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## Genetic etiology of type 2 diabetes

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Stellingen behorende bij het proefschrift:  
**Genetic Etiology of Type 2 Diabetes:  
From Gene Identification to Functional Genomics**

- 1) The multiple rare variant with large effect hypothesis is not applicable to type 2 diabetes. Cirulli, et.al., [Nat Rev Genet](#). 2010 Jun;11(6):415-25.
- 2) Epistatic effects and/or overestimation explain the missing heritability in T2D.
- 3) Type 2 diabetes susceptibility variants demonstrate a large mechanistic heterogeneity. Dimas, et.al., [Diabetes](#). 2014 Jun;63(6):2158-71 (this thesis).
- 4) Functional genetic analysis in *disease specific tissues* is critical to understand a complex disease and study the appropriate gene (this thesis).
- 5) The role of intrauterine environment on development, The Barker Hypothesis known as the thrifty phenotype, contributes to environmental influence on epistatic effects and predisposition to T2D. Hales CN, Barker DJ [Br Med Bull](#). 2001;60:5-20.
- 6) DNA methylation does influence the susceptibility to develop type 2 diabetes Multhaup, ML, et.al., [Cell Metab](#). 2015 Jan 6;21(1):138-49; Dayeh, T, [PLoS Genet](#). 2014 Mar 6;10(3).
- 7) DNA sequencing contributes to the development of therapeutics for type 2 diabetes. Flannick, J, et.al., [Nat Genet](#). 2014 Apr;46(4):357-63.
- 8) Novel strategies for type 2 diabetes therapeutics should focus on prediabetes to prevent it from developing into full blown type 2 diabetes: Diet and exercise, genetic screening, pathway-based approaches and thinking out of the box. Plenge, RM, et.al., [Nat Rev Drug Discov](#). 2013 Aug;12(8):581-94; Fang, S, et.al., [Nat Med](#). 2015 Jan 5.
- 9) Global collaboration and integrative analysis are essential to the understanding of common complex disease.
- 10) “Without music, life would be a mistake” ---Nietzsche
- 11) “If you’re going through hell, keep going” – Winston Churchill
- 12) “It does not matter how slowly you go as long as you do not stop”---Confucius

**Michael Reynolds Erdos**  
Groningen, 18 March 2015