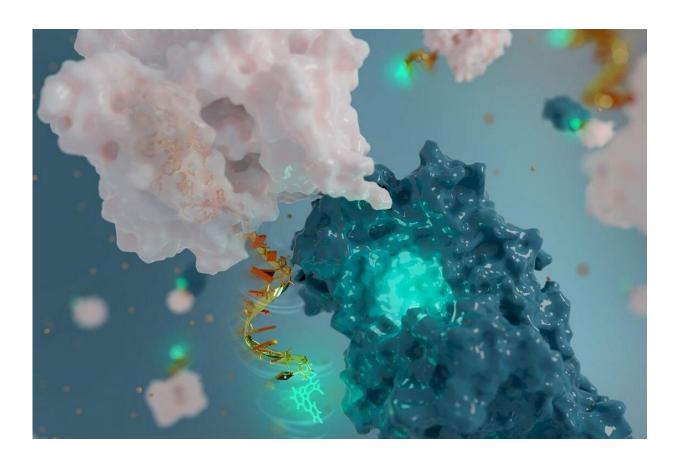


Chemists use DNA to build the world's tiniest antenna

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Like a two-way radio that can both receive and transmit radio waves, the fluorescent nanoantenna designed by Alexis Vallée-Bélisle and his team receives light in one colour and depending on the protein movement it senses, then transmits light back in another colour, which we can detect. One of the main innovations of these nanoantennas is that the receiver part of the antenna (bright green) is also employed to sense the molecular surface of the protein studied via molecular interaction. Credit: Caitlin Monney



Researchers at Université de Montréal have created a nanoantenna to monitor the motions of proteins. Reported this week in *Nature Methods*, the device is a new method to monitor the structural change of proteins over time—and may go a long way to helping scientists better understand natural and human-designed nanotechnologies.

"The results are so exciting that we are currently working on setting up a start-up company to commercialize and make this nanoantenna available to most researchers and the pharmaceutical industry," said UdeM chemistry professor Alexis Vallée-Bélisle, the study's senior author.

An antenna that works like a two-way radio

Over 40 years ago, researchers invented the first DNA synthesizer to create molecules that encode genetic information. "In recent years, chemists have realized that DNA can also be employed to build a variety of nanostructures and nanomachines," added the researcher, who also holds the Canada Research Chair in Bioengineering and Bionanotechnology.

"Inspired by the 'Lego-like' properties of DNA, with <u>building blocks</u> that are typically 20,000 times smaller than a <u>human hair</u>, we have created a DNA-based fluorescent nanoantenna, that can help characterize the function of proteins," he said.

"Like a two-way radio that can both receive and transmit <u>radio waves</u>, the fluorescent nanoantenna receives light in one color, or wavelength, and depending on the <u>protein</u> movement it senses, then transmits light back in another color, which we can detect."

One of the main innovations of these nanoantennae is that the receiver part of the antenna is also employed to sense the molecular surface of the protein studied via molecular interaction.



One of the main advantages of using DNA to engineer these nanoantennas is that DNA chemistry is relatively simple and programmable," said Scott Harroun, an UdeM doctoral student in chemistry and the study's first author.

"The DNA-based nanoantennas can be synthesized with different lengths and flexibilities to optimize their function," he said. "One can easily attach a fluorescent molecule to the DNA, and then attach this fluorescent nanoantenna to a biological nanomachine, such as an enzyme.

"By carefully tuning the <u>nanoantenna</u> design, we have created five nanometer-long antenna that produces a distinct signal when the protein is performing its <u>biological function</u>."

Fluorescent nanoantennas open many exciting avenues in biochemistry and nanotechnology, the scientists believe.

"For example, we were able to detect, in real time and for the first time, the function of the enzyme alkaline phosphatase with a variety of biological molecules and drugs," said Harroun. "This enzyme has been implicated in many diseases, including various cancers and intestinal inflammation.

"In addition to helping us understand how natural nanomachines function or malfunction, consequently leading to disease, this new method can also help chemists identify promising new drugs as well as guide nanoengineers to develop improved nanomachines," added Dominic Lauzon, a co-author of the study doing his Ph.D. in chemistry at UdeM.

One main advance enabled by these nanoantennas is also their ease-ofuse, the scientists said.



"Perhaps what we are most excited by is the realization that many labs around the world, equipped with a conventional spectrofluorometer, could readily employ these nanoantennas to study their favorite protein, such as to identify new drugs or to develop new nanotechnologies," said Vallée-Bélisle.

More information: Scott G. Harroun et al, Monitoring protein conformational changes using fluorescent nanoantennas, *Nature Methods* (2021). DOI: 10.1038/s41592-021-01355-5

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