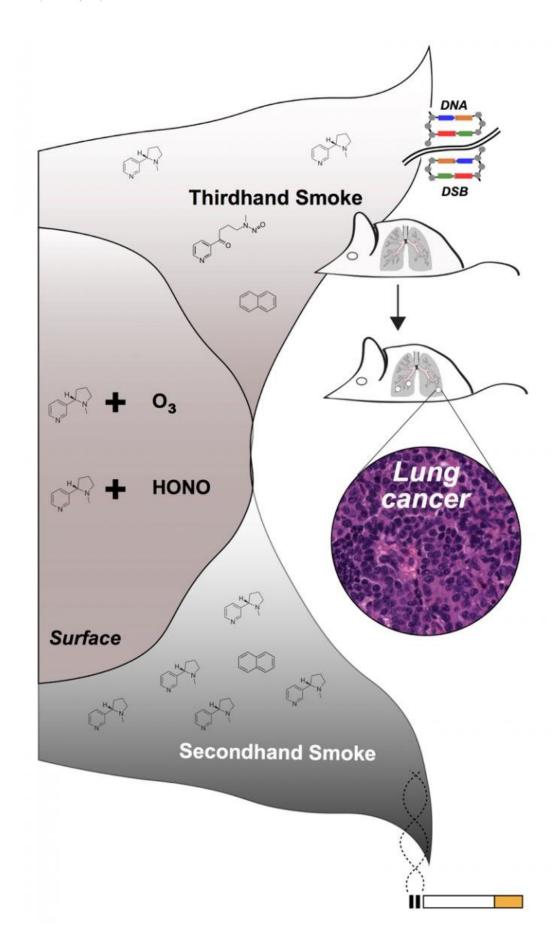


Thirdhand smoke found to increase lung cancer risk in mice

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Thirdhand smoke contains the chemicals in secondhand smoke from a cigarette that are deposited on indoor surfaces. Some of these chemicals interact with molecules from the air to create a toxic mix that includes potentially cancercausing compounds. These compounds induce double-stranded breaks (DSBs) in DNA, which if not repaired correctly, could lead to tumorigenesis in mice. In this study, the researchers have shown for the first time that thirdhand smoke exposure induces lung cancer in A/J mice in early life. Credit: Antoine Snijders, Jian-Hua Mao, and Bo Hang/Berkeley Lab

Researchers at the Department of Energy's Lawrence Berkeley National Laboratory (Berkeley Lab) identified thirdhand smoke, the toxic residues that linger on indoor surfaces and in dust long after a cigarette has been extinguished, as a health hazard nearly 10 years ago. Now a new study has found that it also increases lung cancer risk in mice.

A team led by Antoine Snijders, Jian-Hua Mao, and Bo Hang of Berkeley Lab first reported in 2017 that brief exposure to thirdhand smoke is associated with low body weight and immune changes in juvenile mice. In a follow-up study published recently in *Clinical Science*, the researchers and their team have determined that early thirdhand smoke exposure is also associated with increased incidence and severity of <u>lung cancer</u> in mice.

Field studies in the U.S. and China have confirmed that the presence of thirdhand smoke in indoor environments is widespread, and traditional cleaning methods are not effective at removing it. Because exposure to thirdhand smoke can occur via inhalation, ingestion, or uptake through the skin, young children who crawl and put objects in their mouths are more likely to come in contact with contaminated surfaces, and are therefore the most vulnerable to thirdhand smoke's harmful effects.



In the Berkeley Lab researchers' new study, an experimental cohort of 24 A/J mice (a strain susceptible to spontaneous lung <u>cancer</u> development) was housed with scraps of fabric impregnated with thirdhand smoke from the age of 4 weeks to 7 weeks. The dose the mice received was estimated to be about 77 micrograms per kilogram of body weight per day - comparable to the ingestion exposure of a human toddler living in a home with smokers. Forty weeks after the last exposure, these mice were found to have an increased incidence of lung cancer (adenocarcinoma), larger tumors, and a greater number of tumors, compared to 19 control mice.

Their work also sheds light on what happens on both a molecular and cellular level. If thirdhand smoke toxins damage DNA within cells and the damage is not repaired properly, it can give rise to mutations, which may lead to the cell becoming cancerous. To further investigate how thirdhand smoke exposure promotes tumor formation, the team performed in vitro studies using cultured human lung cancer cells.

These studies indicated that thirdhand smoke exposure induced DNA double-strand breaks and increased cell proliferation and colony formation. In addition, RNA sequencing analysis revealed that thirdhand smoke exposure caused endoplasmic reticulum stress and activated p53 (tumor suppressor) signaling. The physiological, cellular, and molecular data indicate that early exposure to thirdhand smoke is associated with increased lung cancer risk.

More information: Bo Hang et al, Short-term early exposure to thirdhand cigarette smoke increases lung cancer incidence in mice, *Clinical Science* (2018). <u>DOI: 10.1042/CS20171521</u> Bo Hang et al. Short-term early exposure to thirdhand cigarette smoke increases lung cancer incidence in mice, *Clinical Science* (2018). <u>DOI: 10.1042/CS20171521</u>



Provided by Lawrence Berkeley National Laboratory

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