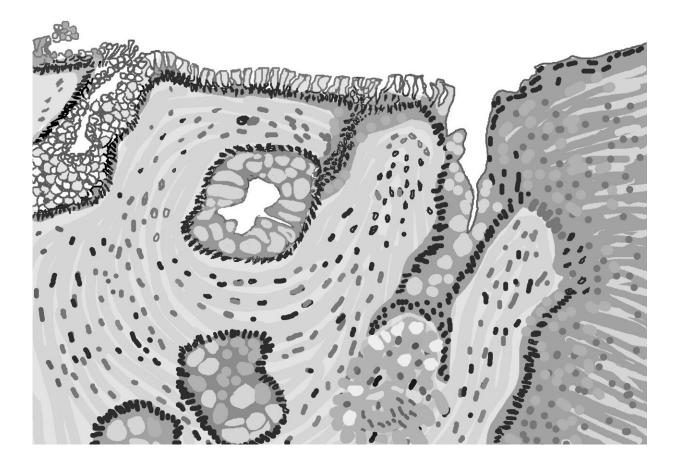


New digital tool to better monitor Barrett's esophagus

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Credit: Oak Ridge National Laboratory

A team including researchers from the Department of Energy's Oak Ridge National Laboratory has developed a digital tool to better monitor



a condition known as Barrett's esophagus, which affects more than 3 million people in the United States. Barrett's occurs when the mucosal lining of the lower esophagus deteriorates, altering its cellular structure, and is most common in those with chronic acid reflux.

Barrett's is considered a premalignant condition because it's a common precursor to esophageal cancer, so monitoring patients is critical. However, the current paradigm for tracking the progress of the disease, known as the Seattle protocol, is invasive, expensive and potentially ineffective.

Per the protocol, Barrett's patients must have their esophagus poked with forceps every few months, an uncomfortable experience that only captures a small percentage of the affected tissue, meaning that despite the invasive procedure problematic tissues can remain undetected.

A research <u>team</u> set out to test the effectiveness of the Seattle protocol by constructing a <u>computational model</u> to represent the human esophagus, using real-world data across a large population of tissues.

The team, which also included researchers from Columbia University's Irving Medical Center, Mount Sinai's Icahn School of Medicine and Massachusetts General Hospital's Institute for Technology Assessment, published their results in the journal *Simulation*.

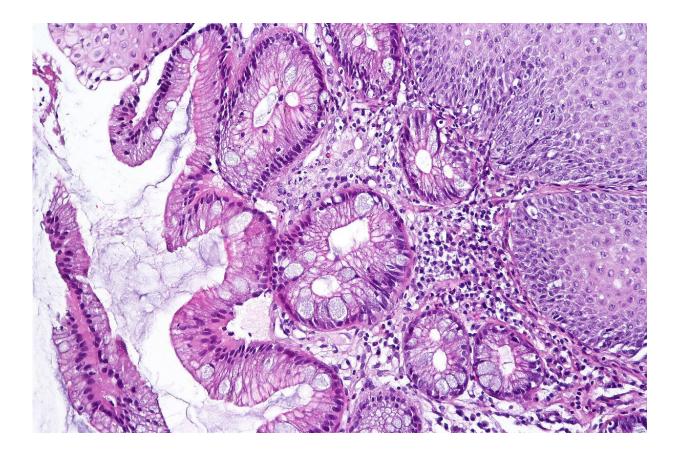
"The ultimate goal was to strike a balance between physically poking a patient and how often you find something that may be concerning," said Jim Nutaro, leader for computational systems engineering at ORNL and a researcher on the project. "It reproduces historical data, and cancers that surface in the simulations are analogous to the real world. These are virtual patients, and we can poke them as much as we want."

The team's end goal is to minimize the invasiveness of tracking the



condition and, by extension, reduce deaths from <u>esophageal cancer</u> by providing a testbed for potential future monitoring regimes.

To examine their digital patients, the researchers first needed to construct a digital esophagus via a computational model. The team drew input data for Barrett's onset and death ages from an actual 1960s cohort and validated the resulting model with real-world data such as detection rates and population statistics.



Biopsy from the tubular esophagus showing incomplete intestinal metaplasia, goblet cells with interposed cells having gastric foveolar-type mucin consistent with Barrett esophagus. Negative for dysplasia. H&E stain. Credit: Wikimedia Commons



"It's classic model construction and development," said Nutaro.

The model was then complemented with simulations of millions of various types of tissues constructed using real-world data, a computationally expensive process that was only possible via parallelization across multicore servers.

"The optimization routine is designed to work on parallel computing platforms and benefits from multicore parallelization to facilitate model calibration in a feasible amount of time," the researchers wrote.

It's a task well within ORNL's wheelhouse, as the laboratory is currently home to the nation's fastest computer, Summit, and hundreds of researchers such as Nutaro and his colleague, ORNL staff scientist and team member Ozgur Ozmen, who work daily to employ such resources in tackling the most complex scientific challenges of our time.

The team's model is in the <u>public domain</u>, available to those who can apply it to the physical treatment of Barrett's patients.

In the near term the researchers are looking to apply it to studying variations in the detection of Barrett's across different timescales and tissue areas. Ultimately, they would like to find partners in the medical community that could use their findings to develop a new protocol and, by extension, improve the lives and life expectancies of patients with Barrett's esophagus.

"We can share the <u>model</u> and work with them to further refine current practices," said Ozmen. "In other words, we can help to optimize future protocols."

More information: Ozgur Ozmen et al, Tissue scale agent-based simulation of premalignant progressions in Barrett's esophagus,



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