

Researchers discover key to mass producing beneficial plant compounds

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Purdue University scientists have discovered the switch in plants that turns off production of terpenoids, compounds that play roles in plant physiology and are used by humans in everything from fragrances and



flavoring to biofuels and pharmaceuticals.

Terpenoids can be used to develop nutritional supplements, <u>natural</u> <u>insecticides</u> and drugs that treat malaria and cancer. The chemotherapy drug Taxol, which is used to treat breast, ovarian, lung, bladder and prostate cancers, is derived from <u>terpenoids</u>. But plants often make them in such low quantities that extracting them for such uses is impractical.

The findings, reported in the journal *Nature Plants* by Natalia Dudareva, Purdue distinguished professor in the Department of Biochemistry and researcher in the Purdue Center for Plant Biology, Laura Henry, a recent doctoral graduate of Dudareva's lab, and colleagues from Purdue and the Salk Institute for Biological Studies, explain how plants control metabolic pathways by regulating the pool of monophosphates used for terpenoid production.

The scientists had earlier determined how plants turn on terpenoid production but having an understanding of both the "on" and "off" switches is essential for terpenoid yield. They also discovered a hidden constraint on flux through the plant terpenoid metabolic pathway.

"This is important basic knowledge that opens new targets for engineering of terpenoid metabolic pathways," Dudareva said. "Plants produce these compounds already, but the amount is small. It might have taken hundreds or thousands of plants to get enough of a compound to use it for something like a pharmaceutical product. This will lead to faster, more efficient ways way to obtain sufficient amounts of these products."

Isopentenyl phosphate kinases (IPK) convert pools of monophosphates into diphosphates that can be converted through downstream processes into terpenes. Dudareva and colleagues determined that two Nudix enzymes are responsible for dephosphorylation—removing a phosphate



group to turn the usable diphosphates back to the inert pool of monophosphates.

"We have shown that IPK and Nudix are working together to regulate downstream product formation," said Henry, who is now an analytical chemist for Heritage Research Group. "Some of these products might be toxic to the plants if the plants make too much of them. This is how the plants regulates their output."

Dudareva's lab is now working on methods to engineer plant metabolic pathways to increase terpenoid production.

"It might be that we have some compounds that aren't naturally in a plant, but we can put a gene of interest in them to create the <u>compounds</u> we want or significantly increase their production," Dudareva said.

More information: Laura K. Henry et al. Contribution of isopentenyl phosphate to plant terpenoid metabolism, *Nature Plants* (2018). <u>DOI:</u> <u>10.1038/s41477-018-0220-z</u>

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