Do monocytes contribute to haemolytic uremic syndrome?

Shiga-toxin (Stx) producing enterohaemorrhagic E.coli (STECs) have been implicated in outbreaks of hemolytic uremic syndrome (HUS). HUS is a life-threatening disease characterized by haemolytic anaemia, thrombocytopenia (low platelet count) and uremia (acute kidney injury/failure). To date no therapy against HUS, highlighting the need for increased research in this area.

Research on HUS has suggested a potential immunopathological contribution by monocyte to HUS pathology. Monocytes subsets in the blood can be distinguished by differential expression of CCR2 and Gr1. Researchers aimed to determine the contribution of Gr1\textsuperscript{high} monocytes during HUS.

Pohl \textit{et al.}, used an an experimental mouse model of HUS. Mice were intravenously injected with Stx and LPS, this resulted in increased blood urea nitrogen (BUN) and other symptoms characteristic of acute kidney injury. Acute HUS (1 day post Stx/LPS injection) resulted in increased
levels of renal monocytes Gr1\textsuperscript{high} expressing high levels of CD11b and were CX\textsubscript{3}CR1\textsuperscript{low}MHC ClassII\textsuperscript{CD64high}. Using non-invasive imaging techniques researchers observed extensive kidney damage which was associated with increased levels of Gr1\textsuperscript{high} monocyte.

HUS has also been associated with increased levels of chemokines CCL2, CCL3 and CCL5, that that recruit monocytes via the chemokine receptor CCR2. High expression of CCR2 on Gr1\textsuperscript{high} monocytes suggested that CCR2 is required for Gr1\textsuperscript{high} monocytes to contribute to immunopathology. Indeed Pohl 	extit{et al.}, demonstrated that in absence of CCR2 there was a reduction on recruitment of Gr1\textsuperscript{high} monocytes, this was associated with reduced kidney damage and increased survival.

The research conducted by Pohl 	extit{et al.}, showing mechanism by which CCL2 via CCR2 contributes to Stx induced HUS immunopathology by recruiting Gr1\textsuperscript{high} monocytes, cells that contribute to HUS immunopathology.

Journal Article: Pohl 	extit{et al.}, 2018. CCR2-dependent Gr1\textsuperscript{high} monocytes promote kidney injury in shiga toxin-induced hemolytic uremic syndrome in mice. European Journal of Immunology.

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