

## Ancient history of Lyme disease revealed with bacterial genomes

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Credit: Yale University

A team of researchers led by the Yale School of Public Health has found that the Lyme disease bacterium is ancient in North America, circulating silently in forests for at least 60,000 years—long before the disease was first described in Lyme, Connecticut, in 1976 and long before the arrival of humans.



For the first time, the full genomes of the Lyme <u>disease bacterium</u>, Borrelia burgdorferi, were sequenced from <u>deer ticks</u> to reconstruct the history of this invading pathogen.

The finding shows that the ongoing Lyme disease epidemic was not sparked by a recent introduction of the bacterium or an evolutionary change—such as a mutation that made the bacterium more readily transmissible. It is tied to the ecological transformation of much of North America. Specifically, forest fragmentation and the population explosion of deer in the last century have created optimal conditions for the spread of ticks and triggered this ongoing epidemic.

Katharine Walter conducted the research while a doctoral student at Yale School of Public Health and is lead author of the study published in *Nature Ecology and Evolution*.

"The Lyme disease bacterium has long been endemic," she said. "But the deforestation and subsequent suburbanization of much of New England and the Midwest created conditions for deer ticks—and the Lyme disease bacterium—to thrive."





Deer ticks are collected on Mashomack Preserve on New York's Shelter Island by dragging a one-meter square section of corduroy across the forest floor. Credit: Katharine Walter

Lyme disease is the most common vector-borne disease in North America. Since it was first described in the 1970s, the disease has rapidly spread across New England and the Midwest. Reported cases of Lyme disease have more than tripled since 1995 and the Centers for Disease Control and Prevention now estimate that more than 300,000 Americans fall ill each year.

The team turned to genomics to reveal the bacterium's origins. By comparing B. burgdorferi genomes collected from different areas and over a 30-year period, the team built an evolutionary tree and reconstructed the history of the pathogen's spread.



Researchers collected deer ticks, vectors of B. burgdorferi, from across New England. They focused sampling efforts in areas predicted to be sources of the epidemic—Cape Cod and areas around Long Island Sound. Over 7,000 <u>tick</u> were collected from these areas during the summer of 2013. To extend the spatial scope of the study, collaborators in the South, Midwest, and across Canada contributed ticks to the team.

Using a method the team previously developed to preferentially sequence bacterial DNA (and avoid sequencing only DNA from the tick), the researchers sequenced 148 B. burgdorferi genomes. Earlier studies of the evolutionary history of B. burgdorferi have relied upon short DNA markers rather than full genomes. Reading the one million letters of the full bacterial genome allowed the team to piece together a more detailed history. The team drew an updated evolutionary tree which showed that the bacterium likely originated in the northeast of the United States and spread south and west across North America to California.

Birds likely transported the pathogen long distances to new regions and small mammals continued its spread. Imprinted on the bacterial genomes was also a signature of dramatic population growth. As it evolved, it seemed to have proliferated.





A tick collecting site on Cuttyhunk Island, one of the Elizabeth Islands in Massachusetts. The earliest records of deer ticks in the northern United States were on the Elizabeth Islands as were some of the first reported cases of Lyme disease. Credit: Katharine Walter

The tree was also far older than the team had expected—at least 60,000 years old. This means that the bacterium existed in North America long before the disease was described by medicine and long before humans first arrived in North America from across the Bering Strait (about 24,000 years ago)

This findings clarify that the bacterium is not a recent invader. Diverse lineages of B. burgdorferi have long existed in North America and the current Lyme disease epidemic is the result of ecological changes that have allowed deer, ticks and, finally, bacterium to invade.



The explosion of deer in the twentieth century into suburban landscapes, free of wolf predators and with strict hunting restrictions, allowed deer ticks to rapidly invade throughout much of New England and the Midwest. Climate change has also contributed. Warmer winters accelerate ticks' life cycles and allow them to survive an estimated 28 miles further north each year.

Ticks expanded into suburbanized landscapes—full of animals like whitefooted mice and robins, excellent hosts for B. burgdorferi. The expansion of ticks into habitats with ideal hosts allowed the bacterium to spread.

**More information:** Katharine S. Walter et al. Genomic insights into the ancient spread of Lyme disease across North America, *Nature Ecology & Evolution* (2017). DOI: 10.1038/s41559-017-0282-8

Provided by Yale University

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