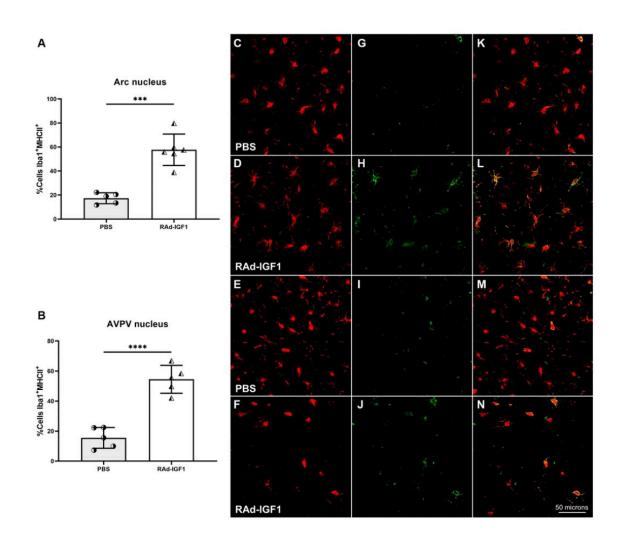


Study proposes IGF1 gene therapy to delay reproductive senescence and potentially optimize women's health and lifespan

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IGF1 gene therapy effect on the activation state of microglia. Credit: 2022



Dolcetti et al.

A new research paper titled "IGF1 gene therapy in middle-aged female rats delays reproductive senescence through its effects on hypothalamic GnRH and kisspeptin neurons" has been published in *Aging*.

The process of aging is the result of progressive loss of homeostasis and functional body impairment, including the central nervous system, where the hypothalamus plays a key role in regulating aging mechanisms. The consequences of aging include a chronic proinflammatory environment in the hypothalamus that leads to decreased secretion of gonadotropin-releasing hormone (GnRH) and impairs kisspeptin neuron functionality.

In this new study, researchers Franco Juan Cruz Dolcetti, Eugenia Falomir-Lockhart, Francisco Acuña, Macarena Lorena Herrera, Sofia Cervellini, Claudio Gustavo Barbeito, Daniela Grassi, Maria-Angeles Arevalo, and María José Bellini from Consejo Nacional de Investigaciones Científicas y Técnicas (UNC-CONICET), Universidad Nacional de La Plata, Autonomous University of Madrid, Instituto Cajal, and Instituto de Salud Carlos III investigated the effect of insulin-like growth factor 1 (IGF1) gene therapy on hypothalamic kisspeptin/GnRH neurons and on microglial cells, that mediate the inflammatory process related with the aging process.

"The aim of the present study is to investigate the effect of IGF1 gene therapy on <u>estrous cycle</u>, kisspeptin and GnRH neurons, and microglial cells in middle-aged female rats," the researchers explain.

The results show that IGF1 rats have higher kisspeptin expression in the anteroventral periventricular (AVPV) nucleus and higher immunoreactivity of GnRH in the arcuate nucleus and median eminence.



In addition, IGF1-treated animals exhibit increased numbers of Iba1+ microglial cells and MHCII+/Iba1+ in the AVPV and arcuate nuclei. In conclusion, IGF1 gene therapy maintains kisspeptin production in the AVPV nucleus, induces GnRH release in the median eminence, and alters the number and reactivity of microglial cells in middle-aged female rats. The researchers suggest that IGF1 gene therapy may have a protective effect against reproductive decline.

"Based on our findings, we propose IGF1 gene therapy to delay reproductive senescence as a potential strategy to optimize lifespan and combat age-related health problems in women," say the researchers.

More information: Franco Juan Cruz Dolcetti et al, IGF1 gene therapy in middle-aged female rats delays reproductive senescence through its effects on hypothalamic GnRH and kisspeptin neurons, *Aging* (2022). DOI: 10.18632/aging.204360

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