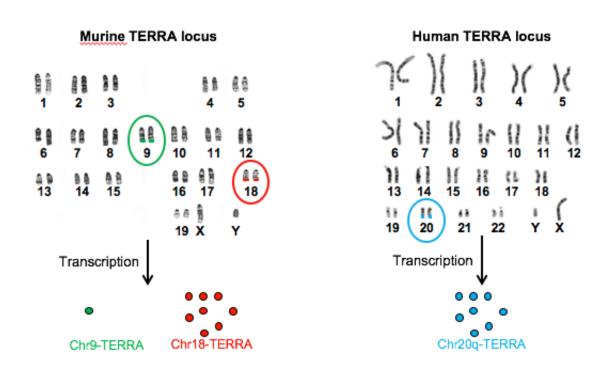


TERRA, the RNAs that protect telomeres

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In murine cells TERRA is transcribed mainly from chromosome 18 and in a less way from chromosome 9, in a similar manner we found that human subtelomere 20q is a TERRA mayor locus. Credit: CNIO

Despite their especially compact and inaccessible structure, telomeres transcribe information like the rest of the DNA. The RNAs resulting from this process are called TERRA, and their function is essential in preserving these protective structures. This is the conclusion of a new study by the Telomere and Telomerase Group at the Spanish National Cancer Research Centre (CNIO), which has also located the part of the



human genome where these molecules are manufactured.

This finding is consistent with the observations made two years ago by the same group, led by Maria A. Blasco. On that occasion, they were working with mouse cells and they observed that the TERRA that protect the 20 chromosomes originated exclusively in pair 18 and, to a lesser extent, in number nine.

The results now published in *Nature Communications* indicate that in humans, these RNAs are transcribed exclusively at one point. The researchers analysed 18 RNAs previously proposed as possible TERRA but only those arising from the long arm of chromosome 20 (20q) and at the short arm of chromosome X (Xp) showed TERRA features.

To demonstrate that the RNAs transcribed at these two points were, indeed, TERRA, the authors genetically removed both loci using the CRISPR-CAS9 technology. They then noted that while the suppression of Xp had no significant consequences, the suppression of 20q had extremely negative effects on <u>telomeres</u>.

This is the first time that the crucial role of TERRA in the preservation of telomeres has been demonstrated in any organism. "These results are striking because they clearly demonstrate that TERRA play an essential role in cell viability as well as in preserving telomeres; they are just as important to the functioning of telomeres as telomerase or shelterins, the proteins that protect the telomeres," explains Blasco.



The discovery of the 20q region as the source of TERRA and the confirmation of the key role played by these RNAs in preserving telomeres opens the door to the study of certain syndromes in which alterations in this chromosomal region have been detected. It also offers a new path for the study of patients with telomeric disorders in which no alterations in telomerase or shelterins have been found.

More information: *Nature Communications*, <u>DOI:</u> 10.1038/NCOMMS12534

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