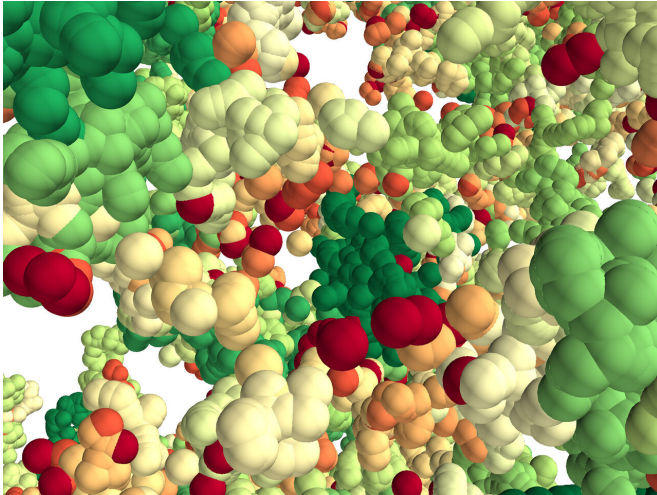


Chromatin organizes itself into 3-D 'forests' in single cells

10 January 2020



A 3D forest of chromatin predicted by the researchers' model. Tree domains are colored from red to green according to their sizes. Each particle represents 2 kilobases of DNA. Credit: Northwestern University

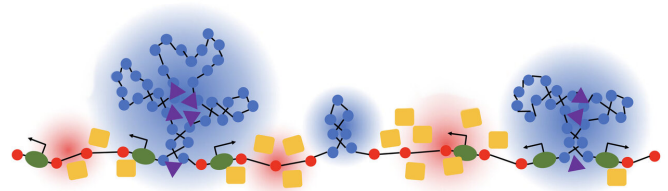
A single cell contains the genetic instructions for an entire organism. This genomic information is managed and processed by the complex machinery of chromatin—a mix of DNA and protein within chromosomes whose function and role in disease are of increasing interest to scientists.

A Northwestern University research team—using mathematical modeling and optical imaging they developed themselves—has discovered how [chromatin](#) folds at the [single-cell](#) level. The researchers found chromatin is folded into a variety of tree-like domains spaced along a chromatin backbone. These small and large areas are like a mixed forest of trees growing from the forest floor. The overall structure is a 3-D forest at microscale.

Chromatin is responsible for packing DNA into the cell nucleus. In humans, that's about six feet of DNA in each cell. The new work suggests that

chromatin is more structured and hierarchical in single [cells](#) than previously thought. Learning how chromatin correctly operates will help scientists understand what goes wrong with it in cancer and other diseases.

"By integrating theoretical and [experimental work](#), we have produced a new chromatin folding picture that helps us see how the 3-D genome is organized at the single-cell level," said Igal Szleifer, the Christina Enroth-Cugell Professor of Biomedical Engineering at Northwestern's McCormick School of Engineering. He co-led the research team with Vadim Backman.



Northwestern University researchers have discovered how chromatin folds at the single-cell level. They found that it folds into a variety of tree-like domains spaced along a chromatin backbone. Credit: Northwestern University

Details of the interdisciplinary study will be published Jan. 10 in the journal *Science Advances*.

"If genes are the hardware, chromatin is the software," said Backman, the Walter Dill Scott Professor of Biomedical Engineering and director of the Center for Physical Genomics and Engineering. "If the structure of chromatin changes, it can alter the processing of the information stored in the genome, but it does not alter the genes themselves. Understanding chromatin folding holds the key to understanding how cells differentiate and

how cancer happens."

Advances in genomic, imaging and information technologies are just beginning to enable scientists to better understand how chromatin works. The Northwestern researchers used a Partial Wave Spectroscopic (PWS) microscope, optical imaging developed by Backman and colleagues, to peer deep into live cells and "sense" alterations in chromatin packing. They also used electron imaging.

"Our paradigm-shifting picture of chromatin folding is an important missing piece in the holistic view of genomic structure," said Kai Huang, the study's first author. Huang is a postdoctoral fellow in Backman's research group. "The results should inspire new strategies to fight cancer."

More information: "Physical and data structure of 3D genome" *Science Advances* (2020).

advances.sciencemag.org/content/6/1/eaay4055

Provided by Northwestern University

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