Stellingen behorende bij het proefschrift

"Tuberous Sclerosis Complex: Mutations, Functions and Phenotypes"

- The majority of sporadic TSC patients are somatic mosaics with a *de novo* mutation in early embryonic development.
  - This thesis

- A negative mutation analysis report does not mean that a definite TSC patient has no mutation. It just means that we have not found it.
  - This thesis

- The hamartin-tuberin complex integrates various stimuli to orchestrate the cellular processes.
  - This thesis, and Inoki, K., T. Zhu, and K.L. Guan. Cell, 2003. 115: p. 577-590

- Functional assays are the ultimate key to determining the effects of amino acid changes and identifying functionally important domains of proteins.
  - This thesis

- Having a mutation in a *TSC* gene does not make one a TSC patient.
  - This thesis

- The primary goals of medicine should be prediction and prevention.
  - Dausset J. J Biomed Biotechnol 2001;1:1-2

- Clinical genetic testing should be performed with the highest possible level of accuracy, since both false positive and false negative results can have a devastating impact on the lives of patients.
  - Strom C.M. Mutation Research 2005; 573:160-167

- Genomic information is uniform among the different cells of a complex organism. It is the epigenome that makes the difference.
  - Fraga M.F. Proc Natl Acad Sci U S A 2005; 102:10604-10609

- The right conclusions can only be reached with the right interpretation of the results.
  - Zeesman S. et al. *Paternal transmission of Fragile X Syndrome.* Am J Med Genet. 2004;129A:184-189 & Steinbach D., Steinbach P. *No evidence of paternal transmission of Fragile X Syndrome.* Am J Med Genet. 2005; 136A:107-108

- The occurrence of transforming proto-oncogene mutations in the germline necessitates redefinition of (proto-)oncogenes.
  - Aoki Y. et al. Nat Genet. 2005 Oct;37:1038-1040

- Only those who risk going too far can possibly find out how far one can go."
  - T.S. Elliot