterol (HDL) determine cognitive function. (This thesis)
- 6. Replication in metabolomics is a crucial step that needs to become standard practice.
- 7. Genetics is to biology what atomic theory is to physics -- *John Stephen Jones*.
- 8. In recent decades rapid technological advances are the drivers of scientific discoveries.
- 9. Genetic studies play a key role in determining causality.
- 10. The genetics of the brain is complex enough to crack our brains for decades to come.
- 11. "Being significantly wise does not imply you are genome-wise significant".

Sven J. van der Lee, Rotterdam, 13 October 2017