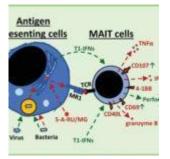
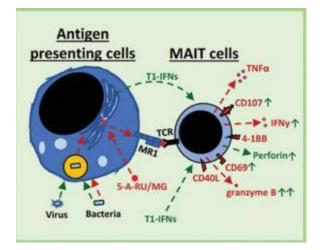
## Type I interferon costimulation of MAIT cell function





Type I interferons enhance activation and effector responses of TCR-stimulated blood and liver-derived MAIT cells. Type I interferons contribute to MAIT cell by riboflavin activation producing bacteria, and act directly on MAIT cells. Together, this suggests type I interferons are important early modulators of the MAIT cell TCR response. (Source: Journal European of

Immunology)

Mucosal-associated invariant T cells (MAITs) are a population of non-conventional innate-like T cells that recognize vitamin B metabolites, primarily riboflavin, via the MHC-related protein-1 (MR1) molecule. MAITs can also be activated in a TCR-independent mechanism primarily via IL-12 and IL-18, and can respond to pathogens that lack a riboflavin pathway such as viruses. Recent evidence suggest that Type I -interferons (T1-IFNs) IFN- $\alpha$  and IFN- $\beta$  can activate of MAITs . However, this mechanism is not fully understood. Researchers from New Zealand (Lamichane *et al.*, 2020) aimed to investigate this and specifically focused on the role of T1-IFN-mediated activation of MAITs in the context of MR1-TCR activation.

Lamichane *et al.*, showed that both IFN- $\alpha$  and IFN- $\beta$  separately enhanced MAIT cell activation and cytokine function only in a co-stimulatory manner. Whereas both T1-IFNs were able to induce expression of cytotoxic molecules Granzyme B, Perforin and CD107 by MAITs in the absence of MR1-TCR activation, albeit at lower proportions compared to TCR-activated MAITs. Researchers further showed that T1-IFNs induced by influenza-A-virus "*enhanced the activation of TCR-stimulated MAIT cells* in a *T1-IFN dependent manner*. *This may have implications for MAIT cell activation during lower respiratory tract infections, where co-infection with bacteria and respiratory viruses are common*." Additionally, these results provide a potential mechanism by which viruses can activate MAIT cells activation.

In summary results by Lamichane *et al.*, establish an important immunomodulatory and co-stimulatory role of T1-IFNs during TCR-mediated MAIT cell activation. T1-IFNs-induced activation could act by either activating antigen-presenting cells to upregulate MR-1 on these, or directly interact with MAITs cells via T1-IFN receptor, which MAITs express at high levels.

Journal Article: Lamichane et al., 2019. Type I interferons

<u>are important co-stimulatory signals during T cell receptor</u> <u>mediated human MAIT cell activation.</u> European Journal of Immunology.

Article by Cheleka AM Mpande