

Journal Pre-proof

Air pollution and age-dependent changes in emotional behavior across early adolescence in the U.S.

Claire E. Campbell, Devyn L. Cotter, Katherine L. Bottenhorn, Elisabeth Burnor, Hedyeh Ahmadi, W. James Gauderman, Carlos Cardenas-Iniguez, Daniel Hackman, Rob McConnell, Kiros Berhane, Joel Schwartz, Jiu-Chiuan Chen, Megan M. Herting

PII: S0013-9351(23)02194-1

DOI: <https://doi.org/10.1016/j.envres.2023.117390>

Reference: YENRS 117390

To appear in: *Environmental Research*

Received Date: 4 April 2023

Revised Date: 24 August 2023

Accepted Date: 11 October 2023

Please cite this article as: Campbell, C.E., Cotter, D.L., Bottenhorn, K.L., Burnor, E., Ahmadi, H., Gauderman, W.J., Cardenas-Iniguez, C., Hackman, D., McConnell, R., Berhane, K., Schwartz, J., Chen, J.-C., Herting, M.M., Air pollution and age-dependent changes in emotional behavior across early adolescence in the U.S., *Environmental Research* (2023), doi: <https://doi.org/10.1016/j.envres.2023.117390>.

This is a PDF file of an article that has undergone enhancements after acceptance, such as the addition of a cover page and metadata, and formatting for readability, but it is not yet the definitive version of record. This version will undergo additional copyediting, typesetting and review before it is published in its final form, but we are providing this version to give early visibility of the article. Please note that, during the production process, errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

© 2023 Published by Elsevier Inc.



Title: Air pollution and age-dependent changes in emotional behavior across early adolescence in the U.S.

Claire E. Campbell^{1,2}, Devyn L. Cotter^{1,2}, Katherine L. Bottenhorn^{1,3}, Elisabeth Burnor¹, Hedyeh Ahmadi¹, W. James Gauderman¹, Carlos Cardenas-Iniguez¹, Daniel Hackman⁴, Rob McConnell¹, Kiros Berhane⁵, Joel Schwartz⁶, Jiu-Chiuan Chen^{1,7}, Megan M. Herting^{1,8}

¹ Department of Population and Public Health Sciences, University of Southern California, Los Angeles, CA, USA

² Neuroscience Graduate Program, University of Southern California, Los Angeles, California, USA 90089-2520

³ Department of Psychology, Florida International University, Miami, FL, USA

⁴ Suzanne Dworak-Peck School of Social Work, University of Southern California, Los Angeles, CA 90089

⁵ Department of Biostatistics, Mailman School of Public Health, Columbia University, New York, NY 10032, USA

⁶ Department of Environmental Health, Harvard T.H. Chan School of Public Health, Boston, MA 02115, USA

⁷ Department of Neurology, Keck School of Medicine of University of Southern California, Los Angeles, CA 90063, USA

⁸ Children's Hospital Los Angeles, Los Angeles, CA 90027, USA

Credit Author Statement

Claire E. Campbell: Project Administration, Methodology, Formal Analysis, Visualization, Writing - Original Draft; Devyn L. Cotter: Methodology, Writing - Original Draft; Katherine L. Bottenhorn: Methodology, Visualization, Writing – Original Draft; Elisabeth Burnor: Methodology, Data Curation, Writing – Review & Editing; Hedyeh Ahmadi: Data Curation, Methodology, Writing – Review & Editing; Carlos Cardenas-Iniguez: Writing - Review & Editing; W. James Gauderman: Methodology, Writing - Review & Editing; Rob McConnell: Methodology, Writing - Review & Editing; Kiros Berhane: Methodology, Writing - Review & Editing; Joel Schwartz: Methodology, Data Curation, Resources, Writing - Review & Editing; Jiu-Chiuan Chen: Conceptualization, Methodology, Writing - Review & Editing; Megan M. Herting: Funding Acquisition, Resources, Conceptualization, Methodology, Supervision, Project Administration, Writing - Original Draft.

1 **Title:** Air pollution and age-dependent changes in emotional behavior across early adolescence in the
2 U.S.

3 Claire E. Campbell^{1,2}, Devyn L. Cotter^{1,2}, Katherine L. Bottenhorn^{1,3}, Elisabeth Burnor¹, Hedyeh Ahmadi¹,
4 W. James Gauderman¹, Carlos Cardenas-Iniguez¹, Daniel Hackman⁴, Rob McConnell¹, Kiros Berhane⁵,
5 Joel Schwartz⁶, Jiu-Chiuan Chen^{1,7}, Megan M. Herting^{1,8}

6 ¹ Department of Population and Public Health Sciences, University of Southern California, Los Angeles,
7 CA, USA

8 ² Neuroscience Graduate Program, University of Southern California, Los Angeles, California,
9 USA 90089-2520

10 ³ Department of Psychology, Florida International University, Miami, FL, USA

11 ⁴ Suzanne Dworak-Peck School of Social Work, University of Southern California, Los Angeles, CA
12 90089

13 ⁵ Department of Biostatistics, Mailman School of Public Health, Columbia University, New York, NY
14 10032, USA

15 ⁶ Department of Environmental Health, Harvard T.H. Chan School of Public Health, Boston, MA 02115,
16 USA

17 ⁷ Department of Neurology, Keck School of Medicine of University of Southern California, Los Angeles,
18 CA 90063, USA

19 ⁸ Children's Hospital Los Angeles, Los Angeles, CA 90027, USA

20 **Corresponding Author**

21 Correspondence to Megan M. Herting. University of Southern California, 1845 N. Soto Rm 225N, Los
22 Angeles, California, 90089.

23 **Abstract**

24 Recent studies have linked air pollution to increased risk for behavioral problems during development,
25 albeit with inconsistent findings. Additional longitudinal studies are needed that consider how emotional
26 behaviors may be affected when exposure coincides with the transition to adolescence – a vulnerable
27 time for developing mental health difficulties. This study investigates if annual average PM_{2.5} and NO₂
28 exposure at ages 9-10 years moderates age-related changes in internalizing and externalizing behaviors
29 over a 2-year follow-up period in a large, nationwide U.S. sample of participants from the Adolescent
30 Brain Cognitive Development (ABCD) Study®. Air pollution exposure was estimated based on the
31 residential address of each participant using an ensemble-based modeling approach. Caregivers
32 answered questions from the Child Behavior Checklist (CBCL) at the baseline, 1-year follow-up, and 2-
33 year follow-up visits, for a total of 3 waves of data; from the CBCL we obtained scores on internalizing
34 and externalizing problems plus 5 syndrome scales (anxious/depressed, withdrawn/depressed, rule-
35 breaking behavior, aggressive behavior, and attention problems). Zero-inflated negative binomial models
36 were used to examine both the main effect of age as well as the interaction of age with each pollutant on
37 behavior while adjusting for various socioeconomic and demographic characteristics. Against our
38 hypothesis, there was no evidence that greater air pollution exposure was related to more behavioral
39 problems with age over time.

40 **Keywords:** air pollution; internalizing; externalizing; adolescence; neurodevelopment

41 1. Introduction

42 Mental health conditions remain a global health challenge for all age groups (World Health Organization,
43 2018), but the risk for onset of psychopathology is highest in childhood and adolescence. Both
44 internalizing and externalizing symptoms typically emerge during adolescence (Achenbach et al., 1991).
45 Moreover, up to approximately 20% of children and adolescents are affected by mental health problems
46 worldwide (Polanczyk et al., 2015) with half of all lifetime mental health conditions diagnosed by age 14
47 years (Kessler et al., 2005). To reduce societal costs and improve quality of life for affected individuals,
48 research on modifiable risk and resilience factors holds the promise to potentially uncover new avenues
49 for early prevention and intervention.

50 Recent evidence indicates that outdoor air pollution may contribute to increased risk for mental health
51 conditions (Braithwaite et al., 2019; Zundel et al., 2022). A growing body of literature has associated
52 ambient exposure to fine particulate matter (PM_{2.5}) and nitrogen dioxide (NO₂) with mental health
53 outcomes in children, adolescents, and adults, including symptoms of anxiety, depression, and
54 aggression in children, as well as increased risk for attention deficit and hyperactivity disorder (ADHD)
55 and delinquency problems (Forns et al., 2016; Margolis et al., 2016; Newman et al., 2013; F. Perera et
56 al., 2016; F. P. Perera et al., 2014; Thygesen et al., 2020; Yorifuji et al., 2016, 2017). However, recent
57 comprehensive reviews of how air pollution relates to anxiety and depression (Zundel et al., 2022) and
58 attention problems (Myhre et al., 2018) highlight a number of inconsistencies and important knowledge
59 gaps in the broader air pollution and mental health literature. For example, while most studies found
60 positive associations between air pollution exposure and anxiety and depression, 25% of studies did not
61 find associations or reported mixed effects, and fewer than 10% examined air pollution exposure during
62 the susceptible window of childhood and adolescence. Even within the developmental literature, there
63 are mixed results: some studies suggest prenatal through adolescent exposure is linked to more
64 internalizing problems (Brokamp et al., 2019; Brunst et al., 2019; B. Fan et al., 2019; Margolis et al.,
65 2016; Rasnick et al., 2021; Yolton et al., 2019), where others have failed to find an association (Jorcano
66 et al., 2019; Zhao et al., 2019). Thus, additional research is needed to clarify the relationship between air
67 pollution exposure and mental health behaviors during the developmental periods of childhood and
68 adolescence.

69 Importantly, most previous studies have been limited to cross-sectional assessment of mental health
70 outcomes and/or are limited in terms of both the geographic and sociodemographic diversity of their study
71 sample. Discrepancies in results may also be due in part to both differences in the window of exposure
72 as well as the timing of the behavioral evaluation, especially given the known developmental patterns in
73 symptom onset. For example, a recent study of 8 European cohorts found that neither prenatal nor early
74 life exposure was related to cross-sectional outcomes of depression, anxiety, or aggression behavior
75 using the Child Behavior Checklist (CBCL) or Strengths and Difficulties Questionnaire (SDQ) when
76 assessed in mid-to-late childhood (Jorcano et al., 2019); however, this study was limited by examining
77 only one time point of internalizing and externalizing behaviors. Given that the frequency and intensity of
78 emotional problems vary across childhood and adolescence (Barch et al., 2021), it is important to
79 consider how air pollution exposure may contribute to mental health problems over time. For example,
80 Roberts and colleagues (Roberts et al., 2019) found that while exposure to higher levels of PM_{2.5} and
81 NO₂ at age 12 years was not associated with concurrent mental health conditions, it did successfully
82 predict a 1.5 fold increased risk for developing major depressive disorder at 18 years of age. Similarly,

83 Reuben and colleagues (2021) found that exposure to nitrous oxides (NO_x) in childhood predicted later
84 onset of internalizing, externalizing, and thought disorder symptoms at age 18 years. The latter two
85 studies suggest that exposure during childhood and adolescence may lead to greater mental health
86 problems overtime, emphasizing the importance in assessing emotional and behavioral problems
87 longitudinally. Thus, additional large-scale longitudinal studies are warranted to further investigate if
88 exposure in late childhood may moderate changes in emotional behavior problems as individuals
89 transition to early adolescence.

90 Beyond the considerations of timing of the exposure and longitudinal assessment of the outcome,
91 questions also remain as to the potential health effects of air pollution below the current air quality
92 standards. That is, despite significant declines in air pollution, recent epidemiological studies continue to
93 find links between adverse health effects and levels of exposure well below the Environmental Protection
94 Agency (EPA)'s specified annual averages for PM_{2.5} ($\leq 12 \mu\text{g}/\text{m}^3$) and NO₂ ($\leq 53 \text{ parts per billion; ppb}$)
95 (Dominici et al., 2019). This emerging body of research showing that negative health effects can be
96 observed at low concentrations of exposure suggests that no observable threshold may be considered
97 "safe" (for review, see Papadogeorgou et al., 2019). As such, the World Health Organization (WHO)
98 updated their regulatory guidelines on ambient air quality in September 2021, recommending that annual
99 averages of PM_{2.5} and NO₂ not exceed $5 \mu\text{g}/\text{m}^3$ and $10 \mu\text{g}/\text{m}^3$ (equivalent to 5.33 ppb) respectively (World
100 Health Organization, 2021). Previous literature linking behavioral problems with ambient air pollution
101 observed relationships with exposure levels that largely exceed both current EPA standards and WHO
102 recommendations. Thus, further research is necessary to investigate to what degree lower levels of
103 exposure seen across the U.S. may influence developmental changes in emotional behavior in today's
104 youth.

105 Leveraging the large (N=11,876), nationwide, and socio-demographically and geographically diverse
106 Adolescent Brain Cognitive Development (ABCD) Study[®] cohort (Jernigan et al., 2018), the current study
107 aimed to examine whether air pollution exposure at ages 9-10 years may relate to longitudinal changes
108 in behavioral and emotional problems over a 2-year follow-up period. The ABCD Study[®] comprises 21
109 study sites across the United States and implements an identical protocol for recruitment and data
110 collection of all participants (Garavan et al., 2018; Lisdahl et al., 2018). Given the extant literature (Cory-
111 Slechta et al., 2023; Reuben et al., 2021; Roberts et al., 2019; Zundel et al., 2022) and also limited
112 availability of ABCD exposure data (C. C. Fan et al., 2021), our study focused on one-year annual
113 average PM_{2.5} and NO₂ exposures at ages 9-10 years. With relatively low concentrations of PM_{2.5} and
114 NO₂ in the ABCD Study (Cserbik et al., 2020; C. C. Fan et al., 2021), the current study aims to address
115 links between air pollution and mental health in those regularly exposed to concentrations largely below
116 EPA standards. Given previous findings (Brokamp et al., 2019; Brunst et al., 2019; B. Fan et al., 2019;
117 Margolis et al., 2016; Rasnick et al., 2021; Yolton et al., 2019), we *a priori* chose to examine internalizing
118 and externalizing summary scores from the Child Behavior Checklist (CBCL), as well as distinct
119 internalizing syndrome subscales of anxious/depressed and withdrawn/depressed, and externalizing
120 syndrome subscales of rule-breaking and aggressive behavior, and the independent subscale of
121 attention, thus addressing a wide range of internalizing and externalizing behaviors. We hypothesized
122 that higher exposure levels during late childhood would predict more emotional problems over the 2-year
123 follow-up period.

124 2. Materials and methods

125 2.1 Study Design and Participants

126 The current study utilized data from the larger ongoing nationwide ABCD Study® (NDA 4.0 data release
127 2021, <https://abcdstudy.org/scientists/data-sharing/>), which enrolled over 11,876 9- and 10-year-old
128 participants across the USA from 2016-2018 with plans to follow subjects for up to 10 years (Garavan et
129 al., 2018; Jernigan et al., 2018; Volkow et al., 2018). The 21 study sites obtained approval from their local
130 Institutional Review Board (IRB) and a centralized IRB approval was obtained from the University of
131 California, San Diego. Written informed consent was provided by each child's parent or legal guardian
132 (hereafter, "caregiver"); each child provided verbal assent. All ethical regulations were complied with
133 during data collection and analysis. Primary inclusion criteria for ABCD Study participants included age
134 (9.0 to 10.99 years at baseline visit), fluency in English, and the ability to complete the baseline visit. For
135 the current analysis, we utilized data from the first 3 waves of annual data collection, with the additional
136 inclusion criteria of having a valid primary residential address at baseline for all subjects. Given both the
137 extant literature on potential neurotoxic and mental health effects (Cory-Slechta et al., 2023; Reuben et
138 al., 2021; Roberts et al., 2019; Zundel et al., 2022) and data availability (C. C. Fan et al., 2021), the
139 current study focused on investigating both PM_{2.5} and NO₂ exposures. Given the distribution of the
140 outcome data and analytic approach required for hypothesis testing (see section 2.5 below), complete
141 predictors were required for each wave of data collection. A flowchart of our target population for our
142 analyses can also be found in **Supplemental Figure 1**. We also selected data collected before March 1,
143 2020, to avoid any potential confounding effects of stress on mental health outcomes related to the
144 COVID-19 pandemic (Hamatani et al., 2022; Kiss et al., 2022; Yip et al., 2022). Lastly, we randomly
145 selected one subject per family to reduce the hierarchical structure of our data from 4 levels (time point,
146 subject, family, site) to 3 levels (time point, subject, site). This resulted in a final sample of 9,273 unique
147 participants: 9,271 for baseline, 8,759 for 1-year follow-up, and 5,827 for 2-year follow-up (**Table 1**). A
148 comparison of the overall ABCD cohort with our final analytical sample at baseline, 1-year, and 2-year
149 follow-up can be found in **Supplemental Table 1**, **Supplemental Table 2**, and **Supplemental Table 3**,
150 respectively. All variable names used in the following analyses are documented in **Supplemental Table**
151 **4**.

152 **Table 1. Demographics**

	Baseline	1-year follow-up	2-year follow-up
N	9271	8759	5827
Sex assigned at birth			
Female	4413 (47.6%)	4154 (47.4%)	2747 (47.1%)
Male	4858 (52.4%)	4605 (52.6%)	3080 (52.9%)
Age at data collection			
Mean (SD)	9.91 (0.62)	10.91 (0.63)	11.92 (0.64)
Range	8.92 - 11.08	9.75 - 12.42	10.58 - 13.67
Race/ethnicity			
Black	1363 (14.7%)	1221 (13.9%)	678 (11.6%)
Hispanic	1958 (21.1%)	1800 (20.6%)	1198 (20.6%)
Other	1191 (12.8%)	1126 (12.9%)	715 (12.3%)
White	4759 (51.3%)	4612 (52.7%)	3236 (55.5%)
Caregiver education			
< HS Diploma	460 (5.0%)	405 (4.6%)	257 (4.4%)
HS Diploma/GED	899 (9.7%)	800 (9.1%)	460 (7.9%)
Some College	2423 (26.1%)	2247 (25.7%)	1466 (25.2%)
Bachelor	2310 (24.9%)	2217 (25.3%)	1551 (26.6%)
Post Graduate Degree	3179 (34.3%)	3090 (35.3%)	2093 (35.9%)
Caregiver employment			
Employed	6444 (69.5%)	6146 (70.2%)	4139 (71.0%)
Stay at Home Parent	1612 (17.4%)	1516 (17.3%)	1002 (17.2%)
Unemployed	539 (5.8%)	481 (5.5%)	299 (5.1%)
Other	676 (7.3%)	616 (7.0%)	387 (6.6%)
Neighborhood safety			
Mean (SD)	3.873 (0.976)	3.884 (0.971)	3.915 (0.948)
Range	1.000 - 5.000	1.000 - 5.000	1.000 - 5.000
Household income			
<\$50k	2564 (27.7%)	2335 (26.7%)	1597 (27.4%)
≥\$50K & <\$100K	2419 (26.1%)	2312 (26.4%)	1597 (27.4%)
≥\$100K	3514 (37.9%)	3418 (39.0%)	2307 (39.6%)
Don't know or refuse	774 (8.3%)	694 (7.9%)	429 (7.4%)

153 Demographic composition of the final sample across three waves of data collection.

154

155 2.2 Estimation of Annual Air Pollution Exposure

156 Details regarding the collection of residential addresses and linkage to one-year annual average ambient
157 PM_{2.5} and NO₂ have been previously published in detail by Fan and colleagues (2021). Briefly, daily
158 pollutant estimates were derived at a 1-km² resolution using hybrid spatiotemporal models that combine
159 satellite-based aerosol optical depth models, land-use regression, and chemical transport models (Di et
160 al., 2019, 2020). The cross-validation of these models with EPA monitored levels across the U.S. were
161 found to perform well, with R² Root Mean Square Error of 0.89 for PM_{2.5} annual averages and 0.84 for
162 NO₂ annual averages (Di et al., 2019, 2020). These daily estimates were then averaged over the 2016
163 calendar year, when the children were aged 9-10 years-of-age and assigned to the geocoded primary
164 residential address at the baseline ABCD study visit. PM_{2.5} is reported in micrograms per meter cubed
165 ($\mu\text{g}/\text{m}^3$) and NO₂ is reported in parts per billion (*ppb*). For subjects who have data indicating their time
166 lived at baseline address (N=9,027), the mean was 5.4 years (standard deviation = 3.75). Average yearly
167 consistency of spatial contrast for each pollutant based on daily estimates at the 1-km² resolution is also
168 presented in the supplement (**Supplemental Figure 2**), as well as the variability in air pollution estimates
169 across ABCD participants by site (**Supplemental Figure 3**).

170 2.3 Emotional Behavior

171 At each annual visit (baseline, 1-year follow-up, 2-year follow-up), the participant's caregiver was asked
172 to report on the child's emotional behavior over the 6 months prior to each study visit using the Child
173 Behavioral Checklist (CBCL) (Achenbach, 2009; Achenbach & Rescorla, 2001). The CBCL within the
174 ABCD Study has 112 different items that each caregiver answers about their child (e.g., "Show little
175 interest in things around him/her") using a 3-point Likert-type scale (0 = Not True, 1 = Somewhat or
176 Sometimes True, 2 = Very True or Often True). These answers are then used to calculate summary
177 scores of internalizing and externalizing behaviors. Based on the prior air pollution and behavioral
178 literature (Brokamp et al., 2019; Brunst et al., 2019; B. Fan et al., 2019; Margolis et al., 2016; Rasnick et
179 al., 2021; Yolton et al., 2019; Zundel et al., 2022), we also chose to examine five additional syndrome
180 subscale scores: anxious/depressed, withdrawn/depressed, rule-breaking behavior, aggressive
181 behavior, and attention problems. Anxious/depressed and withdrawn/depressed subscales fall within the
182 internalizing score, rule-breaking and aggressive behavior subscales fall within the externalizing score,
183 and attention is an independent subscale. Each raw score is a whole number with higher integers
184 indicating increased problem or emotional behaviors, across syndrome scores, such that syndrome score
185 is each on the same scale. While there are age- and sex-normalized scores, we chose to utilize the raw
186 scores to allow us to investigate developmental changes in these behaviors with age, as has been
187 previously done when examining ABCD Study data (Barch et al., 2021). Importantly, the CBCL measures
188 show good test-retest reliability (Pearson's correlations mean = 0.9, min=0.82, max=0.94) and internal
189 consistency was stable over a 12- and 24-month period (Pearson's correlation 12-month mean = 0.74,
190 24-month mean = 0.70) (Achenbach & Rescorla, 2001). The cross-informant agreement between parent
191 and youth using CBCL items has been found to be strong across multi-cultural societies [Q correlations
192 for U.S.= .84] and similar across internalizing and externalizing behaviors (Rescorla et al., 2013). The
193 raw scores have different ranges by subscale: internalizing [0,64], externalizing [0,70],
194 anxious/depressed [0,26], withdrawn/depressed [0,16], rule-breaking [0,34], aggressive [0,36], and
195 attention [0,20], but all subscales use the same Likert scale units, with higher values indicative of greater
196 problems.

197 2.4 Confounders and Covariates

198 We have selected potential confounders based on both prior knowledge and current theories in
199 environmental epidemiology using a directed acyclic graph (Greenland & Brumback, 2002)
200 (**Supplemental Figure 4**). Specifically, we identified confounders that may predict emotional behavior
201 and exposure to ambient air pollutants. All of these variables were reported by the child's caregiver using
202 the PhenX Toolkit (Echeverria et al., 2004; Mujahid et al., 2007). This list includes child's sex,
203 race/ethnicity (non-hispanic white, hispanic, non-hispanic black, other: includes American Indian/Native
204 American, Alaska Native, Native Hawaiian, Guamanian, Samoan, Other Pacific Islander, Asian Indian,
205 Chinese, Filipino, Japanese, Korean, Vietnamese, or Other Race not listed), indicators of family
206 socioeconomic status (e.g., highest caregiver educational attainment, caregiver's employment status,
207 combined total annual household income), as well as perceived neighborhood quality. Highest caregiver
208 educational attainment included <high school diploma, high school diploma or GED, some college,
209 bachelor's degree, or postgraduate degree. Caregiver's employment status included employed (part- or
210 full-time), stay at home parent, unemployed, or other (e.g., temporarily laid off; sick leave; retired;
211 disabled, etc.). Combined total annual household income included less than or equal to \$50,000, greater
212 than \$50,000 but less than \$100,000, greater or equal to \$100,000, or don't know/refuse to answer.
213 Perceived neighborhood quality was an average score of three-items assessing parent perspectives of
214 how safe and free from crime and violence they felt their neighborhood is (Mujahid et al. 2007). Each of
215 these variables' baseline values were used in the model, to align with the timing of the available ambient
216 air pollution estimates. To account for potential confounding of co-exposure, we also included the other
217 air pollutant as an additional variable (i.e., when examining the influence of PM_{2.5}-by-age on CBCL
218 outcomes, NO₂ is added to the model, and vice versa). Importantly, multicollinearity was not an issue in
219 adjusting for the other pollutant in the model as the Pearson correlation coefficient between the baseline
220 annual pollutant concentrations of PM_{2.5} and NO₂ across all sites was low ($r = 0.22$).

221

222 2.5 Analyses

223 All statistical analyses were implemented in R (version 4.1.2) (R Core Team, 2021). Initial descriptive
224 and exploratory analysis were conducted to check all data for potential errors and outliers, and to assess
225 variable distributions required to satisfy modeling assumptions and understand correlations. To
226 investigate how annual PM_{2.5} and NO₂ moderate emotional development of adolescents over 3 visits
227 spaced 1-year apart, we used a multilevel (i.e., mixed effects) modeling approach to account for the
228 repeated measures. We verified our models were appropriate by checking model assumptions post
229 analyses based on prior published methodology (Cameron & Trivedi, 2013; Garay et al., 2011; Hilbe,
230 2011).

231 *2.5.1 Reasoning for Modeling Choice*

232 While previous cross-sectional studies examining CBCL outcomes and air pollution have examined the
233 CBCL t-scores (F. P. Perera et al., 2011, 2012), raw CBCL scores are required to better account for
234 developmental changes in emotional behaviors over time when using a repeated measures design
235 (Barch et al., 2021). Furthermore, when multilevel approaches are required, a common modeling
236 approach is to use a linear mixed-effects model, but since CBCL outcomes are naturally zero-inflated
237 (**Supplemental Figure 5**) and thus over-dispersed (over-dispersion quotient ranges from 2.84-17.29),
238 this can lead to artificial inflation of the coefficients' significance (LAND et al., 1996; Stroup, 2016;

239 Swartout et al., 2015). Thus, we utilized CBCL raw scores as count data and employed a zero-inflated
240 negative binomial (ZINB) model, which adds an extra parameter that accounts for the over-dispersion
241 present (Xu et al., 2017). The *glmm.zinb()* function was used within the NBZIMM package (version 1.0)
242 (<https://github.com/nyiuab/NBZIMM>); a manuscript detailing the development of this package was also
243 published (Zhang & Yi, 2020). This modeling approach has been used in numerous studies with zero-
244 inflated health data (Preisser et al., 2016; Sheu et al., 2004), and specifically when examining mental
245 health outcomes (Kumagai et al., 2021; Vyas et al., 2020). For even further reading on the ZINB
246 approach, we have cited additional readings (Fang et al., 2016; Stroup, 2012; Yau et al., 2003; Young et
247 al., 2022; Zhang & Yi, 2020).

248 2.5.2 Final dataset for ZINB model

249 To implement the ZINB model, complete predictors across timepoints (i.e., no missing values for each
250 subject at each wave of data collection) is required; therefore, listwise deletion was used to remove
251 incomplete data by wave of data collection, making sure at each wave of data collection, each subject
252 had a CBCL outcome score, age at session, race/ethnicity, sex at birth, PM_{2.5} level at year of baseline
253 visit, NO₂ level at year of baseline visit, caregiver's highest level of education at baseline, caregiver's
254 employment status at baseline, perceived neighborhood safety at baseline, and household income at
255 baseline. Since we only had data for our main predictors – PM_{2.5} and NO₂ – at the baseline visit, our
256 environmental covariates and confounders were also only from the baseline visit. Following our initial
257 cleaning steps, creating a dataset across timepoints with complete predictors led to 6%, 5%, and 3% of
258 missing data for the baseline, 1-year, and 2-year follow-up visits, respectively. Bennett (2001) states that
259 greater than 10% missingness could lead to bias within the statistical analysis and prior published
260 literature suggest 5% (on average) missingness is negligible (Jakobsen et al., 2017; Schafer, 1999).
261 Therefore, given the limited amount of missing data, we chose not to perform multiple imputation.

262 2.5.3 Age-only ZINB models

263 For the main analysis, the ZINB model combines two models: 1) zero-inflated model, similar to a logistic
264 regression, that evaluates the likelihood of being in the certain-zero (i.e., no problems) as compared to
265 the non-zero category (i.e., exhibits problems), and 2) count model, assuming a negative binomial
266 distribution, that evaluates the non-zero CBCL subscale scores (i.e., magnitude of problems). Initially,
267 age-only models were performed to establish changes in CBCL outcomes from baseline and two 1-year
268 follow-up periods. These models investigated the main effect of age on each CBCL outcome controlling
269 for necessary covariates in both the zero-inflated and count portions of the model (sex-at-birth,
270 race/ethnicity, highest caregiver educational attainment at baseline, caregiver's employment status at
271 baseline, perceived neighborhood safety at baseline, and the combined total annual household income
272 at baseline). For the random effects within the zero-inflated portion of the model, we only included ABCD
273 site since subjects within the certain-zero group were not strongly clustered by subject (low intraclass
274 correlation coefficients (ICC) for all CBCL outcomes: 0.070-0.107). For the random effects within the
275 count model, subject was nested within a random effect of site to account for the within-subject similarities
276 over time in those with non-zero data (ICCs ranging from 0.506-0.710; this medium-high ICC implies a
277 clustering structure of subject within the non-zero data). For ease of interpretation, age was centered at
278 9 years, the youngest integer age in our cohort.

279 2.5.4 Age-by-air pollutant ZINB models

280 To investigate if air pollution modifies emotional problems over time, we added in an interaction between
281 age and each air pollutant (PM_{2.5} or NO₂). For both the zero-inflated model and the count model we
282 utilized the fixed effects of each pollutant (PM_{2.5} or NO₂), age, pollutant-by-age, while adjusting for the
283 same potential confounders as mentioned above; each pollution-by-age model also corrected for the
284 other pollutant (e.g., for the PM_{2.5}-by-age model, NO₂ was added as a confounder in addition to the
285 previously mentioned covariates, and vice versa for the NO₂-by-age model). For the random effects within
286 the zero-inflated portion of the model, we again only included ABCD site and for the random effects within
287 the count model, again, subject was nested within a random effect of site. PM_{2.5} and NO₂ were centered
288 to the levels recommended by the WHO, 5 µg/m³ and 5.33 ppb, respectively, and age was again centered
289 at 9 years. For models where the interaction term between pollution and age was not significant for both
290 the zero-inflated and count portion of the model, the interaction term was dropped, and the model was
291 run to examine the main effect of pollution.

292 2.5.5 Type-1 error correction

293 For all above models, to avoid type-1 errors, all *p*-values of interest were corrected for multiple
294 comparisons across the same model type using the false-discovery rate of 5% by utilizing the Benjamini-
295 Hochberg procedure ($p_{FDR} < 0.05$) (Benjamini & Hochberg, 1995), which has been used previously with
296 a ZINB modeling approach (Subramaniyam et al., 2019). All model assumptions post analyses were also
297 conducted based on prior published methodology (Cameron & Trivedi, 2013; Garay et al., 2011; Hilbe,
298 2011).

299 2.5.6 Model interpretation

300 In terms of interpreting our PM_{2.5} results, we focused on displaying the predictions of the EPA annual
301 daily standard (PM_{2.5} = 12 µg/m³) as compared to the WHO's recommended target level of 5 µg/m³. For
302 NO₂, our sample's exposure levels were much less than the 53 ppb previously set by EPA in 1971 (US
303 EPA, 2016). Thus, for NO₂ we focused on comparing predictions at 26.1 ppb, based on the 90th percentile
304 of our sample, as compared to the WHO recommended 5.33 ppb. Lastly, given that very large sample
305 sizes tend to identify very small differences as significant, we were sure to also interpret our results in
306 context of effect sizes in order to assess if results were likely to be clinically significant as defined by
307 (Jacobson & Truax, 1991), which requires not only statistical significance, but also a change either in the
308 range of the "dysfunctional population" or "within the range of the functional population".

309 3. Results

310 Descriptives of our analytical dataset separated by baseline, 1-year, and 2-year follow-up can be found
311 in **Table 1**. The mean for PM_{2.5} for the total current sample was 7.706 µg/m³ (range=1.722-15.902
312 SD=1.571) and for NO₂ it was 18.595 ppb (range=0.729-37.940; SD=5.571), which on average falls
313 significantly below the EPA standards (p 's<0.0001) of 12 µg/m³ and 53 ppb, respectively. Furthermore,
314 descriptives of CBCL outcomes across each collection wave are presented in **Supplemental Table 5**.

315 3.1 Internalizing Behavior

316 3.1.1 Changes in internalizing behavior with age

317 There was a significant main effect of age for the zero-inflated portion of the model, demonstrating a 45%
318 increase in the likelihood of having no internalizing problems (i.e., obtaining a true-zero) with increasing
319 age of the child from 9 to 12 years-old (**Figure 1**). For individuals who did experience internalizing
320 symptoms (i.e., modeled by the count portion of the model), there was no significant change in the
321 number of internalizing problems reported from 9 to 12 years of age.

322 3.1.2 Moderating effects of air pollution

323 The aforementioned age effects in internalizing problems from 9 to 12 years-of-age was significantly
324 moderated by PM_{2.5} and NO₂ (**Figure 1, Supplemental Tables 6 and 8**). In contrast to our hypothesis,
325 higher levels of exposure tended to relate to decreases in the probability of exhibiting any problems as
326 well as the number of problems over time between the ages of 9-12 years. Specifically, a PM_{2.5} level of
327 12 µg/m³ (EPA's standard) predicted a 190% increase in the likelihood of having no internalizing
328 problems, as well as a 13% decrease in the number of internalizing problems if problems were present,
329 from 9 to 12 years of age. A similar pattern was also seen for NO₂, with NO₂ levels of 26.1 ppb (90th
330 percentile of sample) relating to a 106% increased likelihood of having no internalizing problems, as well
331 as a 6% decrease in the number of problems, if internalizing behaviors were present, from 9 to 12 years-
332 of-age. Moreover, exposure to lower PM_{2.5} or NO₂ levels (5 µg/m³ and 5.33 ppb based on WHO
333 recommendations) predicted a higher likelihood of having internalizing symptoms (e.g., as seen by a
334 relative decrease in the probability of the caregiver reporting no internalizing problems), as well as an
335 increase in number of internalizing problems, from 9 to 12 years of age. Although these results are
336 counterintuitive in that greater exposure levels were linked with less problems over time, it is important
337 to note the effect sizes of these findings, as the detected changes in probability of exhibiting internalizing
338 problems (i.e., true-zero score) ranged from a 1-4% difference and the magnitude of the number of CBCL
339 internalizing problems was a mere 1-point change.

340 3.1.3 Internalizing subscales: Anxious/Depressed and Withdrawn/Depressed

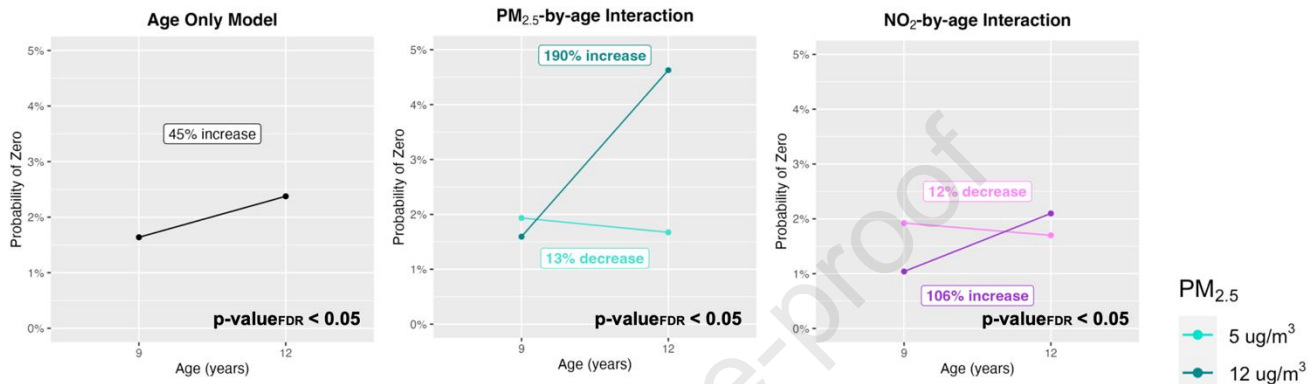
341 These subscales both fall within the internalizing score, therefore, unsurprisingly, a similar pattern was
342 seen for age-only and moderating effects of air pollution for the anxious/depressed problems
343 (**Supplemental Table 6 and 8**). A main effect of age was seen showing an increase of 185% in the
344 likelihood of having no anxious/depressed symptoms (i.e., score of 0), and if symptoms were present, an
345 8% decrease was seen in the number of problems, from 9-12 years-of-age (**Supplemental Figure 6**).
346 PM_{2.5} and NO₂ both moderated these age effects of anxious/depressed symptom scores with similar
347 patterns as seen with overall internalizing problems.

348 A noticeably different pattern was seen for both age-only and moderating effects of air pollution of the
349 withdrawn/depressed subscale. More in line with the literature, the likelihood of having no
350 withdrawn/depressed problems decreased over time, as well as a 34% increase in the number of
351 problems, between the ages of 9-12 years, suggesting a slight increase in the probability of exhibiting
352 withdrawn/depressed problems and a greater number of withdrawn/depressed symptoms with age
353 across early adolescence (**Supplemental Figure 6**). Both PM_{2.5} and NO₂ exposure moderated age-
354 related changes in withdrawn/depressed problems. When examining the probability of having
355 withdrawn/depressed symptoms, higher levels of NO₂ exposure was again associated with a greater
356 likelihood of having no withdrawn/depressed symptoms from ages 9-12 years as compared with lower
357 levels of exposure; PM_{2.5} though, did not moderate this age effect. For individuals who exhibited

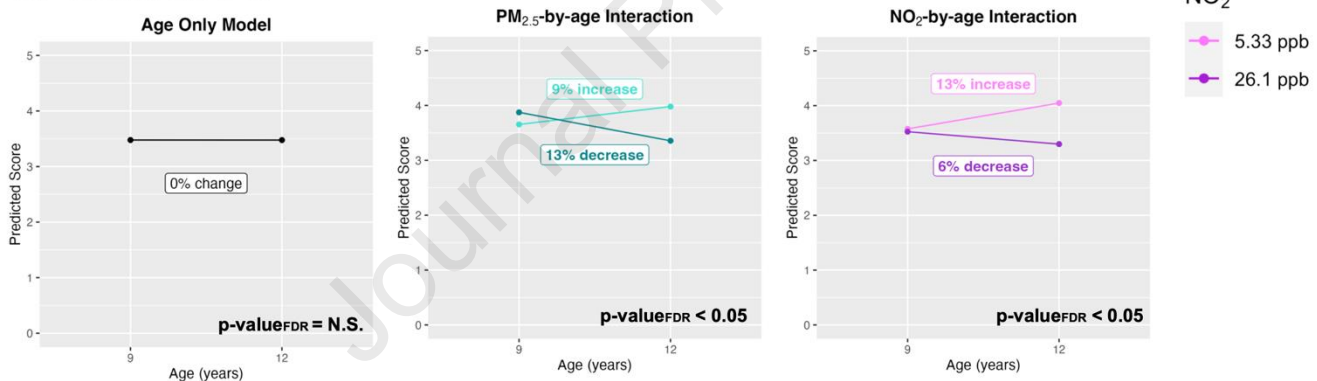
358 withdrawn/depressed problems, both NO_2 and $\text{PM}_{2.5}$ moderated the effect of age, with again lower levels
 359 of NO_2 and $\text{PM}_{2.5}$ predicting greater increases in the number of withdrawn/depressed problems from ages
 360 9-12 years as compared with higher levels of exposure. However, again, the effect sizes for exposure on
 361 both the change in probabilities and the number of problems from 9-12 years-of-age were minimal (i.e.,
 362 1-4% change in probability and 1-point increase in number of problems).

Internalizing

A. Zero-Inflated Models



B. Count Models



363

364 **Figure 1** Results for internalizing behavior. **A)** Displays the estimated probability of being in the absolute zero
 365 category as compared to the non-zero category (i.e., any value for CBCL scores). **B)** Displays the estimated CBCL
 366 score for subjects whose scores were in the non-zero category. Numerous results are presented which include: **1)**
 367 **Age only** which displays the main effects of age excluding air pollution with all other variables held constant from
 368 9 to 12 years-of-age; **2) $\text{PM}_{2.5}$ -by-age interaction** which displays differences in 9 and 12 years-of-age for the WHO
 369 recommended $\text{PM}_{2.5}$ levels - 5 $\mu\text{g}/\text{m}^3$ (light blue) - versus the EPA's - 12 $\mu\text{g}/\text{m}^3$ (dark blue); **3) NO_2 -by-age**
 370 **interaction** which displays differences in 9 and 12 years-of-age for the WHO recommended NO_2 levels - 5.33 ppb
 371 (light purple) - versus the 90th percentile NO_2 level in our sample - 26.1 ppb (The EPA level is 53 ppb
 372 which is outside our sample range). All graphs display percent change with age. All covariates held constant at the
 373 largest N category (sex = "male", race/ethnicity = 'White', caregiver education = 'Post Graduate Degree', caregiver
 374 employment = "Employed", and household income = "≥\$100K"), and mean for neighborhood safety ($\bar{x} = 3.88$); for
 375 interaction models, NO_2 is set to the WHO standard (5.33 ppb) for the $\text{PM}_{2.5}$ -by-age models and $\text{PM}_{2.5}$ is set to the
 376 WHO standard (5 $\mu\text{g}/\text{m}^3$) for the Age-only and NO_2 -by-age models; $p\text{-value}_{\text{FDR}}$ = p-value for predictor graphed once
 377 FDR corrected for multiple comparisons; N.S. = not significant.

378 3.2 Externalizing Behavior

379 *3.1.1 Changes in externalizing behavior with age*

380 There was an 88% increase in the likelihood of having no externalizing problems (i.e., obtaining a true-
381 zero) from 9 to 12 years-old. For individuals who did exhibit externalizing problems (count portion of the
382 model), a 12% decrease in the number of externalizing problems was seen from 9-12 years-of-age
383 (**Figure 2**).

384 *3.1.2 Moderating effects of air pollution*

385 PM_{2.5} did not significantly moderate the aforementioned age effects in externalizing behavior from 9-12
386 years. However, we did find that, regardless of the age of the child, that a main effect of PM_{2.5} was seen,
387 with a 60% increase in the likelihood of no externalizing problems at a PM_{2.5} concentration of 12 as
388 compared to 5 µg/m³ (**Figure 2 and Supplemental Table 7**). A main effect of PM_{2.5}, however, was not
389 seen for the number of externalizing problems. For NO₂, exposure levels did not impact the likelihood of
390 having no externalizing problems (i.e., obtaining a true-zero) from 9 to 12 years-old, but NO₂ did moderate
391 the number of externalizing problems seen with age. Specifically, higher levels of NO₂ were associated
392 with greater decreases in externalizing problems from ages 9-12 years of age as compared to lower
393 levels of NO₂ exposures (**Figure 2 and Supplemental Table 8**). Though, again, the magnitude of these
394 changes equates to less than a 1-point change in the number of problems.

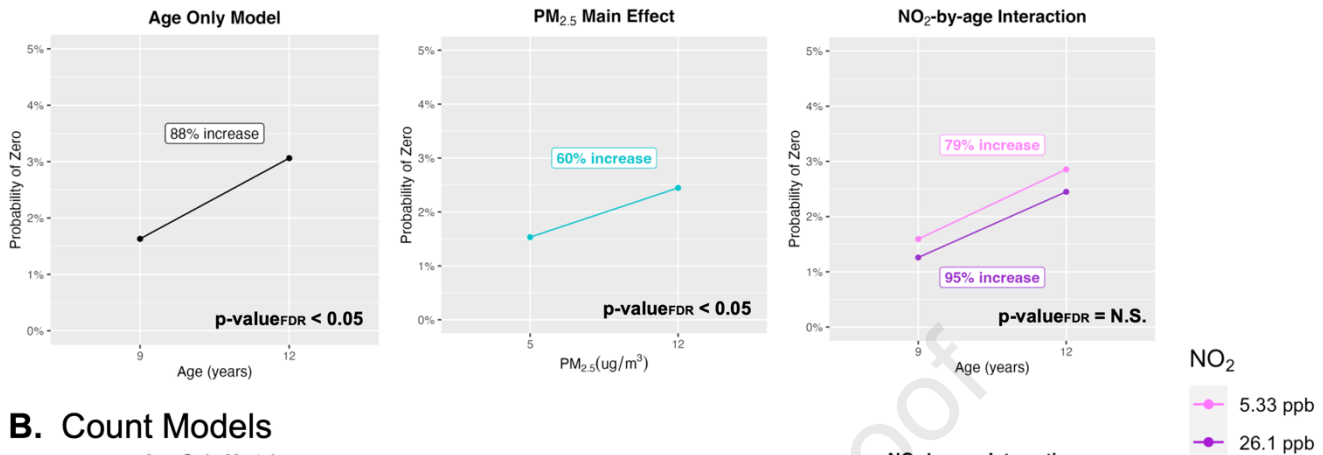
395 *3.1.3 Attention Problems and Externalizing Subscales: Rule-breaking and Aggressive Behavior*

396 There were significant main effects of age for all externalizing subscale behaviors, with an increase in
397 the likelihood of having none of these problems from 9-12 years-of-age. This change in likelihood of
398 problems with age was largest for rule breaking (112%), followed by aggressive behavior (106%), and
399 then attention problems (58%), respectively. For those reporting these problems, a 10% decrease was
400 seen for the number of rule-breaking and attention problems, while a 14% decrease was seen for
401 aggressive behaviors, with age (**Supplemental Figure 7**).

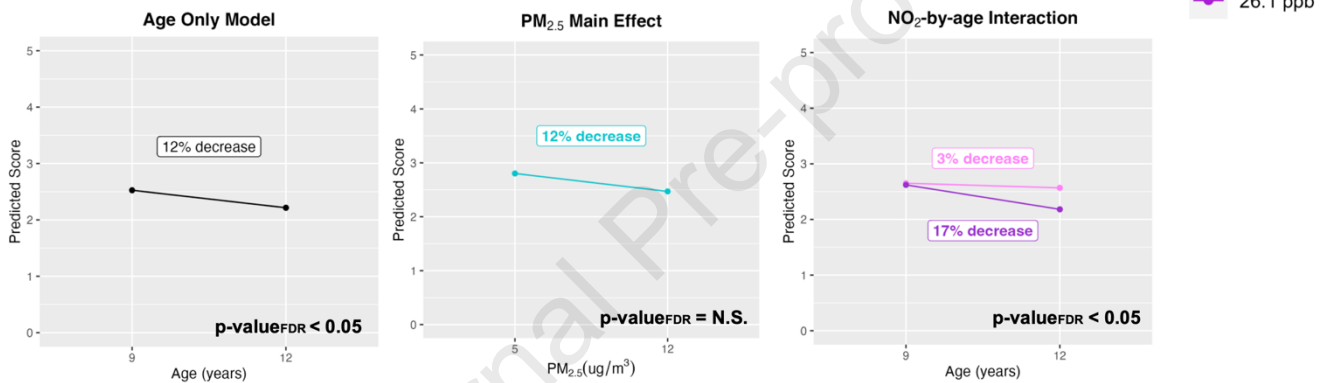
402 Only NO₂ was found to moderate the effects of age on these types of behaviors, albeit slight differences
403 were seen as to the directionality of these effects (**Supplemental Figure 7 and Supplemental Table 8**).
404 Contrary to our hypotheses, higher levels of NO₂ exposure were related to a greater likelihood of having
405 no rule-breaking behavior from 9-12 years as compared to lower levels of NO₂ exposure. Alternatively, in
406 contrast to all other outcomes, but in line with our hypothesis, a greater likelihood of having no attention
407 problems from 9-12 years was seen at lower as compared to higher levels of NO₂ exposure. Again, higher
408 levels of NO₂ exposure were associated with greater decreases in the number of rule-breaking,
409 aggressive, and attention problems from 9-12 years-of-age as compared to lower levels of NO₂ exposure.
410 Since PM_{2.5} did not moderate age-related changes in these behaviors, we investigated the main effect of
411 PM_{2.5} regardless of age. We found greater likelihood of having no aggressive or attention problems with
412 higher as compared to lower levels of PM_{2.5} exposure (**Supplemental Figure 7 and Supplemental Table**
413 **7**). Again, the magnitude of the air pollution effects were marginal, as the differences seen in the
414 probabilities of having problems was on the order of 1% and the number of problems were less than a 1-
415 point change.

Externalizing

A. Zero-Inflated Models



B. Count Models



416

417 **Figure 2** Results for externalizing behavior. **A)** Displays the estimated probability of being in the absolute zero
 418 category as compared to the non-zero category (i.e., any value for CBCL scores). **B)** Displays the estimated CBCL
 419 score for subjects whose scores were in the non-zero category. Numerous results are presented which include: **1)**
 420 **Age only** which displays the main effects of age excluding air pollution with all other variables held constant from
 421 9 to 12 years-of-age; **2) PM_{2.5}-by-age interaction** which displays differences in 9 and 12 years-of-age for the WHO
 422 recommended PM_{2.5} levels - 5 ug/m³ (light blue) - versus the EPA's - 12 ug/m³ (dark blue); **3) NO₂-by-age**
 423 **interaction** which displays differences in 9 and 12 years-of-age for the WHO recommended NO₂ levels - 5.33 ppb
 424 (light purple) - versus the 90th percentile NO₂ level in our sample - 26.1 ppb (dark purple) (The EPA level is 53 ppb
 425 which is outside our sample range). All graphs display percent change with age. All covariates held constant at the
 426 largest N category (sex = "male", race/ethnicity = 'White', caregiver education = 'Post Graduate Degree', caregiver
 427 employment = "Employed", and household income = "≥\$100K"), and mean for neighborhood safety (\bar{x} = 3.88); for
 428 interaction models, NO₂ is set to the WHO standard (5.33 ppb) for the PM_{2.5}-by-age models and PM_{2.5} is set to the
 429 WHO standard (5 ug/m³) for the Age-only and NO₂-by-age models; p-value_{FDR} = p-value for predictor graphed once
 430 FDR corrected for multiple comparisons; N.S. = not significant.

431 Discussion

432 In the current longitudinal study, we leveraged a large, nationwide longitudinal cohort of children to
 433 examine how exposure to both PM_{2.5} and NO₂ at ages 9-10 years affects age-related changes in
 434 behavioral problems as reported on the CBCL over a 2-year follow-up period. To characterize the
 435 developmental trajectory of behavioral problems within our sample, as well as aid interpretation of the
 436 pollutant effects, we first revealed an age-related decrease in the likelihood of having internalizing and

437 externalizing problems as reported on the CBCL from ages 9-12 years-old (in the zero portion of the
438 model), as well as fewer number of internalizing and externalizing problems over time if behaviors were
439 present (in the count portion of the model). Interestingly, we saw the opposite effect in the
440 withdrawn/depressed syndrome scale, where increasing age was related to an increased likelihood of
441 reporting withdrawn/depressed problems, as well as more problems, when present, from ages 9-12 years
442 old. In contrast to our hypothesis, higher levels of PM_{2.5} and NO₂ exposure did not modify these age-
443 related patterns to result in a greater likelihood or frequency in the number of problems over time.
444 Unexpectedly, higher exposure was linked to lower likelihood of having problems as well as slightly fewer
445 problems over time for most CBCL outcomes. In fact, only the association between NO₂ exposure and
446 attention problems was in the expected direction, with lower NO₂ exposure predicting an increased
447 likelihood of zero attention problems with age as compared with higher exposure. While the directions of
448 the relationships between the pollutants and CBCL outcomes are counterintuitive, it is important to
449 consider the magnitude of the effect sizes in such a large sample, rather than the statistical significance
450 of these findings. This is evident by the largest effect we found, which was the effect of PM_{2.5} on the
451 probability of internalizing symptoms arising at age 12. In children with low PM_{2.5} exposure at ages 9-10
452 years, the probability of not having problems at age 12 was 1.7%, while in those exposed to high PM_{2.5}
453 at ages 9-10 years the probability of not having problems at age 12 was 4.6%. Not only is the difference
454 in probability only 2.9%, but the likelihood of having any problems regardless of exposure level falls below
455 5%. Similarly, the effect sizes were extremely small for the quantitative differences in the number of
456 problems, with higher pollution exposure associated with a decrease of less than a single point difference
457 on any given scale. Given that the CBCL uses a 3-point Likert scale (i.e., 0 = Not True, 1 = Somewhat or
458 Sometimes True, 2 = Very True or Often True), a 1-point change is likely clinically negligible, may fall
459 within the range of measurement error, and may not have real-world implications. Thus, against our
460 hypothesis, there was no evidence that low-level exposures to PM_{2.5} and NO₂ at ages 9-10 years resulted
461 in increased emotional problems from ages 9-12 years.

462 Our study focuses on childhood exposure at ages 9-10 years old – a developmental period currently
463 underrepresented in the literature. About 26% of studies on pollution-related differences in mental health
464 problems cover this age range, despite the high incidence of psychiatric diagnoses in early adolescence
465 (Kessler et al., 2005; Solmi et al., 2022; Zundel et al., 2022). Yet even studies focused on linking PM_{2.5}
466 and NO₂ exposure and emotional behaviors in youth have reported mixed findings. Some of the earliest
467 longitudinal research in this area comes from Columbia Center for Children's Environmental Health
468 (CCCEH) longitudinal cohort study of African American and Dominican women in New York City. These
469 essential studies found that prenatal exposure to airborne polycyclic aromatic hydrocarbons (PAHs),
470 which come from fossil fuel combustion, was linked to greater CBCL reported symptoms of
471 anxious/depressed and attention problems at ages 4-5 and 6-7 years-old children (F. P. Perera et al.,
472 2011, 2012). However, in a more recent study that included using either the Strength and Difficulties
473 Questionnaire or the CBCL in 8 European population-based birth cohorts, prenatal and postnatal air
474 pollution, including PM_{2.5} and NO₂ exposure, were not found to relate to the borderline clinical range of
475 depression, anxiety, and aggression in >13,000 children ages 7-11 years-old (Jorcano et al., 2019). In
476 fact, higher postnatal exposure was linked with overall lower odds of having symptoms in the
477 borderline/clinical range when assessed cross-sectionally with the CBCL; albeit the results did not reach
478 statistical significance. Similar findings were also noted when using the quantitative scores of the
479 symptom scales (Jorcano et al., 2019) as implemented in the current analysis. A similar study assessing
480 ADHD symptoms in children 3-10 years-old using these same 8 European birth cohorts also found no

481 association, or even decreased risk, between prenatal air pollution exposure and ADHD (Forns et al.,
482 2018). Given the positive publication bias, it is likely that more evidence of null associations between
483 exposure and behavior problems exists and has been relegated to the so-called file drawer (Mlinarić et
484 al., 2017). Furthermore, a recent study using similar methodology to our own found that higher childhood
485 and prenatal exposure to PM_{2.5} and NO₂, in addition to other pollutants, was associated with fewer
486 internalizing, externalizing, and attention problems in adolescence regardless if CBCL questionnaire was
487 reported by parent or child (Kusters et al., 2022). Thus, our current findings are in line with these more
488 recent multi-research site-based studies. Similar to these studies, it seems very unlikely that the
489 significant effects found in the current study are in fact reflective of a protective effect given both 1) the
490 absence of any postulated mechanism for a protective element of air pollution exposure, as well as 2)
491 the extremely small magnitude of change detected in part to our large sample size and the resulting
492 statistical power. It is feasible that both the previous findings as well as the current results could be due
493 to residual negative confounding (Forns et al., 2018; Jorcano et al., 2019), although it is important to note
494 that in each case the analyses adjusted for many essential sociodemographic variables (i.e., income,
495 caregiver educational attainment, etc.) that are known to be associated with air pollution exposure and
496 mental health in children. Thus, if residual negative confounding is at play, unexplained factor(s) should
497 be explored that may exist across various cities and within various western populations (e.g., U.S.,
498 Germany, Italy, Spain, etc.). Despite the CBCL being a widely used and valid measure in both clinical
499 and research settings (Achenbach & Rescorla, 2001; Wolraich et al., 2008), context and informant
500 differences have been reported in using the CBCL items to assess emotional and behavioral problems
501 in youth (Achenbach et al., 1987). Albeit Kusters et al. (2022) findings suggest air pollution effects on
502 emotional behaviors are consistent regardless of parent or youth (ages 13-16 years) report. Nonetheless,
503 it is feasible that caregiver-report of emotional problems in the current study may contribute to
504 misclassification bias that could contribute towards failing to reject the null hypothesis. Thus, additional
505 studies are warranted using more objective measures, such as clinician-based interviews, of children's
506 mental health and wellbeing.

507 Putting our current results in the larger context of the literature, the importance of windows of exposure
508 and the timing of behavior continue to prevail as to what role air pollution may play in terms of risk for
509 developing mental health problems. That is, while the current study shows a one-year annual average of
510 air pollution exposure during the transition to adolescence does not substantially increase the age-related
511 clinical risk of mental health problems over a 2-year follow-up period, it is still feasible that exposure
512 during this period of development may ultimately predispose an individual to risk for developing
513 psychopathology later in adolescence or early adulthood. Air pollution, then, may influence ongoing brain
514 development and plasticity across adolescence, due to the protracted development of regions and
515 networks associated with mental health conditions and psychopathology (e.g., hippocampus, amygdala,
516 default mode network, frontoparietal network, and salience network) (Menon, 2011, 2013). A number of
517 MRI studies suggest that exposure to ambient air pollution is linked to differences in brain macro- and
518 microarchitecture as well as functional brain network connectivity (Binter et al., 2022; Burnor et al., 2021;
519 Cotter et al., 2023; Essers et al., 2023; Guxens et al., 2018, 2022; Herting et al., 2019; Lubczyńska et al.,
520 2021; Pérez-Crespo et al., 2022; Sukumaran et al., 2023). Thus, it is feasible that these differences may
521 be early neural biomarkers of PM_{2.5} exposure-related risk prior to any overt changes in behavior. As
522 previously mentioned, the idea that exposure during adolescence may ultimately predispose an individual
523 to later develop mental health disorders parallels the findings that higher levels of air pollution during
524 childhood and adolescence predict later onset of major depressive disorder (Roberts et al., 2019) and

525 other internalizing, externalizing, and thought disorder symptoms at age 18 years (Reuben et al., 2021).
526 In fact, the increased incidence of psychopathology and psychiatric diagnoses seen in adolescence
527 typically occurs in mid-adolescence, around age 14 and a half (Solmi et al., 2022), which is above the
528 upper limit of ages included here. However, consortium efforts to eventually estimate lifetime air pollution
529 exposure (C. C. Fan et al., 2021) in the coming years, in addition to active follow-up of ABCD cohort
530 participants through early adulthood, will soon allow researchers to more formally test this hypothesis.
531 Moreover, the results of the current study may also suggest that while PM_{2.5} and NO₂ exposure at 9-10
532 years does not meaningfully impact the age-related relative risk of emotional problems at a population-
533 level, it is feasible that exposure during this time may have harmful effects in children who are more
534 susceptible, due to either genetic risk or due to co-exposure to other adverse environmental threats.
535 Thus, more research is warranted taking a more integrated neural exposome approach to understanding
536 adolescent environmental exposures and risk for psychopathology (Tamiz et al., 2022).

537 The current study has several strengths. Specifically, the statistical approach and data used here
538 contribute to a rigorous assessment of longitudinal, age-related behavioral and emotional problems
539 associated with one-year annual air pollution exposure during the transition to adolescence. While
540 standardized scores are often used to study dimensions of psychopathology and behavior between-
541 subjects, we utilized raw longitudinal CBCL scores in the current study to better capture developmental
542 change (Barch et al., 2021). However, raw CBCL scores are zero-inflated and over-dispersed in
543 normative developmental samples, violating assumptions of general linear models. Our application of a
544 zero-inflated negative binomial (ZINB) model combines the strengths of a logistic regression model with
545 a negative binomial model, allowing robust estimates of associations between air pollution and the
546 emergence of any behavioral or emotional problems (i.e., scores equal to zero vs. scores greater than
547 zero) as children age, and how air pollution is related to the magnitude or number of behavioral or
548 emotional problems (i.e., the range of scores greater than zero). Second, our large, nationwide sample
549 between the ages of 9-12 years provides more geographically diverse estimates of NO₂ and PM_{2.5}. This
550 is an improvement over the smaller, localized samples common to air pollution research that pervade the
551 literature, as sources and concentrations of pollutants vary across locations (Snider et al., 2016) and the
552 health effects of PM_{2.5} vary by source (Holguin, 2008; Sarnat et al., 2008). Although the final sample used
553 here is not fully representative of the larger US population (Garavan et al., 2018), it has greater
554 generalizability compared to smaller scale studies of air pollution and mental health. Further, the models
555 were adjusted for numerous socioeconomic and lifestyle variables that are known to be associated with
556 both exposure and emotional behaviors examined in the current study.

557 A limitation of the current study is that the estimates of air pollution used here only represent a sum
558 across components of PM_{2.5} and capture an average of exposure over one year at the time of study
559 enrollment. Moreover, our study examined exposure levels that are largely below the U.S EPA standards,
560 which may only apply to approximately 50% of high-income countries in North America, Europe, and the
561 Western Pacific, and does not readily apply to existing levels of exposure in many low- and middle-
562 income countries (World Health Organization, 2018). As previously mentioned, different geographical
563 locations have different compositions of PM_{2.5} and the individual components of PM_{2.5} have different
564 effects on human health. It is possible that our results represent an amalgamation of the unique effects
565 of individual components of PM_{2.5} (e.g., elemental carbon, silicon, lead), contributing to our
566 counterintuitive findings. There is also a substantial body of literature quantifying the effects of prenatal
567 air pollution exposure and acute exposure (i.e., days) effects on various mental health outcomes

568 (Braithwaite et al., 2019; Zundel et al., 2022), which are not available in the 4.0 data release of the ABCD
569 Study. Moving forward, incorporating prenatal exposure, as well as acute estimates, could help elucidate
570 potential nuances that exist in the timing of exposure on the emergence of symptomatology across
571 adolescence, in addition to the prevalence of acute mental health crises (for review, see Heo et al., 2021).
572 Another limitation is that the data included here were collected from 2016 until March 2020, at the
573 beginning of the global COVID-19 pandemic. We chose to exclude data collected after March 2020, to
574 avoid the confounding effect of pandemic-induced emotional and behavioral problems in this sample
575 (Hamatani et al., 2022). The onset of the pandemic complicated data collection, as well, and may have
576 contributed to missingness in data collected at later follow-up visits. For example, although sample
577 demographics in the current study were similar to the larger ABCD cohort (Supplemental Tables 1-3)
578 and our overall missingness was small ($\leq 6\%$), we cannot rule out the possibility of selection bias
579 influencing our results. Not all participants had complete data at each wave of data collection, and follow-
580 up waves had slightly higher representation of white children, with greater caregiver educational
581 attainment and household income compared to enrollment at baseline. However, due to this small
582 proportion of missing data, that bias is expected to be small or negligible. Moreover, Asian, American
583 Indian/Alaskan Native, and Native Hawaiian/Pacific Islander populations are underrepresented in the
584 ABCD Study, while families with higher total household incomes and highly educated caregivers are
585 over-represented. Thus, additional studies are needed that include children who may be especially
586 susceptible to air pollution related effects because of potential compounding effects of disadvantage due
587 to poverty and minority-related stressors stemming from racism (Hajat et al., 2015). Although both the
588 exposure models used herein as well as the CBCL questionnaire have shown to have both good validity
589 and reliability and the current study adjusted for key confounders, it is feasible that measurement error
590 or residual confounding may have contributed to the current unexpected findings. Lastly, additional
591 studies are also warranted to examine if annual averages to higher levels of exposure experienced in
592 low- and middle-income countries may influence emotional wellbeing in developing children.

593 **Conclusions**

594 There was no evidence that low-level exposures to $PM_{2.5}$ and NO_2 at ages 9-10 years resulted in greater
595 emotional problems from ages 9-12 years. Future research with additional waves of data extending into
596 late adolescence and early adulthood, as well as incorporating cumulative exposure estimates are
597 necessary to further our understanding between air pollution and mental health during adolescence.

598 **Credit authorship contribution statement**

599 Claire E. Campbell: Project Administration, Methodology, Formal Analysis, Visualization, Writing -
600 Original Draft; Devyn L. Cotter: Methodology, Writing - Original Draft; Katherine L. Bottenhorn:
601 Methodology, Visualization, Writing – Original Draft; Elisabeth Burnor: Methodology, Data Curation,
602 Writing – Review & Editing; Hedyeh Ahmadi: Data Curation, Methodology, Writing – Review & Editing;
603 Carlos Cardenas-Iniguez: Writing - Review & Editing; W. James Gauderman: Methodology, Writing -
604 Review & Editing; Rob McConnell: Methodology, Writing - Review & Editing; Kiros Berhane:
605 Methodology, Writing - Review & Editing; Joel Schwartz: Methodology, Data Curation, Resources, Writing
606 - Review & Editing; Jiu-Chiuan Chen: Conceptualization, Methodology, Writing - Review & Editing; Megan
607 M. Herting: Funding Acquisition, Resources, Conceptualization, Methodology, Supervision, Project
608 Administration, Writing - Original Draft.

609 Acknowledgements

610 A special thank you to all ABCD cohort participants and their families. Portions of the research described
611 in this article were conducted under contract to the HEI, an organization jointly funded by the US
612 Environmental Protection Agency (EPA) and certain motor vehicle 14 and engine manufacturers. The
613 content of this article does not necessarily reflect the views of HEI or its sponsors, nor does it necessarily
614 reflect the views and policies of the US EPA or motor vehicle and engine manufacturers. Additional
615 support for this research was also provided by the National Institutes of Health [NIEHS R01ES032295
616 (Herting), R01ES031074 (Herting), P30ES007048-23S1 (Gilliland), 3P30ES000002-55S1 (Weisskopf),
617 NIHPO1ES022845 (McConnell), T32 ES013678 (Campbell), F31 MH131347 (Campbell)] and EPA grants
618 [RD-83587201 (Schwartz), RD-83544101 (Schwartz)] and the Rose Hills Foundation (Herting). We would
619 also like to acknowledge and thank Shermaine Abad, MPP, for making the maps shown in Supplemental
620 Figure 2.

621 Data used in the preparation of this article were obtained from the Adolescent Brain Cognitive
622 DevelopmentSM (ABCD) Study (<https://abcdstudy.org>), held in the NIMH Data Archive (NDA). This is a
623 multisite, longitudinal study designed to recruit more than 10,000 children aged 9-10 and follow them
624 over 10 years into early adulthood. The ABCD Study® is supported by the National Institutes of Health
625 and additional federal partners under award numbers U01DA041048, U01DA050989, U01DA051016,
626 U01DA041022, U01DA051018, U01DA051037, U01DA050987, U01DA041174, U01DA041106,
627 U01DA041117, U01DA041028, U01DA041134, U01DA050988, U01DA051039, U01DA041156,
628 U01DA041025, U01DA041120, U01DA051038, U01DA041148, U01DA041093, U01DA041089,
629 U24DA041123, U24DA041147. A full list of supporters is available at [https://abcdstudy.org/federal-](https://abcdstudy.org/federal-partners.html)
630 [partners.html](https://abcdstudy.org/federal-partners.html). A listing of participating sites and a complete listing of the study investigators can be found
631 at https://abcdstudy.org/consortium_members/. ABCD consortium investigators designed and
632 implemented the study and/or provided data but did not necessarily participate in the analysis or writing
633 of this report. This manuscript reflects the views of the authors and may not reflect the opinions or views
634 of the NIH or ABCD consortium investigators. Additional support for this work was made possible from
635 NIEHS R01-ES032295 and R01-ES031074.

636 Data and Code Availability Statement

637 Data used in the preparation of this article were obtained from the Adolescent Brain Cognitive
638 Development (ABCD) Study (<https://abcdstudy.org>), held in the NIMH Data Archive (NDA). This is a
639 multisite, longitudinal study designed to recruit more than 10,000 children aged 9-10 and follow them
640 over 10 years into early adulthood. ABCD consortium investigators designed and implemented the study
641 and/or provided data but did not necessarily participate in analysis or writing of this report. This
642 manuscript reflects the views of the authors and may not reflect the opinions or views of the NIH or ABCD
643 consortium investigators. The ABCD data repository grows and changes over time. The ABCD data used
644 in this report came from [10.15154/1523041](https://doi.org/10.15154/1523041).

645
646 R analysis code for this project can be found at [10.5281/zenodo.7787017](https://doi.org/10.5281/zenodo.7787017).

647 **Competing Interests**

648 The authors declare no competing interests.

Journal Pre-proof

649 **References**

- 650 Achenbach, T. M. (2009). *The Achenbach system of empirically based assessment (ASEBA):*
651 *Development, findings, theory, and applications.* University of Vermont, Research Center for
652 Children, Youth, & Families.
- 653 Achenbach, T. M., Howell, C. T., Quay, H. C., Conners, C. K., & Bates, J. E. (1991). National Survey of
654 Problems and Competencies among Four- to Sixteen-Year-Olds: Parents' Reports for Normative
655 and Clinical Samples. *Monographs of the Society for Research in Child Development, 56*(3), i–
656 130. <https://doi.org/10.2307/1166156>
- 657 Achenbach, T. M., McConaughy, S. H., & Howell, C. T. (1987). Child/adolescent behavioral and
658 emotional problems: Implications of cross-informant correlations for situational specificity.
659 *Psychological Bulletin, 101*(2), 213–232.
- 660 Achenbach, T. M., & Rescorla, L. A. (2001). *Manual for the ASEBA School-Age Forms and Profiles.*
661 University of Vermont Research Center for Children, Youth, & Families.
- 662 Barch, D. M., Albaugh, M. D., Baskin-Sommers, A., Bryant, B. E., Clark, D. B., Dick, A. S., Feczko, E.,
663 Foxe, J. J., Gee, D. G., Giedd, J., Glantz, M. D., Hudziak, J. J., Karcher, N. R., LeBlanc, K.,
664 Maddox, M., McGlade, E. C., Mulford, C., Nagel, B. J., Neigh, G., ... Xie, L. (2021). Demographic
665 and mental health assessments in the adolescent brain and cognitive development study:
666 Updates and age-related trajectories. *Developmental Cognitive Neuroscience, 52*, 101031.
667 <https://doi.org/10.1016/j.dcn.2021.101031>
- 668 Benjamini, Y., & Hochberg, Y. (1995). Controlling the False Discovery Rate: A Practical and Powerful
669 Approach to Multiple Testing. *Journal of the Royal Statistical Society: Series B (Methodological),*
670 *57*(1), 289–300. <https://doi.org/10.1111/j.2517-6161.1995.tb02031.x>

- 671 Bennett, D. A. (2001). How can I deal with missing data in my study? *Australian and New Zealand Journal*
672 *of Public Health*, 25(5), 464–469. <https://doi.org/10.1111/j.1467-842X.2001.tb00294.x>
- 673 Binter, A.-C., Kusters, M. S. W., van den Dries, M. A., Alonso, L., Lubczyńska, M. J., Hoek, G., White, T.,
674 Iñiguez, C., Tiemeier, H., & Guxens, M. (2022). Air pollution, white matter microstructure, and
675 brain volumes: Periods of susceptibility from pregnancy to preadolescence. *Environmental*
676 *Pollution (Barking, Essex: 1987)*, 313, 120109. <https://doi.org/10.1016/j.envpol.2022.120109>
- 677 Braithwaite, I., Zhang, S., Kirkbride, J. B., Osborn, D. P. J., & Hayes, J. F. (2019). Air Pollution (Particulate
678 Matter) Exposure and Associations with Depression, Anxiety, Bipolar, Psychosis and Suicide
679 Risk: A Systematic Review and Meta-Analysis. *Environmental Health Perspectives*, 127(12),
680 126002. <https://doi.org/10.1289/EHP4595>
- 681 Brokamp, C., Strawn, J. R., Beck, A. F., & Ryan, P. (2019). Pediatric Psychiatric Emergency Department
682 Utilization and Fine Particulate Matter: A Case-Crossover Study. *Environmental Health*
683 *Perspectives*, 127(9), 97006. <https://doi.org/10.1289/EHP4815>
- 684 Brunst, K. J., Ryan, P. H., Altaye, M., Yolton, K., Maloney, T., Beckwith, T., LeMasters, G., & Cecil, K. M.
685 (2019). Myo-inositol mediates the effects of traffic-related air pollution on generalized anxiety
686 symptoms at age 12 years. *Environmental Research*, 175, 71–78.
687 <https://doi.org/10.1016/j.envres.2019.05.009>
- 688 Burnor, E., Cserbik, D., Cotter, D. L., Palmer, C. E., Ahmadi, H., Eckel, S. P., Berhane, K., McConnell,
689 R., Chen, J.-C., Schwartz, J., Jackson, R., & Herting, M. M. (2021). Association of Outdoor
690 Ambient Fine Particulate Matter With Intracellular White Matter Microstructural Properties Among
691 Children. *JAMA Network Open*, 4(12), e2138300.
692 <https://doi.org/10.1001/jamanetworkopen.2021.38300>

- 693 Cameron, A. C., & Trivedi, P. K. (2013). *Regression analysis of count data* (Vol. 53). Cambridge university
694 press.
- 695 Cory-Slechta, D. A., Merrill, A., & Sobolewski, M. (2023). Air Pollution–Related Neurotoxicity Across the
696 Life Span. *Annual Review of Pharmacology and Toxicology*, 63, 143–163.
697 <https://doi.org/10.1146/annurev-pharmtox-051921-020812>
- 698 Cotter, D. L., Campbell, C. E., Sukumaran, K., McConnell, R., Berhane, K., Schwartz, J., Hackman, D.
699 A., Ahmadi, H., Chen, J.-C., & Herting, M. M. (2023). Effects of ambient fine particulates, nitrogen
700 dioxide, and ozone on maturation of functional brain networks across early adolescence.
701 *Environment International*, 177, 108001. <https://doi.org/10.1016/j.envint.2023.108001>
- 702 Cserbik, D., Chen, J.-C., McConnell, R., Berhane, K., Sowell, E. R., Schwartz, J., Hackman, D. A., Kan,
703 E., Fan, C. C., & Herting, M. M. (2020). Fine particulate matter exposure during childhood relates
704 to hemispheric-specific differences in brain structure. *Environment International*, 143, 105933.
705 <https://doi.org/10.1016/j.envint.2020.105933>
- 706 Di, Q., Amini, H., Shi, L., Kloog, I., Silvern, R., Kelly, J., Sabath, M. B., Choirat, C., Koutrakis, P.,
707 Lyapustin, A., Wang, Y., Mickley, L. J., & Schwartz, J. (2019). An ensemble-based model of
708 PM_{2.5} concentration across the contiguous United States with high spatiotemporal resolution.
709 *Environment International*, 130, 104909. <https://doi.org/10.1016/j.envint.2019.104909>
- 710 Di, Q., Amini, H., Shi, L., Kloog, I., Silvern, R., Kelly, J., Sabath, M. B., Choirat, C., Koutrakis, P.,
711 Lyapustin, A., Wang, Y., Mickley, L. J., & Schwartz, J. (2020). Assessing NO₂ Concentration and
712 Model Uncertainty with High Spatiotemporal Resolution across the Contiguous United States
713 Using Ensemble Model Averaging. *Environmental Science & Technology*, 54(3), 1372–1384.
714 <https://doi.org/10.1021/acs.est.9b03358>

- 715 Dominici, F., Schwartz, J., Di, Q., Braun, D., Choirat, C., & Zanobetti, A. (2019). Assessing Adverse
716 Health Effects of Long-Term Exposure to Low Levels of Ambient Air Pollution: Phase 1. *Research*
717 *Reports: Health Effects Institute, 2019, 200.*
- 718 Echeverria, S. E., Diez-Roux, A. V., & Link, B. G. (2004). Reliability of self-reported neighborhood
719 characteristics. *Journal of Urban Health: Bulletin of the New York Academy of Medicine, 81(4),*
720 *682–701.* <https://doi.org/10.1093/jurban/jth151>
- 721 Essers, E., Binter, A.-C., Neumann, A., White, T., Alemany, S., & Guxens, M. (2023). Air pollution
722 exposure during pregnancy and childhood, APOE ϵ 4 status and Alzheimer polygenic risk score,
723 and brain structural morphology in preadolescents. *Environmental Research, 216(Pt 2), 114595.*
724 <https://doi.org/10.1016/j.envres.2022.114595>
- 725 Fan, B., Wang, T., Wang, W., Zhang, S., Gong, M., Li, W., Lu, C., & Guo, L. (2019). Long-term exposure
726 to ambient fine particulate pollution, sleep disturbance and their interaction effects on suicide
727 attempts among Chinese adolescents. *Journal of Affective Disorders, 258, 89–95.*
728 <https://doi.org/10.1016/j.jad.2019.08.004>
- 729 Fan, C. C., Marshall, A., Smolker, H., Gonzalez, M. R., Tapert, S. F., Barch, D. M., Sowell, E., Dowling,
730 G. J., Cardenas-Iniguez, C., Ross, J., Thompson, W. K., & Herting, M. M. (2021). Adolescent
731 Brain Cognitive Development (ABCD) study Linked External Data (LED): Protocol and practices
732 for geocoding and assignment of environmental data. *Developmental Cognitive Neuroscience,*
733 *52, 101030.* <https://doi.org/10.1016/j.dcn.2021.101030>
- 734 Fang, R., Wagner, B. D., Harris, J. K., & Fillon, S. A. (2016). Zero-inflated negative binomial mixed model:
735 An application to two microbial organisms important in oesophagitis. *Epidemiology & Infection,*
736 *144(11), 2447–2455.*

- 737 Forns, J., Dadvand, P., Foraster, M., Alvarez-Pedrerol, M., Rivas, I., López-Vicente, M., Suades-
738 Gonzalez, E., Garcia-Esteban, R., Esnaola, M., Cirach, M., Grellier, J., Basagaña, X., Querol, X.,
739 Guxens, M., Nieuwenhuijsen, M. J., & Sunyer, J. (2016). Traffic-Related Air Pollution, Noise at
740 School, and Behavioral Problems in Barcelona Schoolchildren: A Cross-Sectional Study.
741 *Environmental Health Perspectives*, *124*(4), 529–535. <https://doi.org/10.1289/ehp.1409449>
- 742 Forns, J., Sunyer, J., Garcia-Esteban, R., Porta, D., Ghasabian, A., Giorgis-Allemand, L., Gong, T.,
743 Gehring, U., Sørensen, M., Standl, M., Sugiri, D., Almqvist, C., Andiaarena, A., Badaloní, C.,
744 Beelen, R., Berdel, D., Cesaroni, G., Charles, M.-A., Eriksen, K. T., ... Guxens, M. (2018). Air
745 Pollution Exposure During Pregnancy and Symptoms of Attention Deficit and Hyperactivity
746 Disorder in Children in Europe. *Epidemiology (Cambridge, Mass.)*, *29*(5), 618–626.
747 <https://doi.org/10.1097/EDE.0000000000000874>
- 748 Garavan, H., Bartsch, H., Conway, K., Decastro, A., Goldstein, R. Z., Heeringa, S., Jernigan, T., Potter,
749 A., Thompson, W., & Zahs, D. (2018). Recruiting the ABCD sample: Design considerations and
750 procedures. *Developmental Cognitive Neuroscience*, *32*, 16–22.
751 <https://doi.org/10.1016/j.dcn.2018.04.004>
- 752 Garay, A. M., Hashimoto, E. M., Ortega, E. M., & Lachos, V. H. (2011). On estimation and influence
753 diagnostics for zero-inflated negative binomial regression models. *Computational Statistics &*
754 *Data Analysis*, *55*(3), 1304–1318.
- 755 Greenland, S., & Brumback, B. (2002). An overview of relations among causal modelling methods.
756 *International Journal of Epidemiology*, *31*(5), 1030–1037. <https://doi.org/10.1093/ije/31.5.1030>
- 757 Guxens, M., Lubczyńska, M. J., Muetzel, R. L., Dalmau-Bueno, A., Jaddoe, V. W. V., Hoek, G., van der
758 Lugt, A., Verhulst, F. C., White, T., Brunekreef, B., Tiemeier, H., & El Marroun, H. (2018). Air
759 Pollution Exposure During Fetal Life, Brain Morphology, and Cognitive Function in School-Age
760 Children. *Biological Psychiatry*, *84*(4), 295–303. <https://doi.org/10.1016/j.biopsych.2018.01.016>

- 761 Guxens, M., Lubczynska, M. J., Perez-Crespo, L., Muetzel, R. L., El Marroun, H., Basagana, X., Hoek,
762 G., & Tiemeier, H. (2022). Associations of Air Pollution on the Brain in Children: A Brain Imaging
763 Study. *Research Report (Health Effects Institute)*, 209, 1–61.
- 764 Hajat, A., Hsia, C., & O'Neill, M. S. (2015). Socioeconomic Disparities and Air Pollution Exposure: A
765 Global Review. *Current Environmental Health Reports*, 2(4), 440–450.
766 <https://doi.org/10.1007/s40572-015-0069-5>
- 767 Hamatani, S., Hiraoka, D., Makita, K., Tomoda, A., & Mizuno, Y. (2022). Longitudinal impact of COVID-
768 19 pandemic on mental health of children in the ABCD study cohort. *Scientific Reports*, 12(1),
769 Article 1. <https://doi.org/10.1038/s41598-022-22694-z>
- 770 Heo, S., Lee, W., & Bell, M. L. (2021). Suicide and Associations with Air Pollution and Ambient
771 Temperature: A Systematic Review and Meta-Analysis. *International Journal of Environmental*
772 *Research and Public Health*, 18(14), 7699. <https://doi.org/10.3390/ijerph18147699>
- 773 Herting, M. M., Younan, D., Campbell, C. E., & Chen, J.-C. (2019). Outdoor Air Pollution and Brain
774 Structure and Function From Across Childhood to Young Adulthood: A Methodological Review of
775 Brain MRI Studies. *Frontiers in Public Health*, 7. <https://doi.org/10.3389/fpubh.2019.00332>
- 776 Hilbe, J. M. (2011). *Negative binomial regression*. Cambridge University Press.
- 777 Holguin, F. (2008). Traffic, Outdoor Air Pollution, and Asthma. *Immunology and Allergy Clinics of North*
778 *America*, 28(3), 577–588. <https://doi.org/10.1016/j.iac.2008.03.008>
- 779 Jacobson, N. S., & Truax, P. (1991). Clinical significance: A statistical approach to defining meaningful
780 change in psychotherapy research. *Journal of Consulting and Clinical Psychology*, 59(1), 12–19.
781 <https://doi.org/10.1037//0022-006x.59.1.12>

- 782 Jakobsen, J. C., Gluud, C., Wetterslev, J., & Winkel, P. (2017). When and how should multiple imputation
783 be used for handling missing data in randomised clinical trials—a practical guide with flowcharts.
784 *BMC Medical Research Methodology*, 17(1), 1–10.
- 785 Jernigan, T. L., Brown, S. A., & Dowling, G. J. (2018). The Adolescent Brain Cognitive Development
786 Study. *Journal of Research on Adolescence: The Official Journal of the Society for Research on*
787 *Adolescence*, 28(1), 154–156. <https://doi.org/10.1111/jora.12374>
- 788 Jorcano, A., Lubczyńska, M. J., Pierotti, L., Altug, H., Ballester, F., Cesaroni, G., El Marroun, H.,
789 Fernández-Somoano, A., Freire, C., Hanke, W., Hoek, G., Ibarluzea, J., Iñiguez, C., Jansen, P.
790 W., Lepeule, J., Markevych, I., Polańska, K., Porta, D., Schikowski, T., ... Guxens, M. (2019).
791 Prenatal and postnatal exposure to air pollution and emotional and aggressive symptoms in
792 children from 8 European birth cohorts. *Environment International*, 131, 104927.
793 <https://doi.org/10.1016/j.envint.2019.104927>
- 794 Kessler, R. C., Berglund, P., Demler, O., Jin, R., Merikangas, K. R., & Walters, E. E. (2005). Lifetime
795 prevalence and age-of-onset distributions of DSM-IV disorders in the National Comorbidity Survey
796 Replication. *Archives of General Psychiatry*, 62(6), 593–602.
797 <https://doi.org/10.1001/archpsyc.62.6.593>
- 798 Kiss, O., Alzueta, E., Yuksel, D., Pohl, K. M., de Zambotti, M., Müller-Oehring, E. M., Prouty, D., Durley,
799 I., Pelham, W. E., McCabe, C. J., Gonzalez, M. R., Brown, S. A., Wade, N. E., Marshall, A. T.,
800 Sowell, E. R., Breslin, F. J., Lisdahl, K. M., Dick, A. S., Sheth, C. S., ... Baker, F. C. (2022). The
801 Pandemic's Toll on Young Adolescents: Prevention and Intervention Targets to Preserve Their
802 Mental Health. *Journal of Adolescent Health*, 70(3), 387–395.
803 <https://doi.org/10.1016/j.jadohealth.2021.11.023>
- 804 Kumagai, N., Tajika, A., Hasegawa, A., Kawanishi, N., Fujita, H., Tsujino, N., Jinnin, R., Uchida, M.,
805 Okamoto, Y., Akechi, T., & Furukawa, T. A. (2021). Assessing recurrence of depression using a

- 806 zero-inflated negative binomial model: A secondary analysis of lifelog data. *Psychiatry Research*,
807 300, 113919. <https://doi.org/10.1016/j.psychres.2021.113919>
- 808 Kusters, M. S. W., Essers, E., Muetzel, R., Ambrós, A., Tiemeier, H., & Guxens, M. (2022). Air pollution
809 exposure during pregnancy and childhood, cognitive function, and emotional and behavioral
810 problems in adolescents. *Environmental Research*, 214(Pt 2), 113891.
811 <https://doi.org/10.1016/j.envres.2022.113891>
- 812 LAND, K. C., McCALL, P. L., & NAGIN, D. S. (1996). A Comparison of Poisson, Negative Binomial, and
813 Semiparametric Mixed Poisson Regression Models: With Empirical Applications to Criminal
814 Careers Data. *Sociological Methods & Research*, 24(4), 387–442.
815 <https://doi.org/10.1177/0049124196024004001>
- 816 Lisdahl, K. M., Sher, K. J., Conway, K. P., Gonzalez, R., Feldstein Ewing, S. W., Nixon, S. J., Tapert, S.,
817 Bartsch, H., Goldstein, R. Z., & Heitzeg, M. (2018). Adolescent brain cognitive development
818 (ABCD) study: Overview of substance use assessment methods. *Developmental Cognitive
819 Neuroscience*, 32, 80–96. <https://doi.org/10.1016/j.dcn.2018.02.007>
- 820 Lubczyńska, M. J., Muetzel, R. L., El Marroun, H., Hoek, G., Kooter, I. M., Thomson, E. M., Hillegers, M.,
821 Vernooij, M. W., White, T., Tiemeier, H., & Guxens, M. (2021). Air pollution exposure during
822 pregnancy and childhood and brain morphology in preadolescents. *Environmental Research*,
823 198, 110446. <https://doi.org/10.1016/j.envres.2020.110446>
- 824 Margolis, A. E., Herbstman, J. B., Davis, K., Thomas, V. K., Tang, D., Wang, Y., Wang, S., Perera, F. P.,
825 Peterson, B. S., & Rauh, V. A. (2016). Longitudinal effects of prenatal exposure to air pollutants
826 on self-regulatory capacities and social competence. *Journal of Child Psychology and Psychiatry,
827 and Allied Disciplines*, 57(7), 851–860. <https://doi.org/10.1111/jcpp.12548>

- 828 Menon, V. (2011). Large-scale brain networks and psychopathology: A unifying triple network model.
829 *Trends in Cognitive Sciences*, 15(10), 483–506. <https://doi.org/10.1016/j.tics.2011.08.003>
- 830 Menon, V. (2013). Developmental pathways to functional brain networks: Emerging principles. *Trends in*
831 *Cognitive Sciences*, 17(12), 627–640. <https://doi.org/10.1016/j.tics.2013.09.015>
- 832 Mlinarić, A., Horvat, M., & Šupak Smolčić, V. (2017). Dealing with the positive publication bias: Why you
833 should really publish your negative results. *Biochemia Medica*, 27(3), 030201.
834 <https://doi.org/10.11613/BM.2017.030201>
- 835 Mujahid, M. S., Diez Roux, A. V., Morenoff, J. D., & Raghunathan, T. (2007). Assessing the measurement
836 properties of neighborhood scales: From psychometrics to ecometrics. *American Journal of*
837 *Epidemiology*, 165(8), 858–867. <https://doi.org/10.1093/aje/kwm040>
- 838 Myhre, O., Låg, M., Villanger, G. D., Oftedal, B., Øvrevik, J., Holme, J. A., Aase, H., Paulsen, R. E., Bal-
839 Price, A., & Dirven, H. (2018). Early life exposure to air pollution particulate matter (PM) as risk
840 factor for attention deficit/hyperactivity disorder (ADHD): Need for novel strategies for
841 mechanisms and causalities. *Toxicology and Applied Pharmacology*, 354, 196–214.
842 <https://doi.org/10.1016/j.taap.2018.03.015>
- 843 Newman, N. C., Ryan, P., Lemasters, G., Levin, L., Bernstein, D., Hershey, G. K. K., Lockey, J. E.,
844 Villareal, M., Reponen, T., Grinshpun, S., Sucharew, H., & Dietrich, K. N. (2013). Traffic-related
845 air pollution exposure in the first year of life and behavioral scores at 7 years of age.
846 *Environmental Health Perspectives*, 121(6), 731–736. <https://doi.org/10.1289/ehp.1205555>
- 847 Papadogeorgou, G., Kioumourtzoglou, M.-A., Braun, D., & Zanobetti, A. (2019). Low Levels of Air
848 Pollution and Health: Effect Estimates, Methodological Challenges, and Future Directions.
849 *Current Environmental Health Reports*, 6(3), 105–115. [https://doi.org/10.1007/s40572-019-](https://doi.org/10.1007/s40572-019-00235-7)
850 00235-7

- 851 Perera, F. P., Chang, H., Tang, D., Roen, E. L., Herbstman, J., Margolis, A., Huang, T.-J., Miller, R. L.,
852 Wang, S., & Rauh, V. (2014). Early-life exposure to polycyclic aromatic hydrocarbons and ADHD
853 behavior problems. *PloS One*, *9*(11), e111670. <https://doi.org/10.1371/journal.pone.0111670>
- 854 Perera, F. P., Tang, D., Wang, S., Vishnevetsky, J., Zhang, B., Diaz, D., Camann, D., & Rauh, V. (2012).
855 Prenatal Polycyclic Aromatic Hydrocarbon (PAH) Exposure and Child Behavior at Age 6–7 Years.
856 *Environmental Health Perspectives*, *120*(6), 921–926. <https://doi.org/10.1289/ehp.11104315>
- 857 Perera, F. P., Wang, S., Vishnevetsky, J., Zhang, B., Cole, K. J., Tang, D., Rauh, V., & Phillips, D. H.
858 (2011). Polycyclic aromatic hydrocarbons-aromatic DNA adducts in cord blood and behavior
859 scores in New York city children. *Environmental Health Perspectives*, *119*(8), 1176–1181.
860 <https://doi.org/10.1289/ehp.1002705>
- 861 Perera, F., Roen Nolte, E. L., Wang, Y., Margolis, A. E., Calafat, A. M., Wang, S., Garcia, W., Hoepner,
862 L. A., Peterson, B. S., Rauh, V., & Herbstman, J. (2016). Bisphenol A Exposure and Symptoms
863 of Anxiety and Depression Among Inner City Children at 10–12 Years of Age. *Environmental*
864 *Research*, *151*, 195–202. <https://doi.org/10.1016/j.envres.2016.07.028>
- 865 Pérez-Crespo, L., Kusters, M. S. W., López-Vicente, M., Lubczyńska, M. J., Foraster, M., White, T., Hoek,
866 G., Tiemeier, H., Muetzel, R. L., & Guxens, M. (2022). Exposure to traffic-related air pollution and
867 noise during pregnancy and childhood, and functional brain connectivity in preadolescents.
868 *Environment International*, *164*, 107275. <https://doi.org/10.1016/j.envint.2022.107275>
- 869 Polanczyk, G. V., Salum, G. A., Sugaya, L. S., Caye, A., & Rohde, L. A. (2015). Annual research review:
870 A meta-analysis of the worldwide prevalence of mental disorders in children and adolescents.
871 *Journal of Child Psychology and Psychiatry, and Allied Disciplines*, *56*(3), 345–365.
872 <https://doi.org/10.1111/jcpp.12381>

- 873 Preisser, J. S., Das, K., Long, D. L., & Divaris, K. (2016). Marginalized zero-inflated negative binomial
874 regression with application to dental caries. *Statistics in Medicine*, 35(10), 1722–1735.
875 <https://doi.org/10.1002/sim.6804>
- 876 R Core Team. (2021). *R: A Language and Environment for Statistical Computing*. R Foundation for
877 Statistical Computing. <https://www.R-project.org/>
- 878 Rasnick, E., Ryan, P. H., Bailer, A. J., Fisher, T., Parsons, P. J., Yolton, K., Newman, N. C., Lanphear,
879 B. P., & Brokamp, C. (2021). Identifying sensitive windows of airborne lead exposure associated
880 with behavioral outcomes at age 12. *Environmental Epidemiology*, 5(2), e144.
881 <https://doi.org/10.1097/EE9.0000000000000144>
- 882 Rescorla, L. A., Ginzburg, S., Achenbach, T. M., Ivanova, M. Y., Almqvist, F., Begovac, I., Bilenberg, N.,
883 Bird, H., Chahed, M., Dobrean, A., Döpfner, M., Erol, N., Hannesdottir, H., Kanbayashi, Y.,
884 Lambert, M. C., Leung, P. W. L., Minaei, A., Novik, T. S., Oh, K.-J., ... Verhulst, F. C. (2013).
885 Cross-Informant Agreement Between Parent-Reported and Adolescent Self-Reported Problems
886 in 25 Societies. *Journal of Clinical Child & Adolescent Psychology*, 42(2), 262–273.
887 <https://doi.org/10.1080/15374416.2012.717870>
- 888 Reuben, A., Arseneault, L., Beddows, A., Beevers, S. D., Moffitt, T. E., Ambler, A., Latham, R. M.,
889 Newbury, J. B., Odgers, C. L., Schaefer, J. D., & Fisher, H. L. (2021). Association of Air Pollution
890 Exposure in Childhood and Adolescence With Psychopathology at the Transition to Adulthood.
891 *JAMA Network Open*, 4(4), e217508–e217508.
892 <https://doi.org/10.1001/jamanetworkopen.2021.7508>
- 893 Roberts, S., Arseneault, L., Barratt, B., Beevers, S., Danese, A., Odgers, C. L., Moffitt, T. E., Reuben, A.,
894 Kelly, F. J., & Fisher, H. L. (2019). Exploration of NO₂ and PM_{2.5} air pollution and mental health
895 problems using high-resolution data in London-based children from a UK longitudinal cohort
896 study. *Psychiatry Research*, 272, 8–17. <https://doi.org/10.1016/j.psychres.2018.12.050>

- 897 Sarnat, J. A., Marmur, A., Klein, M., Kim, E., Russell, A. G., Sarnat, S. E., Mulholland, J. A., Hopke, P.
898 K., & Tolbert, P. E. (2008). Fine Particle Sources and Cardiorespiratory Morbidity: An Application
899 of Chemical Mass Balance and Factor Analytical Source-Appportionment Methods. *Environmental*
900 *Health Perspectives*, 116(4), 459–466. <https://doi.org/10.1289/ehp.10873>
- 901 Schafer, J. L. (1999). Multiple imputation: A primer. *Statistical Methods in Medical Research*, 8(1), 3–15.
- 902 Sheu, M.-L., Hu, T.-W., Keeler, T. E., Ong, M., & Sung, H.-Y. (2004). The effect of a major cigarette price
903 change on smoking behavior in california: A zero-inflated negative binomial model. *Health*
904 *Economics*, 13(8), 781–791. <https://doi.org/10.1002/hec.849>
- 905 Snider, G., Weagle, C. L., Murdymootoo, K. K., Ring, A., Ritchie, Y., Stone, E., Walsh, A., Akoshile, C.,
906 Anh, N. X., & Balasubramanian, R. (2016). Variation in global chemical composition of PM 2.5:
907 Emerging results from SPARTAN. *Atmospheric Chemistry and Physics*, 16(15), 9629–9653.
- 908 Solmi, M., Radua, J., Olivola, M., Croce, E., Soardo, L., Salazar de Pablo, G., Il Shin, J., Kirkbride, J. B.,
909 Jones, P., Kim, J. H., Kim, J. Y., Carvalho, A. F., Seeman, M. V., Correll, C. U., & Fusar-Poli, P.
910 (2022). Age at onset of mental disorders worldwide: Large-scale meta-analysis of 192
911 epidemiological studies. *Molecular Psychiatry*, 27(1), Article 1. [https://doi.org/10.1038/s41380-](https://doi.org/10.1038/s41380-021-01161-7)
912 [021-01161-7](https://doi.org/10.1038/s41380-021-01161-7)
- 913 Stroup, W. W. (2012). *Generalized linear mixed models: Modern concepts, methods and applications*.
914 CRC press.
- 915 Stroup, W. W. (2016). *Generalized Linear Mixed Models: Modern Concepts, Methods and Applications*.
916 CRC Press.
- 917 Subramaniyam, S., DeJesus, M. A., Zaveri, A., Smith, C. M., Baker, R. E., Ehrt, S., Schnappinger, D.,
918 Sassetti, C. M., & Ioerger, T. R. (2019). Statistical analysis of variability in TnSeq data across

- 919 conditions using zero-inflated negative binomial regression. *BMC Bioinformatics*, 20(1), 603.
920 <https://doi.org/10.1186/s12859-019-3156-z>
- 921 Sukumaran, K., Cardenas-Iniguez, C., Burnor, E., Bottenhorn, K., Hackman, D., McConnell, R., Berhane,
922 K., Schwartz, J., Chen, J.-C., & Herting, M. (2023). Ambient fine particulate exposure and
923 subcortical gray matter microarchitecture in 9- and 10-year-olds children across the United States.
924 *IScience*, 106087. <https://doi.org/10.1016/j.isci.2023.106087>
- 925 Swartout, K. M., Thompson, M. P., Koss, M. P., & Su, N. (2015). What is the Best Way to Analyze Less
926 Frequent Forms of Violence? The Case of Sexual Aggression. *Psychology of Violence*, 5(3), 305–
927 313. <https://doi.org/10.1037/a0038316>
- 928 Tamiz, A. P., Koroshetz, W. J., Dhruv, N. T., & Jett, D. A. (2022). A focus on the neural exposome.
929 *Neuron*, 110(8), 1286–1289. <https://doi.org/10.1016/j.neuron.2022.03.019>
- 930 Thygesen, M., Holst, G. J., Hansen, B., Geels, C., Kalkbrenner, A., Schendel, D., Brandt, J., Pedersen,
931 C. B., & Dalsgaard, S. (2020). Exposure to air pollution in early childhood and the association with
932 Attention-Deficit Hyperactivity Disorder. *Environmental Research*, 183, 108930.
933 <https://doi.org/10.1016/j.envres.2019.108930>
- 934 US EPA, O. (2016, July 5). *Fact Sheets and Additional Information Regarding the Primary National*
935 *Ambient Air Quality Standards (NAAQS) for Nitrogen Dioxide (NO2)* [Overviews and Factsheets].
936 [https://www.epa.gov/no2-pollution/fact-sheets-and-additional-information-regarding-primary-](https://www.epa.gov/no2-pollution/fact-sheets-and-additional-information-regarding-primary-national-ambient-air-quality)
937 [national-ambient-air-quality](https://www.epa.gov/no2-pollution/fact-sheets-and-additional-information-regarding-primary-national-ambient-air-quality)
- 938 Volkow, N. D., Koob, G. F., Croyle, R. T., Bianchi, D. W., Gordon, J. A., Koroshetz, W. J., Pérez-Stable,
939 E. J., Riley, W. T., Bloch, M. H., Conway, K., Deeds, B. G., Dowling, G. J., Grant, S., Howlett, K.
940 D., Matochik, J. A., Morgan, G. D., Murray, M. M., Noronha, A., Spong, C. Y., ... Weiss, S. R. B.

- 941 (2018). The conception of the ABCD study: From substance use to a broad NIH collaboration.
942 *Developmental Cognitive Neuroscience*, 32, 4–7. <https://doi.org/10.1016/j.dcn.2017.10.002>
- 943 Vyas, C. M., Donneyong, M., Mischoulon, D., Chang, G., Gibson, H., Cook, N. R., Manson, J. E.,
944 Reynolds, C. F., & Okereke, O. I. (2020). Associations between race and ethnicity and late-life
945 depression severity, symptom burden and care. *JAMA Network Open*, 3(3), e201606.
946 <https://doi.org/10.1001/jamanetworkopen.2020.1606>
- 947 Wolraich, M. L., Drotar, D. D., Dworkin, P. H., & Perrin, E. C. (Eds.). (2008). CHAPTER 18—Internalizing
948 Conditions. In *Developmental-Behavioral Pediatrics* (pp. 627–668). Mosby.
949 <https://doi.org/10.1016/B978-0-323-04025-9.50021-0>
- 950 World Health Organization. (2018). *Mental health atlas 2017*. World Health Organization; WHO IRIS.
951 <https://apps.who.int/iris/handle/10665/272735>
- 952 World Health Organization. (2021). *WHO global air quality guidelines: Particulate matter (PM_{2.5} and
953 PM₁₀), ozone, nitrogen dioxide, sulfur dioxide and carbon monoxide*. World Health Organization.
954 <https://apps.who.int/iris/handle/10665/345329>
- 955 Xu, T., Zhu, G., & Han, S. (2017). Study of depression influencing factors with zero-inflated regression
956 models in a large-scale population survey. *BMJ Open*, 7(11), e016471.
957 <https://doi.org/10.1136/bmjopen-2017-016471>
- 958 Yau, K. K., Wang, K., & Lee, A. H. (2003). Zero-inflated negative binomial mixed regression modeling of
959 over-dispersed count data with extra zeros. *Biometrical Journal: Journal of Mathematical Methods
960 in Biosciences*, 45(4), 437–452.
- 961 Yip, S. W., Jordan, A., Kohler, R. J., Holmes, A., & Bzdok, D. (2022). Multivariate, Transgenerational
962 Associations of the COVID-19 Pandemic Across Minoritized and Marginalized Communities.
963 *JAMA Psychiatry*, 79(4), 350–358. <https://doi.org/10.1001/jamapsychiatry.2021.4331>

- 964 Yolton, K., Khoury, J. C., Burkle, J., LeMasters, G., Cecil, K., & Ryan, P. (2019). Lifetime exposure to
965 traffic-related air pollution and symptoms of depression and anxiety at age 12 years.
966 *Environmental Research*, 173, 199–206. <https://doi.org/10.1016/j.envres.2019.03.005>
- 967 Yorifuji, T., Kashima, S., Diez, M. H., Kado, Y., Sanada, S., & Doi, H. (2017). Prenatal exposure to outdoor
968 air pollution and child behavioral problems at school age in Japan. *Environment International*, 99,
969 192–198. <https://doi.org/10.1016/j.envint.2016.11.016>
- 970 Yorifuji, T., Kashima, S., Higa Diez, M., Kado, Y., Sanada, S., & Doi, H. (2016). Prenatal Exposure to
971 Traffic-related Air Pollution and Child Behavioral Development Milestone Delays in Japan.
972 *Epidemiology (Cambridge, Mass.)*, 27(1), 57–65.
973 <https://doi.org/10.1097/EDE.0000000000000361>
- 974 Young, D. S., Roemmele, E. S., & Yeh, P. (2022). Zero-inflated modeling part I: Traditional zero-inflated
975 count regression models, their applications, and computational tools. *Wiley Interdisciplinary*
976 *Reviews: Computational Statistics*, 14(1), e1541.
- 977 Zhang, X., & Yi, N. (2020). NBZIMM: Negative binomial and zero-inflated mixed models, with application
978 to microbiome/metagenomics data analysis. *BMC Bioinformatics*, 21(1), 488.
979 <https://doi.org/10.1186/s12859-020-03803-z>
- 980 Zhao, T., Markevych, I., Standl, M., Schulte-Körne, G., Schikowski, T., Berdel, D., Koletzko, S., Bauer,
981 C.-P., von Berg, A., Nowak, D., & Heinrich, J. (2019). Ambient ozone exposure and depressive
982 symptoms in adolescents: Results of the GINIplus and LISA birth cohorts. *Environmental*
983 *Research*, 170, 73–81. <https://doi.org/10.1016/j.envres.2018.12.014>
- 984 Zundel, C. G., Ryan, P., Brokamp, C., Heeter, A., Huang, Y., Strawn, J. R., & Marusak, H. A. (2022). Air
985 pollution, depressive and anxiety disorders, and brain effects: A systematic review.
986 *NeuroToxicology*, 93, 272–300. <https://doi.org/10.1016/j.neuro.2022.10.011>

Highlights:

- We examined one-year air pollution exposure on changes in emotion in 9-12 year-olds
- Concentrations of air pollution exposure were below U.S. EPA standards
- Annual measurements of emotional problems were investigated over 3 years
- Overall, less internalizing and externalizing behavior problems seen over time
- Our results do not support the idea that air pollution increases problems over time

Journal Pre-proof

Declaration of interests

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

The authors declare the following financial interests/personal relationships which may be considered as potential competing interests:

Journal Pre-proof