

Biologists discover a key regulator in the pacemakers of our brain and heart

April 25 2014



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The study from the University of Waterloo is published in the *Journal of Biological Chemistry* today and is featured as the "Paper of the Week" for its significance.

The researchers discovered T-type channels in the <u>pond snail</u>, Lymnaea stagnalis, can shift from using calcium ions to using <u>sodium ions</u> to generate the electrical signal because of an outer shield of amino acids called a turret situated above the channel's entrance.

Low voltage T-type channels generate tiny pulses of current at regular intervals by selectively passing positively charged cations across the cell's membrane through a gate-like channel. The channels are normally extremely selective, allowing just one sodium ion to pass for every 10,000 <u>calcium ions</u>.

The resulting rhythmic signals produced by this transfer of cations are what support the synchronous contraction of our heart muscles and neuronal firing in parts of the brain, like the thalamus, which helps regulate our sleep-wake cycle, or circadian rhythm.

In addition to their published findings, the researchers also found the shield-like turrets in pond snails restrict access of therapeutic drugs to the channel.

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Over-active T-type channels are linked to epilepsy, cardiac problems, neuropathic pain, as well as the spreading of several kinds of cancer. Drugs that could quench out-of-control T-type channel activity are unable to bind to the channels themselves.

"We wanted to understand the molecular structures of T-type channels," said Spafford. "How they pass ionic currents to generate electrical activity, and to identify drug binding sites, and the drugs which may block these channels to treat neurological disease or heart complications."

The group is currently investigating how dismantling this extracellular turret will improve drug access and binding in T-type channels.

More information: Paper www.jbc.org/content/289/17/11952

Provided by University of Waterloo

Citation: Biologists discover a key regulator in the pacemakers of our brain and heart (2014, April 25) retrieved 27 April 2024 from <u>https://phys.org/news/2014-04-biologists-key-pacemakers-brain-heart.html</u>

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