Targeted Delivery of siRNA in a Nano-Particle Suppress Glioblastoma Stem Cells

Elham Poonaki1,2, Ali Gorji1,3,4,5, Sajad Sahab Negah1,3*

1Neuroscience Department, Faculty of Medicine, Mashhad University of Medical Sciences, Mashhad, Iran
2Department of Biotechnology, Islamic Azad University, Damghan, Iran
3Shefa Neuroscience Research Center, Khatam Alanbia Hospital, Tehran, Iran
4Department of Neurosurgery, Westfälische Wilhelms-Universität Münster, Münster, Germany
5Epilepsy Research Center, Westfälische Wilhelms-Universität Münster, Robert-Koch-Straße 45, 48149 Münster, Germany

Abstract

Cancer stem cells (CSCs) are suggested as the most dominant causes of recurrence due to their permanent self-renewal and resistance to common cancer treatment in glioblastoma multiform (GBM) which is recognized as the most malignant of brain tumors. It has been indicated that Retinoblastoma-binding protein 5 (RBBP5), a main part of Mixed lineage leukemia protein-1 (MLL1), plays a significant role in cancer stem cell survival. In this study the viability of CSCs derived from human GBM will be evaluated by knocking down the RBBP5 via their SiRNA. To enhance passing the SiRNA thorough the blood-brain barrier, PLGA nanocarrier will be used. CSCs isolated from Human GBM and cultured. To target specific stem cells involved in the growth and spread of cancer cells, CD133+ as a CSCs antigen will be conjugated on the surface of PLGA and then conjugated to SiRNA. The viability, proliferation, apoptosis, and differentiation of CSCs will be performed as a primary outcome. The expression of self-renewal markers such as NANOG, SOX2, CD133+, and Ki67 will be assessed as a secondary outcome. The probable prediction is descending the population of CSCs in the tumors that supposed to recurrence and short life expectancy. In conclusion, the nano drug can decrease the number of CSCs and increase the survival time in the GBM patients.

Keywords: Glioblastoma, Cancer Stem Like Cells, Nanoparticle

*Corresponding Author: Sajad Sahab Negah
Email: sahabnegahs@mums.ac.ir