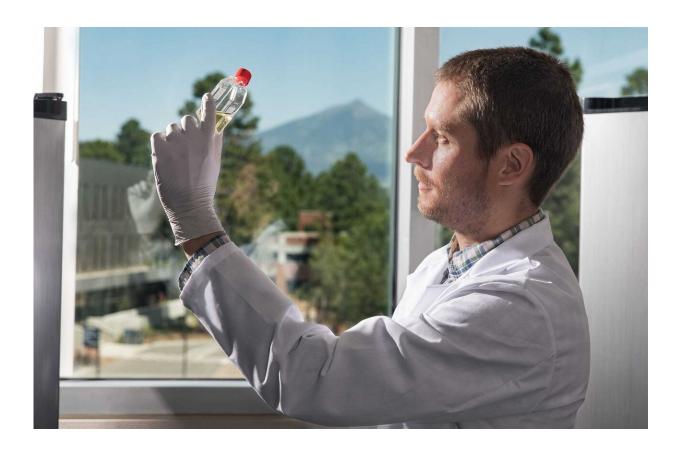


## Shedding light on reptilian health: Researchers investigate origins of snake fungal disease in US

June 29 2022



A team of researchers, including NAU assistant professor Jason Ladner, conducted a genetic study of snake fungal disease that was recently published in *PLOS Biology*. Credit: Northern Arizona University



Although only recently recognized as an issue in wildlife ecology, snake fungal disease (SFD) is of emerging concern in the U.S., with parallels among other better-known wildlife fungal diseases such as white-nose syndrome in bats. SFD can be deadly to snakes, and even in milder cases disrupts an animal's abilities to perform normal biological functions such as hibernation, eating and avoiding predators.

To better understand SFD, a team of researchers, including assistant professor Jason Ladner of Northern Arizona University's Pathogen and Microbiome Institute, conducted a genetic study of the pathogen that was recently published in *PLOS Biology*, "The population genetics of the causative agent of <u>snake</u> fungal <u>disease</u> indicate recent introductions to the U.S.."

Collaborating with study co-author Jeff Lorch of the U.S. Geological Survey (USGS) and other scientists from the USGS, Genencor Technology Center, the University of California-Riverside, Stetson University, the Institute of Zoology, the University of Kentucky and Holyoke Community College, Ladner's goal was to determine whether SFD originated in the U.S. or was introduced from outside the country, which could provide a historical basis for how it emerged—and ultimately inform management of the disease.

"Snake <u>fungal disease</u> first came to be recognized in the U.S. around 2008. There happened to be a well-studied population of rattlesnakes in Illinois that started coming down with some very severe fungal infections. People asked, 'OK, what is this thing? Where is it? What's going on? Is this a new emerging fungal pathogen or not?' What they eventually found was that it was already almost everywhere, at least in the eastern half of the U.S.," Ladner said.

SFD, though seemingly not as deadly as other wildlife <u>fungal diseases</u>, is still a worrying threat to animals that represent an important part of the



ecosystem. "We're very concerned, not just about SFD's effect to drive <u>population declines</u>, but also as a contributing factor amongst many other threats that snakes are already facing, like habitat destruction or over-collection for the pet trade," Lorch said.

Understanding wildlife diseases is critical, both in the context of ecosystem health and in their potential effects on humans. "I have a lot of interest in wildlife diseases, partially because wildlife serves as important reservoirs for diseases that could potentially emerge in humans; SARS-coronavirus-2 is a great example of that. If we want to be prepared for the next emerging infectious disease in humans, we need to better understand the pathogens currently circulating in wildlife populations which may have the potential to be transmitted to humans," Ladner said.

The study presented unique difficulties, however. "For snakes, there's almost no long-term population trend data, especially when we compare snakes to an animal like bats, which have suffered from white-nose syndrome," Lorch said. "In many states, historical data on bat populations exists because they're not generally as difficult to monitor as some other types of wildlife."

Snakes, in contrast, "are pretty secretive animals. They're not something that you probably see on the landscape routinely, unless you're looking for them," Lorch explains. Without a large body of <u>historical data</u> on North American snake populations, "it makes it hard to say what snake populations were doing before SFD was noticed. Long-term trends are really difficult to decipher."

Prior to beginning research, the team had two hypotheses on how the disease originated in the U.S. "One hypothesis was that the fungus that causes this disease may have been introduced only recently into the U.S. and then has been spreading within the past several decades, maybe 100



years. The alternative hypothesis was that this pathogen has been here for a long time and is essentially native to the U.S.; maybe it's been here for thousands of years and has been co-evolving with these snake populations. In the latter case, maybe it seems to be emerging simply because we're looking for it now. Or there's been some type of environmental change, maybe something linked to climate change, that is leading to an increase in the number of cases even though this pathogen has been here all along," Ladner said.

In order to track the disease's evolution, Ladner and Lorch created a "family tree" for strains of the fungus that causes SFD found in the U.S. "One of the ways we could reconstruct the history of the disease was to look at the genetics of the pathogen to get an idea of how long it's been here and how it's changed over time," Lorch said.

Studying the genetics of SFD provided the team with a trail of breadcrumbs, revealing more about its history and throwing light on SFD cases in the U.S. "The reason that genomic data is useful for doing this is because each time this fungus replicates, grows and divides, the polymerase (the molecule that makes the new copy) sometimes makes mistakes. Those mistakes result in mutations. And then those mutations will be passed on through the generations. By looking at those different mutations in the population, we can understand how long certain lineages have existed and have some idea of how the different strains are related to each other. And that can tell us something about how long SFD has been here," Ladner said.

After taking samples from different SFD-affected snakes, the team performed genetic sequencing on 82 strains of the fungus. This included strains of SFD isolated from wild snakes in the U.S. and Europe, as well as captive snakes from three different continents. Based on the genetic similarities and differences among the strains, the team was able to partially reconstruct the evolutionary history of this fungus. "In the U.S.,



we found that there are several divergent lineages of this fungus circulating, but a lack of intermediates between these lineages, which would be expected if they originated in the U.S. Because of that, we think that there were likely multiple, somewhat recent introductions of this fungus to the U.S., and that an unsampled population, somewhere else in the world, acted as a source," Ladner said.

This evidence allowed the team to form conclusions on how SFD arrived in America. "It suggests that this fungus was introduced to the United States through anthropogenic means—humans moving these snakes around. The most likely culprit is the trading of captive <u>snakes</u> as pets: the different clonal lineages that we see in the U.S., we also see represented in captive snake populations," Ladner said.

The study provides guidance for future management of SFD in the U.S., as well as a better understanding of how it was introduced. "If we had caught SFD being introduced very early on, then you can imagine trying to stop the spread of the disease in the U.S. and potentially even eradicate it. I think that's unlikely at this point, given how widespread it is. However, I think it's still helpful to better understand the mechanism for how SFD was introduced, as there's still the potential for new introductions of diverse strains from these source populations. If we know that this fungus was introduced several times over the past several decades through the captive animal trade, then putting more restrictions and controls and testing animals in that process could be important for preventing further spread," Ladner said.

Though their work provides critical insight on SFD, its treatment and movement in the U.S., both scientists stress the need for further research. "What I'm hoping is that this study increases awareness of the disease. I think SFD warrants more of our attention," Lorch said.

More work needs to be done to assess the ecosystem, population and



species effects of SFD. "The broader question of, 'what is going to be the impact of this <u>fungal pathogen</u> on these snake populations?' is a very open question and needs more research," Ladner said.

**More information:** Jason T. Ladner et al, The population genetics of the causative agent of snake fungal disease indicate recent introductions to the USA, *PLOS Biology* (2022). DOI: 10.1371/journal.pbio.3001676

Provided by Northern Arizona University

Citation: Shedding light on reptilian health: Researchers investigate origins of snake fungal disease in US (2022, June 29) retrieved 8 May 2024 from <u>https://phys.org/news/2022-06-reptilian-health-snake-fungal-disease.html</u>

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