Cover Page



Universiteit Leiden



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Title: Identification of therapeutic targets and antisense oligonucleotide mediated exon

skipping based therapies in arthritis

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- 1) Autoantibodies play a role in development of arthritis but they are not sufficient to develop the full-blown disease (This thesis)
- 2) Not all exons are easy to skip by using antisense oligonucleotides (AONs) (This thesis)
- 3) In-vivo delivery of AONs to the target organ/tissue/cell type can be the limiting factor that determines their therapeutic efficacy (This thesis)
- 4) The balance between activating and inhibitory Fc receptors is important and regulates the susceptibility and severity to RA (This thesis).
- 5) Because RA is a multifactorial and heterogeneous disease, an accurate definition of the different sub-forms is required for the development of effective therapies.
- 6) The utility of conditional KO mouse models in the *in vivo* analysis of myeloid cell function is severely hampered by the lack of transgenic mouse lines expressing Cre recombinase exclusively in a particular myeloid subset.
- 7) Modulation of splicing by antisense oligonucleotides has a lot of potential to develop novel therapies for many diseases in the near future, but also as biological research tool.
- 8) Not all the chemical modifications used in antisense oligonucleotides are suitable for in-vivo applications due to their toxicity or other possible side effects.
- 9) "Nothing in life is to be feared, it is only to be understood. Now is the time to understand more, so that we may fear less." MARIE CURIE, *Primary source uncertain, mentioned by Glenn T. Seaborg, 'Need We Fear Our Nuclear Future?' Bulletin of the Atomic Scientists (Jan 1968), 24, No. 1*
- 10) "Be occupied, then, with what you really value and let the thief take something else." MEVLANA JALALUDDIN RUMI, Mathnawi II: 1505-1507 Version by Camille and Kabir Helminski "Rumi: Daylight" (1994)
- 11) "If scientific analysis were conclusively to demonstrate certain claims in Buddhism to be false, then we must accept the findings of science and abandon those claims." Dalai Lama XIV, The Universe in a Single Atom: The Convergence of Science and Spirituality (Jan 2005)