

contain genes that codify proteins. All 30,000 obelisks described so far in the gut and mouth of humans by Stanford's team encode a single major protein known as obulin, and many encode a second smaller obulin. These proteins do not make a shell. Of note, obulins do not share any homology with any other known protein. Hence, there is no clue about their function.

Obelisks are not rare and must be widespread across multiple niches. They were detected in around 7% microbiome datasets from the human gut and 50% of datasets from the human mouth. Different obelisk types were found in different body sites and in distinct donors. Long-term data revealed that people can harbor a single obelisk type for around a year.

Bacteria and fungi are likely hosts of obelisks. At this time, it is unclear whether obelisks may be parasitic and harm cells or they may be beneficial. Hosts may have evolved elaborating defense mechanisms against obelisks or else actively recruit them to gain some unsuspected advantage. If obelisks modulate the human microbiome, this may in turn have implications for human health – they may even have therapeutic potential.

Alternatively, obelisks may cause neither harm nor benefit to their microbial host or humans. Instead, they may simply exist as stealthy evolutionary passengers, silently, and endlessly replicating, like the original “*selfish gene*” (Dawkins R. 1976).

Experiments with obelisks are planned and could reveal truths about the origin of life itself. Because viroids and their relatives are small, simple, and have the capacity to self-replicate, they could be the precursors of all life on Earth. One big question is whether viruses evolved from increasingly complex viroids and obelisks or emerged first and then degenerated into these simpler structures (Penni E. *Science* 2024). The long-term evolution of viruses on Earth starts to slowly emerge.

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## The origin of the four major focus of HTLV-1 in Latin America

HTLV-1 was discovered in 1980 as the first human retrovirus. As zoonosis, HTLV-1 derives from jumps to

humans from simian T-lymphotropic virus naturally infecting monkeys in Central and Western Sub-Saharan Africa for thousands of years. Molecular clock study estimates an ancestor for all human HTLV-1s around 30,000-40,000 years ago<sup>1</sup>.

As expected, Africa accounts for the largest number of HTLV-1 infections worldwide<sup>2</sup>. However, Latin America is the second region endemic for HTLV-1 globally. As shown in figure 1, four areas of high HTLV-1 prevalence can be recognized, namely, at the Caribbean basin, Brazil, Peru, and along the Andes mountains. Phylogenetic studies have examined viral sequences from isolates in all these regions, including the exam of older mummies<sup>3-5</sup>. Based on these data, a reconstruction of the earliest introduction and dissemination of HTLV-1 in America can be postulated.

From its origin in West and Central Africa, HTLV-1 was introduced into Central Asia. During the last Glaciation, human populations migrated through the Bering Strait 35,000-15,000 years ago and introduced HTLV-1 in the Americas. The presence of HTLV-1 in aboriginal populations from Kamchatka at one side of the Bering Strait and in native skimos and Amerindians on the other side confirms this hypothesis<sup>6</sup>. During the warming period that followed, migrations to the South occurred along the long Andean mountains range. This movement carried HTLV-1 south and accounts for the presence of HTLV-1 across distinct native Amerindian tribes from Colombia to Chile<sup>7-11</sup>.

During the XVI to XIX centuries, a second wave of HTLV-1 arrived to the Caribbean basin and Brazil during the colonial times along with the slaves taken in West Africa mostly by Portuguese and British<sup>12-15</sup>. More recently, during the XX century, migrants from Japanese endemic southern areas to Peru and Brazil established a new settlement of HTLV-1 in Latin America.

In Brazil and Peru, both highly endemic countries for HTLV-1 infection, two major distinct sources of the virus can be recognized. In the Amazon tribes of inner Brazil, the ancient HTLV-1 variants infect Amerindians. In the Brazilian coast, HTLV-1 variants that arrived with the slavery trade predominate by far. The arrival of Japanese to large coastal cities of Brazil during the last century added a new variant. In Peru, the newly arrived Japanese HTLV-1 variant has been added to the ongoing circulation of ancient HTLV-1 introduced along the Andean populations thousands of years ago.

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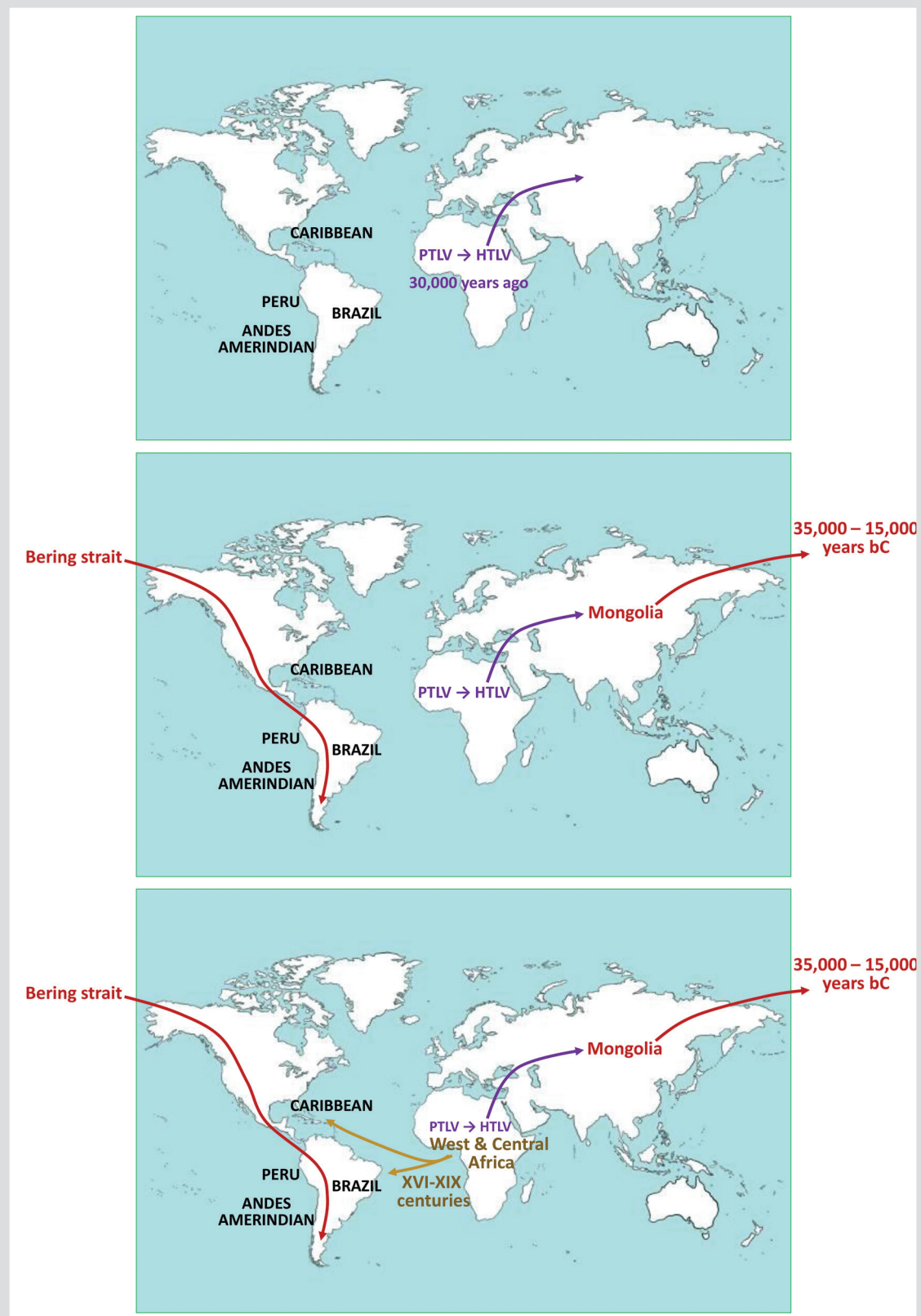


Figure 1. Proposed HTLV-1 origins in Latin America (continues).

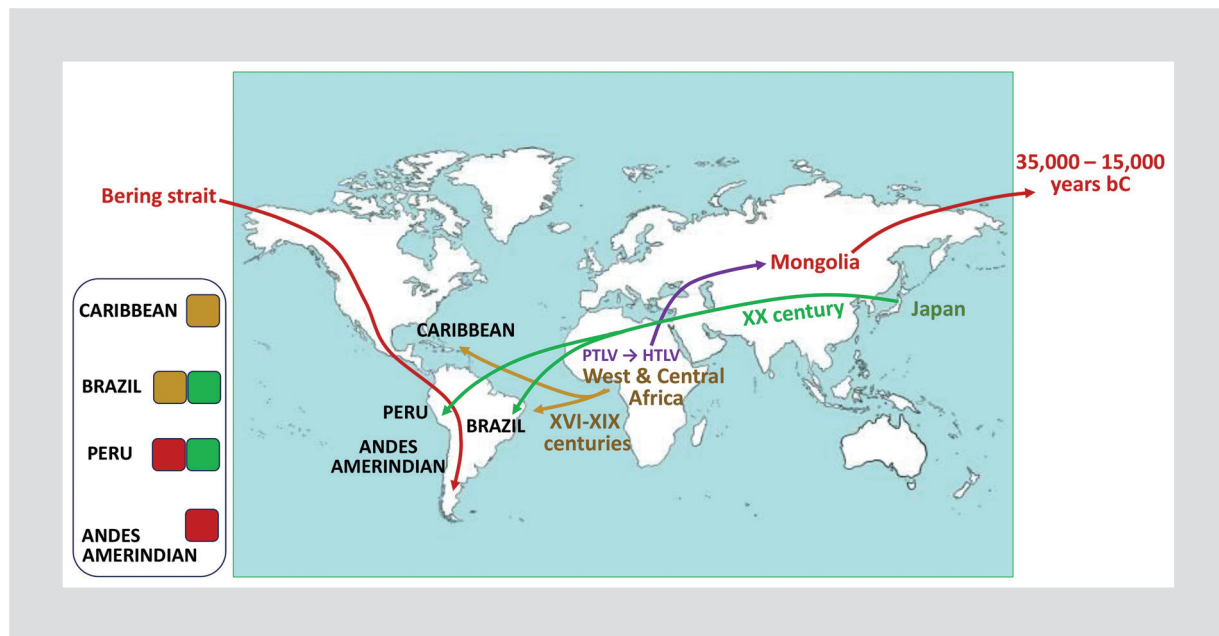


Figure 1. (Continued) Proposed HTLV-1 origins in Latin America.

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