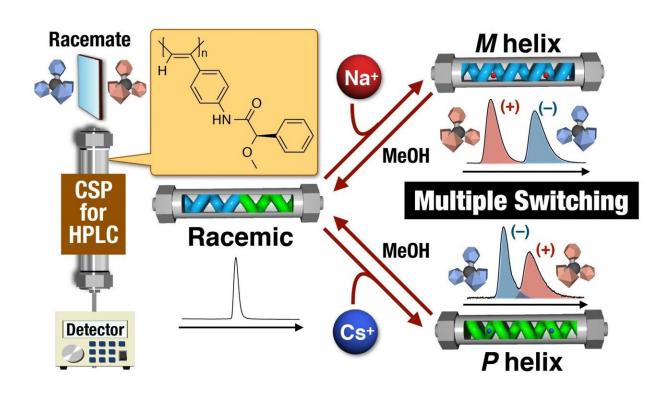


Multi-state switchable stationary phase opens new doors in chiral separation

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Schematic illustration of three-state switchable chiral stationary phase based on macromolecular helicity modulation in a poly(phenylacetylene) derivative using metal cations in the column. Credit: Kanazawa University

The concept of chirality can be challenging for the layperson, with "chemical handedness" seeming a very minor distinction. However, as the consequences of the notorious thalidomide disaster illustrate, understanding chiral materials is a major concern. The continued



development of chiral separation techniques therefore remains a key research area. A team involving researchers from Kanazawa University has reported a three-state switchable chiral stationary phase (CSP) that provides new opportunities in chiral separation. Their findings are published in the *Journal of the American Chemical Society*.

Chiral high performance liquid chromatography (HPLC) remains the most effective method for separating chiral molecules. HPLC involves running samples through a tube—known as a column—containing chiral material (the CSP) that can differentiate between pairs of chiral molecules (enantiomers). However, owing to the numerous enantiomers that cannot be separated using currently available CSPs, research in this area remains ongoing.

The researchers report a CSP based on a helical polymer material containing a chiral pendant group that causes the polymer to adopt different conformations in response to metal ions. When Na^+ ions are present the polymer is forced to adopt a left-handed helix so that the ions can interact with the aromatic component of the group. In contrast, when Cs^+ ions are present the polymer is forced into a right-handed helix to facilitate bonding between the ions and the two oxygen atoms of the group. In the absence of ions the structure is a deactivated mixture of the two helices.

"Our experiments used a CSP based on a poly(phenylacetylene) derivative that can be altered using an achiral external stimulus," study lead author Daisuke Hirose explains. "This allowed us to control the chirality of the column, and hence the retention of enantiomers, simply by introducing metal salts. The <u>metal ions</u> caused the <u>polymer</u> switch to a particular chiral conformation; however, the effect could be reversed by eluting the column with methanol."





An illustration inspired by three-state switchable chiral stationary phase using metal cations. Credit: Kanazawa University

The stability of the ion-induced states was demonstrated over 4 days of



continuous flow through the column, and the separation performance was exhibited by switching between the active and deactivated state numerous times.

"As far as we are aware, our system is the first reported example of a CSP that can be switched between three different recognition states using a stimulus that is not chiral," study corresponding author Katsuhiro Maeda explains. "We hope to build on our findings to extend the range of <u>enantiomers</u> that can be separated, which we believe will benefit numerous research areas such as drug discovery."

More information: Daisuke Hirose et al, Three-State Switchable Chiral Stationary Phase Based on Helicity Control of an Optically Active Poly(phenylacetylene) Derivative by Using Metal Cations in the Solid State, *Journal of the American Chemical Society* (2019). DOI: 10.1021/jacs.9b03177

Provided by Kanazawa University

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