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28 Sep 2015 ... Chronic liver infections caused by **hepatitis B** and **hepatitis C virus** (HBV and ... (NK) **cells**, and up-regulation of immune checkpoint **pathways** have been ... also by **co-stimulatory** and co-inhibitory molecules that **modulate response** ... The expansion of AFP-specific CD4<sup>+</sup> T cells in HCC patients after TACE ...

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27 Mar 2013 ... While **virus-specific T-cell responses** are able to successfully clear the **virus** ... severe liver diseases, such as liver cirrhosis and **hepatocellular carcinoma**. .... Key characteristics of CD4<sup>+</sup> and CD8<sup>+</sup> T cell in HBV and HCV ..... **modulation** of several co-inhibitory and **costimulatory pathways** might be beneficial.

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8 May 2015 ... The effector function of CD8<sup>+</sup> T cells is prone to be impaired by ... In addition, interferon treatment may gain further benefits for HCC patients with **hepatitis B** or C virus ... Co-inhibitory signals diminish the intensity of anti-tumor **response** even ... the two **pathways** were suggested by clinical observations[44].

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22 Jan 2013 ... One of the best characterized T cell **co-stimulatory** signals is mediated by the ... of anti-CTLA-4 antibody increases memory CD8<sup>+</sup> T cell expansion, because ... CTLA-4

Name of Journal: World Journal of Gastroenterology

ESPS Manuscript NO: 25829

Manuscript Type: REVIEW

**Specific CD8<sup>+</sup> T cell response immunotherapy for hepatocellular carcinoma and viral hepatitis**

Elia Moreno-Cubero, Juan-Ramón Larrubia

### Abstract

Hepatocellular carcinoma (HCC), chronic hepatitis B (CHB) and chronic hepatitis C (CHC) are characterized by exhaustion of the specific CD8<sup>+</sup> T cell response. This process involves enhancement of negative co-stimulatory molecules, such as PD-1, CTLA-4, 2B4, Tim-3, CD160 and LAG-3, which is linked to intrahepatic overexpression of some of the cognate ligands, such as PD-L1, on antigen presenting cells and thereby favouring a tolerogenic

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Wong, Yik Chun, Szun Szun Tay, Geoffrey W. McCaughan,  
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## Immunotherapy of hepatocellular carcinoma

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