Stellingen behorende bij het proefschrift

Transcriptional Control During Hematopoietic Development Transcription factor binding and chromatin conformation dynamics

- 1. Intra-aortic hematopoietic clusters (IAHCs) consist of premature hematopoietic stem cells (pre-HSCs) which can mature into true HSCs by the combinatorial actions of a variety of transcription factors. (This Thesis & Taoudi S et al. Cell Stem Cell. 2008;3:99–108)
- Unraveling the fine-scale three-dimensional structure of the genome and its impact on gene regulation is a major goal of current epigenetic research, with direct implications for understanding the molecular mechanisms underlying human phenotypic variation and disease susceptibility. (This thesis)
- 3. Even though genome-wide chromatin conformation analysis approaches appear to be the future in epigenetic research, site-specific approaches like multiplexed 3C-sequencing remain valuable, and are currently still indispensable, for unraveling the fine-scale chromatin structural conformation without the need for substantial computational resources and the high sequencing costs. (This Thesis)
- 4. With the majority of disease-associated genomic alterations located in the non-coding 'regulatory genome', novel enhancer targeting strategies, e.g. transcription factor tethering or genome editing, could become widely applicable therapeutic strategies for molecular diseases. (*This thesis*)
- 5. The erythroid stage-specific enhancer of *Bcl11a* and its conserved enhancer in humans are interesting targets for γ-globin reactivation in therapeutic strategies for β-hemoglobinopathies like sickle cell disease and β-thalassemia. (*This thesis & Bauer D E et al. Science 2013;342:253–257*)
- 6. The statement that "recent findings in metazoan systems have revealed that much of the transcription regulation occurs well after the recruitment of Pol II and the transcription machinery to a gene promoter, through controlling pausing and the efficiency of early elongation. Thus, we are in the midst of a paradigm shift in our understanding of gene regulation as it applies to higher eukaryotic systems. (Adelman, K. & Lis, J.T. Nature Reviews Genetics 2012;13:720-731)" comes a little late. (Frank Grosveld)
- 7. We think in generalities, but we live in detail. (Alfred North Whitehead)
- 8. Research is like top sport; hard work is essential, but success not guaranteed.
- 9. A good work-life balance turns out to be just as hard to achieve as it is essential to have.
- 10. The greatest mistake you can make in life is continually fearing that you'll make one. (Elbert Hubbard)
- 11. Research is what I'm doing when I don't know what I'm doing. (Wernher von Braun, New York Times interview, 16 December 1957)