We declare no competing interests.

© 2018 World Health Organization. Published by Elsevier Ltd. All rights reserved.

- 1 Fine P, Eames K, Heymann DL. "Herd immunity": a rough guide. Clin Infect Dis 2011; **52**: 911–16.
- 2 WHO, Strategic Advisory Group of Experts on Immunization. Assessment report of the Global Vaccine Action Plan. 2017. http://www.who.int/ immunization/web_2017_sage_gvap_assessment_report_en.pdf (accessed March 1, 2018).
- 3 Larson HJ, Jarrett C, Eckersberger E, Smith DM, Paterson P. Understanding vaccine hesitancy around vaccines and vaccination from a global perspective: a systematic review of published literature, 2007–2012. Vaccine 2014; 32: 2150–59.
- 4 Schuster M, Eskola J, Duclos P. Review of vaccine hesitancy: rationale, remit and methods. *Vaccine* 2015; **33**: 4157–60.
- 5 National Vaccine Advisory Committee. Assessing the state of vaccine confidence in the United States: recommendations from the National Vaccine Advisory Committee. *Public Health Rep* 2015; **130**: 573–95.

- 6 Hickler B, MacDonald NE, Senouci K, Schuh HB. Efforts to monitor global progress on individual and community demand for immunization: development of definitions and indicators for the Global Vaccine Action Plan Strategic Objective 2. Vaccine 2017; 35: 3515–19.
- 7 Bedford H, Attwell K, Danchin M, Marshall H, Corben P, Leask J. Vaccine hesitancy, refusal and access barriers: the need for clarity in terminology. *Vaccine* 2018; **36**: 6556–58.
 - MacDonald NE. Vaccine hesitancy: definition, scope and determinants. Vaccine 2015; **33:** 4161–64.
- 9 Peretti-Watel P, Larson HJ, Ward JK, Schulz WS, Verger P. Vaccine hesitancy: clarifying a theoretical framework for an ambiguous notion. PLoS Curr 2015; 7: ecurrents.outbreaks.6844c80ff9f5b273f34c91f71b7 fc289.
- Betsch C, Böhm R, Chapman GB. Using behavioral insights to increase vaccination policy effectiveness. *Policy Insights Behav Brain Sci* 2015; 2: 61–73.

Towards the elimination of HTLV-1 infection in Japan

8

The reported high prevalence of human T-cell leukaemia virus type 1 (HTLV-1) among the Aboriginal population in Australia triggered an open letter to WHO by Fabiola Martin and colleagues, urging action against HTLV-1 infection in May, 2018.¹ Japan is well known for having 1 million people living with HTLV-1 infection, the largest number in the world.^{2,3} HTLV-1 infection is concentrated in the southern islands of Kyushu and Okinawa, but it has also spread to urban areas such as Tokyo and Osaka because of internal migration. Given the large burden of HTLV-1 in Japan, the Government of Japan has maintained its strong commitment towards the elimination of HTLV-1 and has been a leader in implementing various policies to address HTLV-1 infection and related diseases.

Japan has been taking a leading role in advancing science regarding HTLV-1, from the identification of mechanisms of HTLV-1 transmission to the discovery and treatment of HTLV-1-related diseases, including adult T-cell leukaemia-lymphoma, HTLV-1-associated myelopathy and tropical spastic paraparesis (HAM-TSP), and various inflammatory disorders.⁴⁻⁶

Japan was the first country to implement two main strategies for prevention of HTLV-1 transmission. For the prevention of mother-to-child transmission (MTCT) through breastfeeding, antenatal HTLV-1 antibody screening and the recommendation for mothers with positive results to refrain from breastfeeding have been implemented in the Nagasaki prefecture in Kyushu since 1987;⁷ and HTLV-1 antibody screening of all donated blood in Japan has been implemented by the Japanese Red Cross since 1986.³

Efforts towards the elimination of HTLV-1 were further accelerated after the establishment of the HTLV-1 Task Force and endorsement of the Comprehensive Measures for HTLV-1 by the Japanese Government in 2010. The Comprehensive Measures consist of five main pillars. First, routine HTLV-1 antibody testing was incorporated into antenatal pregnancy screening throughout Japan in 2010. Second, counselling was made available for people living with HTLV-1 infection and associated diseases, and training materials and courses for health-care providers and counsellors were organised. Third, the coordination of care for HTLV-1-associated diseases was strengthened, with better coordination between health-care facilities that care for patients with HTLV-1-associated diseases and the development of quidelines for the management of adult T-cell leukaemia-lymphoma and HAM-TSP. Fourth, the website of the Ministry of Health, Labour and Welfare was updated to provide information on HTLV-1 infection and associated diseases, and communication materials were developed and distributed to raise public awareness. Fifth, research on epidemiology, pathophysiology, diagnosis, and therapy was strategically promoted, with JP¥10 billion (approximately US\$9 million) specifically assigned on a recurring annual basis for the research of HTLV-1associated diseases.

To promote these five main pillars of policy, the Comprehensive Measures introduced the establishment of the HTLV-1 Control Promotion Council, consisting of patient groups, scientists, health-care professionals,



and government officials; of public health councils in each prefecture for the prevention of MTCT of HTLV-1; and of conferences to better coordinate projects among the various research groups on HTLV-1 infection and associated diseases.

Further progress after the endorsement of the Comprehensive Measures in 2010 include the following measures. A manual on prevention of MTCT of HTLV-1 was developed, which was revised in 2016 to recommend exclusive formula feeding for all mothers who are positive for HTLV-1.8 Previous recommendations included short-term breastfeeding or frozen-thawed breastmilk feeding, in addition to formula feeding. For pregnant women positive for HTLV-1 antibody, but with an indeterminate result with the western blot test, confirmatory PCR tests are now recommended. 135 health-care facilities provide care for people living with HTLV-1 infection, 146 for people with adult T-cell leukaemia-lymphoma, and 92 for people with HAM-TSP. The Japanese Society of HTLV-1 and Associated Diseases offers accreditation to registered health-care facilities, core hospitals that provide counselling and support services for people living with HTLV-1 and training for health-care workers. This accreditation started in 2018, and six hospitals have been accredited. Lastly, a public awareness campaign, in collaboration with Hataraku Saibou (Cells at Work!; a Japanese animation series), was launched in 2018.

We need to acknowledge further challenges to be tackled. Progress is still needed in basic and clinical research; it is still very difficult to save the life of patients with adult T-cell leukaemia-lymphoma, there is no cure for HAM-TSP, and it is also difficult to prevent the development of HTLV-1-associated disease in people living with HTLV-1 infection. The promotion of preventive measures and awareness of horizontal transmission, mainly sexual transmission, is needed. Routine screening for HTLV-1 in organ donors needs to be considered and more data are needed to assess the prognosis of organ recipients from donors positive for HTLV-1.

Japan continues its strong commitment towards the elimination of HTLV-1. We believe that our progress towards this goal, and our approach to current challenges and future directions, could serve as a good model for other countries in tackling HTLV-1 infection.

*Takeshi Nishijima, Satoshi Shimada, Hiroyuki Noda, Kuniaki Miyake

Infectious Disease Control Division, Health Service Bureau, Ministry of Health, Labour and Welfare, Tokyo 100–8916, Japan nishijimat@who.int

We declare no competing interests. TN is currently on secondment to WHO.

- Martin F, Tagaya Y, Gallo R. Time to eradicate HTLV-1: an open letter to WHO. Lancet 2018; 391: 1893-94.
- 2 Gessain A, Cassar O. Epidemiological aspects and world distribution of HTLV-1 infection. Front Microbiol 2012; **3:** 388.
- 3 Satake M, Iwanaga M, Sagara Y, Watanabe T, Okuma K, Hamaguchi I. Incidence of human T-lymphotropic virus 1 infection in adolescent and adult blood donors in Japan: a nationwide retrospective cohort analysis. Lancet Infect Dis 2016; 16: 1246–54.
- 4 Hinuma Y, Nagata K, Hanaoka M, et al. Adult T-cell leukemia: antigen in an ATL cell line and detection of antibodies to the antigen in human sera. Proc Natl Acad Sci USA 1981; 78: 6476–80.
- 5 Yodoi J, Takatsuki K, Masuda T. Two cases of T-cell chronic lymphocytic leukemia in Japan. N Engl J Med 1974; 290: 572–73.
- 6 Kinoshita K, Yamanouchi K, Ikeda S, et al. Oral infection of a common marmoset with human T-cell leukemia virus type-I (HTLV-I) by inoculating fresh human milk of HTLV-I carrier mothers. Jpn J Cancer Res 1985; 76: 1147–53.
- ⁷ Hino S. Establishment of the milk-borne transmission as a key factor for the peculiar endemicity of human T-lymphotropic virus type 1 (HTLV-1): the ATL Prevention Program Nagasaki. Proc Jpn Acad Ser B Phys Biol Sci 2011; 87: 152–66.
- 8 Ministry of Health, Labour and Welfare. Manual for prevention of motherto-child transmission of HTLV-1. 2016. https://www.mhlw.go.jp/bunya/ kodomo/boshi-hoken16/dl/06.pdf (accessed March 28, 2018, in Japanese).