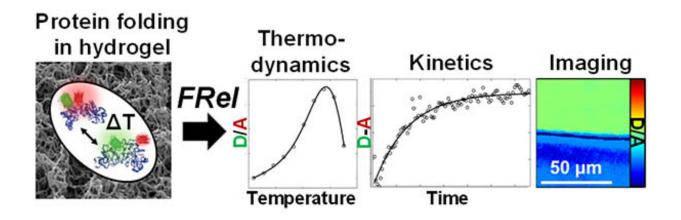


Investigating folding stability and dynamics of proteins

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Researchers at the Beckman Institute at the University of Illinois used Fast Relaxation Imaging (FReI) to investigate the folding stability and dynamics of proteins within polyacrylamide hydrogels. Credit: Lydia Kisley, Beckman Institute for Advanced Science and Technology

Hydrogels are polymer materials that can absorb a large amount of water, making them flexible like human tissue. They are used in a number of medical applications, including contact lenses, wound dressings, and facial reconstruction.

Hydrogels also can be used in drug delivery, for example, as coatings for



drugs. However, questions still remain about the efficacy of the drug delivery as it's not clear how proteins in your body interact with the hydrogels.

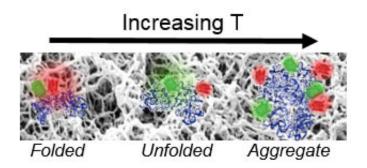
"If you put a hydrogel in your body, there are proteins in your body that can then interact with that material," said Lydia Kisley, a Beckman-Brown Interdisciplinary Postdoctoral Fellow. "Ideally you want proteins to maintain their function and their structure, so there's not any adverse effects."

Kisley, a chemist, and other researchers at the Beckman Institute for Advanced Science and Technology at the University of Illinois recently used Fast Relaxation Imaging (FReI) to investigate the folding stability and dynamics of proteins within polyacrylamide hydrogels. Their findings are published in "Direct Imaging of Protein Stability and Folding Kinetics in Hydrogels" in *ACS Applied Materials and Interfaces*.

"We're trying to understand how proteins behave in these material environments," Kisley said. "There are not many tools available to do that. This paper was the first demonstration of the new imaging technique to understand proteins in a material."

The group used a fluoresence microscope in the lab of Martin Greubele, a professor of chemistry and member of Beckman's Nanoelectronics and Nanomaterials Group. FReI detects protein unfolding in situ by imaging changes in fluorescence resonance energy transfer (FRET) after temperature jump perturbations.





Using Fast Relaxation Imaging (FReI), researchers at the Beckman Institute for Advanced Science and Technology at the University of Illinois investigated the folding stability and dynamics of proteins within polyacrylamide hydrogels. Credit: Lydia Kisley, Beckman Institute for Advanced Science and Technology

"The unique thing that we add here with the microscope is a <u>temperature</u> <u>change</u>. By using this very fast temperature change, we can tell how the protein responds to temperature and how stable it is. And how the confinement and chemistry of the hydrogel changes the protein stability as well," Kisley said.

The function of the protein comes from how it is folded, Kisley explained. In order to understand how proteins that make up some drugs function, it is crucial to understand how they fold.

The group determined that the <u>hydrogel</u> increases protein stability, speeds up folding relaxation, and promotes irreversible binding at the solution-gel interface.

"Although these <u>materials</u> are very compatible, since they have high water content, we found that at low temperatures when the protein is unfolded it's okay, but once the protein starts to unfold a little bit it will start to stick to the material, and may actually promote more proteins to aggregate and stick to it, so it suggests that the protein is destabilized a



bit," Kisley said.

"It was a more complicated system than we were expecting. I thought it would be simple, but that's pretty common in science where things end up being more complicated than you expect."

Kisley's fellowship at the Beckman Institute has allowed her to connect her doctoral research in diffusion and adsorption of proteins in hydrogels to <u>protein</u> folding in hydrogels and additional surface and polymer brush samples.

She plans to investigate if the behaviors we see in a larger 3D gel also happen on a smaller surface, nanoscale environment.

More information: Lydia Kisley et al, Direct Imaging of Protein Stability and Folding Kinetics in Hydrogels, *ACS Applied Materials & Interfaces* (2017). DOI: 10.1021/acsami.7b01371

Provided by Beckman Institute for Advanced Science and Technology

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