

Ancient human DNA hints at why multiple sclerosis affects so many northern Europeans today

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This photo provided by the University of Copenhagen in January 2024 shows the process of ancient DNA extraction at the Lundbeck Foundation GeoGenetics Center in Copenhagen. According to research published Wednesday, Jan. 10, 2024, in the journal *Nature*, ancient DNA helps explain why northern Europeans have a higher risk of multiple sclerosis than other ancestries: It's a genetic legacy of horseback-riding cattle herders who swept into the region about 5,000 years



ago. Credit: Mikal Schlosser/University of Copenhagen via AP

Ancient DNA helps explain why northern Europeans have a higher risk of multiple sclerosis than other ancestries: It's a genetic legacy of horseback-riding cattle herders who swept into the region about 5,000 years ago.

The findings come from a huge project to compare modern DNA with that culled from ancient humans' teeth and bones—allowing scientists to trace both prehistoric migration and disease-linked genes that tagged along.

When a Bronze Age people called the Yamnaya moved from the steppes of what are now Ukraine and Russia into northwestern Europe, they carried gene variants that today are known to increase people's risk of multiple sclerosis, researchers reported Wednesday.

Yet the Yamnaya flourished, widely spreading those variants. Those genes probably also protected the nomadic herders from infections carried by their cattle and sheep, concluded the research published in the journal *Nature*.

"What we found surprised everyone," said study co-author William Barrie, a genetics researcher at the University of Cambridge. "These variants were giving these people an advantage of some kind."

It's one of several findings from a first-of-its-kind gene bank with thousands of samples from <u>early humans</u> in Europe and western Asia, a project headed by Eske Willerslev of Cambridge and the University of Copenhagen who helped pioneer the study of ancient DNA. <u>Similar</u> <u>research</u> has traced even earlier cousins of humans such as Neanderthals.



Using the new gene bank to explore MS was a logical first step. That's because while MS can strike any population, it is most common among white descendants of northern Europeans and scientists have been unable to explain why.

The potentially disabling disease occurs when immune system cells mistakenly attack the protective coating on nerve fibers, gradually eroding them. It causes varying symptoms—numbness and tingling in one person, impaired walking and vision loss in another—that often wax and wane.



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It's not clear what causes MS although a leading theory is that certain infections could trigger it in people who are genetically susceptible. More than 230 genetic variants have been found that can increase someone's risk.

The researchers first examined DNA from about 1,600 ancient Eurasians, mapping some major shifts in northern Europe's population. First, farmers from the Middle East began supplanting <u>hunter-gatherers</u> and then, nearly 5,000 years ago, the Yamnaya began moving in—traveling with horses and wagons as they herded cattle and sheep.

The research team compared the ancient DNA to about 400,000 presentday people stored in a UK gene bank, to see the MS-linked genetic variations persist in the north, the direction the Yamnaya moved, rather than in southern Europe.

In what is now Denmark, the Yamnaya rapidly replaced ancient farmers, making them the closest ancestors of modern Danes, Willerslev said. MS rates are particularly high in Scandinavian countries.

Why would gene variants presumed to have strengthened ancient immunity later play a role in an autoimmune disease? Differences in how <u>modern humans</u> are exposed to animal germs may play a role, knocking the immune system out of balance, said study co-author Dr. Astrid Iversen of Oxford University.

The findings finally offer an explanation for the north-south MS divide



in Europe but more work is needed to confirm the link, cautioned genetic expert Samira Asgari of New York's Mount Sinai School of Medicine, who wasn't involved with the research, in an accompanying commentary.

More information: Elevated Genetic Risk for Multiple Sclerosis Originated in Steppe Pastoralist Populations, *Nature* (2024). <u>DOI:</u> <u>10.1038/s41586-023-06618-z</u>

Samira Asgari et al, Prehistoric events might explain European multiple sclerosis risk, *Nature* (2024). DOI: 10.1038/d41586-023-03977-5, doi.org/10.1038/d41586-023-03977-5

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