

INHIBITION OF PROLINE RACEMASE AFFECTS VERO CELL INVASION BY
Trypanosoma cruzi

Coutinho, L.L.^{1,2}; Alves-Ferreira, M.¹; Berneman, A.³; Cosson, A.³, Chamond, N.³;
Minoprio, P.³; Soeiro M.N.C.² & Degraeve, W.¹

¹ Laboratório de Genômica Funcional e Bioinformática, DBBM, ² DUBC, IOC/FIOCRUZ, Rio de Janeiro, Brazil. ³ Département d'Immunologie, Institut Pasteur, Paris, France.

The kinetoplastid protozoan parasite *Trypanosoma cruzi* is the causal agent of Chagas' disease, a serious public health problem in Latin America. An isoform of the parasite proline racemase enzyme (*TcPRAC*) was described as an effective host B-cell mitogen, contributing to parasite evasion from specific immune responses. *TcPRAC* is encoded by at least two paralogous genes per parasite haploid genome, *TcPRACA* and *TcPRACB*, giving rise to secreted and intracellular protein isoforms. We evaluated parasite infectivity following *in vitro* treatment of cell cultures treated with specific anti-*TcPRAC* polyclonal antibodies or with pyrrole-2-carboxylic acid (PCA), a specific inhibitor of PRAC. Blood trypomastigotes were used to infect Vero cells at 10:1 cell ratio and infectivity was evaluated after 24h. Significant dose dependent decreases in intracellular parasite numbers and in percentage of infected host cells per culture were observed using either anti-*TcPRAC* or PCA. These results substantiate previous observations and put forward a *TcPRAC* role in parasite invasion. Furthermore, our data strengthen *TcPRAC* as a potential target for drug design and immune-modulation.

Supported by: PAPES III and PDTIS – FIOCRUZ; CNPq, FAPERJ, Fonds dédiés Sanofi/Aventis/Ministère de la Recherche/Institut Pasteur.