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In vertebrates, tandem arrays of TTAGGG hexamers are present both at telomeres and at intrachromosomal sites (Interstitial Telomeric Sequence, ITS). An extensive comparative analysis of two primate (human and chimpanzee) and two rodent (mouse and rat) genomes allowed us to describe organization and insertion mechanisms of all the informative ITSs present in the four species. Our results strongly suggest that telomerase was utilized, in some instances, for the repair of DNA double-strand breaks occurring in the genomes of rodents and primates during evolution^{1,2,3}.

Results obtained in collaboration with the ISREC (Switzerland) have demonstrated that mammalian telomeres are actively transcribed into telomeric repeat containing RNA (TERRA)⁴. Using a bio-informatic approach we have identified a putative TERRA promoter; we are now constructing an experimental system to test whether the sequence identified can act as promoter for a reporter gene.

Several ITSs were identified from the now available draft of the horse genome and used for the comparative analysis of the sequence organization of loci orthologous to horse ITSs in different horse breeds and from other equidae species. 9 ITS loci showed either VNTR or presence/absence polymorphism. These loci represent excellent molecular markers for genetic analysis, paternity testing and population genetics^{5,6}. The observation of "empty" ITS loci in the horse population is consistent with the hypothesis that equidae genomes are evolving at a relatively fast rate.

References

- 1. **Nergadze SG**, Santagostino M, Salzano A, Mondello C, Giulotto E. (2007). Contribution of telomerase RNA retrotranscription to DNA double-strand break repair during mammalian genome evolution. Genome Biology. vol. 8, pp. R260.
- 2. Santagostino M., Nergadze SG, Mininni N., Garagna S., Chevret P., Britton-Davidian J., Mondello C., Giulotto E. (2007) Involvement of telomerase in DNA double-strand break repair: an evolutionary approach. FISV 9th Annual Congress. Riva del Garda, 26-29 September. (pp. D07.3).
- 3. **Nergadze SG**, Khoriauli L, Rocchi M, Semino O, Mondello C, Giulotto E. (2007) Evolution and instability of a human interstitial telomeric repeat organized in a head-to-head fashion. FISV 9th Annual Congress. Riva del Garda, Italy, 26-29 September, (pp D07.3).
- 4. Azzalin CM, Reichenbach P, Khoriauli L, Giulotto E, Lingner J. Telomeric repeat containing RNA and RNA surveillance factors at mammalian chromosome ends. Science. 318 (2007) 798.
- 5. **Nergadze SG**, Mondello C., Giulotto E. (2007) Interstitial Telomeric Repeats: A New Class Of Polymorphic Markers In The Horse Genome. Plant & Animal Genomes XV Conference, San Diego, California, January 13-17, (pp. P588).

- 6. **Nergadze SG,** Santagostino M, Mondello C, Giulotto E. (2007) Interstitial Telomeric Repeats in the Horse Genome: Molecular Markers for Population Genetics, Paternity Testing and Phylogenetic Studies. 7th Havemeyer Horse Genome Workshop. Lake Tahoe, CA, August 16-19.
- 7. Alaoui N, Jordana J, Magnani E, **Nergadze SG**, Giulotto E, Ponsa M. New polymorphism in heterochromatin regions of Equus asinus chromosomes detected by horse DNA satellite probes. 6th European Cytogenetic Conference, Istanbul, Turkey, 7-10 July; CHROMOSOME RESEARCH 15 (2007) 130.
- 8. **Nergadze SG**, Santagostino M, Mondello C, Giulotto E. (2007) The Role of Telomeres and Telomerase in Cancer Research. Hyatt Regency San Francisco, San Francisco, California. December 6 9 (pp B31)
- 9. Magnani E., Vidale P., Nergadze SG, Mondello C, Giulotto E. (2008) Immortalization Of Primary Fibroblasts From Equus Species By Ectopic Expression Of Human Telomerase. Plant & Animal Genomes XVI Conference, San Diego, California, January 12-16, (pp. 588).