## Stellingen behorend bij het proefschrift

## Epigenetics in *MLL*-rearranged Infant Acute Lymphoblastic Leukemia

- MLL-rearranged infant acute lymphoblastic leukemia is characterized by pronounced promoter DNA hypermethylation leading to silencing of a substantial number of genes. (this thesis)
- 2. Promoter DNA hypermethylation in t(4;11)-positive infant acute lymphoblastic leukemia encompasses methylation at microRNA loci resulting in aberrant expression of microRNA target genes. (this thesis)
- 3. Clofarabine reverses aberrant epigenetic regulation of tumor suppressor gene activity in *MLL*-rearranged acute lymphoblastic leukemia cells. (*this thesis*)
- 4. Histone deacetylase inhibitors induce leukemic cell death in *MLL*-rearranged infant acute lymphoblastic leukemia *in vitro*, which is in part due to down-regulation of the oncogenic MLL-AF4 fusion. (*this thesis*)
- 5. MLL-rearranged infant acute lymphoblastic leukemia is an epigenetic disease which requires treatment with epigenetic drugs. (this thesis)
- 6. The inherently Darwinian character of cancer is the primary reason for therapeutic failure, but it may also hold the key to more effective control. (*Mel Greaves, Nature, 2012*)
- 7. Genetic changes occur in the development of cancers, but there also are epigenetic changes, and those come first. (*Andrew P. Feinberg, Nature Reviews Genetics, 2006*)
- 8. Epigenomics is where genomics was 30 years ago, when everyone was working on part of the puzzle. Nevertheless, the field is currently moving forward at high speed. (Peter A. Jones, Nature, 2006, and Stephen B. Baylin, Nature Reviews Cancer, 2011)
- 9. The important thing in science is not so much to obtain new facts as to discover new ways of thinking about them. (William L. Bragg, youngest Nobel Prize Winner)
- 10. It takes two to tango in translational research: close collaboration of medical doctors and basic biologists is indispensable.
- 11. Vectatio iterque et mutata regio vigorem dant (Journeys and change of venues impart new vigor to the mind). (*Lucius A. Seneca*)