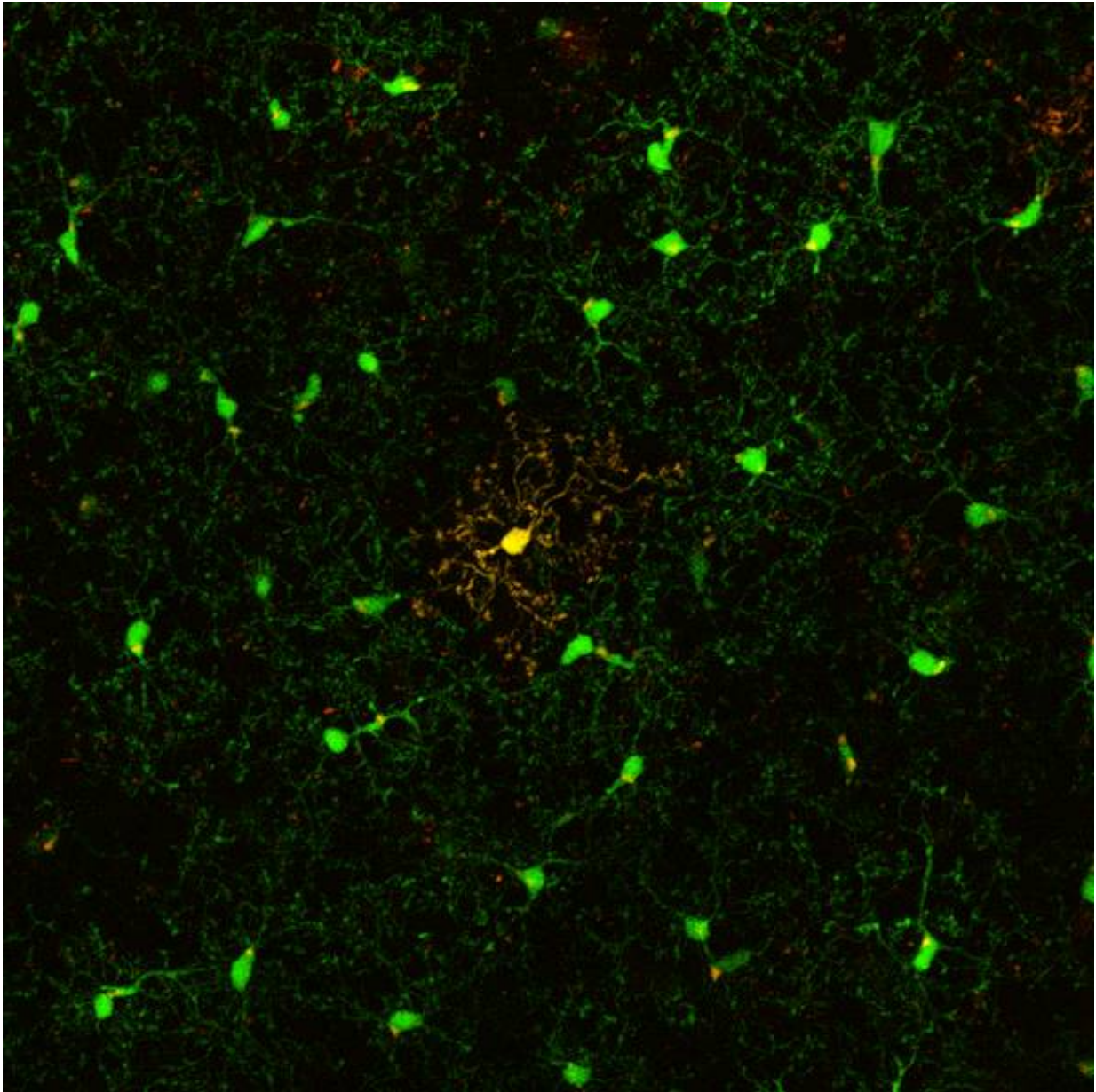


Brain defense cells live longer than expected

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The scientists labelled individual microglial cells (one cell marked in yellow).

This allowed them to differentiate the cells from other microglia (in green) and to follow them over their lifespan in vivo. Credit: University of Tübingen

Eliminating pathogens and cellular waste is an important task of microglia, the immune cells of the brain. They are among the group of non-neural brain cells that support the normal function of nerve cells. A new study now shows that in mice, these scavenger cells may live as long as the rodents themselves. The new finding comes from scientists at the Hertie Institute for Clinical Brain Research, the University of Tübingen, and the German Center for Neurodegenerative Diseases (DZNE).

In an advance online publication of the journal *Nature Neuroscience*, the scientists followed individual microglial [cells](#) under the microscope in vivo. The unexpectedly long lifespan of [microglia](#) indicates further functions these cells may perform: "Their longevity allows them to learn and to age," explains Professor Dr. Mathias Jucker. "This may enable them to form an immunological memory and contribute to the development of [neurodegenerative diseases](#)."

Until now, it has been unclear whether microglial cells can create a memory for pathogens similar to the one developed by [immune cells](#) in the rest of the body. This function ensures a faster and more efficient activation of the defense cells when a second contact is made. "If microglial cells lived only briefly, an immune memory would not make much sense. Now that we know this is not the case, the idea becomes conceivable," says co-corresponding author Dr. Angelos Skodras. In fact, there are first indications that an early stimulation of the [brain's](#) immune system permanently changes the activity of the microglial cells.

Microglia have long been suspected of playing a role in the development of age-related neurological diseases. "An amazing finding in recent years

is that most risk factors for Alzheimer's disease map to changes in genes that are active in [microglial cells](#)," Jucker says. It remains unclear how the cells contribute to the development of the disease. However: "Aging and senescence of microglia may play a role—this requires a long life of the cells."

In the healthy brain, the number of microglia remains more or less constant. However, until now, scientists have wondered whether microglia are short-lived cells that rapidly proliferate, or whether they are long-lived cells that rarely divide. Previous measurements were done only indirectly and led to contradictory results.

In the present study, first author Dr. Petra Fügen genetically labeled individual microglia in mice and directly observed the turnover of these cells using 2-photon imaging over many months in the mouse brain. "As the outcome was completely open, we had a bet running in the department with predictions that ranged from a couple of months to more than a year," Jucker recalls. In the end, half of the cells studied showed a calculated lifespan of up to 28 months, which corresponds to a mouse's lifetime. "In our study, we were finally able to prove the longevity of microglia," the authors conclude.

More information: Petra Fügen et al. Microglia turnover with aging and in an Alzheimer's model via long-term in vivo single-cell imaging, *Nature Neuroscience* (2017). [DOI: 10.1038/nn.4631](https://doi.org/10.1038/nn.4631)

Provided by University of Tübingen

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