

## Alternative end-joining of DNA breaks

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#### Citation

Schendel, R. van. (2016, December 15). *Alternative end-joining of DNA breaks*. Retrieved from https://hdl.handle.net/1887/45030

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Author: Schendel, Robin van Title: Alternative end-joining of DNA breaks Issue Date: 2016-12-15

### Stellingen behorende bij het proefschrift:

### Alternative end-joining of DNA breaks

- 1. DNA double-strand breaks in *C. elegans* germ cells are not repaired by NHEJ, but by POLQ-mediated end joining. (this thesis)
- 2. The A-family polymerase POLQ has the unique ability to join two break-ends by extending from single nucleotide homology, minimizing the loss of genetic information. (this thesis)
- 3. Overcoming replication blocking lesions by translesion synthesis prevents large genomic rearrangements. (this thesis)
- 4. POLQ-mediated repair allows for genome diversification. (this thesis)
- POLQ-mediated alt-EJ is an important pathway to repair DNA damage generated during the rapid cleavage divisions of early embryogenesis. (Thyme et al., Cell reports, 2016)
- POLQ is upregulated in HR-deficient ovarian and breast cancers, suggesting that alt-EJ can serve as a backup pathway for the repair of DSBs when HR is defective. (Ceccaldi et al., Trends in Cell Biology, 2016)
- 7. The location of a DSB within the cellular genome and the chromatin context are suspected to influence the facility with which the cell will process a given break and the repair pathway it will choose for this purpose. (Mladenov et al., Seminars in Cancer Biology, 2016)
- 8. Mutational signatures can be used as a physiological readout of the biological history of a cancer and also have potential use for discerning ongoing mutational processes from historical ones, thus possibly revealing new targets for anticancer therapies. (Helleday et al., Nature Review Genetics, 2014)
- 9. Three things cannot be long hidden: the sun, the moon, and the truth. (Buddha, 450 BC)
- 10. We have evolved from worm to man, but much within us is still worm. (Friedrich Nietzsche, Thus Spoke Zarathustra, 1885)
- 11. You fall down. You get up. Life goes on.
- 12. Understanding computers is easy: if you rule out 1, it has to be 0, understanding biology is hard: if you rule out 1, it can be any of the other 1000 ways you did not think of.