Mesenchymal Stem Cells Encapsulated in a Self-Assembling Nanopeptide Scaffold Attenuate Neuroinflammation and Behavioral Function in a Model of Traumatic Brain Injury in Rats

Mohammad Moein Shirzad¹, Farzin Naseri¹, Ghazale Biglari¹, Ali Hasani Dooghabadi¹, Ali Jahanbazi Jahan Abad², Arezou Es’haghabadi², Ali Gorji³,⁴, Sajad Sahab Negah⁵*

¹Neuroscience Department, Faculty of Medicine, Mashhad University of Medical Sciences, Mashhad, Iran
²Shefa Neuroscience Research Center, Khatam Alanbia Hospital, Tehran, Iran
³Department of Neurosurgery, Westfälische Wilhelms-Universität Münster, Münster, Germany
⁴Epilepsy Research Center, Westfälische Wilhelms-Universität Münster, Robert-Koch-Straße 45, 48149 Münster, Germany

Abstract

Traumatic brain injury is one of the major causes of brain function impairments and surgery is involved in the treatment program of many cases although it cannot rescue the brain functions completely and is confined to reduce the second injury. In this study, we aimed to investigate the effects of mesenchymal stem cells encapsulated in RADA1-GGS IKVAV, surgically injected into the lesion site. 36 male Wistar rats underwent an acute model of TBI. Subjects were divided into 5 groups, each consisting of 6 to 9 rats: Sham (receiving no treatment), PBS, GSIKVAV, Mesenchymal Stem Cells (MSCs), GSIKVAV+MSCs. MSCs were stained with BrdU. Flow cytometry was done to characterize MSCs. Several Behavioral tests were conducted: Open Field (OF) and Elevated Plus Maze (EPM) to assess anxiety-like behavior and modified Neurological Severity Score (mNSS) to evaluate the sensory-motor function. Subjects were euthanized at day 30. IHC was carried out to measure MSCs’ viability and differentiation. Also, Western blotting was performed to check for inflammatory factors including toll like receptor3, 4, tumor necrosis factor α and glial fibrillar acidic protein. There was a significant decrease ($P < 0.05$) between MSCs and MSCs + GSIKVAV groups in mNSS score. In addition, the number of entries to the open arm in the EPM test and total distance in OF test was significantly higher ($P < 0.05$). Our data suggest that using MSCs in combination with GSIKVAV can rescue cognitive function. These findings suggest that new assembly peptides can be a new and potential therapy for TBI patients.

Keywords: Tissue Engineering, Neuroinflammation, Functional Recovery, Traumatic Brain Injury

*Corresponding Author: Sajad Sahab Negah
E-mail: sahabnegahs@mums.ac.ir