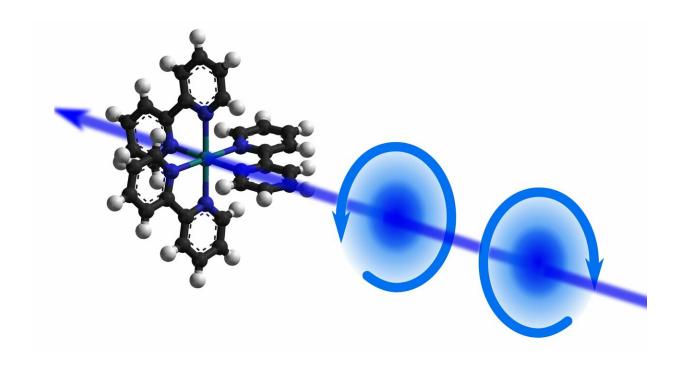


Chirality in real time

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An illustration of chirality in a molecule. Credit: M. Oppermann, EPFL

Distinguishing between left-handed and right-handed (chiral) molecules is crucial in chemistry and the life sciences, and is commonly achieved using a method called circular dichroism. However, during biochemical reactions, the chiral character of molecules may change. EPFL scientists have now developed a method that uses ultrashort, deep-ultraviolet pulses to accurately probe such changes in real-time in biomolecular systems.



In nature, certain molecules with the same <u>chemical composition</u> can exist in two mirrored configurations, much like human hands. This property is known as "chirality," and molecules with different chirality are called enantiomers. Enantiomers can exhibit entirely different chemical or biological properties, and separating them is a major issue in drug development and in medicine.

The method commonly used to detect enantiomers is circular dichroism (CD) spectroscopy. It exploits the fact that light polarized into a circular wave (like a whirlpool) is absorbed differently by left-handed and right-handed enantiomers. Steady-state CD spectroscopy is a major structural tool in (bio)chemical analysis.

While functioning, biomolecules undergo structural changes that affect their chiral properties. Probing these in real-time (i.e. between one picosecond and one nanosecond) provides a view of their biological function, but this has been challenging in the deep-UV spectrum (wavelengths below 300 nm) where most biologically relevant molecules such as <u>amino acids</u>, DNA and peptide helices absorb light.

The limitations are due to the lack of adequate sources of pulsed light and of sensitive detection schemes. But now, the group of Majed Chergui at the Lausanne Centre for Ultrafast Science (EPFL) has developed a setup to visualize the chiral response of (bio)molecules by CD spectroscopy with a resolution of 0.5 picoseconds.

The setup uses a photoelastic modulator, which is an optical device that can control the polarization of light. In this system, the modulator permits shot-to-shot polarization switching of a 20 kHz femtosecond pulse train in the deep-UV range (250-370 nm). It is then possible to record changes in the <u>chirality</u> of <u>molecules</u> at variable time-delays after they are excited with a short laser pulse.



"Amino acid residues and DNA bases absorb light below 300 nm," says Malte Oppermann, the paper's first author. "This set-up is the first to cover this region, and we successfully tested it on a model molecular system. Our next aim is to move on to larger biosystems, like DNA oligomers."

More information: Malte Oppermann et al, Ultrafast broadband circular dichroism in the deep ultraviolet, *Optica* (2019). <u>DOI:</u> 10.1364/OPTICA.6.000056

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