

Failed antibiotic could be a game-changing weed killer for farmers

May 24 2023



The monomeric crystal structure of AtDHDPR1 (cyan, PDB ID: 7T34) aligned with chain C of the dimeric AtDHDPR2 crystal structure (green, PDB ID: 5UA0). The inset depicts the proximity of the conserved 'latch and catch' residues of AtDHDPR1 (blue) and AtDHDPR2 (pink), allowing for hydrogen bonding, which stabilizes the enzyme's closed conformation. Credit: *Communications Biology* (2023). DOI: 10.1038/s42003-023-04895-y

Weed killers of the future could soon be based on failed antibiotics.

A molecule that was initially developed to treat tuberculosis but failed to progress out of the lab as an antibiotic is now showing promise as a powerful foe for <u>weeds</u> that invade our gardens and cost farmers billions



of dollars each year.

While the failed antibiotic wasn't fit for its original purpose, scientists at the University of Adelaide discovered that by tweaking its structure, the molecule became effective at killing two of the most problematic weeds in Australia, <u>annual ryegrass</u> and wild radish, without harming bacterial and human cells. This research has been published in the journal *Communications Biology*.

"This discovery is a potential game changer for the agricultural industry. Many weeds are now resistant to the existing herbicides on the market, costing farmers billions of dollars each year," said lead researcher Dr. Tatiana Soares da Costa from the University of Adelaide's Waite Research Institute.

"Using failed antibiotics as herbicides provides a short-cut for faster development of new, more effective weed killers that target damaging and <u>invasive weeds</u> that farmers find hard to control."

Researchers at the University's Herbicide and Antibiotic Innovation Lab discovered there were similarities between bacterial superbugs and weeds at a molecular level.

They exploited these similarities, and by chemically modifying the structure of a failed antibiotic, they were able to block the production of amino acid lysine, which is essential for weed growth.





(From left) Emily Mackie, Dr Andrew Barrow and Dr Tatiana Soares da Costa. Credit: University of Adelaide

"There are no commercially available herbicides on the market that work in this way. In fact, in the past 40 years, there have been hardly any new herbicides with new mechanisms of action that have entered the market," said Dr. Andrew Barrow, a postdoctoral researcher in Dr. Soares da Costa's team at the University of Adelaide's Waite Research Institute.

It's estimated that weeds cost the Australian agriculture industry more than \$5 billion each year.



Annual ryegrass in particular is one of the most serious and costly weeds in southern Australia.

"The short-cut strategy saves valuable time and resources, and therefore could expedite the commercialization of much needed new herbicides," said Dr. Soares da Costa.

"It's also important to note that using failed <u>antibiotics</u> won't drive <u>antibiotic resistance</u> because the herbicidal <u>molecules</u> we discovered don't kill bacteria. They specifically target weeds, with no effects on <u>human cells</u>," she said.

It's not just farmers who could reap the benefits of this discovery. Researchers say it could also lead to the development of new weed killers to target pesky weeds growing in our backyards and driveways.

"Our re-purposing approach has the potential to discover herbicides with broad applications that can kill a variety of weeds," said Dr. Barrow.

More information: Emily R. R. Mackie et al, Repurposed inhibitor of bacterial dihydrodipicolinate reductase exhibits effective herbicidal activity, *Communications Biology* (2023). DOI: 10.1038/s42003-023-04895-y

Provided by University of Adelaide

Citation: Failed antibiotic could be a game-changing weed killer for farmers (2023, May 24) retrieved 27 June 2024 from <u>https://phys.org/news/2023-05-antibiotic-game-changing-weed-killer-farmers.html</u>

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