

## New isomer separation method a boon for research on protein oxidation

April 9 2020



Chemists at UC Santa Cruz reported a new method for separation of methionine sulfoxide diastereomers that opens up new opportunities for studying their roles in biological processes. Credit: Cover art by J. Raskatov

Proteins are made up of long chains of amino acids, and most proteins have one or more of the sulfur-containing amino acid methionine. Oxidation of the sulfur atom in methionine is an important biomolecular



reaction that can have a wide range of biological consequences depending on the context and the protein involved.

Interestingly, the sulfur <u>oxidation</u> of methionine can give rise to two different versions of the oxidized molecule (methionine sulfoxide) differing only in the 3-dimensional spatial arrangement of its atoms. The two versions are called stereoisomers (or, more precisely, diastereomers, which are stereoisomers that are not mirror images). Researchers studying the biological effects of sulfur oxidation would like to be able to separate the two stereoisomers, but this has been extremely difficult to do.

Now, chemists at UC Santa Cruz have reported a new method for separation of methionine sulfoxide diastereomers that opens up new opportunities for studying their roles in biological processes. In a paper published April 6 in *Chemistry, A European Journal*, they reported obtaining both stereoisomers in purities exceeding 99%.

"The field of methionine oxidation has been hampered for decades by a lack of robust access to these reagents, and we believe this will be a tremendous boost," said first author Jevgenij Raskatov, assistant professor of chemistry and biochemistry at UC Santa Cruz.

Raskatov's lab collaborated with researchers at the California Institute of Technology on the paper, which was featured on the cover of the journal. Raskatov said he has already heard from other researchers in the field who are interested in the new method.

Raskatov's team used an advanced chromatography technique called supercritical fluid chromatography to purify the <u>methionine</u> sulfoxide stereoisomers. Supercritical fluids have the properties of both liquids and gases, which can be very useful in chromatography. The researchers also analyzed the structures of the purified stereoisomers using X-ray



crystallography and confirmed their remarkable stereochemical stability.

In addition to Raskatov, the coauthors of the paper include Scott Virgil and Lawrence Henling at Caltech and Hsiau-Wei Lee, Ka Chan, Ariel Kuhn, and Alejandro Foley at UC Santa Cruz.

**More information:** Jevgenij A. Raskatov et al, A Facile Method for the Separation of Methionine Sulfoxide Diastereomers, Structural Assignment, and DFT Analysis, *Chemistry – A European Journal* (2019). DOI: 10.1002/chem.201904848

## Provided by University of California - Santa Cruz

Citation: New isomer separation method a boon for research on protein oxidation (2020, April 9) retrieved 24 May 2024 from <u>https://phys.org/news/2020-04-isomer-method-boon-protein-oxidation.html</u>

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