

'Instant acid' method offers new insight into nanoparticle dispersal in the environment and the body

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Successive test runs at NIST show how clumping of typical nanoparticles in a solution depends on changes in acidity. Time after acidity jump is shown on the horizontal axis, while the vertical axis is a measure of the size of the nanoparticle aggregates. As pH goes down (and acidity up), both rate of aggregation and size of clumps of nanoparticles goes up. Credit: R. Murphy/NIST

Using a chemical trick that allows them to change the acidity of a solution almost instantly, a team at the National Institute of Standards and Technology has demonstrated a simple and effective technique for quantifying how the stability of nanoparticle solutions change when the acidity of their environment suddenly changes. The measurement method and the problem studied are part of a broader effort at NIST to understand the environmental, health and safety implications of nanoparticles.

Any change in nanoparticle solubility with local acidity (pH) ultimately affects how they are distributed in the environment as well as their



potential for uptake into organisms. This is crucial when designing nanoparticles for use in medicine, explains NIST chemical engineer Vivek Prabhu. "Cells in the body are very compartmentalized. There are places within the cell that have vastly different pH. For instance, in the sea of the cell, the cytosol, pH is regulated to be about 7.2, which is slightly basic. But within the <u>lysosome</u>, which is where things go to get broken down, the pH is about 4.5, so it's very acidic."

Nanoparticles designed for use in drug therapy or as contrast agents for medical imaging typically are coated with molecules to prevent the <u>particles</u> from clumping together, which would reduce their effectiveness. But the efficacy of the anti-clumping coating often depends on the pH of the environment. According to the NIST team, while it's relatively easy to put nanoparticles in a solution at a particular pH and to study the stability of the suspension over long times, it is difficult to tell what happens when the particles are suddenly exposed to a different level of acidity as often occurs in environmental and application contexts. How long does it take them to react to this change and how?

"Our idea borrows some of the materials used in <u>photolithography</u> to make microcircuits," says Prabhu. "There are molecules that become acids when you shine a light on them—photo acid generators. So instead of manually pouring acid into a solution and stirring it around, you start with a solution in which these molecules already are mixed and dissolved. Once you shine light on it ...bam! Photolysis occurs and it becomes acidic." The acidity of the solution can be made to jump a major step—an amount chosen by the experimenter—without needing to wait for mixing or disturbing the solution. "It gives you a way to probe the nanoparticle solution dynamics at much shorter timescales than before," says Prabhu.

Using their "instant <u>acid</u>" technique and light scattering instruments to



monitor the aggregation of nanoparticles, the NIST team followed the growth of clusters of chemically stabilized latex nanoparticles for the first few seconds after inducing the pH transition with light. Their results demonstrate that under certain conditions, the stability of the nanoparticles—their tendency to resist clumping—becomes very sensitive to pH. Studies such as these could provide a stronger foundation to design nanoparticles for applications such as targeting tumor cells that have levels of acidity markedly different from normal cells.

More information: R.J. Murphy, D. Pristinski, K. Migler, J.F. Douglas and V.M. Prabhu. Dynamic light scattering investigations of nanoparticle aggregation following a light-induced pH jump. Journal of Chemical Physics. 132, 194903 (2010) <u>doi:10.1063/1.3425883</u>

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