

Alkylphenol Binding Site Identified in Human HCN1 Channels

In a just published study in the journal Nature, three research teams identify the binding site in human HCN1 (hHCN1) ion channel.

NEW YORK, NY, USA, August 20, 2024 /EINPresswire.com/ -- In a just published study in the journal Nature, research teams at Weill Cornell Medicine, the University of Miami, and Linköping University, Sweden, report having identified the binding site for the alkylphenol general anesthetic 2,6-di-iso-propylphenol (a.k.a., propofol) on the human HCN1 (hHCN1) ion channel.

The study's lead authors are Dr. Crina Nimigean, professor of physiology and biophysics in anesthesiology at Weill



Cornell Medicine, and Dr. Peter Larsson, a professor in the department of biomedical and clinical sciences at Linköping University.

Study co-author Dr. Peter Goldstein and his lab have previously demonstrated 2,6-di-iso-propylphenol, as well as its non-anesthetic congener 2,6-di-tert-buylphenol (2,6-DTBP), effectively relieves neuropathic pain resulting from peripheral nerve injury. 2,6-DTBP is the basis of Akelos's peripherally-restricted non-narcotic antihyperalgesic lead molecule BP4L-18:1:1.

"Using cryoelectron microscopy along with other high-resolution approaches, the study identified the binding site for a clinically important class of molecules that target the human HCN1 ion channel. The findings will guide efforts for novel drug development for poorly treated neurologic disorders in which HCN1 channels play a role. By further defining the underlying mechanism of action of propofol, these results will also support Akelos' efforts to submit an IND application to the FDA for first-in-human trials with AKE-1018 with the aim of treating

neuropathic pain," says Dr. Goldstein, who is scientific co-founder and a scientific advisory board member of Akelos, Inc. He is also an inventor on patents related to the use of 2,6-DTBP and its derivatives to treat neuropathic pain, which have been licensed by Cornell University to Akelos, Inc.

Says Akelos founder Dr. Steven Fox, "BP4L-18:1:1 effectively relieves peripheral neuropathic pain and does so with an excellent safety profile. The study's findings strengthen our ability to introduce it into clinical practice. The authors of the Nature paper are to be congratulated on an outstanding achievement."

Forward-Looking Statement

This press release contains forward-looking statements. These forward-looking statements are based on management's expectations and are subject to certain factors, risks and uncertainties that may cause actual results, outcomes of events, timing and performance to differ materially from those expressed or implied by such statements.

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