





International PhD Program in Experimental Medicine

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Immune Surveillance of the liver

Kupffer cells (KCs) are highly abundant, intravascular, liver-resident macrophages known for their scavenger and phagocytic functions. KCs can also present antigens to CD8+ T cells and promote either tolerance or effector differentiation, but the mechanisms underlying these discrepant outcomes are poorly understood. Here, we used a mouse model of hepatitis B virus (HBV) infection – where HBV-specific naïve CD8+ T cells recognizing hepatocellular antigens are driven into a state of immune dysfunction – to identify a subset of KCs (referred to as KC2) that cross-presents hepatocellular antigens upon interleukin-2 (IL-2) administration, thus improving the antiviral function of T cells.

Removing MHC-I from all KCs – including KC2 – or selectively depleting KC2 Impaired the capacity of IL-2 to revert the T cell dysfunction induced by intrahepatic priming.

In summary, by sensing IL-2 and cross-presenting hepatocellular antigens, KC2 overcome the tolerogenic potential of the hepatic microenvironment, suggesting new strategies for boosting hepatic T cell immunity.

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