Stellingen behorende bij het proefschrift

“TCR TRANSLOCATIONS AT THE NORMAL-MALIGNANT T CELL INTERFACE”

Rotterdam, 11 september 2013

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1. The V(D)J recombination machinery is not the driver, but rather the facilitator in the formation of TCR translocations and non-TCR chromosomal aberrations in T-ALL. (This thesis)

2. DNA double strand breaks in breakpoint cluster regions are primarily induced in the presence of a genetic element of sequence vulnerability, while breaks at single breakpoint sites lacking such elements of vulnerability are more likely induced by chance or perhaps due to a patient specific genetic vulnerability. (This thesis)

3. The time frame of the V(D)J recombination process at the TCR locus primarily determines the “window of opportunity” for a TCR translocation to occur. (This thesis)

4. Nuclear spatial organization of TCR and oncogene loci in thymocytes is random. (This thesis)

5. Genetic vulnerability to double stand break induction primarily determines the frequency of TCR translocation formation, but the positive selection by the oncogenic effect resulting from the translocation determines the frequency of oncogene involvement in T-ALL. (This thesis)

6. Genetic changes occur in development of cancers, but there also are epigenetic changes and those come first. (Adapted from A. Feinberg M.D., Press release Johns Hopkins Medicine [online], 2005)
7. If there is no innovative fundamental research, there will be nothing to translate. (D. Vance, Editorial. Biochimica et Biophysica Acta Newsletter [online], 2009)

8. A new era of cancer research emerges in which patients may be stratified for appropriate therapy on the basis of the DNA damage response status of their tumor, rather than on the tissue of origin. (Adapted from N. J. Curtin, Nature Reviews Cancer, 2012)

9. DNA neither knows nor cares. DNA just is. And we dance to its music. (M. K. Cunningham, God and Evolution: A Reader, page 191, 2007)

10. It is a capital mistake to theorize before one has data. Insensibily one begins to twist facts to suit theories, instead of theories to suit the facts. (Arthur Conan Doyle, Sherlock Holmes, A Scandal in Bohemia, page 163, 1891)

11. Eindelijk mijn persoonlijk bewijs dat “Antilianen” niet lui zijn!