

Characterization and administration of bone marrow-derived mesenchymal stem cells in an animal model of congenital hydrocephalus

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Bone marrow-derived mesenchymal stem cells (BM-MSK) are considered as a potential therapeutic tool in neurodegenerative diseases, due to their ability to migrate to degenerated tissues and the production of growth factors. Congenital hydrocephalus is a disorder characterized by a degeneration of the periventricular cerebral parenchyma and the white matter. In the present study, using an animal model of congenital hydrocephalus, the *hyh* mouse, it has been studied the capacity of the BM-MSK to reach the degenerated regions exhibiting glial reactions and their probable neuroprotector effects.

The BM-MSK were isolated from two sources: a) transgenic mice expressing the monomeric red fluorescent protein (mRFP1); b) wild type mice. In the second case, the BM-MSK were labelled *in vitro* using bromodeoxyuridine, a fluorescent cell tracker and the lipophilic DiI. Before application, the cells were analysed using flow cytometry and immunofluorescence. The BM-MSK were injected into the retro-orbital sinus or into the lateral ventricle of *hyh* mice. After 24/96 hours of administration, they were detected under light, confocal and electron microscopes.

The injected BM-MSK reached the degenerated periventricular regions and the disrupted neurogenic niches. They were detected in the periventricular parenchyma, around periventricular blood vessels and in the ventral meninges. Most of the applied BM-MSK expressed the glial cell-derived neurotrophic factor (GDNF), in the same way as the periventricular reactive astrocytes, suggesting a possible neuroprotector effect.

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