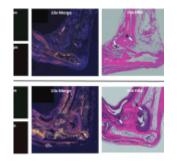
Novel protein linked to rheumatoid arthritis pathogenesis



In a recent paper, researchers have discovered and described a protein known as sulfatase-2 that may be a key role player in the pathology of <u>rheumatoid arthritis</u> (RA) (Figure 1). This finding helps researchers understand the <u>mechanisms</u> underlying the inflammation seeing in RA. Currently there is no cure, but these findings may lead to improved treatments of the disease.

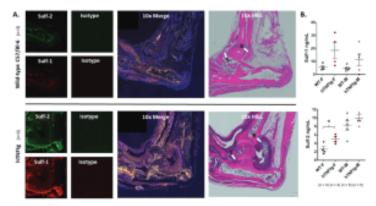


Figure 1: Extracellular sulfatase levels are elevated in inflamed joint tissues and serum of a human TNF-transgenic mouse model of RA. Extracellular sulfatase protein levels were compared in human TNFtransgenic mice of the 3647 line (hTNFtg) and littermate wild-type (WT) C57BL/6 mice. A Murine Sulf-1

and Sulf-2 proteins were аt qualitatively higher levels in ankle sections of hTNFtg mice as determined by fluorescence IHC. Images are shown with isotypematched controls on the same tissue and slide. H&E images illustrate synovial hyperplasia and leukocyte infiltration, which are characteristic of hTNFtg mice. Bone (B) and synovial tissues (S) are labeled. B Murine Sulf-1 and Sulf-2 proteins in serum samples from hTNFtg and littermate WT mice were quantitated by ELISA. Data are presented as the mean \pm SEM. *p < 0.05

RA is a chronic disease in which the immune system attacks the body's own joint tissues. Tumor necrosis factor-alpha (TNF- α) is a common and major inflammatory marker driving RA and is the target of many therapies. Patients can develop resistant to treatments overtime; therefore new treatments are needed.

Investigating sulfatase-2 in synovial fibroblasts (cells which line the joints and lubricate them), the researchers removed the protein from one group of cells and stimulated the cells with TNF- α . They were able to show that the cells without sulfatase-2 showed reduced inflammation. This could open the field to new research into using the inhibition of sulfatase-2 to reduce RA symptoms.

Journal article: Siegel, R.J., et al., 2022. Extracellular sulfatase-2 is overexpressed in rheumatoid arthritis and mediates the TNF- α -induced inflammatory activation of synovial fibroblasts. Cellular & Molecular Immunology.

Summary by Stefan Botha