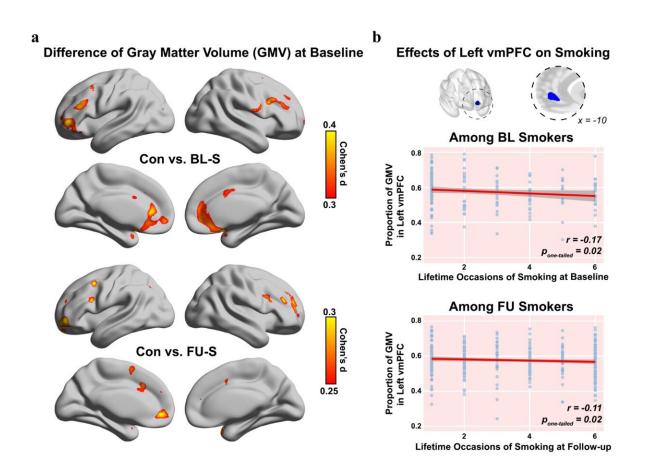


## Reduced gray matter in frontal lobes linked to teenage smoking and nicotine addiction: Study

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Smaller Left vmPFC was associated with future smoking. Credit: *Nature Communications* (2023). DOI: 10.1038/s41467-023-40079-2



Levels of gray matter in two parts of the brain may be linked to a desire to start smoking during adolescence and the strengthening of nicotine addiction, a new study has shown.

A team of scientists, led by the universities of Cambridge and Warwick in the UK and Fudan University in China, analyzed brain imaging and behavioral data of over 800 young people at the ages of 14, 19 and 23.

They found that, on average, teenagers who started <u>smoking</u> by 14 years of age had markedly less gray matter in a section of the left frontal lobe linked to decision-making and rule-breaking.

Gray matter is the <u>brain tissue</u> that processes information, and contains all of the organ's neurons. While <u>brain development</u> continues into adulthood, gray matter growth peaks before adolescence.

Low gray matter volume in the left side of the <u>ventromedial prefrontal</u> <u>cortex</u> may be an "inheritable biomarker" for nicotine addiction, say researchers—with implications for prevention and treatment.

In addition, the scientists found that the opposite, right part of the same brain region also had less gray matter in smokers.

Importantly, loss of gray matter in the right prefrontal <u>cortex</u> appears to speed up only after someone has started smoking. This region is linked to the seeking of sensations.

The team argue that less gray matter in the left forebrain could lower cognitive function and lead to "disinhibition": impulsive, rule-breaking behavior arising from a limited ability to consider consequences. This may increase the chances of smoking at a young age.

Once a nicotine habit takes hold, gray matter in the right frontal lobe



shrinks, which may weaken control over smoking by affecting "hedonic motivation": the way pleasure is sought and managed. Excessive loss of gray matter in the right brain was also linked to binge drinking and marijuana use.

Taken together, the findings point to a damaged "neurobehavioural mechanism" that can lead to nicotine use starting early and becoming locked into long-term addiction, say researchers. The study used data from the IMAGEN project and is published in the journal *Nature Communications*.

"Smoking is perhaps the most common addictive behavior in the world, and a leading cause of adult mortality," said Prof Trevor Robbins, cosenior author from Cambridge's Department of Psychology.

"The initiation of a smoking habit is most likely to occur during adolescence. Any way of detecting an increased chance of this, so we can target interventions, could help save millions of lives."

Annual deaths from cigarettes are expected to reach eight million worldwide by the end of the decade. Currently, one in five adult deaths each year are attributed to smoking in the US alone.

"In our study, reduced gray matter in the left prefrontal cortex is associated with increased rule-breaking behavior as well as early smoking experiences. It could be that this rule-breaking leads to the violation of anti-smoking norms," said Robbins.

Co-author Prof Barbara Sahakian from Cambridge's Department of Psychiatry said, "The ventromedial prefrontal cortex is a key region for dopamine, the brain's pleasure chemical. As well as a role in rewarding experiences, dopamine has long been believed to affect self-control.



"Less gray matter across this brain region may limit cognitive function, leading to lower self-control and a propensity for risky behavior, such as smoking."

The study used data gathered by the IMAGEN project from sites in four European countries: UK, Germany, France and Ireland. The researchers compared brain imaging data for those who had smoked by age 14 with those who had not, and repeated this for the same participants at ages 19 and 23.

Those with smoking experience by 14 years of age had significantly less gray matter in the left prefrontal cortex, on average. Additionally, those who started smoking by age 19 also had less gray matter in their left prefrontal cortex at 14, indicating a potential causal influence.

The scientists also looked at the right ventromedial prefrontal cortex. Gray matter loss occurs in everyone as they age. However, those who smoked from age 14 as well as those smoking from age 19 both ended up with excessive gray matter loss in the right frontal lobe.

For the right prefrontal cortex, 19-year-old smokers who did not start during adolescence had similar gray matter levels at age 14 to those who never smoked at all. This suggests a rapid reduction in the right ventromedial prefrontal cortex only begins with the onset of smoking.

Data at age 23 showed that gray matter volume in the right prefrontal cortex shrank at a faster pace in those who continued to smoke, suggesting an influence of smoking itself on prefrontal function.

Researchers also analyzed data from two questionnaires completed by participants to investigate the personality traits of novelty seeking and sensation seeking.



"Both questionnaires examine the pursuit of thrilling experiences, but they measure distinct behaviors," said Robbins. "The sensation seeking scale focuses on pleasurable experiences, while the novelty seeking questionnaire includes items on impulsiveness and rule-breaking."

Less gray matter in the left prefrontal cortex was associated with novelty seeking, particularly disorderly and rule-breaking behavior, while reduced gray matter volume in the right prefrontal cortex was linked to sensation seeking only.

Lead author Prof Tianye Jia from Fudan University added, "Less gray <u>matter</u> in the left frontal lobes is linked to behaviors that increase the likelihood of smoking in adolescence.

"Smokers then experience excessive loss of <u>gray matter</u> in the right frontal lobes, which is linked to behaviors that reinforce substance use. This may provide a causal account of how smoking is initiated in young people, and how it turns into dependence."

**More information:** Tianye Jia, Association between vmPFC gray matter volume and smoking initiation in adolescents, *Nature Communications* (2023). DOI: 10.1038/s41467-023-40079-2. www.nature.com/articles/s41467-023-40079-2

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