

pandemics of the 21st century, including severe acute respiratory syndrome, ebola, avian and swine flu, dengue, chikungunya, zika, monkeypox, and above all, COVID-19, that have raised unprecedented awareness on global health. They are all zoonoses or use an animal vector.

In 2018, four international organizations (WHO, Food and Agriculture Organization, UN, and World Organization for Animal Health [WOAH]) signed an agreement for collaboration and joint promotion of “*One health*”, highlighting the close relationship and interdependence between human, animal, and environmental health (Fig. 1). It is a comprehensive and unifying approach to balancing and optimizing the health of people, animals, and ecosystems. It addresses the full spectrum of disease control, from prevention to diagnosis, management, and treatment, contributing to global health security.

According to the WHO, the “*One health*” concept refers to the global goal of increasing interdisciplinary collaboration (public health, medicine, sanitary, veterinary, research, environmental sciences, etc.) in the care of health of people, animals, and the environment, to develop and implement programs, policies, and laws in favor of improving global health.

The relationship between humans, other living beings, and ecosystems is dynamic. With a holistic vision, “*One health*” seeks to adapt to changing relationships that, in recent decades, are especially significant due to phenomena such as globalization, human travel and migration, changes in the geographical distribution of different animal species, climate change, deforestation, intensive livestock farming, new animal migratory routes, environmental pollution, etc. All this has favored the transmission of diseases, with jumps of microbes from animals to people (zoonoses), as new opportunities for contact between humans and animals have arisen, in altered ecosystems.

According to the WOAHA, 60% of known human infectious diseases and more than 75% of emerging infectious diseases are of animal origin, either domestic or wild animals. Therefore, animal health is essential for the maintenance of public health. This integration of the health of humans, animals, and ecosystems must be reflected in the new policies of all countries. Areas where “*One health's*” focus is particularly needed are food safety, zoonosis control, and the fight against antimicrobial resistance.

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HTLV-1 in Brazil: epidemiological scenario in the highest endemic country in the world

HTLV-1 infection is widely disseminated, mainly in developing countries, such as South America, and the Caribbean basin. Japan is also considered an important place for endemicity¹. Since HTLV-1 is responsible for developing inflammatory diseases in almost 30% of infected people² and adult T-cell leukemia/lymphoma in 5-10% of carriers³, it is important to implement measures to block virus transmission in a long fashion.

Brazil implemented a national HTLV screening in blood banks in 1993⁴. In 2023, discussions on universal maternal screening nationwide are taking place, after the publication of a cost-effectiveness study to implement this procedure in the Brazilian population⁵. Japan implemented national screening for HTLV-1 to prevent mother-to-child transmission (MTCT) in 2011, due to similar high prevalence of infection and vertical transmission⁶.

A recent Japanese study evaluated a prefecture-wide antenatal adult T-cell leukemia prevention program in Nagasaki, where 57,323 pregnant women were screened for anti-HTLV-1 antibodies during 2011-2018. The results showed that 133 (0.79%) pregnant women were HTLV-1-positive during their first pregnancy and 9 (0.05%) seroconverted before or during subsequent pregnancies⁷. These data reinforce the importance of the implementation of programs to prevent MTCT of HTLV-1 in endemic countries, such as Brazil.

Since both strategies, inhibition of breast-feeding from HTLV-infected women and blood bank screening for HTLV antibodies showed important benefits for controlling and reduce new HTLV-1 infections, in Japan⁸, similar initiatives in other endemic countries would perform equally well. These strategies should also be considered by health authorities in countries that receive migrants from areas known to have high HTLV-1 endemicity.

In Brazil, as the country with the largest number of HTLV-1 carriers in Latin America, the estimated number of infected persons approaches one million⁹. Paradoxically, HTLV screening is only recommended in blood banks since 1993⁴, and public health policies miss other key populations, such as pregnant women, organ transplant donors, and clinics for sexually transmitted infections. Furthermore, there is a lack of infrastructure for caring with good clinical support to people already diagnosed with HTLV-1 infection.



Figure 1. Estimated number of HTLV-1 infected people in Brazil (215 millions in year 2023).

The most recent survey of HTLV prevalence in Brazil showed 10-year results (2007-2016) in first-time blood donors. HTLV-1/2 infection predominated in women, donors over 50 years of age, black skin color, and low education level. The rate of HTLV-1/2 infection remained stable during the study period, but a trend was noticed for an increased rate among the youngest ($p < 0.001$), males ($p = 0.049$), whites ($p < 0.001$), and higher education profile ($p = 0.014$)¹⁰.

Considering the prevalence of HTLV in blood banks for the entire country, and despite the study being published in the mid 90's, estimates around 0.5% for the general population can be made¹¹. Using prediction models such as those reported by Catalan-Soares¹¹ and the Instituto Brasileiro de Geografia e Estatística (IBGE)¹², nationwide estimates can be inferred (Fig. 1).

Approximately 1,150,000 individuals with HTLV-1 infection are currently living in Brazil, with the highest prevalence being noticed in the North and Northeast states. Accordingly, a recent study conducted in Belém city, Pará state (North Brazil), identified 2% of HTLV-1 infection in the general population, which could account for around 15,000 cases of myelopathy associated with HTLV-1 (HAM) in the region¹³.

Based on this scenario, it would be naïve to plan or claim for HTLV elimination programs in the absence of effective measures to unveil people unaware of the infection and stop HTLV transmission. At this time, HTLV-1 transmission routes are clear¹⁴⁻¹⁸, the risk of

clinical development is well established^{2,3,19-26} and death rates well figured out²⁷.

Given the lack of specific treatment and/or vaccine, any control of the HTLV-1 pandemic worldwide should follow the Japanese strategy, which relies in universal blood donor screening and avoidance of breastfeeding in HTLV-1+ pregnant women.

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References

- Gessain A, Cassar O. Epidemiological aspects and world distribution of HTLV-1 infection. *Front Microbiol.* 2012;3:388.
- Haziot ME, Gascon MR, Assone T, Fonseca LA, Luiz OD, Smid J, et al. Detection of clinical and neurological signs in apparently asymptomatic HTLV-1 infected carriers: association with high proviral load. *PLoS Negl Trop. Dis.* 2019;13:e0006967.
- Rosadas C, Puccioni-Sohler M, Oliveira A, Casseb J, Sousa M, Taylor G. Adult T-cell leukaemia/lymphoma in Brazil: a rare disease or rarely diagnosed? *Br J Haematol.* 2020;188:e46-e49.
- Ministério da Saúde, PORTARIA No 1.376. Available from: <https://Chrome-extension://efaidnbmnndpcajpcglclefindmkaj>; https://redsang.ial.sp.gov.br/site/docs_leis/ps/ps29.pdf [Last accessed on 1993 Nov 19].
- Rosadas C, Senna K, Da Costa M, Assone T, Casseb J, Nukui Y. Economic analysis of antenatal screening for HTLV Type 1 in Brazil: an open access cost-utility model. *Lancet Glob Health.* 2023;11:e781-90.
- Yonemoto N, Suzuki S, Sekizawa A, Hoshi S, Sagara Y, Itabashi K. Implementation of nationwide screening of pregnant women for HTLV-1 infection in Japan: Analysis of a repeated cross-sectional study. *BMC Public Health.* 2020;20:1150.
- Komatsu N, Iwanaga M, Hasegawa Y, Miura S, Fuchi N, Moriuchi H, et al. Frequency of HTLV-1 seroconversion between pregnancies in Nagasaki, Japan, 2011-2018. *Front Microbiol.* 2022;13:1036955.
- Satake M, Sagara Y, Hamaguchi I. Lower prevalence of anti-HTLV-1 as expected by previous models among first-time blood donors in Japan. *J Med Virol.* 2023;95:e28606.
- Rosadas C, Miranda AE, Gonçalves DU, Caterino-de-Araujo A, Assone T, Ishak R. Coordenação-Geral de Vigilância das Infecções Sexualmente Transmissíveis (CGIST/DCCI/SVS). Prevalência da infecção por HTLV-1/2 no Brasil-Boletim Epidemiológico. SVS/MS. 51 2020.
- Miranda C, Utsch-Gonçalves D, Piassi F, Loureiro P, Gomes I, Ribeiro MA, et al. Prevalence and risk factors for human T-Cell lymphotropic virus (HTLV) in blood donors in Brazil-A 10-year study (2007-2016). *Front Med.* 2022;9:844265.
- Catalan-Soares B, Carneiro-Proietti AB, Proietti F. Interdisciplinary HTLV Research Group. Heterogeneous geographic distribution of human T-cell lymphotropic viruses I and II (HTLV-I/II): serological screening prevalence rates in blood donors from large urban areas in Brazil. *Cad Saude Publica.* 2005;21:926-31.
- IBGE. Portal do IBGE. Available from: <https://www.ibge.gov.br> [Last accessed 2023 Nov 28].
- Silva I, Pinheiro B, Nobre A, Coelho JL, Pereira CC, Ferreira LS, et al. Moderate endemicity of the human T-lymphotropic virus infection in the metropolitan region of Belém, Pará, Brazil. *Rev Bras Epidemiol Braz J Epidemiol.* 2018;21:e180018.
- Hino S. Establishment of the milk-borne transmission as a key factor for the peculiar endemicity of HTLV-1: the ATL prevention program Nagasaki. *Proc Jpn Acad Ser B Phys Biol Sci.* 2011;87:152-66.

15. Paiva A, Assone T, Haziot ME, Smid J, Fonseca LA, Luiz OD, et al. Risk factors associated with HTLV-1 vertical transmission in Brazil: longer breastfeeding, higher maternal proviral load and previous HTLV-1-infected offspring. *Sci Rep*. 2018;8:7742.
16. Prates G, Paiva A, Haziot ME, Fonseca LA, Smid J, Marcusso RM, et al. Could cesarean delivery (C-section) help prevent mother-to-child transmission of HTLV-1? *J Infect Dis*. 2023 Jun 30:jjad219. doi: 10.1093/infdis/jiad219. Online ahead of print.
17. Rosadas C, Taylor G. Mother-to-child HTLV-1 transmission: unmet research needs. *Front Microbiol*. 2019;10:999.
18. Tezuka K, Fuchi N, Okuma K, Tsukiyama T, Miura S, Hasegawa Y, et al. HTLV-1 targets human placental trophoblasts in seropositive pregnant women. *J Clin Invest*. 2020;130:6171-86.
19. Araujo A, Hall WW. Human T-lymphotropic virus type II and neurological disease. *Ann Neurol*. 2004;56:10-9.
20. Araujo A, Bangham C, Casseb J, Gotuzzo E, Jacobson S, Martin F, et al. Management of HAM/TSP: systematic review and consensus-based recommendations 2019. *Neurol Clin Pract* 2021;11:49-56.
21. Dantas L, Netto E, Glesby M, Carvalho E, Machado P. Dermatological manifestations of individuals infected with HTLV-1. *Int J Dermatol*. 2014;53:1098-102.
22. De Mendoza C, Pérez L, Rando A, Reina G, Aguilera A, Benito R, et al. Spanish HTLV Network. HTLV-1-associated myelopathy in Spain. *J Clin Virol*. 2023;169:105619.
23. Izumo S, Umehara F, Kashio N, Kubota R, Sato E, Osame M. Neuropathology of HTLV-1-associated myelopathy (HAM/TSP). *Leukemia*. 1997;11(Suppl 3):82-4.
24. Martin F, Taylor GP, Jacobson S. Inflammatory manifestations of HTLV-1 and their therapeutic options. *Expert Rev Clin Immunol* 2014;10:1531-46.
25. Tanajura D, Castro N, Oliveira P, Neto A, Muniz A, Carvalho NB, et al. Neurological manifestations in human T-Cell lymphotropic virus type 1 (HTLV-1)-infected individuals without HTLV-1-associated myelopathy/tropical spastic paraparesis: a longitudinal cohort study. *Clin Infect Dis*. 2015;61:49-56.
26. Tattermusch S, Bangham C. HTLV-1 infection: what determines the risk of inflammatory disease? *Trends Microbiol*. 2012;20:494-500.
27. Marcusso RM, Van Weyenbergh J, de Moura J, Dahy FE, De Moura Brasil Matos A, Haziot ME, et al. Dichotomy in fatal outcomes in a large cohort of people living with HTLV-1 in São Paulo, Brazil. *Pathogens*. 2019;9:E25.