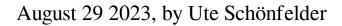
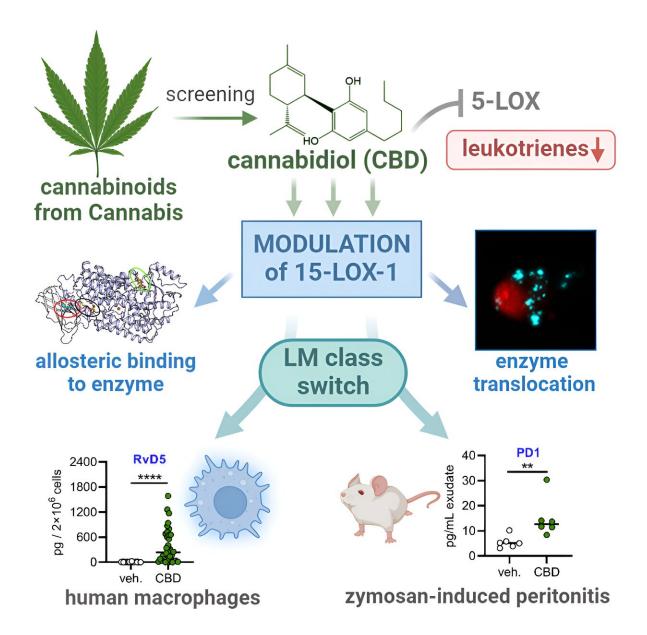


Research team clarifies mode of action of cannabinoids in inflammation







Graphical abstract. Credit: *Cell Chemical Biology* (2023). DOI: 10.1016/j.chembiol.2023.08.001

While the German government is planning to relax legislation on the use of cannabis, researchers from the Friedrich Schiller University Jena, together with colleagues from Italy, Austria and the U.S., have identified the mode of action underlying anti-inflammatory effects demonstrated by cannabinoids.

A few days ago, the <u>federal government</u> took the <u>controversial decision</u> to make the acquisition and possession of small amounts of cannabis exempt from punishment. Provided the German parliament approves the draft bill, the "Cannabis Act" will come into force next year. While some consider this move to be long overdue, others continue to warn strongly against the health risks of cannabis use.

The Jena researchers and their colleagues are now taking a different look at cannabis—at the traditional medicinal plant—with a study published in the journal *Cell Chemical Biology*. The team from the Institute of Pharmacy investigated how certain ingredients from the cannabis plant counteract inflammation. It was already known from previous studies that cannabis is not only an analgesic and an antispasmodic, but also has an anti-inflammatory effect.

"However, the reason for the anti-inflammatory effect was largely unclear until now," says Dr. Paul Mike Jordan, who led the study together with Prof. Oliver Werz.

The researchers studied how different <u>cannabinoids</u>, including the psychoactive THC (tetrahydrocannabinol) and CBD (cannabidiol), which is already found in freely available products today, act on human



immune cells. "We found that all eight cannabinoids we studied had antiinflammatory effects," says Lukas Peltner, doctoral student and first author of the study. "All the compounds we studied were found to inhibit the formation of pro-inflammatory messenger substances in cells while enhancing the formation of inflammation-resolving substances."

CBD induces a switch in immune cells

CBD in particular proved to be highly effective and the team investigated it in more detail with regard to its mode of action. The researchers were able to determine that CBD activates the 15-lipoxygenase-1 enzyme, which triggers the production of inflammation-resolving messenger substances that subsequently cause the inflammation to subside.

"CBD thus induces a switch in the affected cells, so to speak, which steers the <u>inflammatory process</u> from the promoting to the inhibiting side," explains Dr. Jordan. The researchers were also able to confirm these results, which were obtained in <u>cell cultures</u>, in animal experiments on mice.

In the long term, the insights gained could lead to <u>new therapeutic</u> <u>strategies</u> for treating <u>inflammatory diseases</u>, the researchers conclude. The focus should be on CBD, which was the most effective cannabinoid in the study. Previously approved preparations with cannabinoids contain CBD, "but also the psychoactive THC, which can be associated with a variety of side effects," notes Dr. Jordan. Therapeutics containing only CBD would reduce this problem.

More information: Lukas K. Peltner et al, Cannabidiol acts as molecular switch in innate immune cells to promote the biosynthesis of inflammation-resolving lipid mediators, *Cell Chemical Biology* (2023). DOI: 10.1016/j.chembiol.2023.08.001



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