

## Discovery may lead to lower doses of chemotherapy

October 9 2014



Ph.D. candidate Vajira Weerasekara and Professor Josh Andersen.

No matter what type of chemotherapy you attack a tumor with, many cancer cells resort to the same survival tactic: They start eating themselves.

Scientists at Brigham Young University discovered the two proteins that



pair up and switch on this process – known as <u>autophagy</u>.

"This gives us a therapeutic avenue to target autophagy in tumors," said Josh Andersen, a BYU chemistry professor. "The idea would be to make tumors more chemo-sensitive. You could target these proteins and the mechanism of this switch to block autophagy, which would allow for lower doses of chemotherapy while hopefully improving <u>patient</u> <u>outcomes</u>."

Lower doses would mean milder side effects. The prospect spurred an international hunt for this switch. For good reason, several other labs started with a <u>protein</u> called ATG9 as their prime suspect and then looked for its accomplice among thousands of other proteins.

But the BYU team, comprised mainly of undergraduate students, stumbled into the race unexpectedly, coming at it from a different direction. They wanted to know why <u>cancer cells</u> made a surplus of protein called 14-3-3 zeta.

Using breast cancer tissue in the lab, they forced <u>tumor cells</u> to undergo autophagy by depriving them of oxygen and glucose. A comparison with a control group let them see that the two proteins hook up only when under attack. That's because stress makes Atg9 undergo a modification that enables 14-3-3 zeta to bind with it and switch the cancer cells to survival mode.

"This unique approach we used, partially by luck, gave us an advantage," Andersen said. "I don't think we would have discovered this through more conventional approaches."

The study results appear in the journal Molecular and Cellular Biology.

Andersen notes that several medicines already exist that could block



autophagy and make <u>chemotherapy</u> more effective. One of them is called chloroquine, an anti-malarial drug invented in 1934. In the event that this and other existing inhibitors don't cross-over safely or effectively, the study offers a blueprint for development of a drug specific to the task.

More information: <u>mcb.asm.org/content/early/2014</u>... 740-14.full.pdf+html

Provided by Brigham Young University

Citation: Discovery may lead to lower doses of chemotherapy (2014, October 9) retrieved 3 May 2024 from <u>https://medicalxpress.com/news/2014-10-discovery-doses-chemotherapy.html</u>

This document is subject to copyright. Apart from any fair dealing for the purpose of private study or research, no part may be reproduced without the written permission. The content is provided for information purposes only.