

Study finds possibility of new ways to treat, manage epilepsy seizures

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Credit: University of Kentucky

New findings from the University of Kentucky published in the *Journal* of Neuroscience demonstrate that there may be ways to address bloodbrain barrier dysfunction in epilepsy.



Epilepsy is one of the most common neurological disorders and around one-third of <u>epilepsy</u> patients do not respond well to anti-<u>seizure</u> drugs. Until now, it was believed that the cause and effect of epilepsy was merely based on a dysfunction in the brain's neurons. However, recent findings suggest that epilepsy can be caused by many other factors, including a dysfunctional <u>blood-brain barrier</u>. Essentially, seizures erode the lining of capillaries in the brain which plays a role in letting nutrients in and keeping toxins out. This can result in a "leaky" blood-brain barrier, which leads to more seizures, resulting in epilepsy progression.

Björn Bauer's lab at the UK College of Pharmacy collaborated with Sanders-Brown Center on Aging scientists to conduct research focused on this barrier leakage. Bauer and colleagues hypothesized that glutamate, released during seizures, mediates an increase in certain enzymes and activity levels, thereby contributing to barrier leakage.

Through their research, they found that the neurotransmitter glutamate, released during seizures, increased the activity of two types of enzymes, which increased barrier leakage. They also found that blocking the enzyme cPLA2 and genetically deleting cPLA2 may prevent the changes mentioned and the associated leakage. This suggests that cPLA2 is responsible for barrier leakage.

Since 30 percent of people with epilepsy do not respond well to current anti-seizure medications, these findings demonstrate there could be new ways to treat and manage seizures that currently do not respond well to medication.

The data gathered implies that cPLA2 could be a pharmaceutical target to repair and normalize barrier dysfunction and improve the treatment of epilepsy and potentially other neurological disorders that are accompanied by blood-brain barrier leakage These strategies to repair barrier dysfunction could be valuable add-on treatments to existing



pharmacotherapy.

More information: Ralf G. Rempe et al. Matrix Metalloproteinase-Mediated Blood-Brain Barrier Dysfunction in Epilepsy, *The Journal of Neuroscience* (2018). DOI: 10.1523/JNEUROSCI.2751-17.2018

Provided by University of Kentucky

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