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Name of Journal: *World Journal of Stem Cells*

ESPS Manuscript NO: 28727

Manuscript Type: Review

Immunomodulatory oligonucleotide IMT504: Effects on mesenchymal stem cells as a first-in-class immunoprotective/immunoregenerative therapy

11
Jorge Zorzopulos, Steven M Opal, Andrés Hernando-Insúa, Juan M Rodriguez, Fernanda Elias, Juan Fló, Ricardo A López, Norma A Chasseing, Victoria A Lux-Lantos, Maria F Coronel, Raul Franco, Alejandro D Montaner and David L Horn

Abstract

The immune responses of humans and animals to insults (*i.e.* infections, traumas, tumoral transformation and radiation) are based on an intricate network of cells and chemical messengers. Abnormally high inflammation immediately after insult or abnormally prolonged pro-inflammatory stimuli bringing about chronic inflammation can lead to life-threatening or severely debilitating diseases. Mesenchymal stem cell (MSC) transplant has proved to be an effective therapy in preclinical studies which evaluated a vast diversity of inflammatory conditions. MSCs lead to resolution of inflammation, preparation for regeneration and actual regeneration, and then ultimate return to normal baseline or homeostasis. However, in clinical trials of transplanted MSCs, the expectations of great medical benefit have not yet been fulfilled. As a practical alternative to MSC transplant, a synthetic drug with the capacity to boost endogenous MSC expansion and/or activation may also be effective. Regarding this, IMT504, the prototype of a major class of immunomodulatory oligonucleotides, induces

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