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Telomeres





Table of Contents

1 Termination of DNA Replication

² Function of Telomeres

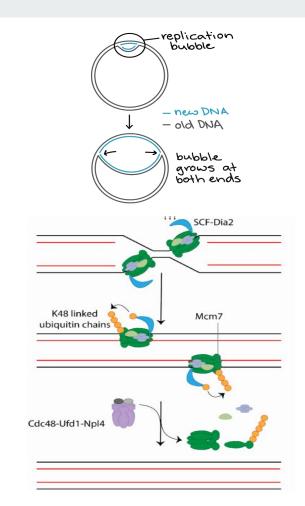
Termination of DNA replication

Prokaryotes

- DNA replication begins at a single origin
- A special DNA sequence slows down the movement of each fork causing them to disassemble

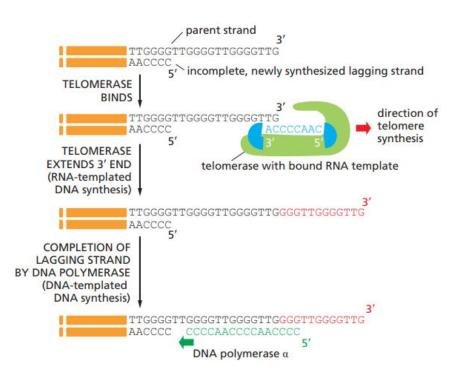
Eukaryotes

- Fork encounters cause termination
- CMG helicase is modified by ubiquitin, causing disassembly



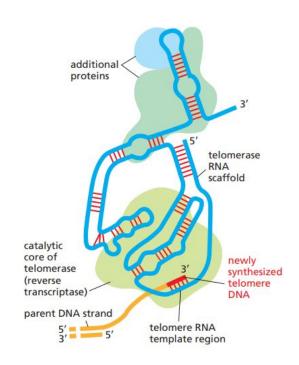
Telomeres and Telomerase

- The final RNA primer at the end of the lagging strand cannot be replaced by DNA polymerase
- Telomeres are special nucleotide sequences at the end of the eukaryotic chromosome
- Telomerase replenishes telomeres after each cell division



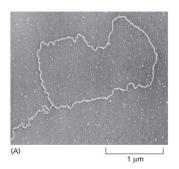
Structure of Telomerase

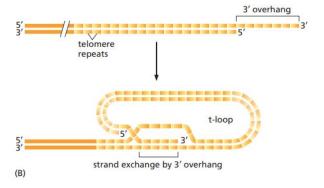
- Telomerase provides an RNA scaffold that provides the template of telomere repeat synthesis
- Contains reverse transcriptase domain
- In some organisms, telomerases have proteins that guide the telomerase to the ends of the chromosome



Packaging of Telomeres

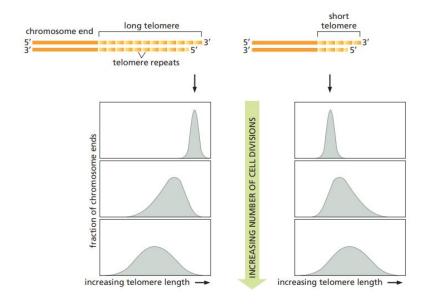
- Telomeres protect chromosome ends preventing them from being mistaken for accidental DNA breaks
- GGGTTA repeats form a complex with caps the DNA
- A nuclease degrades 5' end leaving a 3' single strand overhang
- Forms a t-loop which conceals chromosome ends





Telomere Length is Regulated

- Telomere length shorten in most somatic cell division, acting as a biological clock
- Telomere shortening triggers replicative senescence protecting against cancer
- Most cancer cells reactivate telomerase enabling indefinite growth
- Telomere shortening protects against cancer however can lead to aging-related diseases-dyskeratosis congenital



Thank you

