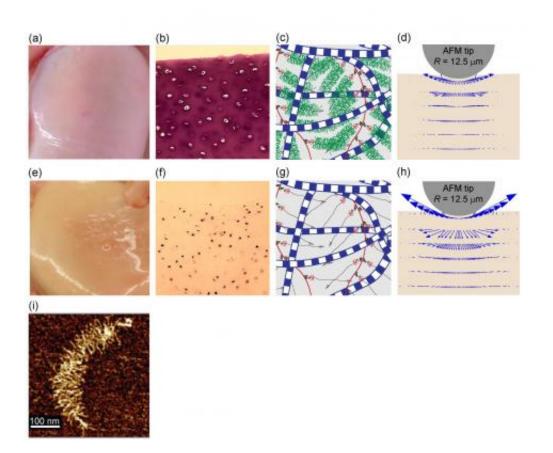


Cartilage damaged from exercise may aid in early osteoarthritis detection

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These are images of (a) normal human joint cartilage, and (e) cartilage with very early matrix GAG loss. (b) Histologic image of a normal cartilage stained with Toluidine Blue to visualize content of GAGs. (f) Histologic image of a GAGdepleted cartilage (c) Schematic of extracellular matrix of normal cartilage composed mainly of collagen fibers and aggrecan (g) Schematic of matrix of GAG-depleted cartilage (d,h) Schematic of AFM-based dynamic compression of normal (d) and GAG-depleted (h) cartilage, which results in intra-tissue fluid flow velocity depicted by the blue arrows (from Finite element model computer



simulations): loss of GAG (h) enables more fluid to flow out of the cartilage at high loading rates. (i) Human aggrecan imaged by Atomic Force Microscopy (AFM). Credit: *Biophysical Journal*, Nia et al.

Osteoarthritis is the most common joint disorder, affecting about onethird of older adults, and currently there is no cure. A study published by Cell Press April 2nd in the *Biophysical Journal* reveals how the nanoscale biomechanical properties of cartilage at joints change at the earliest stages of osteoarthritis, making the tissue more prone to damage during fast physical activities. The findings could improve early detection of the disease as well as tissue engineering strategies to repair damaged cartilage in patients.

"Our techniques enable detection of the earliest loss of mechanical function associated with daily activities involving high loading rates, such as running and jumping," says senior study author Alan Grodzinsky of the Massachusetts Institute of Technology. "The findings can also be used to evaluate replacement tissue to ensure that it can survive these daily activities."

Osteoarthritis is a painful condition marked by the deterioration of cartilage—firm, rubbery tissue that cushions bones and prevents them from rubbing together. At the earliest stages of the disease, cartilage loses molecules called glycosaminoglycans (GAGs), which reduces the ability of the tissue to resist impact caused by physical activity. But until now, little was known about how GAG loss affects the functioning of cartilage across a wide spectrum of activities, from walking to running and jumping.

To address this question, Grodzinsky and his team developed a new system to measure the biomechanical properties of cartilage in response



to cyclic compression forces that simulated a range of physical activities, each occurring at a different timescale. GAG-depleted cartilage was less capable of increasing its stiffness to deal with forces associated with high-rate activities such as running, when compared with normal tissue. Moreover, GAG loss resulted in a dramatic increase in the ability of fluids to flow out of cartilage, which is expected to diminish protection against impact caused by fast activities.

Together, the findings show how GAG depletion at the earliest disease stages could affect the <u>nanoscale</u> properties of cartilage, reducing the ability of this tissue to withstand high-rate activities. "We discovered that GAG-depleted tissue is most vulnerable to high rates of loading and not just the magnitude of the load," Grodzinsky says. "This finding suggests that people with early degradation of <u>cartilage</u>, even before such changes would be felt as pain, should be careful of dynamic activities such as running or jumping."

More information: *Biophysical Journal*, Nia et al.: "High-Bandwidth AFM-Based Rheology Reveals that Cartilage is Most Sensitive to High Loading Rates at Early Stages of Impairment." <u>dx.doi.org/10.1016/j.bpj.2013.02.048</u>

Provided by Cell Press

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