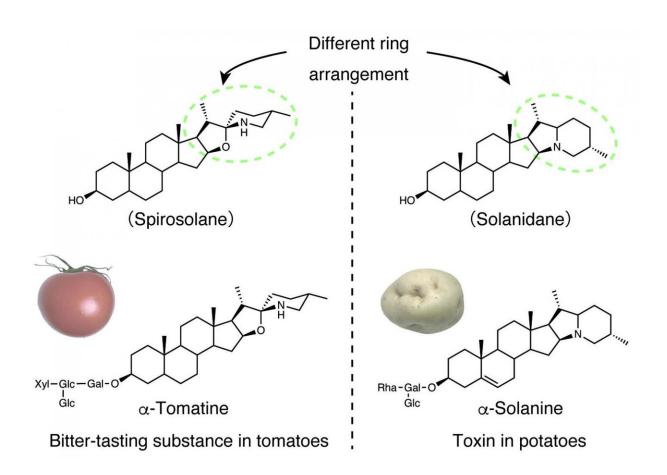


Toxin in potatoes evolved from a bittertasting compound in tomatoes

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The chemical structures of SGAs found in tomatoes and potatoes. Credit: Ryota Akiyama/Masaharu Mizutani

A multi-institutional collaboration has revealed that α -solanine, a toxic compound found in potato plants, is a divergent of the bitter-tasting α -



tomatine, which is found in tomato plants. The research group included Associate Professor Mizutani Masaharu and Researcher Akiyama Ryota et al. of Kobe University's Graduate School of Agricultural Science, Assistant Professor Watanabe Bunta of Kyoto University's Institute for Chemical Research, Senior Research Scientist Umemoto Naoyuki of the RIKEN Center for Sustainable Resource Science, and Professor Muranaka Toshiya of Osaka University's Graduate School of Engineering.

It is hoped that these research results can be used in <u>potato breeding</u> as a basis for suppressing the synthetization of poisonous compounds.

These research results were published in the international academic journal *Nature Communications* on February 26.

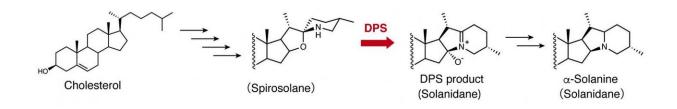
Main Points

- α -solanine is a toxic steroidal glycoalkaloid (SGA) found in potatoes.
- Tomato's α -tomatine is astringent-tasting SGA that accumulates inside unripe fruits.
- Based on their chemical structures, SGAs can be divided into two general classes, solanidanes e.g.α-solanine and spirosolanes, e.g. α-tomatine.
- The research group revealed that the toxic α -solanine in potatoes is biosynthesized from spirosolane.
- They discovered that the dioxygenase DPS is the key enzyme for this catalytic conversion.
- It was also revealed that the α -solanine biosynthesis pathway in potatoes diverged from the spirosolane biosynthesis pathway due to the evolution of DPS.



Research Background

 α -solanine is a type of toxic steroidal glycoalkaloid (SGA), which accumulates in the green skin on <u>potato</u> tubers and in tuber sprouts. SGA is not only found in potatoes but also in other plants of the Solanaceae family, including crops like tomatoes and eggplants. These substances are poisonous to many living things and serve as one of the plants' natural defenses. Low concentrations of SGA in potatoes cause a bitter taste and larger amounts can cause food poisoning. For this reason, biosynthesis research has been conducted with the aim of controlling the accumulation of SGA in potatoes.



The research group discovered that the enzyme DPS is a catalyst for this reaction (indicated by the red arrow). Credit: Ryota Akiyama/Masaharu Mizutani

Based on their skeletal chemical structures, SGAs can be divided into two general classes, solanidanes and spirosolanes. The potato toxin α solanine is an example of a solanidane, whereas α -tomatine, which accumulates inside unripe tomatoes, is a spirosolane. It is known that both classes of SGA are biosynthesized from cholesterol. Up until now, several genes that encode the catalytic enzymes in SGA biosynthesis have been discovered and potato and <u>tomato plants</u> share these enzymes in the common pathway of SGA biosynthesis. However, the steps and enzymes involved in the metabolic branch point between solanidane-



skeleton and spirosolane-skeleton formation remains an unsolved mystery.

This research group showed that the potato $toxin\alpha$ -solanine is biosynthesized from spirosolane. In a world first, they discovered that the dioxygenase DPS is the key to this conversion.

Research Findings

Potatoes contain the toxic solanidanes α -solanine and α -chaconine. The research group investigated the α -solanine biosynthesis pathway in <u>potato</u> plants. Using genome editing, they disrupted the biosynthetic enzyme gene in potato so that it was unable to produce α -solanine. Feeding α -tomatine (a spirosolane found in tomatoes) to the disruptant resulted in a metabolic conversion to the corresponding solanidane compound. In addition, it was found that this metabolic alteration could be suppressed with a 2-oxoglutarate dependent dioxygenase inhibitor, revealing that a dioxygenase is responsible for the oxidation reaction that synthesizes solanidanes from spirosolanes.

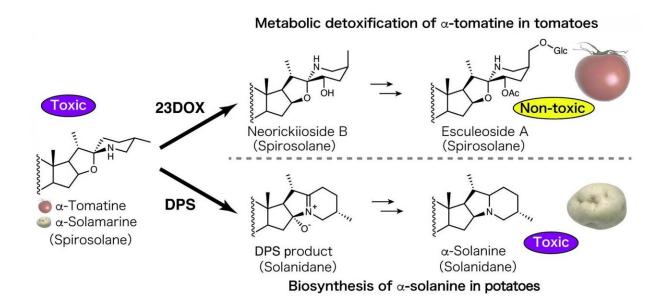
The researchers singled out a 2-oxoglutarate dependent dioxygenase (DPS) gene that was expressed in potato during α -solanine synthesis. To investigate this further, the researchers generated modified plants in which DPS gene expression was suppressed via RNA interference. The solanidane concentrations in these modified potato plants were far lower than in the unmodified group, and spirosolanes accumulated inside the plants in place of solanidanes. Next, the researchers measured the enzymatic activity of DPS by recombining the proteins and expressing them in E. coli. The results revealed the unique catalytic role of DPS in spirosolane's conversion into solanidane. This proved that DPS is the key enzyme responsible for this conversion.

This research revealed that the potato's ability to produce α -solanine



came about due to the evolution of DPS, which is responsible for metabolically converting spirosolanes (e.g. α -tomatine) into solanidanes. It is known that tomatoes also have an enzyme for metabolizing spirosolanes. The bitter-tasting α -tomatine is found in unripe tomatoes but is metabolized into the tasteless, non-toxic esculeoside A as the fruits ripen. The catalyst for this reaction is 23DOX, which is also a dioxygenase.

From the relationship between their chromosomal positions and phylogenetic analysis, it was revealed that the α -solanine biosynthase DPS has evolved from the same precursor gene as the non-toxic α -tomatine's catalytic enzyme 23DOX. Thus, it is believed that the evolution of the dioxygenase gene that metabolizes spirosolanes is one of the main drivers of the development of structural and functional variation in SGAs.



Metabolic reactions of spirosolane in tomatoes and potatoes. Credit: Ryota Akiyama/Masaharu Mizutani



Further Developments

Potatoes have been termed a potentially dangerous food because large concentrations of toxic α -solanine can cause food poisoning. It is hoped that these research results can provide a basis for future potato varieties in which the biosynthesis of toxic compounds is suppressed by targeting the DPS gene.

As shown in this research, the evolutionary origins of the structural diversity of SGAs provide clues towards discovering unknown SGA synthesis enzymes involved in biological functions in various plants. Illuminating these functions could pave the way for the molecular breeding of plant varieties that are able to adapt to different stressful environments.

More information: Ryota Akiyama et al, The biosynthetic pathway of potato solanidanes diverged from that of spirosolanes due to evolution of a dioxygenase, *Nature Communications* (2021). DOI: 10.1038/s41467-021-21546-0

Provided by Kobe University

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