

Evaluating the safety profile of defibrotide in sickle cell disease: an *in vitro* study

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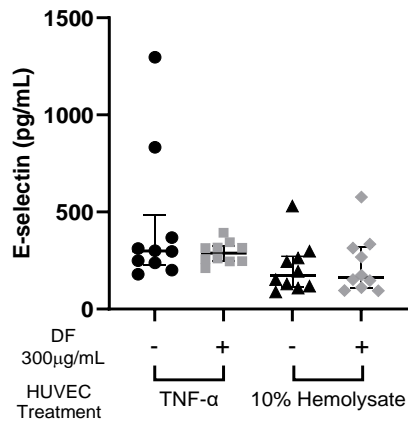
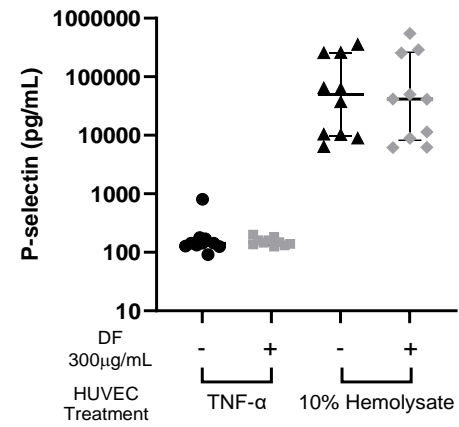
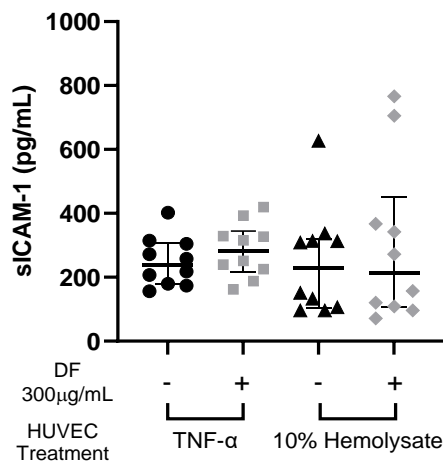
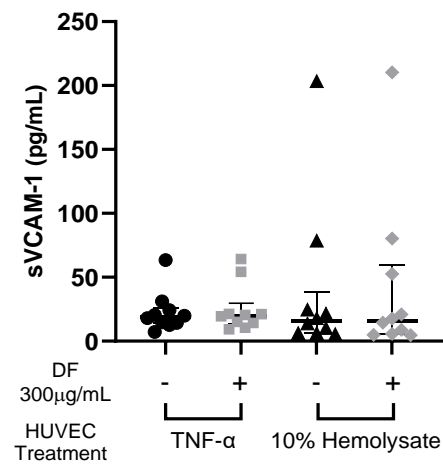
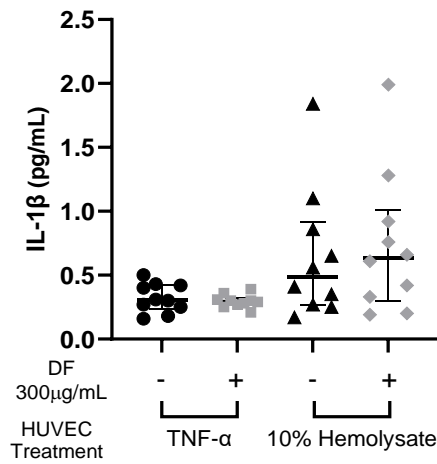
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Supplemental Figure legend:

Figure S1. Defibrotide does not modify the release of soluble E-selectin, P-selectin, ICAM-1, VCAM-1, IL-1 β and vWF by endothelial cells. After 16-40 hours of culture under flow conditions (using KIMA pumps, Cellix) in μ -slides (IBIDI) coated with fibronectin, human umbilical vein endothelial cells (HUVEC) were perfused for 4 hours with culture medium containing 20 ng/mL TNF- α , with or without 300 μ g/mL Defibrotide (DF) or 10% hemolysate +/- 300 μ g/mL DF. (A) Soluble E-selectin, (B) P-selectin, (C) ICAM-1, (D) VCAM-1, (E) IL-1 β , and (F) von Willebrand factor (vWF) concentrations were measured by Luminex (MagPix, Thermofisher) and by ELISA for vWF (Thermofisher) in HUVEC supernatant collected after the 4-hour treatment with TNF- α or hemolysate (samples treated with DF in grey). Data are presented as median with interquartile range and were analyzed using the Wilcoxon test (comparisons to the 0 μ g/mL condition) on GraphPad Prism 10. All p-values were > 0.05.

A**Soluble [E-selectin] in HUVEC supernatant****B****Soluble [P-selectin] in HUVEC supernatant****C****Soluble [ICAM-1] in HUVEC supernatant****D****Soluble [VCAM-1] in HUVEC supernatant****E****[IL-1 β] in HUVEC supernatant****F****[vWF] in HUVEC supernatant**