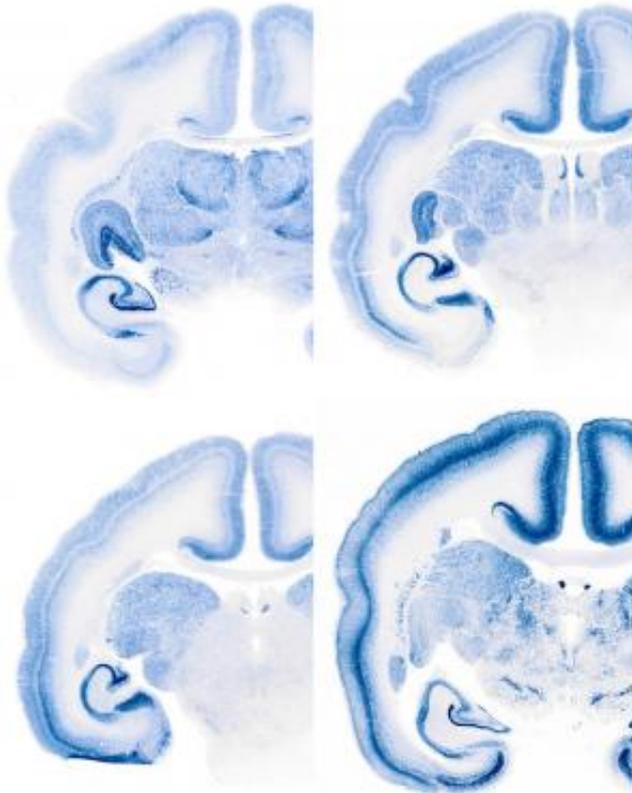


Two heads are better than one: Gene expression reveals molecular mechanisms underlying evolution of cerebral cortex

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Gene expression patterns in the brain of the newborn common marmoset revealed by in situ hybridization. Credit: 2012 Tomomi Shimogori, RIKEN Brain Science Institute

Dramatic expansion of the human cerebral cortex, over the course of evolution, accommodated new areas for specialized cognitive function, including language. Understanding the genetic mechanisms underlying these changes, however, remains a challenge to neuroscientists.

A team of researchers in Japan has now elucidated the mechanisms of cortical evolution. They used molecular techniques to compare the

[gene expression patterns](#) in mouse and monkey brains.

Using the technique called in situ hybridization to visualize the distribution of mRNA transcripts, Okano, Shimogori and their colleagues examined the expression patterns of genes that are known to regulate development of the [mouse brain](#). They compared these patterns to those of the same genes in the brain of the common marmoset. They found that most of the genes had similar expression patterns in mice and marmosets, but that some had strikingly different patterns between the two species. Notably, some areas of the visual and prefrontal [cortices](#) showed expression patterns that were unique to marmosets.

The researchers observed that the *Btbd3* gene, for example, which encodes a transcription factor that regulates the expression of other genes, was expressed throughout the visual cortex of the mouse but restricted to layer 4 of the [primary visual cortex](#), or the V1 area, of the marmoset. Similarly, the gene encoding connective tissue growth factor (CTGF) was expressed throughout the mouse cortex in layer 5, but was restricted to layer 4 of area V1 in the marmoset.

Some of the genes that are expressed widely throughout the mouse prefrontal cortex were likewise restricted to specific layers and sub-regions in the marmoset. Okano, Shimogori and colleagues also noted differences in expression patterns in the subplate region of the developing cortex, which contains the first neurons to receive inputs from the thalamus, a deep [brain structure](#) that relays sensory information to the cortex.

The researchers also found differences in gene expression within regions that connect the prefrontal cortex and hippocampus, a structure that

is critical for learning and memory.

"The distinct gene expression patterns and anatomical differences between marmosets and mice provide enormous insights into the evolution of the brain," says Okano. "We are interested in characterizing the functions of genes that could act as driving forces of brain evolution and have started to investigate several candidate genes. Such approaches will eventually lead to a better understanding of brain function and mental disorders."

More information: Mashiko, H., Yoshida, A.C., Kikuchi, S.S., Niimi, K., Takahashi, E., Aruga, J., Okano, H. & Shimogori, T. Comparative anatomy of marmoset and mouse cortex from genomic expression. *Journal of Neuroscience* 32, 5039–5053 (2012).
www.jneurosci.org/content/32/15/5039.short

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