



Genome Sequence of Trypanosoma cruzi Strain Bug2148

Francisco Callejas-Hernández,^a DNúria Gironès,^{a,b} Manuel Fresno^{a,b}

^aCentro de Biología Molecular Severo Ochoa, Consejo Superior de Investigaciones Científicas, Universidad Autónoma de Madrid, Cantoblanco, Madrid, Spain ^bInstituto Sanitario de Investigación Princesa, Madrid, Spain

ABSTRACT *Trypanosoma cruzi* belongs to the group of mitochondrion-containing eukaryotes and has a highly plastic genome, unusual gene organization, and complex mechanisms for gene expression (polycistronic transcription). We report here the genome sequence of strain Bug2148, the first genomic sequence belonging to cluster TcV, which has been related to vertical transmission.

Typanosoma cruzi is a highly polymorphic parasite that belongs to the *Kinetoplastidae* order and is the causative agent of Chagas disease, also known as American trypanosomiasis, a chronic illness and one of the most neglected tropical diseases (1). Chagas disease is endemic in Latin America, but due to the migration of infected people, this disease has been extended to countries that are nonendemic for the disease, such as those in the European Union, making Chagas disease a serious public health problem (2, 3). Seven to 10 million people are chronically infected with this disease, and 10,000 to 14,000 deaths per year are caused by it (4). There are thousands of different strains of this parasite, but in 2009, a classification based on the genetic structure was proposed, establishing the existence of six separate clusters or discrete typing units (DTUs), named Tcl to TcVI (5).

The complete genome of *Trypanosoma cruzi*, predominantly described as diploid, has been predicted to be around 105 Mb in length, distributed across 20 to 46 chromosomes; however, the total genome size can vary extensively among strains even of the same DTU, mainly due to aneuploidies and variations in gene copy number (6–8). This complex genetic content has been related to evolution, genetic conservation, and variability processes, but its marked differential behavior in *in vitro* and *in vivo* models proposes also its relationship with infectivity and disease development (9, 10).

To date, there are available public genomes of some strains belonging to DTUs I, II, and VI. We have sequenced genomic DNA from metacyclic trypomastigotes cultured in Vero cells and RPMI medium supplemented with 5% fetal bovine serum (FBS) at 37°C; strain Bug2148, belonging to DTU TcV, was sequenced by Pacific Biosciences technology (8-kb to 15-kb read length) and assembled with HGAP version 3 (11), obtaining 55.22 Mb distributed in 934 contigs, with 68× coverage, corresponding to 100% of its haploid estimated genome. Contigs with coverage lower than 15× and without any predicted gene were filtered from the assemblies, and as was expected for this kinetoplastid, the G+C content was around 50% (51.63%). About 91% of its complete predicted genes showed BLASTN similarities to available *Trypanosoma cruzi* predicted genes (including hypothetical genes and pseudogenes [http://tritrypdb.org/tritrypdb/]), in agreement with previous results (12).

Accession number(s). The complete genome sequence of Bug2148 has been deposited in GenBank under accession number NMZN00000000.

ACKNOWLEDGMENTS

This work was supported by the Consejo Nacional de Ciencia y Tecnología (CONACYT, Mexico) through Ph.D. studentship number 411595 to F.C.-H. and the

Received 4 December 2017 Accepted 6

December 2017 **Published** 18 January 2018 **Citation** Callejas-Hernández F, Gironès N, Fresno M. 2018. Genome sequence of *Trypanosoma cruzi* strain Bug2148. Genome Announc 6:e01497-17. https://doi.org/10.1128/ genomeA.01497-17.

Copyright © 2018 Callejas-Hernández et al. This is an open-access article distributed under the terms of the Creative Commons Attribution 4.0 International license.

Address correspondence to Núria Gironès, ngirones@cbm.csic.es. N.G. and M.F. contributed equally to this work. Consejo de Ciência, Tecnología e Innovación de Hidalgo (CITNOVA, Mexico); by Ministerio de Economía y Competitividad grant SAF2015-63868-R (MINECO/FEDER) to N.G. and SAF2016-75988-R (MINECO/FEDER) to M.F.; by Red de Investigación Colaborativa en Enfermedades Tropicales (RICET grant RD12/0018/0004) to M.F.; by the European Union (grant HEALTH-FE-2008-22303, ChagasEpiNet) to M.F.; and by the Comunidad de Madrid (grant S-2010/BMD-2332) and institutional grants from Fundación Ramón Areces.

The technical and scientific assistance provided by Maria Chorro de Villa-Ceballos, Maria C. Maza Moreno, and Alberto Rastrojo Lastras is greatly appreciated.

REFERENCES

- 1. World Health Organization. 2017. Neglected tropical diseases 2017. World Health Organization, Geneva, Switzerland. www.who.int/neglected _______diseases/en/.
- 2. Schmunis GA, Yadon ZE. 2010. Chagas disease: a Latin American health problem becoming a world health problem. Acta Trop 115:14–21. https://doi.org/10.1016/j.actatropica.2009.11.003.
- Bern C, Montgomery SP, Herwaldt BL, Rassi A, Marin-Neto JA, Dantas RO, Maguire JH, Acquatella H, Morillo C, Kirchhoff LV, Gilman RH, Reyes PA, Salvatella R, Moore AC. 2007. Evaluation and treatment of Chagas disease in the United States: a systematic review. JAMA 298:2171–2181. https://doi.org/10.1001/jama.298.18.2171.
- Rassi A, Rassi A, Marcondes de Rezende J. 2012. American trypanosomiasis (Chagas disease). Infect Dis Clin North Am 26:275–291. https://doi .org/10.1016/j.idc.2012.03.002.
- Zingales B, Andrade S, Briones M, Campbell D, Chiari E, Fernandes O, Guhl F, Lages-Silva E, Macedo A, Machado C, Miles M, Romanha A, Sturm N, Tibayrenc M, Schijman A. 2009. A new consensus for *Trypanosoma cruzi* intraspecific nomenclature: second revision meeting recommends Tcl to TcVI. Mem Inst Oswaldo Cruz 104:1051–1054. https://doi.org/10 .1590/S0074-02762009000700021.
- Souza RT, Lima FM, Barros RM, Cortez DR, Santos MF, Cordero EM, Ruiz JC, Goldenberg S, Teixeira MMG, da Silveira JF. 2011. Genome size, karyotype polymorphism and chromosomal evolution in *Trypanosoma cruzi*. PLoS One 6:e23042. https://doi.org/10.1371/journal.pone.0023042.
- 7. Henriksson J, Åslund L, Pettersson U. 1996. Karyotype variability in Trypano-

soma cruzi. Parasitol Today 12:108-114. https://doi.org/10.1016/0169 -4758(96)80670-3.

- Lewis MD, Llewellyn MS, Gaunt MW, Yeo M, Carrasco HJ, Miles MA. 2009. Flow cytometric analysis and microsatellite genotyping reveal extensive DNA content variation in *Trypanosoma cruzi* populations and expose contrasts between natural and experimental hybrids. Int J Parasitol 39:1305–1317. https://doi.org/10.1016/j.ijpara.2009.04.001.
- Rodriguez HO, Guerrero NA, Fortes A, Santi-Rocca J, Gironès N, Fresno M. 2014. *Trypanosoma cruzi* strains cause different myocarditis patterns in infected mice. Acta Trop 139:57–66. https://doi.org/10.1016/j.actatropica .2014.07.005.
- Sanoja C, Carbajosa S, Fresno M, Ria Gironè SN. 2013. Analysis of the dynamics of infiltrating CD4⁺ T cell subsets in the heart during experimental *Trypanosoma cruzi* infection. PLoS One 8:e65820. https://doi.org/ 10.1371/journal.pone.0065820.
- Chin CS, Alexander DH, Marks P, Klammer AA, Drake J, Heiner C, Clum A, Copeland A, Huddleston J, Eichler EE, Turner SW, Korlach J. 2013. Nonhybrid, finished microbial genome assemblies from long-read SMRT sequencing data. Nat Methods 10:563–569. https://doi.org/10.1038/nmeth .2474.
- Franzén O, Ochaya S, Sherwood E, Lewis MD, Llewellyn MS, Miles MA, Andersson B. 2011. Shotgun sequencing analysis of *Trypanosoma cruzi* I Sylvio X10/1 and comparison with *T. cruzi* VI CL Brener. PLoS Negl Trop Dis 5:1–9. https://doi.org/10.1371/journal.pntd.0000984.