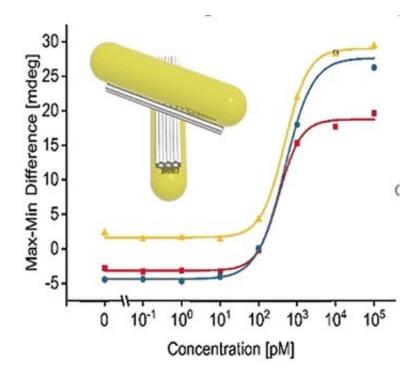


Optical detection of picomolar concentrations of RNA using switches in plasmonic chirality

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Even tiny amounts of viruses can have disastrous consequences. RNA identification can reveal the type of virus present. A fast and sensitive technique based on optical detection has now been outlined in the journal *Angewandte Chemie*. Scientists from Germany and Finland have demonstrated the binding of an RNA target to a probe made of gold nanorods and a DNA origami structure. Chirality switches triggered by binding can be measured by circular dichroism spectroscopy.



Identifying the pathogen—often a virus—that is troubling a patient is among the biggest challenges in healthcare. Viruses responsible for Zika fever, AIDS, and hepatitis C contain mutating RNA sequences. Physicians need to know quickly which type of virus their patients have acquired, but current techniques based on multiplying RNA are costly and time-consuming. Now, Tim Liedl from Ludwigs-Maximilians-Universität in Munich, Germany, and his colleagues, have developed a fast detection strategy based on nanoplasmonics, DNA origami, and an optical readout.

Light can induce plasmonic waves in nanosized metal structures smaller than the wavelength of the incident light. This resonance may lead to strongly enhanced light emission even from nanoscopic structures—a feature that is highly interesting for biosensing applications. Liedl and colleagues have created a nanosized sensing probe for RNA molecules.

The probe, a nanosized apparatus made of DNA and <u>gold nanorods</u>, was assembled by the so-called DNA origami technique, which exploits the specific interactions of the DNA bases to fold and glue together single strands in any desired form. The authors constructed two bars of parallel DNA helices loosely connected through a hinge in the middle of the bars. Gold nanorods were placed on top of each of the crossed bars. Both crossing arms were supplied with functionality at their ends: the scientists attached one single DNA sequence complemented with a blocking strand to one arm, and the complementing DNA sequence to the other. In the presence of target RNA, which could be a typical viral RNA sequence, the blocking strand would leave its DNA in favor of RNA hybridization, and both single DNA sequences would complementarily form a double strand whereby the two arms of the cross are pulled together. This structural change introduces chirality to the probe.

Chirality can be detected with circular dichroism. And indeed, the



structural changes triggered by the RNA binding induced a circular dichroism signal detectable with a CD spectrometer. Concentrations as low as 100 picomolar of the target RNA were recognized, according to the authors. The scientists hope to establish this technique in lab-on-achip systems where few steps are required for sample preparation and low-cost miniature devices lead to sensitive results. Preliminary results on serum from blood with added viral RNA were promising.

The authors admit that the detection limits are still not low enough to be clinically relevant. However, they believe improvements should be possible; including, better protection of the nanosensors from serum proteins, a change to better resonating plasmonic metals, and expansion of RNA recognition sites. This could make the technique a promising diagnostic tool that is not necessarily limited to viral RNA.

More information: Timon Funck et al, Sensing Picomolar Concentrations of RNA Using Switchable Plasmonic Chirality, *Angewandte Chemie International Edition* (2018). DOI: 10.1002/anie.201807029

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