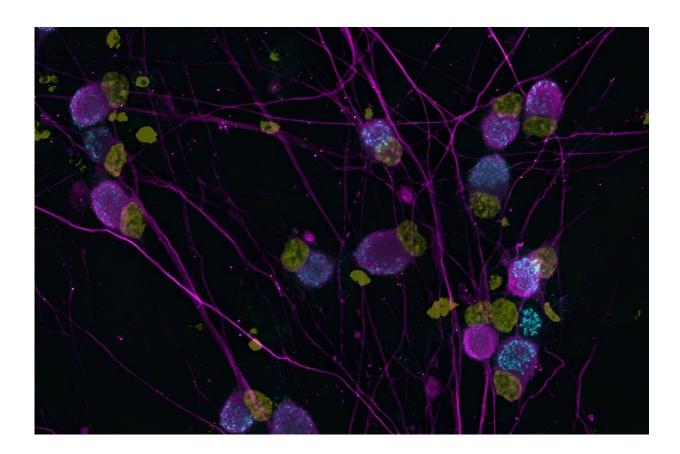


A new ion channel that plays a vital role in touch perception discovered

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Induced human sensory neurons with the ion channel Elkin1 (cyan), nucleus (yellow) and Neurofilament 200 (magenta). Credit: Amy Hulme, University of Wollongong

Every hug, every handshake, every dexterous act engages and requires touch perception. Therefore, it is essential to understand the molecular



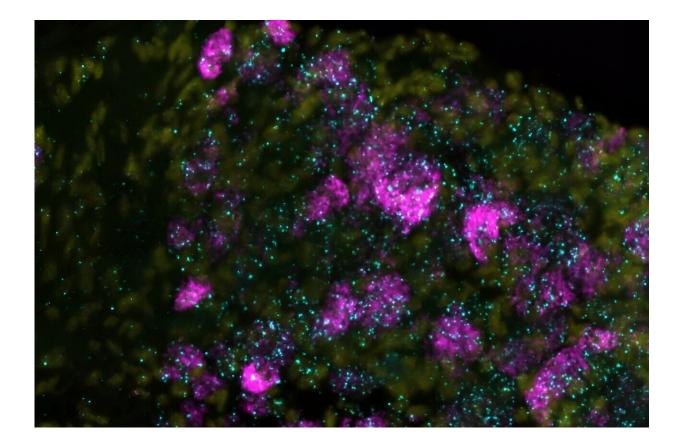
basis of touch. "Until now, we had known that the ion channel—Piezo2—is required for touch perception, but it was clear that this protein alone cannot explain the entirety of touch sensation," says Professor Gary Lewin, head of the Molecular Physiology of Somatic Sensation Lab at the Max Delbrück Center.

For over 20 years Lewin has been studying the molecular basis of the sensation of touch. He and his team have now discovered a new ion channel, named Elkin1, that plays a vital role in touch perception. This is only the second ion channel implicated in the touch perception. It is likely that the protein is directly involved in converting a mechanical stimulus, such as light touch, into an <u>electrical signal</u>.

When Elkin1 is present, the receptors in the skin can transmit the touch signals via <u>nerve fibers</u>, to the central nervous system and brain. The researchers have published their <u>findings</u> in the journal *Science*.

Lewin's team came across Elkin1 a few years ago while investigating a malignant melanoma cell line. The researchers had found that the protein is required for sensing mechanical forces by these highly motile cancer cells. "Now we wanted to determine whether the same protein also plays a role in touch sensation," says Lewin.





Mouse neurons with the new ion channel Elkin1 (cyan), which is responsible for touch sensation, nucleus (yellow) and the already known ion channel Piezo2 (magenta). Credit: Sampurna Chakrabarti, Max Delbrück Center

Lack of Elkin1 reduces touch sensitivity

The researchers bred genetically-modified mice that lacked the Elkin1 gene. They then conducted simple behavioral experiments that involved lightly brushing a cotton swab against the rodents' hind paws.

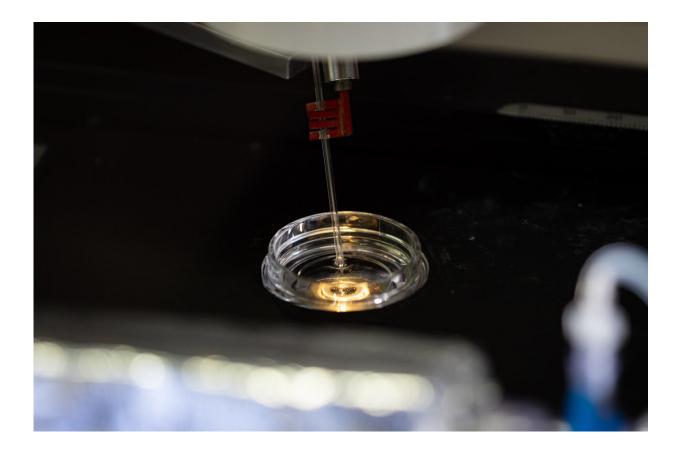
"Usually, normal mice react to the cotton swab 90% of the time," says Lewin. "In contrast, mice lacking Elkin1 only reacted half of the time, indicating touch insensitivity." Importantly, the rodents' reaction to nonmechanical stimuli like temperature was not affected.



At the neuronal level, Dr. Sampurna Chakrabarti, a scientist in Lewin's team, used the patch clamp method to record the electrical activity of sensory neurons in response to poking of the neuronal membrane.

"Around half of the neurons in genetically modified mice lacking Elkin1 didn't respond to mechanical stimuli, and no signal transmission occurred," says Chakrabarti.

Further experiments confirmed that there were no signals relayed from the neuron's receptor ending in the skin, on the first leg of the signals journey from skin to the spinal cord and brain.



Petri dish with sensory neurons that are poked with the glass electrode. Credit: Felix Petermann, MDC





Dr. Sampurna Chakrabati at the microscope in the lab of Professor Gary Lewin from the Max Delbrück Center. Credit: Katharina Bohm, Max Delbrück Center

Furthermore, their Australian collaborators in the lab of Professor Mirella Dottori in the University of Wollongong tested whether Elkin1 in necessary for touch transduction in human sensory neurons grown in a petri dish from stem cells. Their findings also strongly suggest that Elkin1 could play a major role in human touch perception.

The researchers assume that during normal signal transmission, Elkin1 and Piezo2 share roles in touch perception. They have also found evidence that Elkin1 may play a part in the transmission of painful mechanical stimuli.



"If this is confirmed to be the case, we will have not only identified a second ion channel with an indispensable role in normal touch <u>perception</u>, but also a new potential target for treating <u>chronic pain</u>," says Lewin.

More information: Sampurna Chakrabarti et al, Touch sensation requires the mechanically-gated ion channel ELKIN1, *Science* (2024). DOI: 10.1126/science.adl0495. www.science.org/doi/10.1126/science.adl0495

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