

New method detects telomere length for research into cancer, aging

29 June 2016



Dr. Jerry Shay. Credit: UT Southwestern

UT Southwestern Medical Center cell biologists have identified a new method for determining the length of telomeres, the endcaps of chromosomes, which can influence cancer progression and aging.

In broad terms, telomeres help determine whether a cell reproduces accurately. As the cell divides, these endcaps degrade, causing the cell to age - and it is believed this degradation may lead to some aspects of aging in humans. UT Southwestern scientists hope to use knowledge of telomeres to slow or stop cells from aging, or to potentially help stop the uncontrolled growth of cancer cells that maintain their telomeres.

Finding the best and most sensitive methods for measuring telomeres is an important initial step that could eventually enable scientists to encourage healthy cell growth or, in the case of cancer, limit or halt the cell growth.

"Probes generated using this new approach significantly enhance the sensitivity of telomere length measurements," explained Dr. Jerry Shay, Professor of Cell Biology, who along with his postdoctoral fellow, Dr. Tsung-Po Lai, developed the new method together with Dr. Woodring Wright, Professor of Cell Biology and Internal Medicine. Dr. Shay and Dr. Wright, who hold The Southland Financial Corporation Distinguished Chair in Geriatrics, are members of UT Southwestern's Harold C. Simmons Comprehensive Cancer Center, and together direct a joint laboratory studying telomere biology.

Telomeres are the ends of chromosomes, which package genetic information to pass from parents to children. As the cell divides, these endcaps help keep the genetic information stable and prevent the chromosomes from fusing together. But as cells divide, the telomeres shorten and, eventually, trigger DNA damage that in conjunction with other factors can lead to progression of aging and an increased chance of cancer emerging.

Scientists are interesting in understanding this cellular aging process to create an "immortal cell," which could continue to create new cells without degrading and that could have important implications for diseases related to aging. In reverse, controlling the cellular aging process could accelerate the mortality of cancer cells. If telomeres in cancer cells shorten, this action could prevent the cancer cells from dividing uncontrollably in their early stages of development, slowing or halting cancer progression. In the event that a tumor has already developed, anti-telomerase therapy might be administered to prevent relapse.

The first step is developing the ability to accurately measure telomeres. Radioactive probes often used for measuring lengths are less stable, require long exposure times, and result in hazardous waste



disposal requirements and other potential safety issues. A popular nonradioactive alternative called DIG (digoxigenin) probes requires relatively large amounts of genomic DNA and is less sensitive. The new nonradioactive method developed by the researchers here uses multiple DIG molecules, which increases both the sensitivity and stability of telomere detection.

Provided by UT Southwestern Medical Center APA citation: New method detects telomere length for research into cancer, aging (2016, June 29) retrieved 4 May 2021 from https://medicalxpress.com/news/2016-06-method-telomere-length-cancer-aging.html

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