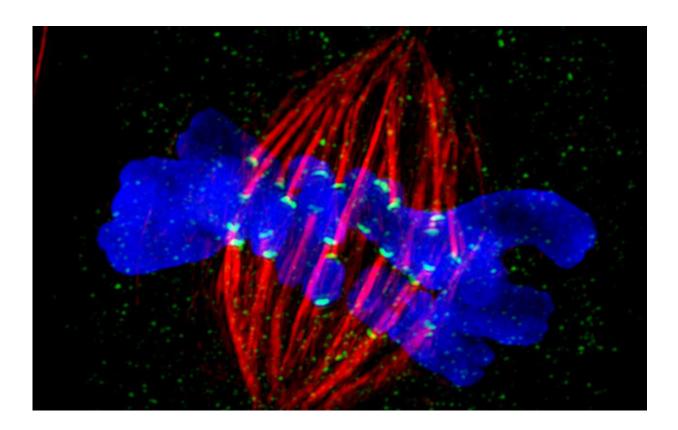


Machinery used in basic cell division does double duty as builder of neurons

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PtK2 cell undergoing mitosis or cell division. Microtubules are shown in red with kinetochores in green and DNA stained blue. Credit: Cell Image Library

Researchers at the San Diego branch of the Ludwig Institute for Cancer Research at University of California San Diego have identified an entirely new mechanism underlying the development and structure of the



nervous system during embryogenesis.

The findings, publishing in the February 28, 2019 issue of *Developmental Cell*, focus on the dynamic coupling of microtubule ends to kinetochores, built on the centromeres of chromosomes, to direct chromosome segregation during cell division. The work was conducted using *Caenorhabditis elegans*, a species of nematode, as the animal model.

During <u>cell division</u> or mitosis, the centromere regions of chromosomes assemble large protein machines called kinetochores to connect chromosomes to microtubules, which the chromosomes then use to separate to opposite ends of the cell. The microtubule-based physical separation of chromosomes ensures that the two new <u>cells</u> born after division inherit a complete genome.

In their new paper, Arshad Desai, Ph.D., professor in the Department of Cellular and Molecular Medicine at UC San Diego School of Medicine, and colleagues report that the evolutionarily ancient kinetochoremicrotubule coupling machine, called the KMN network, plays a critical role in neuronal morphogenesis —the <u>biological process</u> that causes a cell or organism to develop its shape.

"This is an entirely new discovery," said Desai. "We've found molecular similarity between the movement of <u>chromosomes</u> in dividing cells and the formation of neuronal projections, both of which involve microtubule polymers that dynamically grow and shorten."

Desai said the findings may help illuminate the underlying pathology of some neurological conditions. For example, he said, mutations in one of the components of the KMN network have been associated with microcephaly, a condition in which the brain does not develop properly, resulting in a smaller than normal head and numerous health and



cognitive issues.

"Our work suggests a potential explanation for why that happens," said Desai.

In related work, researchers at Harvard Medical School report similar findings, working primarily with a fruit fly model.

More information: *Developmental Cell* (2019). <u>DOI:</u> <u>10.1016/j.devcel.2019.02.002</u>

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