Compounds from Tropical Seaweed May be Promising Anti-malarial Drugs

Georgia Tech Institute of Technology

A group of chemical compounds used by a species of tropical seaweed to ward off fungus attacks may have promising anti-malarial properties for humans. The compounds are part of a unique chemical signaling system that seaweeds use to battle enemies -- and that may provide a wealth of potential new pharmaceutical compounds.

Using a novel analytical process, researchers at the Georgia Institute of Technology found that the complex antifungal molecules are not distributed evenly across the seaweed surfaces, but instead appear to be concentrated at specific locations – possibly where an injury increases the risk of fungal infection.

A Georgia Tech scientist reported on the class of compounds, known as bromophycolides, at the annual meeting of the American Association for the Advancement of Science (AAAS) Feb. 21, 2011 in Washington, D.C. The research, supported by the National Institutes of Health, is part of a long-term study of chemical signaling among organisms that are part of coral reef communities.

"The language of chemistry in the natural world has been around for billions of years, and it is crucial for the survival of these species," said Julia Kubanek, an associate professor in Georgia Techs School of Biology and School of Chemistry and Biochemistry. "We can co-opt these chemical processes for human benefit in the form of new treatments for diseases that affect us."

More than a million people die each year from malaria, which is caused by the parasite *Plasmodium falciparum*. The parasite has developed resistance to many anti-malarial drugs and has begun to show resistance to artemisinin -- today's most important anti-malarial drug. The stakes are high because half the world's population is at risk for the disease.

"These molecules are promising leads for the treatment of malaria, and they operate through an interesting mechanism that we are studying," Kubanek explained. "There are only a couple of drugs left that are effective against malaria in all areas of the world, so we are hopeful that these molecules will continue to show promise as we develop them further as pharmaceutical leads."

In laboratory studies led by Georgia Tech student Paige Stout from Kubaneks lab -- and in collaboration with California scientists -- the lead molecule has shown promising activity against malaria, and the next step will be to test it in a mouse model of the disease. As with other potential drug compounds, however, the likelihood that this molecule will have just the right chemistry to be useful in humans is relatively small.

Other Georgia Tech researchers have begun research on synthesizing the compound in the laboratory. Beyond producing quantities sufficient for testing, laboratory synthesis may be able to modify the compound to improve its activity -- or to lessen any side effects. Ultimately, yeast or another microorganism may be able to be modified genetically to grow large amounts of bromophycolide.

The researchers found the anti-fungal compounds associated with light-colored patches on the surface of the *Callophycus serratus* seaweed using a new analytical technique known as desorption electrospray ionization mass spectrometry (DESI-MS). The technique was developed in the laboratory of Facundo Fernandez, an associate professor in Georgia Tech's School of Chemistry and Biochemistry. DESI-MS allowed researchers for the first time to study the unique chemical activity taking place on the surfaces of the seaweeds.

As part of the project, Georgia Tech scientists have been cataloging and analyzing natural compounds from more than 800 species found in the waters surrounding the Fiji Islands. They were interested in *Callophycus serratus* because it seemed particularly adept at fighting off microbial infections.

Using the DESI-MS technique, researchers Leonard Nyadong and Asiri Galhena analyzed samples of the seaweed and found groups of potent anti-fungal compounds. In laboratory testing, graduate student Amy Lane found that these bromophycolide compounds effectively inhibited the growth of *Lindra thalassiae*, a common marine fungus.

"The alga is marshalling its defenses and displaying them in a way that blocks the entry points for microbes that might invade and cause disease," Kubanek said. "Seaweeds don't have immune responses like humans do. But instead, they have some chemical compounds in their tissues to protect them."

Though all the seaweed they studied was from a single species, the researchers were surprised to find two distinct groups of anti-fungal chemicals. From one seaweed sub-population, dubbed the "bushy" type for its appearance, 23 different anti-fungal compounds were identified. In a second group of seaweed, the researchers found 10 different anti-fungal compounds - all different from the ones seen in the first group.

In the DESI-MS technique, a charged stream of polar solvent is directed at the surface of a sample under study at ambient pressure and temperature. The spray desorbs molecules, which are then ionized and delivered to the mass spectrometer for analysis.

"Our collaborative team of researchers from the Department of Biomedical Engineering and the College of Sciences has worked within the newly-formed Bioimaging Mass Spectrometry Center at Georgia Tech to better understand the mechanisms of chemical defenses in marine organisms," said Fernandez. "This is an example of cross-cutting interdisciplinary research that characterizes our institute."

Kubanek is hopeful that other useful compounds will emerge from the study of signaling compounds in the coral reef community.

"In the natural world, we have seaweed that is making these molecules and we have fungi that are trying to colonize, infect and perhaps use the seaweed as a substrate for its own growth," Kubanek said. "The seaweed uses these molecules to try to prevent the fungus from doing this, so there is an interaction between the seaweed and the fungus. These molecules function like words in a language, communicating between the seaweed and the fungus."

Research News & Publications Office
Georgia Institute of Technology
75 Fifth Street, N.W., Suite 314
Atlanta, Georgia 30308 USA

Media Relations Contacts: John Toon (404-894-6986)(jtoon@gatech.edu [1]) or Abby Robinson (404-385-3364)(abby@innovate.gatech.edu [2]).

Writer: John Toon

SOURCE [3]

Source URL (retrieved on *07/10/2014 - 8:27am*):
http://www.mdtmag.com/news/2011/02/compounds-tropical-seaweed-may-be-promising-anti-malarial-drugs

Links:
[1] mailto://www.gatech.edu/newsroom/jtoon@gatech.edu
[2] mailto://www.gatech.edu/newsroom/abby@innovate.gatech.edu
[3] http://www.gatech.edu/newsroom/release.html?nid=64407