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Home > Chemistry > Analytical Chemistry > February 20, 2018

## Microanalysis of biological samples for early disease detection

February 20, 2018, Osaka University

Fig.1. A schematic drawing of solution measurement by using fabricated terahertz microfluidic chip. The chip consists of a local THz radiation point source, a single microchannel and a few arrays of split ring resonators. The THz waves are ... more ▾

The use of terahertz (THz) waves for biosensing is currently receiving considerable attention. THz waves are able to detect molecular vibrations and rotations, without using labels that can affect the properties of the substances of interest.

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However, until now, the diffraction limit of THz waves and their strong absorption by water have constrained this technique. Microfluidic devices are also promising analytical systems because of the low sample volumes needed for sample measurement. A group of researchers from Osaka University has now developed a nonlinear optical crystal (NLOC) chip, which combines THz waves with a microfluidic device, utilizing the close proximity of the THz wave source and the solution of interest in a microchannel. Their work was published in *APL Photonics*.

"Using our technique, we have been able to detect solution concentrations of several femtomoles in volumes of less than a nanoliter," corresponding author Masayoshi Tonouchi says. "Such high-sensitivity detection without the need for labelling moieties has great potential for future low-invasivity clinical techniques."

Early and rapid detection of a number of common diseases is expected to be one of the major applications of the technique. Cancer, diabetes, and the influenza virus could potentially be detected with only very small volumes of bodily fluid, reducing the pain and discomfort of numerous exploratory procedures for patients. In addition, the technique allows living cells to be analyzed in a non-destructive way, which has numerous potential benefits in research.

Mineral Concentration (mg/L)	Frequency Shift (THz)
10	~0.004
40	~0.007
200	~0.014
600	~0.015
0	~0.005
50	~0.006
100	~0.009
150	~0.010

Fig.2. Plots of the resonance frequency shift with respect to the mineral concentration in the actual amount of 318 picoliters of mineral water. By observing the amount of shift from the resonance frequency of pure water, it is found that ... more ▾

The developed NLOC chip is able to locally generate the THz radiation in close proximity to the single microchannel device, improving efficiency. The sensor chip was used to analyze mineral concentrations by comparing frequency shifts resulting from the presence of ions to those of pure water. Using this technique they determined a sensitivity of 31.8 femtomoles.

"Achieving high sensitivity without the need for a high-power optical or THz source, near-field probes or prisms, opens up a number of possibilities," lead author Kazunori Serita says. "We are very excited about the potential of our findings to lead to rapid detection and compact device design. In particular, we see our results accelerating the development of THz lab-on-a-chip devices."

This highly adaptable technology is likely to ripple out into many areas of analytical and biochemistry, as well as cell biology, and clinical medicine.

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More information: Kazunori Serita et al. Invited Article: Terahertz microfluidic chips sensitivity-enhanced with a few arrays of meta-atoms. *APL Photonics* (2018). DOI: 10.1063/1.5007681

Provided by: Osaka University

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