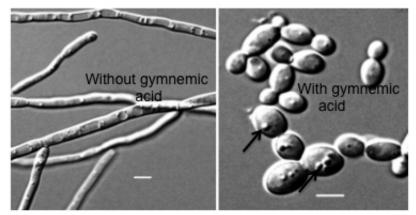


Research treats the fungus among us with nontoxic medicinal compound

September 12 2013, by Jennifer Tidball



Gymnemic acids prevent *C. albicans* yeast-to-hyphal transition and hyphal growths. Scale bars, 5µm.

Credit: Kansas State University

A Kansas State University microbiologist has found a breakthrough herbal medicine treatment for a common human fungal pathogen that lives in almost 80 percent of people.

Govindsamy Vediyappan, assistant professor of biology, noticed that diabetic people in developing countries use a medicinal herb called *Gymnema slyvestre* to help control sugar levels. He decided to study the microbiological use of *Gymnema slyvestre*—a tropical vine plant found in India, China and Australia—to see if it could treat a common human <u>fungal pathogen</u> called *Candida albicans*.



The investigation was successful on two levels: Vediyappan's research team found the medicinal compound is both nontoxic and blocks the virulence properties of the <u>fungus</u> so that it is more treatable. The results are important for human health, <u>biomedical applications</u> and potential drug development.

"We have shown that this compound is safe to use because it doesn't hurt our body cells, yet it blocks the virulence of this fungus under in vitro conditions," Vediyappan said. "Taking the medicine could potentially help patients control the invasive growth of the fungus and also help bring their sugar levels down."

Candida albicans is one of the major fungal pathogens in humans because it lives in oral and intestinal areas as a normal flora, Vediyappan said. But the fungus can overgrow and can cause oral, intestinal and genital infections. The fungus kills almost 30 percent of people who have it and it is a concern among cancer patients—especially patients with neck or <u>oral cancer</u>—HIV patients, organ transplant patients and other people with compromised immune systems.

The fungus can grow in two forms: a treatable yeast and a difficult-to-treat hyphal form. Once the fungus transforms from a yeast to a hyphal growth it becomes difficult to treat because the hyphal growth has long filament-like structures that can spread into various organs. Vediyappan's study aimed to block the hyphal growth form.

"Once it gets into the tissue, it spreads like roots and is difficult to contain by our immune system," Vediyappan said.

If the fungus remains in yeast form, it is easy to manage and does not invade tissues. Vediyappan's research team purified gymnemic acid compounds that prevented the transition stage from occurring and stopped the fungus spread. The gymnemic acids come from the leaves of



Gymnema sylvestre, a traditional medicinal plant.

The research appears in the peer-reviewed journal *PLOS ONE* in an article titled "Gymnemic acids inhibit hyphal growth and virulence in *Candida albicans.*"

Gymnema extract is commonly used to treat diabetes and other ailments because it is a cost-effective treatment, Vediyappan said. Often, people drink the extract to control their sugar levels or to lose weight.

Although Vediyappan's research team is not the first to discover gymnemic acid compounds, the team is the first to discover that the compounds block the fungal transition. The researchers found that the compounds work quickly, too, which was an important characteristic. The treatable fungal yeast can transition to a hyphal growth within 30 minutes of an infection. When the hyphal transition has occurred, it will grow into branched filaments.

The gymnemic acid compounds are nontoxic, which is especially important for cancer patients and other immunocompromised patients. The gymnemic acids can stop the unwanted invasive infection while preserving important healthy cells.

The *Candida albicans* fungus also makes a biofilm, which is a fungal cell collection that can be difficult to treat. The researchers found that the gymnemic acid compounds converted the biofilm back to treatable yeast cells.

"This compound prevents the biofilm formation because hyphae are the major builders of biofilms and biofilms are resistant to antifungals," Vediyappan said. "Yeast cells by themselves cannot make biofilms and are sensitive to antifungal treatments."



Another interesting aspect: The gymnemic acid compounds also stopped the growth of *Aspergillus*, another fungal pathogen that can affect heart transplant patients and leukemia patients.

Vediyappan plans future studies to research mode of action, potential drug development, diabetes applications and other ways to improve treatment for *Candida albicans* and other fungal pathogens.

More information: dx.plos.org/10.1371/journal.pone.0074189

Provided by Kansas State University

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