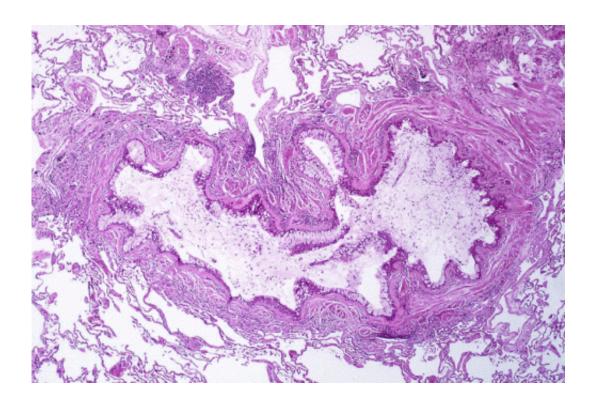


Genetic damage caused by asthma shows up in circulating blood stream, too

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Obstruction of the lumen of a bronchiole by mucoid exudate, goblet cell metaplasia, and epithelial basement membrane thickening in a person with asthma. Credit: Yale Rosen/Wikipedia/CC BY-SA 2.0

Asthma may be more harmful than was previously thought, according to UCLA researchers who found that genetic damage is present in circulating, or peripheral, blood. Doctors previously thought that the genetic damage it caused was limited to the lungs.



In the study, researchers looked for the overexpression of a cytokine called interleukin 13 (IL-13), which is known to mediate inflammation, a critical problem for people with asthma.

The study, which was conducted in an animal model that mimicked human asthma, was the first to assess the role of IL-13 in genetic damage to cells, or genotoxicity, said its senior author, Robert Schiestl, a professor of pathology and radiation oncology at the David Geffen School of Medicine at UCLA.

"Asthma is a very widespread disease, and we show for the first time an association between asthma and genotoxicity in peripheral blood," said Schiestl, who also is a professor of <u>environmental health sciences</u> at the Fielding School of Public Health at UCLA. "This is important because it shows a whole-body effect from asthma, not just damage in the lungs."

The findings were published today in the peer-reviewed journal *Mutation Research/Fundamental and Molecular Mechanisms of Mutagenesis*.

Schiestl said it appears that IL-13 increases important elements of the inflammatory response, including reactive oxygen species molecules—ions or very small molecules that include free radicals. His research team found that ROS-derived oxidative stress induced genetic damage with four types of systemic effects in the peripheral blood:

- Oxidative DNA damage.
- Single and double DNA strand breaks.
- Micronucleus formation.
- Protein damage.

Schiestl said all four effects causes the chromosomes to become unstable, which could result in a variety of other diseases.



"We found four different markers of DNA damage and one marker of protein damage in <u>blood cells</u> in the body periphery, which was very surprising," Schiestl said. "This could indicate that other organs in asthmatics have a higher risk of developing disease."



Robert Schiestl, University of California - Los Angeles Health Sciences. Credit: UCLA

Schiestl and his team will next attempt to use chemicals that help repair the DNA of damaged cells. Their goal is to determine whether doing so can make asthma less damaging by reducing genetic instability in the peripheral blood supply.

Asthma, a chronic disease that inflames and narrows the airways of the lungs, affects more than 150 million individuals worldwide. It causes recurring periods of wheezing, chest tightness, shortness of breath and



coughing, and the symptoms can worsen at any time, making breathing difficult. There is no cure.

Asthma affects people of all ages, but it most often starts during childhood. More than 25 million Americans are known to have the disease, including about 7 million children. There are two types of asthma, allergic and non-allergic asthma, which account for 70 and 30 percent of cases respectively. Non-allergic asthmatics experience more severe and more frequent symptoms.

Provided by University of California, Los Angeles

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