# Cryptococcus gattii Emerges in Pacific Northwest

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ATLANTA — *Cryptococcus gattii*, a fungal pathogen previously found only in tropical and subtropical areas, is emerging as a serious infection in the U.S. Pacific Northwest, according to the Centers for Disease Control and Prevention.

It has infected at least 60 people and 52 animals in Washington, Oregon, Idaho, and California, according to Julie R. Harris, Ph.D., a CDC epidemiologist. This is "one of the most interesting emerging infectious diseases in the United States today," she said.

In September 2009, the CDC and various organizations in the affected states began passive surveillance for human and veterinary cases in the Pacific North-

'Patients might require prolonged therapy with antifungal drugs, or special procedures and follow-up that normally aren't warranted with *C. neoformans* infections.'

west to understand the extent of the pathogen's spread and its epidemiology.

Twenty cases were identified in 2009, and she expects that even more cases will be identified this year.

Of the 39 patients on which researchers have data, 89% were hospitalized; 33% died of or with the *C. gattii* infection, she said. The mean age of infected patients is 52 years (range, 15-95 years); *C. gattii* appears to affect men and women equally. It's rarely a pediatric disease. Moreover, it appears to infect both immunocompetent and immunocompromised patients, most of whom were not infected with HIV.

The most common symptoms are cough, shortness of breath, headache, nausea and vomiting, fever, and weight loss. The patients developed pneumonia (57%), meningitis (44%), encephalitis (21%), and cryptococcomas in the lung (34%) and the brain (25%).

Emilio DeBess, D.M.V., of the Oregon Department of Human Services, noted that *C. gattii* spores appear to live in association with certain trees and the soil around those trees. Humans can become infected by inhalation of the airborne fungi. "As far as we know, there is no human-to-human or animal-to-human transmission," he said.

Before 1999, reports of infection were generally limited to tropical and subtropical areas and were most prevalent in Australia. In 1999, it was identified in Vancouver Island, B.C., and then jumped to the mainland in 2004, and spread to the U.S. Pacific Northwest. Most cases are in Washington and Oregon, but reports of infection also have been identified in Idaho and California, Dr. DeBess said.

One explanation for the recent transmissions is that the spores are carried by trucks on the I-5 corridor, which stretches from British Columbia to San Diego. "There are a lot of theories out there. That's the one we are working with at this point," he said. "Marion County [Oregon] along the I-5 corridor is the Petri dish of *C. gattii*."

Then again, Dr. DeBess acknowledged that "it's hard to say" whether *C. gattii* is

truly spreading or if more cases are being identified as a result of more intense surveillance.

*C. gattii* may be underdiagnosed and, as a result, the mortality rate may be inflated, he added. Both he and Dr. Harris noted the U.S. mortality rate is higher than that reported in British Columbia. He hopes to improve surveillance and promote awareness among physicians and veterinarians.

One of the clinical challenges, Dr. Harris noted, is that most labs cannot distinguish between *Cryptococcus neoformans* and *C. gattii*.

"We want all physicians to be aware that this is a new possible etiology of cryptococcal infections in the United States, and if infected with *C. gattii*, their patients might require prolonged therapy with antifungal drugs, or special procedures and follow-up that normally

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# Important Safety Information for Lantus®

#### **Contraindications**

Lantus® is contraindicated in patients hypersensitive to insulin glargine or one of its excipients.

#### Warnings and precautions

Monitor blood glucose in all patients treated with insulin. Insulin regimens should be modified cautiously and only under medical supervision. Changes in insulin strength, manufacturer, type, or method of administration may result in the need for a change in insulin dose or an adjustment in concomitant oral antidiabetic treatment.

Do not dilute or mix Lantus® with any other insulin or solution. If mixed or diluted, the solution may become cloudy, and the onset of action/time to peak effect may be altered in an unpredictable manner. Do not administer Lantus® via an insulin pump or intravenously because severe hypoglycemia can occur. Insulin devices and needles must not be shared between patients.

Hypoglycemia is the most common adverse reaction of insulin therapy, including Lantus<sup>®</sup>, and may be life-threatening

Severe life-threatening, generalized allergy, including anaphylaxis, can occur.

A reduction in the Lantus® dose may be required in patients with renal or hepatic impairment.

#### **Drug interactions**

Certain drugs may affect glucose metabolism, requiring insulin dose adjustment and close monitoring of blood glucose. The signs of hypoglycemia may be reduced in patients taking anti-adrenergic drugs (e.g., beta-blockers, clonidine, guanethidine, and reserpine).

#### **Adverse reactions**

Other adverse reactions commonly associated with Lantus® are injection site reaction, lipodystrophy, pruritus, and rash.

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Lantus® is a long-acting insulin analog indicated to improve glycemic control in adults and children (6 years and older) with type 1 diabetes mellitus and in adults with type 2 diabetes mellitus. Lantus® should be administered once a day at the same time every day.

Important Limitations of Use: Lantus® is not recommended for the treatment of diabetic ketoacidosis. Use intravenous short-acting insulin instead.

Lantus® SoloSTAR® is a disposable prefilled insulin pen.

Please see brief summary of full prescribing information for Lantus® on the next page.

References: 1. Data on file, sanofi-aventis U.S. LLC. 2. Lantus Prescribing Information. September 2009.

aren't warranted with *C. neoformans* infections," she said in an interview.

"Although most patients either live in or have visited the Pacific Northwest, we would encourage all physicians to keep *Cryptococcus gattii* in mind when they see an atypical patient with cryptococcal infection, as we don't yet know the extent of the endemic area," she said.

Because *C. neoformans* is so much more common than *C. gattii*, at this point she is not suggesting submission and speciation of every *Cryptococcus* isolate. "However, for patients with suspected *C. gattii* infection, we encourage

clinicians to report the cases to their states and submit isolates as requested for further speciation," she said.

Risk factors are poorly defined at this point, she added. "We know anecdotally that many patients live in or near wooded areas or have occupations that involve contact with soil or wood, but we don't have a good group with which to compare these types of exposures. We do plan to conduct a study on this in the near future."

**Disclosures:** Dr. DeBess and Dr. Harris reported that they had no conflicts.



