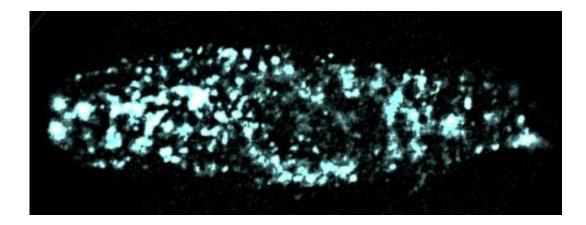


Scientists gain a clear picture of E1 protein

March 4 2014, by Leah Burrows



E1 protein, on the surface of a mammalian cell, illuminated with a teal fluorescent protein and viewed through a laser mounted total internal reflection microscope.

(Medical Xpress)—The heart's cellular blueprint has been the subject of scientific debate for decades, with no component causing more trouble the KCNE1.

This little protein, E1 for short, is one of two subunits in a <u>potassium ion</u> <u>channel</u> that regulates heart contractions. Understanding the construction of this channel is key to understanding life-threatening <u>heart conditions</u>, such as arrhythmias, and developing drugs to threat those conditions.

For years, scientists have debated how many E1 proteins are required to build one of these channels, theorizing anywhere between one and 14. Now, Brandeis University researchers found that these channels are built



with two E1s.

Leigh Plant, assistant research professor of biochemistry, along with postdoctoral fellows Dazhi Xiong, Hui Dai and provost and professor of biochemistry Steve Goldstein, published their findings in the *Proceedings of the National Academy of Sciences* on Monday, March 3.

This report challenges a previous study—the findings of which are currently being used in drug development trials and animal models—that anywhere between one and four E1s are required per channel. Brandeis researchers hope their new findings may help create more effective models to study heart conditions and their treatment.

A single heartbeat is the slow expanding and contracting of the heart muscle. It is controlled, in part, by a series of channels on the surface of heart cells that regulate the movement of different ions into and out of the cells. The potassium ion <u>channel</u> is critical to ending each <u>heart</u> contraction and is made up of the proteins Q1 and E1. Q1s create the pore that the potassium flows through and the E1s control how slowly that pore opens and closes, how many channels are on the cell surface of each cell and how they are regulated by drugs.

Goldstein's team observed E1 in live, mammalian cells at remarkable sensitivity, counting the proteins in individual channels, something that had never been done before in this area of research. Because this mechanism has been so widely debated, Goldstein and his team used three different means to count E1—including tagging them with different fluorescent colors and using a scorpion toxin to bind to Q1. Each time, the team got the same results.

While there is always room for debate in science, Goldstein and his team said they hope these findings will give researchers a quintessential key to unlocking the intricacies of the heartbeat.



More information: Individual IKs channels at the surface of mammalian cells contain two KCNE1 accessory subunits, <u>www.pnas.org/cgi/doi/10.1073/pnas.1323548111</u>

Provided by Brandeis University

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