

Temporal Changes of Recovery-related Transcriptome in Stroke

In-Ae Choi^{2*}, Cheol Soon Lee², Nayeon Ko¹, Dong-Hee Choi², Jongmin Lee^{1,2†}

Konkuk University School of Medicine and Konkuk University Medical Center, Department of Rehabilitation Medicine¹, Konkuk University, Center for Neuroscience Research, Institute of Biomedical Science and Technology²

Objective

The endogenous remodeling post stroke, such as angiogenesis, neurogenesis and axonal sprouting is restrictive to induce complete restoration of neurological function. To identify the therapeutic targets to evoke endogenous restorative mechanism for stroke recovery, we investigated temporal alteration of endogenous recovery-related genes at different time points post stroke.

Methods

Photothrombotic stroke was induced in wistar rats. Peri-infarct tissues were collected at different time points (1w, 4w and 8w) and RNA sequencing was conducted. Kyoto Encyclopedia of Genes and Genomes (KEGG) pathway analysis and Gene Ontology (GO) analysis were conducted to analyze the profiles of the genes involved in axonal sprouting, growth factor and angiogenesis categories (FDR<0.05 and |fc|>2)

Results

KEGG pathway analysis revealed that inflammatory response was dominant at 1 week and downregulated afterwards. ECM-receptor interaction was facilitated to support recovery until late stages. GO analysis discovered that endogenous recovery process was active until 4 weeks and the change was not completed even at 8 weeks, *Nefh*, *Syt2* and *Robo3* were downregulated and conversely, *C3*, *Gfap*, *Igf2*, *Aqp1*, *Spp1*, *Anaxa1*, *Cd44*, *Bmp6*, *Dcn*, *Crabp2*, *Dab2*, *Vim*, *Lgals1*, *Lgals3*, *Aprnr*, *Col1a1*, *Col3a1*, *Foxd1*, *Foxc2*, *Efemp1*, *sfrp1*, *Aldh1a2*, *Cdkn1c* were upregulated.

Conclusion

These gene profile alterations may identify new therapeutic target to prompt recovery post stroke at various time points.

Acknowledgment

This research was supported by Basic Science Research Program through the National Research Foundation of Korea (NRF) funded by the Ministry of Science, Information and Communications Technology (ICT) and Future Planning (NRF-2014R1A2A1A11050248 and NRF-2017R1A2B4004837).