Network Open.

Original Investigation | Psychiatry Use of Tobacco Products and Suicide Attempts Among Elementary School-Aged Children

Phil H. Lee, PhD; Brenden Tervo-Clemmens, PhD; Richard T. Liu, PhD; Maia B. Gersten, BS; Jae-Yoon Jung, PhD; Amy C. Janes, PhD; Jodi Gilman, PhD

Abstract

IMPORTANCE The use of tobacco products, including e-cigarettes and vaping, has rapidly increased among children. However, despite consistent associations found between smoking cigarettes and suicidal behaviors among adolescents and adults, there are limited data on associations between emerging tobacco products and suicidal behaviors, especially among preadolescent children.

OBJECTIVE To examine whether the use of tobacco products is associated with nonsuicidal selfinjury (NSSI), suicidal ideation (SI), and suicide attempts (SAs) among preadolescent children.

DESIGN, SETTING, AND PARTICIPANTS This cohort study, conducted from September 1, 2022, to September 5, 2023, included participants in the Adolescent Brain Cognitive Development study, a population-based cohort of 11 868 US children enrolled at 9 and 10 years of age. The cross-sectional investigation focused on 3-year periods starting from the baseline to year 2 of follow-up. Statistical analysis was performed from October 1, 2022, to June 30, 2023.

MAIN OUTCOMES AND MEASURES Children's use of tobacco products was assessed based on youth reports, including lifetime experiences of various nicotine-related products, supplemented with hair toxicologic tests. Main outcomes were children's lifetime experiences of NSSI, SI, and SAs, assessed using the K-SADS-5 (Kiddie Schedule for Affective Disorders and Schizophrenia for the *DSM-5*). Multivariate logistic regression was conducted to examine the associations of the use of tobacco products with NSSI, SI, and SAs among the study participants. Sociodemographic, familial, and children's behavioral, temperamental, and clinical outcomes were adjusted in the analyses.

RESULTS Of 8988 unrelated study participants (median age, 9.8 years [range, 8.9-11.0 years]; 4301 girls [47.9%]), 101 children (1.1%) and 151 children (1.7%) acknowledged lifetime use of tobacco products at baseline and at 18-month follow-up, respectively. After accounting for various suicide risk factors and potential confounders, children reporting use of tobacco products were at a 3 to 5 times increased risk of SAs (baseline: n = 153 [adjusted odds ratio (OR), 4.67; 95% CI, 2.35-9.28; false discovery rate (FDR)-corrected P < .001]; year 1: n = 227 [adjusted OR, 4.25; 95% CI, 2.33-7.74; FDR-corrected P < .001]; and year 2: n = 321 [adjusted OR, 2.85; 95% CI, 1.58-5.13; FDR-corrected P = .001]). Of all facets of impulsivity measures that were significant correlates of use of tobacco products, negative urgency was the only independent risk factor for SAs (adjusted OR, 1.52 [95% CI, 1.31-1.78]; FDR-corrected P < .001). In contrast, children's alcohol, cannabis, and prescription drug use were not associated with SAs.

CONCLUSIONS AND RELEVANCE This study of US children suggests that the increased risk of SAs, consistently reported for adolescents and adults who smoke cigarettes, extends to a range of emerging tobacco products and manifests among elementary school-aged children. Further

(continued)

Den Access. This is an open access article distributed under the terms of the CC-BY License.

JAMA Network Open. 2024;7(2):e240376. doi:10.1001/jamanetworkopen.2024.0376

Key Points

Question Is the use of tobacco products associated with an increased risk of self-injurious thoughts and behaviors among children?

Findings This cohort study of 8988 preadolescent children enrolled in the Adolescent Brain Cognitive Development study found statistically significant associations between children's use of tobacco products and suicide attempts. These associations were partially related to increased negative urgency and remained significant after accounting for various demographic, socioeconomic, familial, and clinical risk factors of suicide.

Meaning The findings suggest that smoking tobacco products may be a modifiable risk factor that can be addressed in suicide prevention efforts, especially among children.

Supplemental content

Author affiliations and article information are listed at the end of this article.

Abstract (continued)

investigations are imperative to clarify the underlying mechanisms and to implement effective preventive policies for children.

JAMA Network Open. 2024;7(2):e240376. doi:10.1001/jamanetworkopen.2024.0376

Introduction

The increasing use of various tobacco products, including e-cigarettes, vapes, and hookahs, among children and adolescents is a critical public health problem worldwide.¹ Smoking, especially when started at a young age, has been associated with lifelong negative mental health outcomes, ranging from diminished neurocognition²⁻⁴ to neurodevelopmental alterations⁵⁻¹⁰ to increased risk of delinquent behaviors and addiction¹¹ in later life.¹²⁻¹⁴ Multiple population-based studies have reported that adolescent smokers exhibit a 2 to 5 times greater risk of self-injurious thoughts and behaviors (SITBs), even after adjusting for their mental health problems.¹⁵⁻¹⁷ Higher rates of suicide risk are consistently reported among current smokers compared with individuals who quit,^{16,18-21} underscoring that further investigations are necessary to clarify whether smoking could be a major preventable risk factor associated with suicidal behaviors.²²

Compared with adolescents¹⁵⁻¹⁷ and adults,^{21,23} however, there is a dearth of information available on the association between smoking and suicidal behaviors among preadolescent children. Many observational studies focused on uncovering risk factors of childhood smoking initiation, including prenatal exposure,²⁴ psychiatric disorders,²⁵ family conflict,^{24,26} and peer influence.⁵ Several groups examined brain structures and functional networks for biomarkers of early-onset smoking.⁵ Others inspected genome-wide data to elucidate the genetic effects shared between smoking and childhood psychopathologic problems.²⁷ Some of these same risk factors (eg, psychiatric disorders, family conflict or dysfunction, peer relationships) have similarly been associated with preadolescent suicidal thoughts and/or behavior.²⁸ However, it remains an open question whether associations between smoking and suicidal behaviors, consistently observed among adolescents and adults who use combustible cigarettes, also manifest among preadolescent children,²⁹ an increasing number of whom are exposed to emerging tobacco products.

The primary aim of this study was to fill in this major knowledge gap. We analyzed data from the Adolescent Brain Cognitive Neurodevelopment (ABCD) study, a US population-based cohort of 11 878 elementary school-aged children (enrolled at 9-10 years of age). The ABCD data consist of various social, familial, mental, and physical well-being and behavioral measures assessed from baseline and the first 2 years of follow-up (release version 4.0). Using these data, we examined the following 3 questions: (1) Is the use of tobacco products associated with SITBs among children, while accounting for demographic, familial, and socioeconomic confounders? (2) If so, are children's cognition, temperament, and psychopathologic outcomes associated with the link between smoking and SITBs? (3) Are these findings unique to the use of tobacco products or shared with other substance use problems, such as alcohol, cannabis, and prescription drugs use, among children? This research should have an effect on the prioritization of smoking prevention and cessation efforts in health care policies, especially concerning suicide prevention among children and adolescents.

Methods

Study Participants

The ABCD study, an ongoing longitudinal study that began in 2017, enrolled a total of 11868 participants aged 9 and 10 years in 22 US study sites (eTable 1 in Supplement 1). We downloaded ABCD study data, version 4.0, from September 1, 2022, to September 5, 2023, from the National Institute of Mental Health Data Archive. Comprehensive information regarding sample collection,

survey measures, and study protocols is available elsewhere.³⁰⁻³² The ABCD study was approved by a central institutional review board at the University of California, San Diego. All ABCD participants and their caregivers provided written informed assent and consent for human research. This study is the secondary analysis of deidentified ABCD participants and is exempt from informed consent by the Massachusetts General Hospital institutional review board. This report followed the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) reporting guideline.

Measures of the Use of Tobacco Products

Children's substance use data were assessed at baseline using the ABCD Youth Substance Use Interview.²⁹ The ABCD Youth Mid-Year Phone Interview for Substance Use was conducted at the 6-month follow-up and then on a yearly basis, asking whether the child has smoked tobacco products at any time in the past 6 months until the interview date. We assembled hair toxicology screening test results from the ABCD Youth Hair Results. Using these data, we generated 3 binary variables for lifetime use of tobacco products corresponding to baseline, 6-month follow-up, and 18-month follow-up. Participants were classified as cases if they ever reported the use of tobacco products in the surveys or had positive test results for cotinine in the hair toxicology tests (eTable 2 in Supplement 1).

Measures of Suicidal Behaviors

Data on SITBs of the participants were obtained from the youth and caregiver reports of the K-SADS-5 (Kiddie Schedule for Affective Disorders and Schizophrenia for the *DSM-5*). Lifetime nonsuicidal self-injuries (NSSIs), suicide attempts (SAs), and suicidal ideation (SI) were assessed based on the K-SADS-5 suicide survey module (eTable 3 in Supplement 1). Considering the young age of the ABCD study participants, SA cases included participants who reported attempted, interrupted, and/or aborted SA actions with intention to die, while SI cases were participants with passive or active thoughts of suicide but without any SA actions. Nonsuicidal self-injury cases were participants who reported NSSIs without reporting any SA or SI experience (eFigure 1 in Supplement 1). Response rates for the use of tobacco products and SITB outcomes are provided in eTable 4 in Supplement 1.

Additional Measures on Family and Children

To identify and control for potential confounding factors, we examined demographic characteristics, socioeconomic characteristics, family history, and prenatal exposure of ABCD study participants stratified by their use of tobacco products status using the ABCD Parent Demographics Survey, Parent Family History Summary, and Developmental History Questionnaire (eTable 5 in Supplement 1). Parent-reported race and ethnicity information was obtained from the ABCD Parent Demographics Survey. For children's cognition, temperament, and psychopathology outcomes, we analyzed data from the ABCD Child Behavior Checklist, National Institutes of Health Toolbox Summary Scores, Pearson Scores, Little Man Task Summary Scores, Mental Health Youth Sum Scores, and Early Adolescent Temperament Questionnaire Parent data (eTable 6 in Supplement 1). We generated ever-use experiences of each participant's alcohol sipping or tasting, alcohol drinking (ie, at least a full cup of drinking), cannabis use, and prescription use at baseline, 6-month follow-up, and 18-month follow-up (eTable 7 in Supplement 1).

Statistical Analysis

Statistical analysis was performed from October 1, 2022, to June 30, 2023. Bivariate analyses were conducted for summarizing the characteristics of the study participants stratified by their use of tobacco product status. We examined demographic characteristics (age, sex, parent-reported race and ethnicity [Asian, Black, Hispanic, White, and other (American Indian or Alaska Native, Native Hawaiian or Other Pacific Islander, multiple races, or other in the survey)]), socioeconomic factors (parental education, household income, parental marital status), parental history (depression,

JAMA Network Open. 2024;7(2):e240376. doi:10.1001/jamanetworkopen.2024.0376

alcohol abuse, drug use, behavioral troubles, suicide), and prenatal exposure to smoking. Race and ethnicity information was included because smoking behaviors are influenced by cultural and social norms, which may vary among different racial and ethnic groups. Unpaired *t* tests were used for continuous variables, and χ^2 tests were used for categorical variables. False discovery rates (FDRs) were calculated to ensure a proper type I error rate at 5% for all analyses involving multiple testing.

Multivariate logistic regression was conducted to examine the association of the use of tobacco products with NSSI, SI, and SAs. First, we examined a basic regression model, in which the use of tobacco products and each SITB outcome were tested as an independent and a dependent variable, respectively, while controlling for age, sex, and parent-reported race and ethnicity. In the following full covariate-adjusted model, we additionally included sociodemographic, family history, and prenatal substance use variables that were identified as significant correlates of the use of tobacco products in the preceding bivariate analyses. We also examined whether specific children's cognition, temperaments, and psychopathology outcomes were significant correlates of the use of tobacco products using multivariate logistic regression. All child behavioral outcomes with significant correlation with the use of tobacco products were added in the full covariate-adjusted regression model to estimate the independent association of the use of tobacco products with SITBs above and beyond these potential confounders. Last, we evaluated the specificity of the association between the use of tobacco products and the risk of SITB outcomes by comparing the analysis results with other types of substance use. Statistical analysis was conducted in R, version 4.2.1 (R Project for Statistical Computing). All P values were from 2-sided tests, and results were deemed statistically significant at an FDR-corrected P < .05.

Results

Study Sample

Table 1 summarizes the major characteristics of ABCD participants stratified by their lifetime use of tobacco products at baseline. Of the 8988 unrelated youths (median age, 9.8 years [8.9-11.0]; 4301 girls [47.9%]) included in this study (eFigure 2 in Supplement 1), 101 (1.1%) reported ever using tobacco products at baseline. Of these participants, 42 girls (41.6%) reported the use of tobacco products (median age, 10.1 years [range, 8.2-11.0 years]). A total of 8887 youths (98.9%; 4259 girls [47.9%]; median age, 9.8 years [range, 8.9-11.0 years]) did not report use of tobacco products. Prevalence of smoking experience was not statistically different in either sex or across different races and ethnicities. However, children in the case group were exposed to more disadvantaged socioeconomic environments and smoking during pregnancy and were more likely to have parents with depression, alcohol problems, behavioral troubles, and suicide-related incidences compared with children in the control group. Although the prevalence of tobacco product use increased significantly from 1.1% (n = 101) at baseline to 1.7% (n = 151) at 18-month follow-up ($\chi^2 = 9.66$; *P* = .002), all major characteristics of the group that used tobacco products observed at baseline remained consistent in the follow-up years (eTables 8 and 9 in Supplement 1).

Association of Smoking and SITBs

First, we examined whether the use of tobacco products by children was associated with SITBs. Of 8988 youths, the lifetime prevalence of SITBs at baseline, year 1, and year 2, respectively, were as follows: NSSI, 4.8% (n = 428), 5.7% (n = 508), and 6.2% (n = 560); SI, 12.8% (n = 1151), 16.2% (n = 1454), and 20.6% (n = 1853); and SAs, 1.7% (n = 153), 2.5% (n = 227), and 3.6% (n = 321) (eFigure 1 in Supplement 1). In the basic model, we found statistically significant associations of the use of tobacco products with SAs and SI at all assessed time points (eFigure 3 and eTable 10 in Supplement 1). In the full covariate-adjusted model, the effect size of the use of tobacco products was attenuated, yet it remained independent and statistically significant for both SAs and SI (Figure 1; eTable 11 in Supplement 1). The use of tobacco products assessed at baseline was associated with SAs and SI at baseline, as well as with the outcomes reported in year 1 and year 2.

Children reporting use of tobacco products were at a 3 to 5 times increased risk of SAs (baseline: n = 153 [adjusted odds ratio (OR), 4.67; 95% CI, 2.35-9.28; false discovery rate (FDR)-corrected P < .001]; year 1: n = 227 [adjusted OR, 4.25; 95% CI, 2.33-7.74; FDR-corrected P < .001]; and year 2: n = 321 [adjusted OR, 2.85; 95% CI, 1.58-5.13; FDR-corrected P = .001]). Similarly, lifetime use of tobacco products assessed at 6-month and 18-month follow-up was associated with the 2 SITB outcomes subsequently measured in year 1 and year 2. In all cases, the estimated effect sizes of the use of tobacco products for SAs were approximately 3 times larger compared with those for SI (Figure 1). Sensitivity analyses confirmed consistent findings across various measures of SITB outcomes (eTables 12-14 in Supplement 1). Despite limited power, we also found prospective associations of the use of tobacco products assessed at baseline and new SA cases reported in follow-up years 1 and 2 (ie, someone who did not report SAs at baseline but did in year 1 and year 2) (eTable 15 in Supplement 1). Conversely, we found no associations of the use of tobacco products with NSSI at all assessed times (Figure 1).

Table 1. Major Characteristics of Adolescent Brain Cognitive Development Study Participants Based on Youth-Reported Ever Use of Tobacco Products in the Baseline

	Ever use of tobacco products (baseline) ^a				
Characteristic ^b	Cases (n = 101 [1.1%])	Controls (n = 8887 [98.9%])	P value ^c		
Age, median (range), y ^d	10.1 (8.9-11.0)	9.8 (8.9-11.0)	.09		
Sex, No. (%)					
Male	59 (58.4)	4628 (52.1)	24		
Female	42 (41.6)	4259 (47.9)	.24		
Race and ethnicity, No. (%)					
Asian	1 (1.0)	210 (2.4)			
Black	16 (15.8)	1221 (13.7)			
Hispanic	15 (14.9)	1752 (19.7)	.009		
White	48 (47.5)	4785 (51.3)			
Other ^e	21 (20.8)	919 (10.3)			
Parental education, No. (%)					
<high certification<="" diploma="" ged="" or="" school="" td=""><td>6 (5.9)</td><td>345 (3.9)</td><td colspan="2"></td></high>	6 (5.9)	345 (3.9)			
High school diploma or GED certification	16 (15.8)	759 (8.5)			
Some college	43 (42.6)	2269 (25.5)	<.001 [†]		
Bachelor's degree	20 (19.8)	2301 (25.9)			
Postgraduate degree	16 (15.8)	3213 (36.3)			
Household income, No. (%)					
≤\$50 000	45 (44.6)	2078 (23.4)			
>\$50 000 and ≤\$100 000	30 (29.7)	2328 (26.2)	<.001 ^f		
>\$100 000	26 (25.7)	4481 (50.4)			
Marital status of parents, No. (%)					
Not married	55 (54.5)	2758 (31.0)	. 001f		
Married	46 (45.5)	6129 (69.0)	<.001		
Parental history, No. (%)					
Alcohol problem	35 (34.7)	1282 (14.4)	<.001 ^f		
Drug problem	48 (47.5)	2685 (30.2)	<.001		
Depression	33 (32.7)	945 (10.6)	<.001 ^f		
Behavioral troubles	33 (32.7)	1141 (12.8)	<.001 ^f		
Mental disorders	52 (51.5)	3519 (39.6)	.02		
Suicide	17 (16.8)	477 (5.4)	<.001 ^f		
Prenatal smoking, No. (%)					
Before acknowledging pregnancy	45 (44.6)	1187 (13.4)	<.001 ^f		
After acknowledging pregnancy	25 (24.8)	459 (5.2)	<.001 ^f		

Abbreviation: GED, General Educational Development.

- ^a Cases represent children who acknowledged the use of tobacco products in the past or at present or had positive cotinine results from hair toxicology tests, while controls did not.
- ^b Information about the demographic, socioeconomic, and family history was obtained from the Adolescent Brain Cognitive Development Parent Demographic survey data.
- ^c *P* values were from χ^2 tests for categorical variables (eg, sex) and unpaired *t* test for quantitative measures (eg, age).
- ^d Age in years represents the age in months of study participants at the time of the interview divided by 12.
- Indicates study participants who selected either
 American Indian or Alaska Native, Native Hawaiian or
 Other Pacific, multiple races, or others in the
 Adolescent Brain Cognitive Development Parent
 Demographic survey.

^f Significant after multiple testing correction.

Investigation of Children's Cognition and Behavioral Measures

Next, we examined whether the associations between the use of tobacco products and increased risk of SAs and SI are due to children's cognition, temperament, and psychopathology outcomes. Of the 48 measures that we examined in year 2 (eTable 6 in Supplement 1), 13 were significant correlates of the use of tobacco products after multiple testing correction (**Figure 2**). Children who had reported the use of tobacco products at 18 months were more likely to have conduct disorder and oppositional defiant disorder compared with controls in year 2. These children also exhibited increased levels of rule-breaking behaviors, aggressive behaviors, and social problems relative to controls. In addition, there were notable differences in several impulsivity traits, including a lack of planning, positive urgency, and negative urgency.

However, both bivariate and multivariate regression analyses, which examined the 13 correlates of the use of tobacco products in the same model, found that the association between the use of tobacco products and SAs was independent of all childhood behavioral or psychopathology outcomes (OR, 3.09 [95% CI, 1.70-5.65]; P < .001; likelihood ratio test FDR-corrected P < .001) (**Table 2**). Negative urgency was the only correlate of the use of tobacco products that retained a significant association with SAs independent from other variables (OR, 1.52 [95% CI, 1.31-1.78]; P < .001; FDR-corrected P < .001). Smoking data assessed at baseline and 6-month follow-up presented similar findings (eTables 16 and 17 in Supplement 1). In contrast, the association between the use of tobacco products and SI dissipated once childhood behavioral measures were accounted for (eTables 18-20 in Supplement 1). Interaction analyses of smoking with childhood outcome measures did not reveal any significant associations with suicide risk outcomes (eTable 21 in Supplement 1).

Variable/ outcome	Odds ratio (95% CI)	Does not favor outcome	Favors outcome				FDR-corrected
Smoking (baselin	ie)						P value
NSSI							
Baseline	0.89 (0.32-2.48)						.93
Year 1	0.84 (0.30-2.37)						.93
Year 2	0.96 (0.38-2.44)		<u> </u>				.94
SA							
Baseline	4.67 (2.35-9.27)			a			<.001
Year 1	4.25 (2.33-7.74)			a			<.001
Year 2	2.85 (1.58-5.12)						.001
SI							
Baseline	2.03 (1.25-3.29)		D				.009
Year 1	2.00 (1.25-3.21)		b				.009
Year 2	1.63 (1.03-2.59)						.10
Smoking (6 mo)							
NSSI							
Year 1	1.05 (0.45-2.44)						.94
Year 2	1.09 (0.49-2.42)	—					.93
SA							
Year 1	4.24 (2.42-7.43)			a			<.001
Year 2	2.83 (1.63-4.90)						<.001
SI							
Year 1	1.87 (1.21-2.87)		D				.009
Year 2	1.49 (0.98-2.28)						.10
Smoking (18 mo)						
NSSI							
Year 2	1.32 (0.67-2.59)	_	-				.58
SA							
Year 2	3.17 (1.95-5.17)			a			<.001
SI							
Year 2	1.46 (0.99-2.14)		-				.09
		0	2	4	6	8	「 10
				Odds ratio (95	% CI)		

In multivariate logistic regression, lifetime use of tobacco products was used as an independent variable, along with other covariates, to measure associations with 3 suicide risk outcome measures: nonsuicidal self-injury (NSSI), suicide attempts (SAs), and suicidal ideation without SAs (SI). Fixed-effects covariates included age, sex, race and ethnicity, parents' highest educational level, parents' marital status, household income, prenatal exposure to smoking, parental history of depression, alcohol problems, behavioral troubles, and suicide.

^a False discovery rate (FDR)-corrected *P* < .001.
 ^b FDR-corrected *P* < .01.

Comparison With Other Substance Use Data

To evaluate the specificity of the association between the use of tobacco products and increased risk of SAs, we examined the association of SAs and smoking with other types of substance use data. Along with alcohol, cannabis, and prescription drug use, significant associations between SAs and use of tobacco products remained robust (use of tobacco products and alcohol sipping: OR, 2.70

Figure 2. Associations of Children's Personality, Cognition, and Psychology Outcomes With the Use of Tobacco Products

Variable/ outcome	β (95% CI)	Does not favor outcome	Favors outcome	FDR-c P valu	orrected:
Psychopathology (B)		-			
Conduct disorder	1.57 (0.78 to 2.36)			a <.001	
Oppositional defiant disorder	1.42 (0.72 to 2.12)			<.001	
ADHD	0.89 (0.15 to 1.64)		_	.06	
Anxiety disorder	0.47 (-0.34 to 1.29)			.51	
Depression	0.32 (-0.49 to 1.14)			.64	
Psychopathology					
Rule-breaking behaviors	0.66 (0.47 to 0.85)		a	<.001	
Aggressive behaviors	0.48 (0.29 to 0.67)		a	<.001	
Social problems	0.31 (0.12 to 0.50)		b	.009	
Attention problems	0.28 (0.08 to 0.47)		C	.02	
Total psychosis symptoms	0.20 (0.03 to 0.37)		- - -	.06	
Thought problems	0.16 (-0.03 to 0.36)	-		.28	
Anxious or depressed	0 15 (-0 05 to 0 34)	-		34	
Psychosis severity	0.13(-0.05 to 0.30)			35	
Somatic symptoms	0.11 (-0.09 to 0.30)	_		.55	
Withdrawn or depressed	0.07 (-0.12 to 0.27)			.51	
Mania	0.07(-0.12 to 0.27)	_		.04	
Porconality	0.07 (=0.11 (0 0.24)		-	.04	
Lack of planning	0 E7 (0 40 to 0 7E)		<u>a</u>	< 001	
	0.37 (0.40 to 0.73)		b	<.001	
Positive urgency	0.30 (0.12 to 0.47)		b	.006	
	0.28 (0.11 to 0.45)		 C	.009	
Sensation seeking	0.26 (0.09 to 0.44)		C	.02	
Aggression	0.25 (0.07 to 0.43)		<u> </u>	.02	
Negative urgency	0.24 (0.07 to 0.42)			.03	
Depressive mood	0.22 (0.04 to 0.40)			.049	
Fun seeking	0.11 (-0.07 to 0.28)			.49	
High-intensity pleasure	0.11 (-0.07 to 0.28)			.49	
Drive	0.09 (-0.08 to 0.26)			.55	
Frustration	0.09 (-0.09 to 0.26)			.58	
Fear	0.08 (-0.09 to 0.25)			.63	
Shyness	0.05 (-0.12 to 0.22)	_		.69	
Inhibitory control	0.05 (-0.13 to 0.23)			.72	
Attention	-0.01 (-0.19 to 0.16)	<u> </u>		.93	
Affiliation	-0.03 (-0.20 to 0.14)		—	.86	
Behavioral inhibition	-0.07 (-0.24 to 0.11)			.64	
Reward responsiveness	-0.13 (-0.30 to 0.05)) —	-	.35	
Activation control	-0.16 (-0.33 to 0.02)			.24	
Cognition					
Processing speed	0.06 (-0.13 to 0.26)			.68	
Short delay recall	0.04 (-0.13 to 0.22)		-	.72	
Attention	0.04 (-0.15 to 0.23)	_	-	.77	
Crystallized cognition	0.02 (-0.15 to 0.19)			.89	
Visuospatial accuracy	0.00 (-0.18 to 0.18)	- -		.10	
Visuospatial reaction time	-0.01 (-0.19 to 0.18)) —		.94	
Long delay recall	-0.02 (-0.19 to 0.16)) —		.90	
Episodic memory	-0.06 (-0.23 to 0.11)) —	_	.65	
Vocabulary	-0.06 (-0.21 to 0.09)) —	-	.64	
Reading	-0.07 (-0.23 to 0.09)) —	_	.64	
		-0.5	0 0.5 1.0 1. β (95% CI)	5 2.0 2.5	

JAMA Network Open. 2024;7(2):e240376. doi:10.1001/jamanetworkopen.2024.0376

The use of tobacco products was assessed in 18-month follow-up, while the assessed personality, cognition, and psychopathology outcomes were assessed in year 2. Effect size indicates the regression coefficient $\boldsymbol{\beta}$ of the exposure (ie, ever use of tobacco products) in multivariate regression, where each child outcome was used as a dependent variable. Additional covariates included age, sex, parent-reported race and ethnicity, parents' highest educational level, household income, parents' marital status, prenatal exposure to tobacco, number of substances used during pregnancy by mother, parental history of depression, alcohol problem, and behavioral troubles. The category Psychopathology (B) includes binary variables for DSM-5--oriented Adolescent Brain Cognitive Development study Child Behavior Checklist measures. ADHD indicates attention-deficit/ hyperactivity disorder.

^a False discovery rate (FDR)-corrected P < .001.

^b FDR-corrected P < .01.

^c FDR-corrected P < .05.

Table 2. Multivariate Logistic Regression Results in Year 2

	Multivariate logistic regression		Likelihoo		
Independent variable	OR (95% CI)	P value	R ² , %	P value	FDR-corrected P value
Demographic characteristics					
Age	1.13 (0.98-1.32)	.10	0.20	.10	.41
Female	0.91 (0.67-1.22)	.05	0.03	.52	.88
Race and ethnicity					
Asian	1.50 (0.52-4.34)	.04	0.00	>.99	>.99
Black	1.47 (0.91-2.36)	.12	0.00	>.99	>.99
Hispanic	1.40 (0.94-2.09)	.10	0.00	>.99	>.99
Other ^b	1.32 (0.84-2.07)	.24	0.00	>.99	>.99
Socioeconomic factors					
Income	1.06 (0.87-1.29)	.58	0.02	.06	.90
Married	0.59 (0.42-0.83)	.003	0.65	.003 ^c	.03
Parental education					
High school diploma or GED certification	1.57 (0.67-3.67)	.30	0.00	>.99	>.99
Some college	1.76 (0.81-3.83)	.15	0.00	>.99	>.99
Bachelor's degree	1.45 (0.63-3.36)	.38	0.00	>.99	>.99
Postgraduate	1.27 (0.53-3.03)	.59	0.00	>.99	>.99
Prenatal substance exposure					
Early tobacco exposure	0.89 (0.48-1.63)	.70	0.01	.70	.97
No early tobacco exposure	1.08 (0.88-1.32)	.44	0.04	.44	.85
Late tobacco exposure	1.37 (0.54-3.45)	.51	0.03	.51	.88
No late tobacco exposure	0.96 (0.78-1.19)	.72	0.01	.72	.97
Parental history					
Alcohol problems	0.86 (0.57-1.28)	.45	0.04	.45	.85
Depression	1.05 (0.67-1.67)	.82	0.00	.82	>.99
Behavioral troubles	1.14 (0.74-1.76)	.54	0.03	.55	.90
Suicide	1.72 (1.07-2.76)	.02	0.34	.03 ^c	.27
Child psychopathology					
Rule-breaking behaviors	1.17 (1.00-1.37)	.05	0.27	.05	.34
Social problems	1.13 (0.98-1.32)	.10	0.19	.10	.41
Aggressive behaviors	1.10 (0.92-1.33)	.30	0.08	.30	.74
Attention problems	1.03 (0.89-1.19)	.73	0.01	.73	.97
Anxious or depressed	1.11 (0.94-1.30)	.21	0.11	.21	.59
Withdrawn or depressed	1.08 (0.93-1.25)	.30	0.07	.31	.74
DSM-5 disorders					
Conduct disorder	0.72 (0.31-1.65)	.44	0.04	.44	.85
Oppositional defiant disorder	0.55 (0.26-1.18)	.12	0.18	.12	.42
Temperament					
Negative urgency	1.52 (1.31-1.78)	<.001	1.99	<.001 ^c	<.001
Lack of planning	1.17 (1.00-1.37)	.05	0.26	.06	.34
Positive urgency	1.13 (0.98-1.32)	.10	0.19	.10	.41
Lack of perseverance	1.07 (0.92-1.25)	.37	0.06	.38	.84
Sensation seeking	0.90 (0.77-1.06)	.20	0.12	.20	.59
Aggression	0.97 (0.84-1.12)	.69	0.01	.69	.97
Depressive mood	0.90 (0.78-1.05)	.19	0.12	.19	.59
Substance use					
Use of tobacco products	3.09 (1.70-5.65)	<.001	0.83	<.001 ^c	.011

Abbreviations: *DSM-5*, *Diagnostic and Statistical Manual of Mental Disorders* (Fifth Edition); FDR, false discovery rate; GED, General Educational Development; OR, odds ratio.

- ^a All listed independent variables were examined simultaneously in the regression model using suicide attempts as a dependent variable. Likelihood ratio tests were conducted by comparing the full regression model with the same model without the corresponding independent variable.
- ^b Indicates study participants who selected either
 American Indian or Alaska Native, Native Hawaiian or
 Other Pacific Islander, multiple races, or other in the
 Adolescent Brain Cognitive Development Parent
 Demographics survey.
- ^c Likelihood ratio test FDR less than 5%.

[95% Cl, 1.46-4.99]; FDR-corrected P = .003; use of tobacco products and alcohol drinking: OR, 3.26 [95% Cl, 1.77-5.98]; FDR-corrected P = .008; use of tobacco products and cannabis use: OR, 2.78 [95% Cl, 1.50-5.16]; FDR-corrected P = .04; and use of tobacco products and prescription drug use: OR, 2.89 [95% Cl, 1.56-5.35]; FDR-corrected P = .03) (eTable 22 in Supplement 1). Of other substance use data, only experiences of alcohol sipping were associated with SAs independent of other variables (OR, 1.55 [95% Cl, 1.14-2.13]; FDR-corrected P = .007).

Discussion

Despite a rapid increase in the use of various tobacco products among youths, there have been limited data on its associations with mental health and suicidal behaviors among younger children. Using 3-year assessment data from a population-based sample of 8988 US preadolescent children, our study provides robust evidence that the use of tobacco products is associated with a 3 to 5 times increased risk of SAs during preadolescence. The heightened risk was detected among children as young as 9 and 10 years and manifested consistently in 2 subsequent years (aged <13 years).This association remained independent of well-established suicide risk factors, including sociodemographic, familial, and children's psychopathology outcomes, as well as other types of substance use (including alcohol, cannabis, and prescription drugs).

Our study advances the field of pediatric suicide research in several important ways. First, to our knowledge, our findings provide the first empirical evidence that an increased risk of SAs, consistently reported for cigarette smokers,¹⁵⁻¹⁷ extends to a range of emerging tobacco products and was observable among preadolescent children. In recent years, e-cigarettes, vapes, and flavored nicotine products have gained wide popularity, in part because they are often marketed as safe alternatives to conventional cigarettes.^{18,33,34} However, several nationwide surveys have reported an increased risk of SAs among middle school and high school students with a lifetime use of e-cigarettes^{16,35,36} and vaping devices.³⁷⁻⁴⁰ Additional work suggests that the association between smoking and SAs may be largest for smoking initiation,⁴¹ which is consistent with the relatively early developmental patterns observed in the present work.

Second, our data extend prior evidence that use of tobacco products by children is a risk factor specifically associated with SAs but not with NSSI or SI. Using data from a community sample of 1458 youths aged 9 to 17 years, Wu et al¹⁷ reported significant associations between cigarette smoking and SAs after controlling for depression, while the previously significant association with SI disappeared. A study by Orri et al⁴² reported an association of prolonged exposure to maternal smoking during pregnancy and late childhood with increased risk of children's SAs after accounting for various socioeconomic factors and parents' mental health problems. Likewise, Orri et al⁴² did not find associations of maternal smoking with SI. The present study provides additional evidence for this distinction because the association between the use of tobacco products and children's SAs remained robust after accounting for various confounders (including children's own externalizing and internalizing problems), while the association with SI vanished. Our findings thus suggest that information on the use of tobacco products may help distinguish individuals who have SI from those who may act on their thoughts.^{20,43,44}

Third, our data suggest that use of tobacco products was correlated specifically with externalizing problems and temperament issues rather than internalizing problems among this age group. For adolescents and adults, there have been mixed findings about the associations between suicidal behaviors, substance abuse, and psychiatric disorders. Several previous studies noted that smoking among youths with suicidal behaviors is likely to be secondary to affective disorders.^{45,46} Other groups reported that the associations between smoking and SAs were independent of depression and excessive alcohol use among adolescents and adults.^{21,47,48} Additional work has suggested a particular association with impulsive SAs that is more specific to externalizing symptoms.^{49,50}

Our findings provide evidence that negative urgency is associated with both smoking exposure and increased risk of SAs among preadolescent children. We found several childhood behavioral outcomes that were significant correlates of the use of tobacco products, many of which, if not all, are established risk factors for substance use problems during adolescence.⁵¹ However, negative urgency was the only correlate of the use of tobacco products that remained an independent correlate of SAs after controlling for various suicide risk factors. The robust association between negative urgency and SAs is notable for several reasons. First, this association may be particularly relevant to this age group. This hypothesis is supported by age-related decreases in impulsivity from youth to adulthood^{52,53} and evidence that the association between impulsivity and suicide is stronger among youths than adults.^{54,55} Second, the finding that negative urgency specifically, rather than trait impulsivity more broadly, was a correlate of SAs is consistent with current conceptualizations of suicide risk. Several models of suicide^{56,57} posit that proximal risk for this outcome often occurs within circumscribed periods of high arousal. Further work is required to understand the neural mechanisms contributing to the association between smoking, negative urgency, and SAs.

Limitations

Our study should be interpreted with several limitations in mind. First, although the ABCD study enrolled participants from various sites across the US, representing diverse geographical regions and demographic groups, the sample may not fully represent the entire US population.³⁰⁻³² Findings from our study thus may not be generalizable to the entire US population of preadolescent children. Second, while we have identified significant associations between the use of tobacco products and SAs both cross-sectionally (at baseline) and prospectively (between different assessment events), our observational study design precludes the determination of causal relationships. For our additional thoughts on potential underlying mechanisms between the use of tobacco products and SAs, please refer to the eAppendix in Supplement 1). Third, we used hair toxicology tests to assess tobacco product use, which, while valuable, has limitations in detecting certain nicotine substances and may not capture the full range of tobacco exposure. Environmental factors, particularly exposure to secondhand smoke, might also be associated with the concentrations of substances detected in hair samples, therefore falsely increasing the prevalence of smoking among the study participants. Fourth, given the young age of the study participants, the prevalence of the use of tobacco products was less than 2%, limiting the statistical power to perform fine-grained secondary analyses, such as investigating the effect of individual types of tobacco products, the role of substance use patterns (eg, quantity, frequency, concentrations, administration routes), and proximal associations between newly developed use of tobacco products and SITBs between various assessment periods. As the ABCD study continues to collect data from adolescents over time, we anticipate that future datasets may offer improved statistical power to enhance our understanding of these critical aspects. Fifth, despite our efforts to control for various confounding factors, including sociodemographic variables, familial influences, and children's psychopathology, the possibility of unmeasured confounders remains. which could influence both tobacco use and suicidal behaviors.

Conclusions

In this study, children reporting tobacco use had an increased risk of SAs but not NSSI or SI. The findings suggest that smoking tobacco products may be a modifiable risk factor for suicide. Taken together, our findings suggest that the presumption that noncombustible tobacco products, including e-cigarettes and vapes, are safer alternatives to conventional cigarettes needs to be more thoroughly investigated, ⁵⁸ especially in the context of their potential effect on the mental health and suicidal behaviors of children and adolescents. Further research using neurobiology, genomics, and neuroimaging data is warranted to clarify the causal mechanisms underlying the association between the use of tobacco products and the increased risk of SAs through a developmental perspective.⁵⁹

Until we have a clear understanding of the role of smoking and associated neural mechanisms, the use of tobacco products, particularly among children and adolescents, should not be overlooked. In conjunction with more active smoking prevention and intervention effort, we call for routine screening of the use of tobacco products among children and adolescents, especially when assessing suicide risk.

ARTICLE INFORMATION

Accepted for Publication: January 7, 2024.

Published: February 26, 2024. doi:10.1001/jamanetworkopen.2024.0376

Open Access: This is an open access article distributed under the terms of the CC-BY License. © 2024 Lee PH et al. *JAMA Network Open*.

Corresponding Author: Phil H. Lee, PhD, Center for Genomic Medicine, Massachusetts General Hospital, Simches Research Building, 185 Cambridge St, Boston, MA 02114 (pleeO@mgh.harvard.edu).

Author Affiliations: Department of Psychiatry, Massachusetts General Hospital and Harvard Medical School, Boston (Lee, Tervo-Clemmens, Liu, Gersten, Gilman); Center for Genomic Medicine, Massachusetts General Hospital, Boston (Lee); Department of Psychiatry and Behavioral Sciences, University of Minnesota, Minneapolis (Tervo-Clemmens); Center for Addiction Medicine, Massachusetts General Hospital, Boston (Tervo-Clemmens, Gilman); Depression Clinical and Research Program, Massachusetts General Hospital, Boston (Liu); Division of Neuropsychiatry, Massachusetts General Hospital, Boston (Gersten); Department of Pediatrics, Stanford University, Stanford, California (Jung); Cognitive and Pharmacological Neuroimaging Unit, National Institute on Drug Abuse, Biomedical Research Center, Baltimore, Maryland (Janes).

Author Contributions: Dr Lee had full access to all of the data in the study and takes responsibility for the integrity of the data and the accuracy of the data analysis.

Concept and design: Lee, Gersten, Jung, Gilman.

Acquisition, analysis, or interpretation of data: Lee, Tervo-Clemmens, Liu, Janes.

Drafting of the manuscript: Lee, Liu, Jung, Gilman.

Critical review of the manuscript for important intellectual content: Lee, Tervo-Clemmens, Liu, Gersten, Janes.

Statistical analysis: Lee, Jung.

Obtained funding: Lee.

Administrative, technical, or material support: Lee, Tervo-Clemmens, Gersten, Jung.

Supervision: Lee, Gilman.

Conflict of Interest Disclosures: Dr Liu reported receiving personal fees from Relmada Therapeutics, the American Psychological Association, the Association for Behavioral and Cognitive Therapies, and the American Foundation for Suicide Prevention outside the submitted work. No other disclosures were reported.

Funding/Support: This research was supported by grants R01 MH119243 (Dr Lee) and RF1MH120830, R01MH115905, R01MH124899 and R21MH130767 (Dr Liu) from the National Institute of Mental Health; grant K02 DA052684-01A1 from the National Institute of Drug Abuse (Dr Gilman); and the intramural research program of the National Institute of Drug Abuse (Dr Janes). Statistical analyses were carried out on the Partner's Research Computing Cluster servers and high-performance computing clusters hosted by the Broad Institute of MIT and Harvard. The Adolescent Brain Cognitive Development (ABCD) study was supported by the National Institutes of Health under award numbers: U01DA041022, U01DA041025, U01DA041028, U01DA041048, U01DA041089, U01DA041093, U01DA041106, U01DA041117, U01DA041120, U01DA041134, U01DA041148, U01DA041156, U01DA041174, U24DA041123, and U24DA041147. A full list of supporters is available at https://abcdstudy.org/federalpartners.html. A listing of participating sites and a complete listing of the study investigators can be found at https://abcdstudy.org. ABCD consortium investigators designed the study and provided data through the National Institute of Mental Health Data Archive (https://nda.nih.gov), but did not participate in data analysis or writing of this study.

Role of the Funder/Sponsor: The funding sources had no role in the design and conduct of the study; collection, management, analysis, and interpretation of the data; preparation, review, or approval of the manuscript; and decision to submit the manuscript for publication.

Data Sharing Statement: See Supplement 2.

Additional Contributions: We thank the ABCD study research team for their great efforts in collecting data. We also thank the ABCD study research participants and their families for their continued support of the ABCD study. We are deeply indebted to the investigators for their dedications in the open data science policy.

REFERENCES

1. Coutts J, Langley RJ. Toxic and addictive effects of nicotine on children and adolescents: are we sleepwalking into a public health disaster? *Arch Dis Child*. 2023;108(9):691-692. doi:10.1136/archdischild-2022-323891

2. Counotte DS, Spijker S, Van de Burgwal LH, et al. Long-lasting cognitive deficits resulting from adolescent nicotine exposure in rats. *Neuropsychopharmacology*. 2009;34(2):299-306. doi:10.1038/npp.2008.96

3. Mahedy L, Wootton R, Suddell S, et al. Testing the association between tobacco and cannabis use and cognitive functioning: findings from an observational and Mendelian randomization study. *Drug Alcohol Depend*. 2021;221: 108591. doi:10.1016/j.drugalcdep.2021.108591

4. Fluharty ME, Heron J, Munafò MR. Longitudinal associations of social cognition and substance use in childhood and early adolescence: findings from the Avon Longitudinal Study of Parents and Children. *Eur Child Adolesc Psychiatry*. 2018;27(6):739-752. doi:10.1007/s00787-017-1068-x

5. Dai HD, Doucet GE, Wang Y, et al. Longitudinal assessments of neurocognitive performance and brain structure associated with initiation of tobacco use in children, 2016 to 2021. *JAMA Netw Open*. 2022;5(8):e2225991. doi: 10.1001/jamanetworkopen.2022.25991

6. Jobson CLM, Renard J, Szkudlarek H, et al. Adolescent nicotine exposure induces dysregulation of mesocorticolimbic activity states and depressive and anxiety-like prefrontal cortical molecular phenotypes persisting into adulthood. *Cereb Cortex*. 2019;29(7):3140-3153. doi:10.1093/cercor/bhy179

7. Akkermans SEA, van Rooij D, Rommelse N, et al. Effect of tobacco smoking on frontal cortical thickness development: a longitudinal study in a mixed cohort of ADHD-affected and -unaffected youth. *Eur Neuropsychopharmacol.* 2017;27(10):1022-1031. doi:10.1016/j.euroneuro.2017.07.007

8. Gazula H, Holla B, Zhang Z, et al. Decentralized multisite VBM analysis during adolescence shows structural changes linked to age, body mass index, and smoking: a COINSTAC analysis. *Neuroinformatics*. 2021;19(4): 553-566. doi:10.1007/s12021-020-09502-7

9. Chaarani B, Kan KJ, Mackey S, et al; IMAGEN Consortium. Low smoking exposure, the adolescent brain, and the modulating role of *CHRNA5* polymorphisms. *Biol Psychiatry Cogn Neurosci Neuroimaging*. 2019;4(7):672-679. doi: 10.1016/j.bpsc.2019.02.006

10. Cheng W, Rolls ET, Robbins TW, et al. Decreased brain connectivity in smoking contrasts with increased connectivity in drinking. *eLife*. 2019;8:e40765. doi:10.7554/eLife.40765

11. Romoli B, Lozada AF, Sandoval IM, et al. Neonatal nicotine exposure primes midbrain neurons to a dopaminergic phenotype and increases adult drug consumption. *Biol Psychiatry*. 2019;86(5):344-355. doi:10. 1016/j.biopsych.2019.04.019

12. Lees B, Meredith LR, Kirkland AE, Bryant BE, Squeglia LM. Effect of alcohol use on the adolescent brain and behavior. *Pharmacol Biochem Behav.* 2020;192:172906. doi:10.1016/j.pbb.2020.172906

13. El Marroun H, Klapwijk ET, Koevoets M, et al. Alcohol use and brain morphology in adolescence: a longitudinal study in three different cohorts. *Eur J Neurosci*. 2021;54(6):6012-6026. doi:10.1111/ejn.15411

14. Degenhardt L, Stockings E, Patton G, Hall WD, Lynskey M. The increasing global health priority of substance use in young people. *Lancet Psychiatry*. 2016;3(3):251-264. doi:10.1016/S2215-0366(15)00508-8

15. Lange S, Koyanagi A, Rehm J, Roerecke M, Carvalho AF. Association of tobacco use and exposure to secondhand smoke with suicide attempts among adolescents: findings from 33 countries. *Nicotine Tob Res.* 2020; 22(8):1322-1329. doi:10.1093/ntr/ntz172

16. Huh Y, Cho HJ. Associations between the type of tobacco products and suicidal behaviors: a nationwide population-based study among Korean adolescents. *Int J Environ Res Public Health*. 2021;18(2):367. doi:10.3390/ ijerph18020367

17. Wu P, Hoven CW, Liu X, Cohen P, Fuller CJ, Shaffer D. Substance use, suicidal ideation and attempts in children and adolescents. *Suicide Life Threat Behav*. 2004;34(4):408-420. doi:10.1521/suli.34.4.408.53733

18. Kim JS, Kim K. Electronic cigarette use and suicidal behaviors among adolescents. *J Public Health (Oxf)*. 2021; 43(2):274-280. doi:10.1093/pubmed/fdz086

19. Mullins N, Kang J, Campos AI, et al; Major Depressive Disorder Working Group of the Psychiatric Genomics Consortium; Bipolar Disorder Working Group of the Psychiatric Genomics Consortium; Eating Disorders Working Group of the Psychiatric Genomics Consortium; German Borderline Genomics Consortium; MVP Suicide Exemplar Workgroup; VA Million Veteran Program. Dissecting the shared genetic architecture of suicide attempt, psychiatric disorders, and known risk factors. *Biol Psychiatry*. 2022;91(3):313-327. doi:10.1016/j.biopsych.2021.05.029

20. Mars B, Heron J, Klonsky ED, et al. What distinguishes adolescents with suicidal thoughts from those who have attempted suicide? a population-based birth cohort study. *J Child Psychol Psychiatry*. 2019;60(1):91-99. doi: 10.1111/jcpp.12878

21. Evins AE, Korhonen T, Kinnunen TH, Kaprio J. Prospective association between tobacco smoking and death by suicide: a competing risks hazard analysis in a large twin cohort with 35-year follow-up. *Psychol Med*. 2017;47(12): 2143-2154. doi:10.1017/S0033291717000587

22. Franklin JC, Ribeiro JD, Fox KR, et al. Risk factors for suicidal thoughts and behaviors: a meta-analysis of 50 years of research. *Psychol Bull*. 2017;143(2):187-232. doi:10.1037/bul0000084

23. Poorolajal J, Darvishi N. Smoking and suicide: a meta-analysis. *PLoS One*. 2016;11(7):e0156348. doi:10.1371/journal.pone.0156348

24. Goldschmidt L, Cornelius MD, Day NL. Prenatal cigarette smoke exposure and early initiation of multiple substance use. *Nicotine Tob Res.* 2012;14(6):694-702. doi:10.1093/ntr/ntr280

25. De Angelis F, Wendt FR, Pathak GA, et al. Drinking and smoking polygenic risk is associated with childhood and early-adulthood psychiatric and behavioral traits independently of substance use and psychiatric genetic risk. *Transl Psychiatry*. 2021;11(1):586. doi:10.1038/s41398-021-01713-z

26. Luk TT, Wang MP, Leung LT, et al. Associations of perceived interparental relationship, family harmony and family happiness with smoking intention in never-smoking Chinese children and adolescents: a cross-sectional study. *BMJ Open*. 2017;7(10):e017523. doi:10.1136/bmjopen-2017-017523

27. Barkhuizen W, Dudbridge F, Ronald A. Genetic overlap and causal associations between smoking behaviours and mental health. *Sci Rep.* 2021;11(1):14871. doi:10.1038/s41598-021-93962-7

28. Liu RT, Walsh RFL, Sheehan AE, Cheek SM, Sanzari CM. Prevalence and correlates of suicide and nonsuicidal self-injury in children: a systematic review and meta-analysis. *JAMA Psychiatry*. 2022;79(7):718-726. doi:10.1001/jamapsychiatry.2022.1256

29. Lisdahl KM, Sher KJ, Conway KP, et al. Adolescent Brain Cognitive Development (ABCD) study: overview of substance use assessment methods. *Dev Cogn Neurosci.* 2018;32:80-96. doi:10.1016/j.dcn.2018.02.007

30. Zucker RA, Gonzalez R, Feldstein Ewing SW, et al. Assessment of culture and environment in the Adolescent Brain and Cognitive Development study: rationale, description of measures, and early data. *Dev Cogn Neurosci*. 2018;32:107-120. doi:10.1016/j.dcn.2018.03.004

31. Barch DM, Albaugh MD, Avenevoli S, et al. Demographic, physical and mental health assessments in the Adolescent Brain and Cognitive Development study: rationale and description. *Dev Cogn Neurosci*. 2018; 32:55-66. doi:10.1016/j.dcn.2017.10.010

32. Luciana M, Bjork JM, Nagel BJ, et al. Adolescent neurocognitive development and impacts of substance use: overview of the Adolescent Brain Cognitive Development (ABCD) baseline neurocognition battery. *Dev Cogn Neurosci.* 2018;32:67-79. doi:10.1016/j.dcn.2018.02.006

33. Moustafa AF, Testa S, Rodriguez D, Pianin S, Audrain-McGovern J. Adolescent depression symptoms and e-cigarette progression. *Drug Alcohol Depend*. 2021;228:109072. doi:10.1016/j.drugalcdep.2021.109072

34. Wang G, Liu W, Song W. Toxicity assessment of electronic cigarettes. *Inhal Toxicol*. 2019;31(7):259-273. doi:10. 1080/08958378.2019.1671558

35. Park S, Lee KS. Association of heated tobacco product use and secondhand smoke exposure with suicidal ideation, suicide plans and suicide attempts among Korean adolescents: a 2019 national survey. *Tob Induc Dis.* 2021;19:72. doi:10.18332/tid/140824

36. Kim SH, Jeong SH, Park EC, Jang SI. Association of cigarette type initially smoked with suicidal behaviors among adolescents in Korea from 2015 to 2018. *JAMA Netw Open*. 2021;4(4):e218803. doi:10.1001/jamanetworkopen.2021.8803

37. Striley CW, Nutley SK, Hoeflich CC. E-cigarettes and non-suicidal self-injury: prevalence of risk behavior and variation by substance inhaled. *Front Psychiatry*. 2022;13:911136. doi:10.3389/fpsyt.2022.911136

38. Javed S, Usmani S, Sarfraz Z, et al. A scoping review of vaping, e-cigarettes and mental health impact: depression and suicidality. *J Community Hosp Intern Med Perspect*. 2022;12(3):33-39. doi:10.55729/2000-9666.1053

39. Jacobs W, Orozco G, Villanueva G, Merianos AL. E-cigarette and cannabis use patterns, depression, and suicide behaviors among US youth: analysis of 2019 Youth Risk Behavior Survey data. *Am J Health Promot*. 2023;37 (1):77-83. doi:10.1177/08901171221112927

40. Swann AC, Graham DP, Wilkinson AV, Kosten TR. Nicotine inhalation and suicide: clinical correlates and behavioral mechanisms. *Am J Addict*. 2021;30(4):316-329. doi:10.1111/ajad.13171

41. Harrison R, Munafò MR, Davey Smith G, Wootton RE. Examining the effect of smoking on suicidal ideation and attempts: triangulation of epidemiological approaches. *Br J Psychiatry*. 2020;217(6):701-707. doi:10.1192/bjp. 2020.68

42. Orri M, Chadi N, Ahun MN, et al. Suicidal ideation and attempt in adolescents exposed to maternal smoking across pregnancy and childhood: a 20-year prospective cohort study. *J Affect Disord*. 2021;286:10-18. doi:10.1016/j.jad.2021.02.047

43. O'Connor RC, Kirtley OJ. The integrated motivational-volitional model of suicidal behaviour. *Philos Trans R Soc Lond B Biol Sci.* 2018;373(1754):20170268. doi:10.1098/rstb.2017.0268

44. Klonsky ED, Saffer BY, Bryan CJ. Ideation-to-action theories of suicide: a conceptual and empirical update. *Curr Opin Psychol.* 2018;22:38-43. doi:10.1016/j.copsyc.2017.07.020

45. Park S, Romer D. Associations between smoking and depression in adolescence: an integrative review. *J Korean Acad Nurs*. 2007;37(2):227-241. doi:10.4040/jkan.2007.37.2.227

46. Paperwalla KN, Levin TT, Weiner J, Saravay SM. Smoking and depression. *Med Clin North Am*. 2004;88(6): 1483-1494, x-xi. doi:10.1016/j.mcna.2004.06.007

47. Schneider B, Lukaschek K, Baumert J, Meisinger C, Erazo N, Ladwig KH. Living alone, obesity, and smoking increase risk for suicide independently of depressive mood findings from the population-based MONICA/KORA Augsburg cohort study. *J Affect Disord*. 2014;152-154:416-421. doi:10.1016/j.jad.2013.10.007

48. Lucas M, O'Reilly EJ, Mirzaei F, et al. Cigarette smoking and completed suicide: results from 3 prospective cohorts of American adults. *J Affect Disord*. 2013;151(3):1053-1058. doi:10.1016/j.jad.2013.08.033

49. Orri M, Geoffroy MC, Turecki G, et al. Contribution of genes and environment to the longitudinal association between childhood impulsive-aggression and suicidality in adolescence. *J Child Psychol Psychiatry*. 2020;61(6): 711-720. doi:10.1111/jcpp.13163

50. Esposito-Smythers C, Spirito A. Adolescent substance use and suicidal behavior: a review with implications for treatment research. *Alcohol Clin Exp Res*. 2004;28(5)(suppl):77S-88S. doi:10.1097/01.ALC.0000127417.99752.87

51. Micalizzi L, Knopik VS. Maternal smoking during pregnancy and offspring executive function: what do we know and what are the next steps? *Dev Psychopathol*. 2018;30(4):1333-1354. doi:10.1017/S0954579417001687

52. Harden KP, Tucker-Drob EM. Individual differences in the development of sensation seeking and impulsivity during adolescence: further evidence for a dual systems model. *Dev Psychol*. 2011;47(3):739-746. doi:10.1037/a0023279

53. Steinberg L, Graham S, O'Brien L, Woolard J, Cauffman E, Banich M. Age differences in future orientation and delay discounting. *Child Dev.* 2009;80(1):28-44. doi:10.1111/j.1467-8624.2008.01244.x

54. Kasen S, Cohen P, Chen H. Developmental course of impulsivity and capability from age 10 to age 25 as related to trajectory of suicide attempt in a community cohort. *Suicide Life Threat Behav.* 2011;41(2):180-192. doi:10.1111/j. 1943-278X.2011.00017.x

55. McGirr A, Renaud J, Bureau A, Seguin M, Lesage A, Turecki G. Impulsive-aggressive behaviours and completed suicide across the life cycle: a predisposition for younger age of suicide. *Psychol Med.* 2008;38(3):407-417. doi:10. 1017/S0033291707001419

56. Miller AB, Prinstein MJ. Adolescent suicide as a failure of acute stress-response systems. *Annu Rev Clin Psychol*. 2019;15:425-450. doi:10.1146/annurev-clinpsy-050718-095625

57. Rudd MD. Fluid vulnerability theory: a cognitive approach to understanding the process of acute and chronic suicide risk. In: Ellis TE, ed. *Cognition and Suicide: Theory, Research, and Therapy*. American Psychological Association; 2006:355-368. doi:10.1037/11377-016

58. Bals R, Boyd J, Esposito S, et al. Electronic cigarettes: a task force report from the European Respiratory Society. *Eur Respir J*. 2019;53(2):1801151. doi:10.1183/13993003.01151-2018

59. Orri M, Séguin JR, Castellanos-Ryan N, et al. A genetically informed study on the association of cannabis, alcohol, and tobacco smoking with suicide attempt. *Mol Psychiatry*. 2021;26(9):5061-5070. doi:10.1038/s41380-020-0785-6

SUPPLEMENT 1.

eAppendix. Discussion of Hypothesized Neurobiological Pathways and Shared Risk Factors Linking Smoking and Suicidal Behaviors

eFigure 1. Number of Study Participants who Endorsed Suicide Attempts (SAs), Ideation (SI), and Non-Suicidal

Self-Injuries (NSSI) from the KSADS-5, and Case Group Classifications Used in the Present Study

eFigure 2. Flowchart for Study Participant Selection

eFigure 3. Association Between the Use of Tobacco Products (UTPs) and Suicide Risk Outcomes Based on the Basic Model

eTable 1. List of 22 ABCD Study Enrollment Sites

eTable 2. Details of Children's Tobacco Use Outcome Measures

eTable 3. Details of KSADS-5 Variables Used for Constructing Lifetime Suicide Risk and Self-Injury Outcome Measures

eTable 4. Response Rates for UTP and SITB Measures

eTable 5. Data on Sociodemographics, Family History, and Prenatal Exposure to Substance Use

eTable 6. Child Behavioral Outcome Measures

eTable 7. Details of Children's Substance Use Measures

eTable 8. Major Characteristics of ABCD Study Participants Based on Youth-Reported Ever Use of Tobacco Products in the 6-Month Follow-Up

eTable 9. Major Characteristics of ABCD Study Participants Based on Youth-Reported Ever Use of Tobacco Products in the 18-Month Follow-Up

eTable 10. Association Between UTPs and Suicide Risk Outcomes Based on the Basic Model

eTable 11. Association Between UTPs and Suicide Risk Outcomes Based on the All Covariate-Adjusted Model

eTable 12. Association Between UTPs and KSADS-5 Youth-Report-Based Suicide Risk Outcomes Based on the All Covariate-Adjusted Model

eTable 13. Association Between UTPs and KSADS-5 Parent-Report-Based Suicide Risk Outcomes Based on the All Covariate-Adjusted Model

eTable 14. Association Between UTPs and Suicide Risk Outcome Measures Concordant Between KSAS-5 Youthand Parent-Reports Assessed in the All Covariate-Adjusted Model

eTable 15. Associations of the Use of Tobacco Products (UTPs) and New SA Cases, While Adjusting for Various Confounding Factors

eTable 16. Multivariate Logistic Regression Results for Predicting Suicide Attempts Using UTPs (Both Lifetime Measures Assessed in the Baseline)

eTable 17. Multivariate Logistic Regression Results for Predicting Suicide Attempts Using UTPs (UTPs Assessed in 6-Month Follow-Up, SA Assessed in Year 1 Follow-Up)

eTable 18. Multivariate Logistic Regression Results for Predicting Suicidal Ideation Using UTPs (UTPs and SI Both Assessed in Baseline)

eTable 19. Multivariate Logistic Regression Results for Predicting Suicidal Ideation Using UTPs (UTPs Assessed in 6-Month Follow-Up, SI Assessed in Year 1 Follow-Up)

eTable 20. Multivariate Logistic Regression Results for Predicting Suicidal Ideation using UTPs (UTPs Assessed in 18-Month Follow-Up, SI Assessed in Year 2 Follow-Up)

eTable 21. Interaction Analysis Results of UTP at Baseline with Children's Cognition, Temperament/Personality, and Psychopathology Measures on SA.

eTable 22. Multivariate Logistic Regression Results for Predicting Suicide Attempts (SAs) While Integrating Multiple Types of Substance Use Data in Addition to Smoking Tobacco Products

SUPPLEMENT 2.

Data Sharing Statement