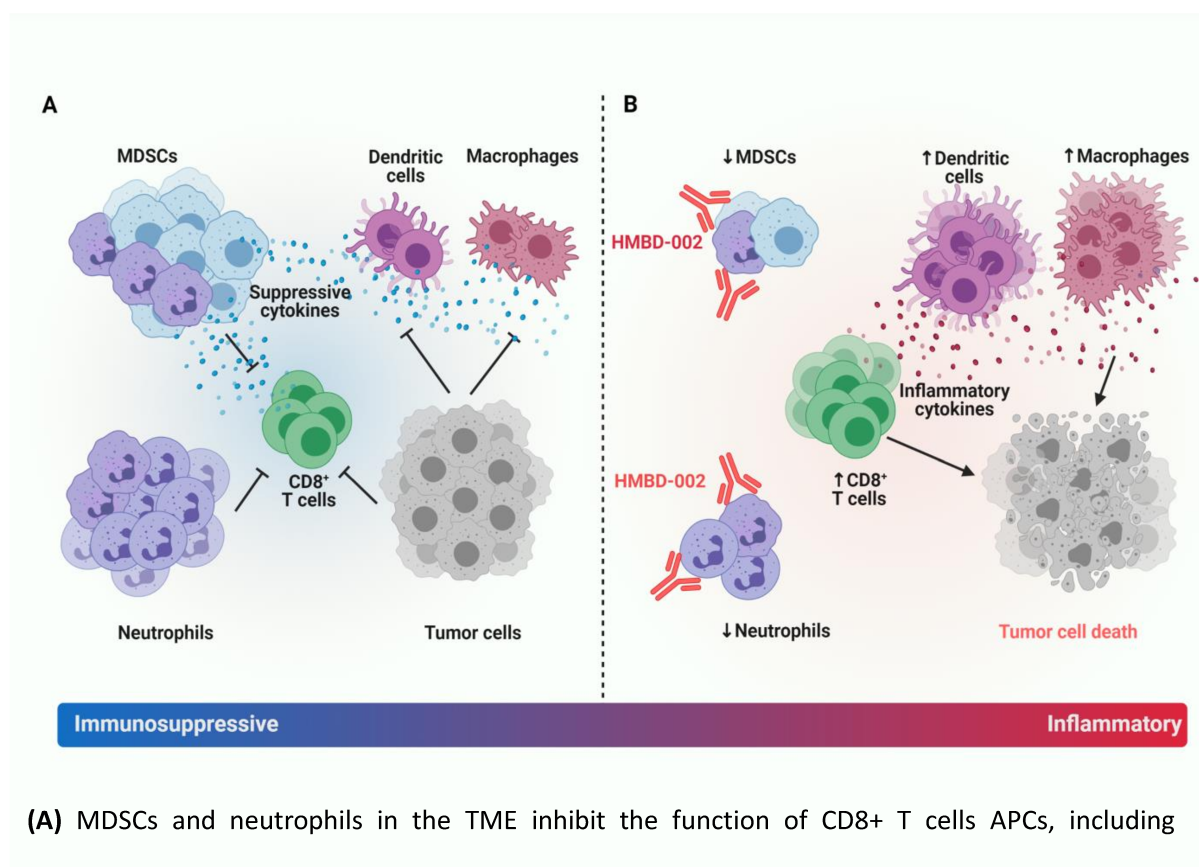


**Title: A rationally targeted anti-VISTA antibody that blockades the C-C' loop region can reverse VISTA immune suppression and remodel the immune microenvironment to potently inhibit tumor growth in an Fc independent manner**

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**(A)** MDSCs and neutrophils in the TME inhibit the function of CD8+ T cells APCs, including macrophages and dendritic cells leading to an immunosuppressive TME permissive of tumor growth. **(B)** HMBD-002 binds to and blocks VISTA on neutrophils and MDSCs, releasing their immunosuppressive effects and shifting the TME to a pro-inflammatory anti-tumorigenic phenotype, characterised by increased inflammatory cytokine release and increased proliferation and activation of CD8+ T cells and APCs.