Changing fate: Reprogramming inflammatory cells to induced neurons

Negar Mahmoudi ^{*a, b, c**}, Alan R Harvey ^{*d*}, Clare L. Parish ^{*e*}, Richard J. Williams ^{*f*}, and David R. Nisbet ^{*a, b, g, h, i*}

^a ACRF Department of Cancer Biology and Therapeutics, The John Curtin School of Medical Research, ANU College of Health & Medicine, Australia

^bResearch School of Chemistry, ANU College of Science, Australia

^c ANU College of Engineering & Computer Science, ACT 2601, Australia

^d School of Human Sciences, The University of Western Australia, and Perron Institute for Neurological and Translational Science, Perth, WA 6009, Australia

^e The Florey Institute of Neuroscience and Mental Health, The University of Melbourne, Parkville, Melbourne, VIC 3010, Australia

^f iMPACT, School of Medicine, Deakin University, Waurn Ponds, VIC 3216, Australia[,] Federation Uni

^g The Graeme Clark Institute, The University of Melbourne, Melbourne, Australia

^h Melbourne Medical School, Faculty of Medicine, Dentistry and Health Science, The University of Melbourne, Melbourne Australia

ⁱ Department of Biomedical Engineering, Faculty of Engineering and Information Technology, The University of Melbourne, Melbourne, Australia

* Presenter's email: david.nisbet@unimelb.edu.au

Abstract

The incapacity of the central nervous system (CNS) to regenerate is a barrier to the effective treatment of neurodegenerative diseases and traumatic injuries. Following CNS injury, astrocytes undergo reactive gliosis and create a glial scar that cordons off the injury that restricts the extension of regenerating axons through the injury site. Reversing this natural biological response to CNS injury is challenging using existing treatment approaches. Therefore, converting glial cells into functional neurons *in situ* provides an efficient way to obtain desirable endogenous neurons from a large cellular pool for "on-site" brain repair.

Here, we examined the ability of adeno-associated viral vector (AAV) encoded with genes relevant for reprogramming astrocytes into functional neurons. Our *in vitro* results demonstrated the successful conversion reactive astrocytes to neurons through direct and indirect reprogramming.