

Propositions *belonging to this thesis*

The Toll of too much Interferon:

The systemic Interferon signature in the pathogenesis of Sjögren's syndrome

1. The Interferon type I signature in primary Sjögren's syndrome identifies a subgroup of patients with higher disease activity as assessed by the EULAR Sjögren's Syndrome Disease Activity Index. (*this thesis*)
2. MxA is a clinically applicable biomarker for measuring systemic Interferon type I activity in primary Sjögren's syndrome, systemic lupus erythematosus and systemic sclerosis. (*this thesis*)
3. Triggering of the ribonucleic acid sensing Toll-like receptor 7 results in the upregulation of the cytosolic RIG-I like receptors RIG-I and MDA5, and downregulation of Toll-like receptor 9. (*this thesis*)
4. Regulatory T cells are positively correlated with IDO and IFN type I activity in patients with primary Sjögren's syndrome. (*this thesis*)
5. Patients with primary Sjögren's syndrome will benefit from optimising therapies tailored to specific subgroups of the disease, according to their distinct Interferon signatures. (*this thesis*)
6. Innate immunity exhibits enhanced innate immune responses after exposure to secondary infections, giving rise to innate immunological memory, termed "trained immunity". (*Netea M.G. et al, Cell Host Microbe 2011, 9:355-361*)
7. Together with the bacterial microbiome, the retrovirome shapes both the magnitude and the quality of the immune response. (*Grasset E.K., Cerutti A., Science 2014, 346:1454-1455*)
8. The discovery of the central nervous system lymphatic system calls for a reassessment of basic assumptions in neuroimmunology and sheds new light on the aetiology of neuroinflammatory and neurodegenerative diseases associated with immune system dysfunction. (*Louveau A. et al. Nature 2015, 523:337-341*).
9. Metastatic behaviour of tumor cells can be phenocopied through extracellular vesicle exchange between cells. (*Zomer A. et al. Cell 2015, 161:1046-1057*)
10. Brief, repeated exposures to sunlight are more efficient in boosting the body's vitamin D supply compared to longer exposures, that additionally cause skin damage and increase the risk of skin cancer. (*Mead. M.N. Environ Health Perspect 2008, 116:A160-167*)
11. "If you can't fly, then run, if you can't run, then walk, If you can't walk, then crawl, but whatever you do, you have to keep moving forward " – *Martin Luther King*