

The open chromatin profile changes at genome compartments and at tDNA loci in *Trypanosoma cruzi* life forms

Alex Ranieri Jerônimo Lima^{1,2}, Saloe Bispo Poubel^{1,2}, Juliana Nunes Rosón^{1,2,3}, Loyze Paola Oliveira de Lima^{1,2}, Héllida Marina Costa-Silva^{1,2}, Herbert Guimarães de Sousa Silva^{1,2,3}, Pedro Leonardo Lima, Camila Silva Gonçalves^{4,5}, Pedro A. F. Galante⁶, Fabiola Holetz⁷, Maria Cristina Machado M. Motta^{4,5}, Ariel M. Silber⁸, M. Carolina Elias^{1,2}, Julia Pinheiro Chagas da Cunha^{1,2#}

Abstract

Genomic organization and gene expression regulation in trypanosomes are remarkable because protein-coding genes are organized into codirectional gene clusters with unrelated functions. Moreover, there is no dedicated promoter for each gene, resulting in polycistronic gene transcription, with posttranscriptional control playing a major role. Nonetheless, these parasites harbor epigenetic modifications at critical regulatory genome features that dynamically change among parasite stages, which are not fully understood. Here, we investigated the impact of chromatin changes in a scenario commanded by posttranscriptional control exploring the parasite *Trypanosoma cruzi* and its differentiation program using genome-wide approaches supported by transmission electron microscopy. The integration of FAIRE and MNase-seq data, two complementary epigenomic approaches, enabled us to identify differences in *T. cruzi* genome compartments, putative transcriptional start regions, and virulence factors. In addition, we also detected developmental chromatin regulation at tRNA loci (tDNA), which seems to be linked to the translation regulatory mechanism required for parasite differentiation. Strikingly, a positive correlation was observed between active chromatin and steady-state and nascent transcription levels. Taken together, our results indicate that chromatin changes reflect the unusual gene expression regulation of trypanosomes and the differences among parasite developmental stages, even in the context of a lack of canonical transcriptional control of protein-coding genes. Supported by FAPESP, Serrapilheira and CAPES.