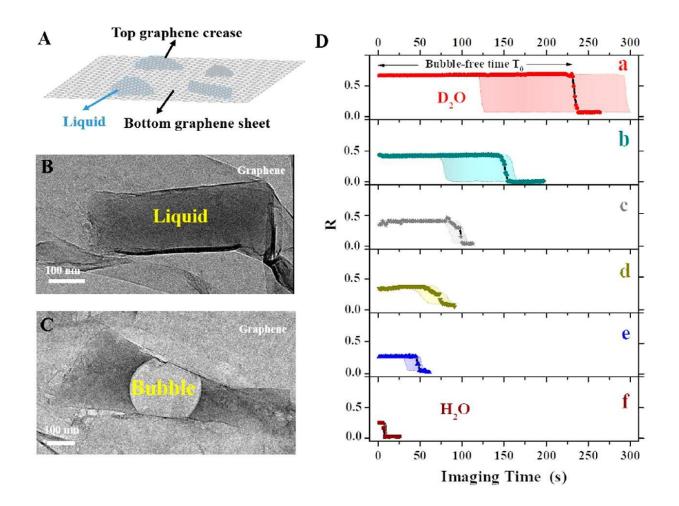


## **Sunscreen for dancing molecules**

## August 1 2018



(A) Graphene pocket protects the sample contained in the liquid. (B) Photo of the graphene pocket. (C) As a result of the incident electron beam, an air bubble fills most of the pocket after a short period of time. (D) Bubble forming time, depending on the type of solution filling the pocket. The longest bubble-free time is obtained when the pocket is filled with a) heavy water (D2O), followed by water (H2O) with b) 50% glycerol, c) 100 mM NaCl, d) 10 mM n-propyl gallate, e) 3% glycerol, and f) water only. Credit: Institute for Basic Science



Since life is mostly based on water, our molecules are moving, vibrating and somersaulting in a liquid environment. But electron microscopy—a technique to study a static version of this nanoworld—has been almost impossible to use to see moving molecules, because the incident electron beam damages the samples. Scientists at the Center for Soft and Living Matter, within the Institute for Basic Science (IBS), report a major improvement in this area.

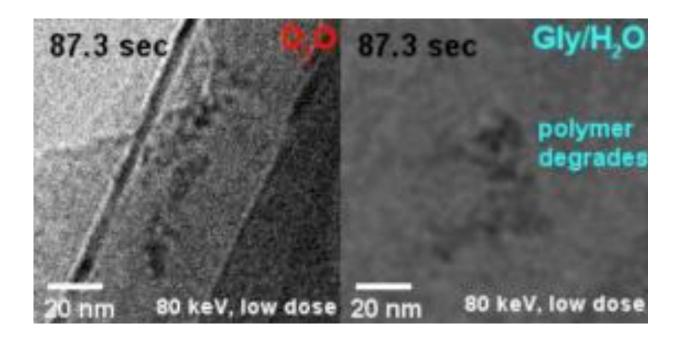
This study, published in *ACS Nano*, is the first to use heavy water  $(D_2O)$ —a form of water that contains <u>deuterium</u> (D) instead of hydrogen—in the field of transmission <u>electron microscopy</u> (TEM). This approach significantly delays sample damage, which is one of the major impediments for broader application of liquid-phase TEM to fragile biological samples.

In electron microscopy, electrons emitted against the sample have a much shorter wavelength than light, so they are better suited to provide information about single molecules. On the other hand, the electron beam is extremely powerful and risks damaging the specimen due to its high energy, which generates an electric charge and breaks the chemical bonds.

IBS researchers used a tiny pocket filled with liquid sandwiched between atomically-thin sheets of graphene, within which the sample molecules are free to move and are protected from electrical charging, and tested several types of liquids to find the one that preserves the sample longer. "In contrast to the common approach of reducing the energy of the electron beam to delay sample damage, we focused on tuning the environment—the water in which the molecules of interest are dissolved," says Huan Wang, co-author of the study.



IBS scientists have shown that use of <u>heavy water</u> has several advantages over competing methods.  $D_2O$  most effectively delays not only the formation of gas bubbles, but also structural damage of individual polymer molecules. Compared to  $H_2O$ ,  $D_2O$  has one more neutron, which means that it is heavier, thus more difficult to dissociate into radicals, and less reactive in the subsequent damaging process.



When exposed to the electron beam, polymer molecules (polystyrene sulfonate) are degraded more rapidly in the glycerol solution than in D2O. Credit: Institute for Basic Science

"Heavy water outperforms the competing methods by a factor of two to five at least," said Kandula Hima Nagamanasa, co-author of the study. "Since bubble formation is delayed and the molecules were visible for twice as long."



An equally important advantage is that  $D^2O$  is a harmless sunscreen. The sample, a polymer of polystyrene sulfonate in this case, showed the same pattern of dynamics and similar contrast in  $D_2O$  and in water.

"In the future, we plan to extend this study to more complex macromolecules, like DNA and proteins," explained Steve Granick, director of the IBS center and corresponding author of the study. "Moreover, the study opens avenues to observe long-term phenomena in other related microscopy techniques, like cryoEM (cryogenic electron microscopy), and to get more statistical information about complex phenomena, like self-assembly of single molecules into more complex biological structures."

**More information:** Huan Wang et al. Longer-Lasting Electron-Based Microscopy of Single Molecules in Aqueous Medium, *ACS Nano* (2018). DOI: 10.1021/acsnano.8b04190

Provided by Institute for Basic Science

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