

Genetic Quality Control During Meiosis

Infertility and birth defects often arise due to improper genetic quality control during meiosis. Robert Weiss, Biomedical Sciences, and Marcus Smolka, Molecular Biology and Genetics, are resolving how the cellular DNA damage response ensures genetic quality control during meiosis and enables the efficient and accurate production of gametes.

To ensure the production of gametes without genetic defects, the genome of meiocytes is monitored by a set of evolutionarily conserved kinases, known as checkpoint kinases. These kinases sense damage to DNA or problems in chromosome pairing and, upon activation, can block meiotic progression and induce cell death.

How meiotic checkpoint signaling is regulated, however, is not understood, especially in mammals. Little is known about how checkpoint kinases can act as essential regulators of normal meiotic progression as well as effectors of quality control mechanisms that lead to cell death. This knowledge gap poses a major barrier for understanding the determinants of genetic quality control and how misregulation of checkpoint signaling may promote infertility.

The same pathways also mediate fundamental DNA repair and checkpoint functions in mitotic cells, sometimes deregulated in cancers. Researchers are targeting these pathways as an emerging strategy for cancer clinical treatment.

The findings will provide fundamental knowledge about DNA repair and DNA damage signaling mechanisms that have key roles in organismal development, cancer, and reproduction.

NIH Award Number: 1R01HD095296-01



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Funding Received

\$1.2 Million
spanning 5 years

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Eunice Kennedy Shriver National Institute
of Child Health and Human Development

Other Research Sponsored by

National Institutes of Health, National Institute of Child Health and Human
Development





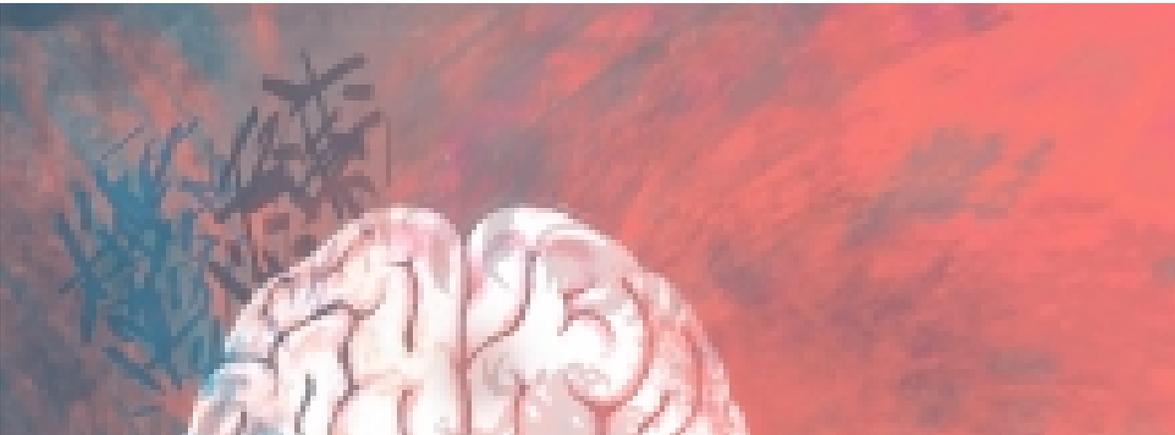
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