Graphical Abstract

Monocyte

B cell

Active CD Healthy/non-active CD Deep Ulcers/ Lamina propria **MALT** Inflammation Macroscopic Inflammation GO-pathways: **ER Stress Epithelial Differentiation** Lymphocyte Differentiation Inflammatory Cytokine Macroautophagy Neural Signal mRNA Processing Immune Cell Differentiation Ion Transport Translation **Endothelial Activation** Cell Division CCL2/3, CXCL1/2, CD14, IL6, OSM, PTGS2, IL1A/B Anti-TNF non-response Anti-TNF response Aquired and genetic IL-10 non-response **TNF** IL-10 IL-23 IL-21 **TNF** IL-17A/F IL-22 IFN-γ TNF IL-17A/F IL-22 TNF Potential therapeutic Macrophage CD4⁺ T cell Intestinal Epithelial Cells implications: **Dendritic Cell** IL-1 targeting FOXP3+ regulatory T cell Intestinal microbes Anti-IL-23p19