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## Familial Confounding of the Association between Maternal Smoking during Pregnancy and Offspring Criminality: A Population-Based Study in Sweden

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### Abstract

**Context**—The association between maternal smoking during pregnancy (SDP) and offspring disruptive behaviors has been well documented, but it is unclear whether exposure to SDP or the effects of factors correlated with SDP account for the increased risk.

**Objective**—To test whether the association between SDP and offspring criminal convictions was consistent with a causal connection or due to familial background factors by controlling for measured covariates and using a quasi-experimental approach.

**Design**—We used a population-based study of children born in Sweden from 1983–1989 (N=609,372) to examine the association between SDP and offspring criminal convictions, while controlling for measured traits of both parents. We also compared siblings differentially exposed to SDP (N=50,339) to account for unmeasured familial factors that could account for the association.

**Setting**—Population-based study of all children born in Sweden from 1983–1989 with information on maternal SDP and offspring criminal convictions, based on national registries collected by the Swedish government.

**Main Outcome Measures**—Violent and nonviolent convictions, based on the Swedish National Crime Register, a register with detailed information on all convictions in the country.

**Results**—Moderate (HR=2.47, CI=2.34–2.60) and high levels (HR=3.43, CI=3.25–3.63) of maternal SDP was associated with an increased risk for offspring violent convictions, even when controlling for maternal and paternal traits. There was no association between SDP and violent convictions, however, when comparing differentially exposed siblings (HR<sub>moderate</sub>=1.02, CI=0.79–1.30; HR<sub>high</sub>=1.03, CI=0.78–1.37). SDP also was associated with nonviolent convictions in the entire population (HR<sub>moderate</sub>=1.62, CI=1.58–1.66; HR<sub>high</sub>=1.87, CI=1.82–1.92) and when

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The primary investigator had full access to all of the data in the study and takes responsibility for the integrity of the data and the accuracy of the data analysis.

controlling for covariates. But, there was no association when comparing siblings who were differentially exposed ( $HR_{\text{moderate}}=0.89$ ,  $CI=0.78-1.01$ ;  $HR_{\text{high}}=0.89$ ,  $CI=0.78-1.02$ ).

**Conclusions**—The results suggest that familial background factors account for the association between maternal SDP and criminal convictions, not the specific exposure to SDP.

Maternal smoking during pregnancy (SDP) has consistently been linked with disruptive behaviors in offspring,<sup>1-4</sup> including parent-reported conduct problems,<sup>5, 6</sup> diagnoses of Oppositional Defiant<sup>7</sup> and Conduct Disorder,<sup>8-10</sup> and arrest history from national crime registries.<sup>11, 12</sup> Reviews have noted that the associations are consistent with a causal connection because they are specific to disruptive behaviors (and related constructs), have been found across diverse samples and using different measures, demonstrate a dose-response relationship, and are commensurate with findings from basic research, including animal studies.<sup>1, 2, 13</sup>

Many researchers, however, have noted that causal interpretations are impossible to prove because the association between SDP and offspring behavior may be due to risk factors associated with SDP rather than the influence of prenatal nicotine exposure.<sup>1, 14, 15</sup> SDP is correlated with many parental and family environmental factors that also predict criminal and psychiatric problems in children, such as low socioeconomic status,<sup>16, 17</sup> young age of childbearing,<sup>2, 17</sup> history of maternal<sup>14</sup> and paternal antisocial behavior,<sup>18</sup> and poor family functioning.<sup>19</sup> Genetic factors could also explain the relation between SDP and offspring disruptive behaviors.<sup>13, 14, 20-23</sup> SDP has been shown to be influenced by genetic factors.<sup>20, 24</sup> Mothers who smoke during pregnancy, therefore, could pass down genetic risk for disruptive behaviors to their offspring.<sup>25</sup>

A number of studies have utilized numerous quasi-experimental designs, approaches that use natural experiments, to explore the mechanisms through which SDP is associated with offspring development.<sup>4</sup> Recent sibling comparison<sup>5, 6, 23, 26, 27</sup>, children of twins<sup>28</sup>, and prenatal in vitro fertilization (IVF) cross-fostering<sup>29, 30</sup> studies suggest the associations between SDP and offspring conduct problems, academic achievement, intellectual abilities, and Attention Deficit Hyperactivity Disorder (ADHD) are better explained by background familial factors. The use of quasi-experimental approaches, therefore, suggests that later neurocognitive and behavioral problems that are associated with SDP may actually be due to background familial factors rather than the teratogenic effects of SDP.

The quasi-experimental studies of the association between SDP and offspring disruptive behaviors are limited, however. The extant sibling comparison studies<sup>5, 6</sup> are based on behaviors in young children. Using IVF to study SDP is a novel approach for disentangling genetic confounds from prenatal risks,<sup>29, 30</sup> but the findings for SDP need to be replicated, as the results were based on parent report of behavior problems during childhood from a small study of children conceived through IVF. And, many studies are confounded by the fact that mothers reported on both SDP and offspring behaviors.<sup>31</sup> Certainly, causal inferences can only be drawn based on converging evidence from multiple studies using various designs, each with their own strengths and weaknesses.<sup>32</sup>

The current manuscript explores the association between maternal SDP and offspring criminal behavior using a population study of all individuals born in Sweden during a span of seven years. The study statistically controlled for measured covariates from both mothers and fathers, a limitation of many existent studies of SDP,<sup>18</sup> and compared siblings differentially exposed to SDP. The comparison of siblings is a powerful approach for testing causal inferences because the design rules out most genetic factors that can confound the association between SDP and offspring criminality, as well as all environmental factors that make siblings similar.<sup>31, 33, 34</sup> The assessments of criminal behavior were based on

government records of convictions, which assesses serious violations during late adolescence and early adulthood that are not based on parental report. The study also included assessments of the offspring that were highly associated with criminal convictions, including an independent evaluation of all males in the country as part of the mandatory armed forces evaluation at age 18 and school grades at age 15, which gives the opportunity provide converging evidence. As such, the current study provides a rigorous test of the underlying etiological factors that increase the risk for serious criminal behavior associated with maternal SDP.

## METHODS

### Sample

The sample was based on merging Swedish population registries that are maintained by government agencies in Sweden. The current study is based on a subset of offspring with information on criminal convictions from a larger sample with reports of maternal SDP.<sup>23</sup>

The Swedish Medical Birth Registry (MBR), which is maintained by the Swedish Centre for Epidemiology at the National Board of Health and Welfare, includes information on over 99% of all births in Sweden since 1973.<sup>35</sup> The information is gathered during pregnancy and delivery using standardized assessments and records. Beginning in 1983 the register included assessment of maternal smoking during pregnancy at the first antenatal visit. The MBR also includes assessments of demographics, family structure, maternal age, birth order, physical attributes of the children, and complications.

The National Crime Register (NCR) is a register maintained by the Swedish National Council for Crime Prevention that includes all registered convictions in Sweden (from 1973–2004). The register includes information about every conviction, including the type of offense and sentence type. Merging the dataset in the study provided criminal background information starting at age 15 (the age of criminal responsibility in Sweden) for the offspring, as well as the criminal records for their mothers and fathers.

The Conscript Registry includes all 18-year old males in Sweden since 1970 (tested for compulsory military service) with systematic data on cognitive, psychological and physical performance.<sup>36</sup> The Education Register is maintained by Statistics Sweden and includes information about the highest obtained level of education and type of degree for each individual. We used the highest level of education in 2004 for the mothers and fathers of the target children in the study. The Register also includes assessments of achievement, including summary grades for all students at the end of grade nine.<sup>23, 26</sup> The 1990 Census Swedish Population and Housing Census was an assessment of individuals in each household in Sweden, which included measures of employment, occupation, and income. The Cause of Death Register contains data on the date and cause of deaths, and the Migration Register includes the dates of migration in and out of Sweden.

The Multi-Generation Register is a large database maintained by Statistics Sweden, which contains each individual's unique identifier, as well as information on the identity of the parents of each child born in Sweden since 1932, or who immigrated and became Swedish citizens before age 18.<sup>37</sup> The information allowed us to link all children to their biological fathers (based on maternal reports) and identify all siblings. Because of the focus on maternal SDP all siblings in the current study share the same mother (i.e. they are either full siblings or maternal half-siblings).<sup>23</sup>

Between 1983 and 1989, 709,125 children were included in the MBR. Children who had serious malformations at birth (N=12,220), were from multiple births (e.g. twins and

triplets) (N=14,375), died (N=6,262), or emigrated from the country (N=24,482) were not eligible for inclusion in the analyses, leaving 654,954 offspring. Offspring who were missing the identification number of their mother (N=745) or who were missing data on smoking during pregnancy (N=50,660) were then excluded from the sample, resulting in a sample of 609,372 offspring (93% of the targeted population and 86% of all births).

## Measures

**Offspring Outcomes**—Violent crime is defined according to the Swedish Penal Code as attempted or completed murder, manslaughter, and filicide (offence codes 3:1-3:3), aggravated assault (3:5-3:6), gross violation of a person's integrity (4a), kidnapping and illegal restraint (4:1-4:2), illegal coercion and threats (4:4-4:5), harassment (4:7), aggravated robbery (8:5-8:6), aggravated arson (13:1-13:2), and/or threats or violence against an officer (17:1). Any sexual offense was also included [rape, sexual coercion, child molestation, and sexual harassment including indecent exposure] (6:1-6:10, 6:12). This corresponds to a wider definition of violent offending that was used in earlier scientific reports on this sample.<sup>38, 39</sup> Analyses are only based on first conviction for repeat offenders. Kaplan Meier estimates of conviction for a violent crime in the sample indicate that 2.3% of individuals were convicted by 21.9 years of age.

Non-violent crime is defined as all other convictions, including crimes according to the Narcotic Drugs Act and Traffic Offences Act. Kaplan Meier estimates from the current sample indicate a higher prevalence of non-violent crime (10.1%) by 21.9 years.

The sample included 93,675 males with information from the Conscript Registry. The registry includes an evaluation of military officer suitability, which is an independent evaluation of the ability of each male to hold positions of greater responsibility in the military based on two days of interviews and standardized assessments, including a psychiatric evaluation.<sup>40</sup> Previous research has shown that the scale is the most powerful predictor of later functioning in the military when compared to the other assessments included in the registry.<sup>40</sup> The measure of low military officer suitability, which was based on the lowest two stanines (6.24%), was highly associated with convictions for later violent (Hazard Rate=3.98,  $b_{\text{logit}}=1.38$ ,  $SE=0.09$ ,  $p<.0001$ ) and nonviolent crimes (Hazard Rate=1.78,  $b_{\text{logit}}=0.57$ ,  $SE=0.04$ ,  $p<.0001$ ) in the current sample.

The study also included assessment of performance in grade 9 for 605,294 offspring; 15.1% were identified as having low academic performance, commensurate with failing.<sup>26</sup> Low academic performance was highly associated with later violent (Odds Ratio=11.35, CI=10.81–11.93) and nonviolent crimes (OR=3.69, CI=3.61–3.78), in addition to low military officer suitability (OR=3.99, CI=3.69–4.31).

**Risk Factors**—SDP was assessed at the first antenatal visit using three responses: no-smoking, 1–9 cigarettes per day (moderate smoking), and 10+ cigarettes per day (high). Analyses based on the collapsing the moderate and high levels were commensurate with those presented here (results available upon request). Previous studies found that only 6% of self-reported nonsmokers in Sweden had serum cotinine levels indicating active smoking.<sup>41</sup> Studies in the US<sup>42</sup> and Sweden<sup>43</sup> also suggest the validity of self-reported smoking has not changed over time. In the current sample 28.8% of the offspring were exposed to SDP; documentation of decline in SDP during this time span is available elsewhere.<sup>2</sup>

The MBR also included maternal age at birth, the birth order, and the gestational age of each child. Family structure at the time of birth was also assessed. Women reported either cohabiting or not cohabiting with the father of the child at childbirth.

A standard measure of socioeconomic status of both the mothers and fathers, as assessed by the Census, was also included in the analyses.<sup>44</sup> The groupings include (a) unskilled blue collar worker, (b) skilled blue collar worker, (c) low-level white collar worker, (d) intermediate-level white collar worker, (e) high-level white collar worker, (f) self-employed, (g) employed but uncategorized, and (h) not working/missing. The final category included individuals too young to be assessed (below the age of 15) and individuals that were not working at the time of the census (because they were students or unemployed). Because previous research has shown that the groupings cannot be ordered,<sup>45</sup> dummy codes were created to compare each occupational group to the unskilled blue collar worker category.

Maternal and paternal highest level of education was based on seven categories: (a) less than 9 years of primary and lower secondary education, (b) 9 years of primary and lower secondary education, (c) 1–2 years of upper secondary education, (d) 3 years of upper secondary education, (e) less than 3 years of post-secondary education, (f) more than 3 years of post-secondary education, and (g) postgraduate education. Dummy codes were used to compare higher levels of education to having completed less than 9 years of primary education. Maternal and Paternal history of criminal behavior for mothers and fathers was based on the presence or absence of a conviction for any criminal offense in the NCR through 2004.

Demographic characteristics are presented in Table 1. Previous analyses using the current sample have found that maternal SDP is associated with all of the covariates.<sup>23, 27</sup> SDP is associated with earlier age at childbearing for mothers and fathers, earlier gestational age, greater likelihood the mother is not cohabiting with the biological father at childbirth; lower maternal and paternal occupation status, lower maternal and paternal socioeconomic status, and greater risk of maternal and paternal history of criminal convictions.

## Statistical Analyses

**Estimates of Risks Comparing Unrelated Individuals and Differentially Exposed Siblings**—For descriptive purposes, estimates of risk for violent and nonviolent convictions based on Kaplan Meier Estimates were calculated separately for unexposed and exposed individuals (collapsing the moderate and high SDP categories) in two samples. The first sample was based on all offspring and, thus, reflected the increased risk of criminal convictions associated with exposure to SDP in the total population. The second sample was based solely on differentially exposed siblings (N=29,842). The comparison of risk for convictions in differentially exposed siblings provides an estimate of the risk associated with SDP while controlling for background familial factors. If SDP caused convictions, siblings exposed to SDP would have greater risk for criminality than their siblings who were not exposed.<sup>31, 33, 34</sup>

**Survival Analyses**—Formal statistical analyses of the association between the levels of SDP and criminality were conducted using survival analyses because the measures of convictions were right censored (i.e. each offspring had not lived through risk period). The models used a sandwich estimator to take into account clustering of individuals in extended families. Both moderate and high levels of SDP were compared to no SDP during pregnancy in each model by using dummy codes. The first model regressed each outcome on maternal smoking during pregnancy using the entire sample, controlling for offspring sex and birth order.

The second model explored the association between SDP and criminality while controlling for measured covariates. The model included offspring-specific covariates, maternal traits, and paternal traits. The models were based on subsets of the sample with complete data (pairwise deletion); the models were fit with 566,183 individuals (93%).



The third model examined the association between SDP and each outcome comparing siblings who were differentially exposed when considering moderate and high levels of SDP (N=50,339). We used standard Fixed Effects Models<sup>46</sup> for survival analyses that fit separate hazard functions for each nuclear family.<sup>47</sup> The analyses controls for all factors shared by siblings, which estimates the association between SDP and convictions using within-mother variation in SDP. The models, therefore, compare siblings who were differentially exposed.<sup>48</sup> The analyses also controlled for offspring sex and birth order, which is important when conducting sibling comparison models.<sup>34</sup>

**Sensitivity Analyses**—Because of the size of the data and complexity of the analyses, more advanced approaches of handling missing values<sup>49</sup> in model two were difficult to fit. The first set of sensitivity analyses, however, tested how robust the results were to the use of pairwise deletion. Two additional models were run for each measure of criminality, a worst-case and a best-case scenario. The worst-case scenario replaced each missing value with the value conferring the greater risk for the outcome (e.g. parents were assigned a positive history of criminal convictions), whereas the best-case scenario assigned values associated with a decreased risk of each outcome (e.g. parents were assigned a negative history of criminal convictions). Comparing the size of the SDP parameter in these two conditions with the value from model two provided an estimate of the possible bias in the parameter values due to pairwise deletion.

Additional models testing the association between SDP and low military officer suitability were also conducted to see whether the results replicate the findings for criminality. Comparisons of unrelated individuals and differentially exposed siblings were conducted using logistic regressions. Previous studies have already reported on sibling comparison analyses of low academic achievement at the end of grade 9, suggesting familial confounding of the association between SDP and grades.<sup>23, 26</sup>

Finally, we tested the assumption inherent in the sibling comparison design that the results from women who varied their smoking over across pregnancies to women who did not. One way to test the assumption is to assess whether the association between SDP and each outcome among *unrelated* individuals is comparable in two groups: (a) a sample of women who varied their smoking across pregnancies and (b) a sample of women with more than one child who did not vary their smoking across pregnancies (the women never smoked or smoked the same in each pregnancy).<sup>4</sup> Survival models assessed the between-family association between SDP and each outcome in the subset of offspring from the first group (N=50,339) and second group (N=337,168) separately. We then used interaction effects to test whether the magnitude of the associations between the two groups were statistically significant in a subset of the data that included all offspring of women with more than one child (combining the two groups described above). The analyses controlled for offspring sex, birth order, and number of children in each family.

## Results

**Estimates of Risks Comparing Unrelated Individuals and Differentially Exposed Siblings**—Kaplan Meier Estimates of risk for violent criminal convictions are presented graphically in Figure 1. The first panel graphs the increasing risk of being convicted of a violent offense across the ages of 15–21 years of age for the entire sample, depending on exposure to maternal SDP. As the graph illustrates, maternal SDP was associated with roughly a three-fold increased in criminal convictions of a violent nature. The second panel presents the risk for violent offenses in the sample of differentially exposed siblings. The figure indicates that exposed and unexposed siblings had similar rates of convictions of violent crime.

Figure 2 presents the Kaplan Meier Estimates for convictions for nonviolent offenses. The first panel, based on the entire sample, suggests offspring exposed to SDP were at increased risk for being convicted of a nonviolent offense. The second panel, based on the subset of differentially exposed siblings, however, indicates that exposed and unexposed siblings had equivalent rates of nonviolent convictions. The figures suggest that the association between SDP and criminal convictions was due to background familial risks.<sup>33, 34</sup>

**Survival Analyses**—The results of the survival analyses predicting violent offenses are presented in Table 2. Model one, which estimates the raw association with SDP, found that offspring exposed to moderate (HR=2.47, CI=2.34–2.60) and high levels (HR=3.43, CI=3.25–3.63) of SDP were more likely to be convicted of a violent offense. The inclusion of the measured covariates in model two reduced the associations, but SDP still uniquely predicted violent convictions (HR<sub>moderate</sub>=1.57, CI=1.48–1.67; HR<sub>high</sub>=1.87, CI=1.42–1.99). Model three compared differentially exposed siblings using Fixed Effects Models. The results indicate that SDP was not associated with violent offenses when comparing differentially exposed siblings (HR<sub>moderate</sub>=1.02, CI=0.79–1.30; HR<sub>high</sub>=1.03, CI=0.78–1.37), signifying that SDP was not associated with violent offenses when controlling for background familial factors.

The parameter estimates of the survival models predicting offspring nonviolent offenses are presented in Table 3. Model one indicated that SDP is associated with an increased risk of nonviolent offenses (HR<sub>moderate</sub>=1.62, CI=1.58–1.66; HR<sub>high</sub>=1.87, CI=1.82–1.92) in the entire population. Model two indicated that SDP still robustly predicted nonviolent offenses when statistically controlling for the measured covariates (HR<sub>moderate</sub>=1.31, CI=1.27–1.34; HR<sub>high</sub>=1.38, CI=1.34–1.42). Model three revealed no associations between levels of SDP and nonviolent convictions (HR<sub>moderate</sub>=0.89, CI=0.78–1.01; HR<sub>high</sub>=0.89, CI=0.78–1.02) when comparing differentially exposed siblings, findings consistent with the results from the models predicting violent convictions. Survival analyses on subsets of the data based on year of birth (1983–1986 and 1987–1989) were consistent with those presented in the text (results available upon request).

**Sensitivity Analyses**—Two additional models, a worst- and best-case scenario, were run to test whether the results from model two were robust to the assumptions of generalizability given pairwise deletion. For each measure of criminality the first two decimal places for each unstandardized parameter estimate (logits) did not change (full results available upon request). The sensitivity analyses, therefore, indicate that the results of the models that controlled for measured covariates were quite robust.

SDP was associated with increased risk for low military officer suitability in the entire population (OR<sub>moderate</sub>=1.34, CI=1.25–1.44; OR<sub>high</sub>=1.65, CI=1.52–1.46), even when controlling for the measured covariates (OR<sub>moderate</sub>=1.16, CI=1.08–1.25; OR<sub>high</sub>=1.33, CI=1.21–1.46). But, the comparison of differentially exposed siblings using Fixed Effects Models (OR<sub>moderate</sub>=0.83, CI=0.61–1.13; OR<sub>high</sub>=0.93, CI=0.73–1.19) suggests no association, which parallels the findings for criminal convictions.

We tested whether the association between SDP and convictions using *unrelated* individuals differed for mothers who did and did not vary their smoking across multiple pregnancies. The association between SDP and violent criminal offenses in unrelated individuals in the subset of women with variation in SDP across pregnancies (HR<sub>moderate</sub>=2.29, CI=1.62–3.24; HR<sub>high</sub>=3.63, CI=2.82–4.66) was somewhat smaller than in women who did not vary their smoking (HR<sub>moderate</sub>=2.77, CI=2.55–3.01; HR<sub>high</sub>=4.10, CI=3.77–4.46), but the differences were small and not statistically significant ( $b_{\text{logits-moderate}}=-0.18$ ,  $SE=0.18$ ,  $p=0.31$ ;  $b_{\text{logits-high}}=-0.13$ ,  $SE=0.13$ ,  $p=0.33$ ). The findings for nonviolent convictions were similar.

The associations in women without variation in SDP ( $HR_{\text{moderate}}=1.44$ ,  $CI=1.21-1.73$ ;  $HR_{\text{high}}=1.77$ ,  $CI=1.55-2.01$ ) were generally consistent with the association in women who did not vary their smoking ( $HR_{\text{moderate}}=1.65$ ,  $CI=1.58-1.71$ ;  $HR_{\text{high}}=2.04$ ,  $CI=1.95-2.13$ ), and the differences were not statistically significant ( $b_{\text{logits-moderate}}=-0.13$ ,  $SE=0.09$ ,  $p=0.16$ ;  $b_{\text{logits-high}}=-0.13$ ,  $SE=0.07$ ,  $p=0.08$ ). The sensitivity analyses, thus, indicate the findings from the Fixed Effects Models are not due to the differences between women with and without variation in their SDP across their pregnancies.

## Comment

The current study, which was based on a population-based study of offspring born in Sweden over seven years, explored the risk associated with SDP for violent and nonviolent convictions, acts associated with enormous societal and personal costs. Although the associations between SDP and violent and nonviolent criminality showed a dose-dependent response and were robust when controlling for measured traits of both mothers and fathers, differentially exposed siblings had the same risk for being convicted. The results suggest that the increased risk of convictions associated with SDP is due to familial background factors, not the putative biological effects of SDP.

The results for criminality support and extend the findings of recent sibling comparison<sup>5, 6</sup> and IVF<sup>29</sup> studies of SDP that were based on smaller samples and assessed conduct problem during childhood.<sup>31</sup> The findings are also commensurate with previous quasi-experimental studies of traits associated with delinquency, such as poor academic achievement,<sup>23, 26</sup> low intellectual abilities,<sup>27</sup> and ADHD.<sup>28, 30</sup> It is important to note the findings for convictions were replicated using highly correlated assessments based on independent evaluations (military officer suitability), school grades<sup>23, 26</sup>, and standardized measures of achievement<sup>23</sup> in the same sample. The use of quasi-experimental studies of SDP as a whole<sup>4</sup>, therefore, seriously calls into question causal inferences concerning the teratogenic effects of SDP for certain later neurocognitive and psychiatric problems in offspring.

Most research on SDP has typically found robust statistical associations between SDP and disruptive behaviors while controlling for correlated risks,<sup>2, 13</sup> as was the case in the current study. The results of the sibling comparison analyses, therefore, imply that familial factors that are not frequently measured (or measured well) in research protocols are actually responsible for the increased risk in offspring whose mothers smoke during pregnancy. Future research will need to include better assessment possible environmental confounds (e.g., maternal report of conduct disorder as a teenager<sup>21</sup> and antisocial behavior of both parents<sup>18</sup>) and use designs that can examine whether shared genetic liability accounts for the statistical associations.<sup>23, 29</sup> Designs that can further elucidate the underlying causal mechanisms include randomized controlled interventions targeting SDP<sup>50</sup>, studies exploring variation in SDP within pregnancies<sup>51</sup>, different patterns of SDP across pregnancies (comparing women who stop smoking versus those who begin smoking over time)<sup>52</sup>, adoptions studies<sup>53</sup>, and the children of twins/siblings<sup>5, 20, 23, 54</sup>, to name a few possibilities.<sup>4, 32</sup>

The study also has some limitations which must be considered. Using criminal convictions as a measure of delinquency can limit the interpretation of the results. Children less than 15 years of age cannot be convicted, and certain crimes (e.g., homicide) have a higher probability of leading to a verdict. The use of convictions inevitably covers less than the true prevalence of criminal behavior, but this will not necessarily affect the magnitude of the association with SDP because findings for criminality based on self report and official records are often similar across different exposures.<sup>55</sup> The assessment of convictions also was only available until the offspring reached 21.9 years old, which does not cover the entire risk period for convictions. Future research will need to study the association between SDP



and criminality throughout adulthood and specifically study whether SDP is independently associated with other outcomes, such as drug and alcohol problems.<sup>56</sup> The findings for convictions, however, were consistent with those for low military officer suitability, a measure available on all males in Sweden at the age of 18, and other traits in the entire Swedish sample (e.g., academic achievement<sup>23, 26</sup> and intellectual abilities<sup>27</sup>) that are highly associated with criminality.

The assessment of SDP is based on maternal report, which is not as accurate as using biological assays to measure extent of smoking, and the measure only differentiated between moderate and high levels of SDP. SDP was only assessed during the first trimester of pregnancy, so we were also unable to examine possible differences in the timing or pattern of use. Such assessment were included in the Medical Birth Registry in later years, which will allow researchers to examine such questions once those offspring reach late adolescence.

The sibling comparison design also has a number of inherent assumptions.<sup>4, 34</sup> We examined whether women who varied their smoking across multiple pregnancies were comparable to women who did not. The results suggest that the association between SDP and criminality was slightly less in women who altered their smoking, but the differences were small and not statistically significant. The design can only account for environmental factors that influence siblings similarly and also assumes that traits of the offspring do not influence maternal SDP.<sup>23</sup>

Furthermore, we did not explore moderators of the association between SDP and offspring criminality (e.g., pregnancy complications<sup>11</sup> or genetic risk measured by candidate genes<sup>4</sup>). The current findings, however, have implications for such research. SDP may only have influence offspring who are particularly at risk, or the identified moderators could be interacting with background familial factors.

Given the limitations of the current study and of previous quasi-experimental research, definitive conclusions concerning SDP can only be drawn from considering all research on the topic.<sup>4, 32</sup> The challenge for the research field is to incorporate the recent quasi-experimental findings, which implicate the importance of familial confounding, with the results from the existent traditional family studies that support the causal hypothesis. More translational research is also needed to understand how these findings fit with animal studies, which some researchers suggest are not conclusive about the association between SDP and aggressive behavior.<sup>30, 57</sup> Certainly, more interdisciplinary research is needed to truly understand mechanisms underlying the association between SDP and offspring disruptive problems. The current study, as well as other recent reports, suggest that quasi-experimental studies will play an important role in the endeavor.<sup>4</sup>

It is important to note, however, that the results of the current study do not indicate that SDP is benign. Recent quasi-experimental studies suggest that SDP causes pregnancy-related problems. Children of twins<sup>20</sup>, sibling comparison<sup>5, 6</sup> (including the current sample<sup>23</sup>), prenatal IVF cross-fostering,<sup>29</sup> and intervention studies<sup>58</sup> indicate that the association between SDP and offspring birth weight is consistent with a causal effect. A recent study from Sweden that compared consecutive births also found that SDP is independently associated with infant mortality.<sup>59</sup> Reducing SDP, therefore, remains an important public health issue.

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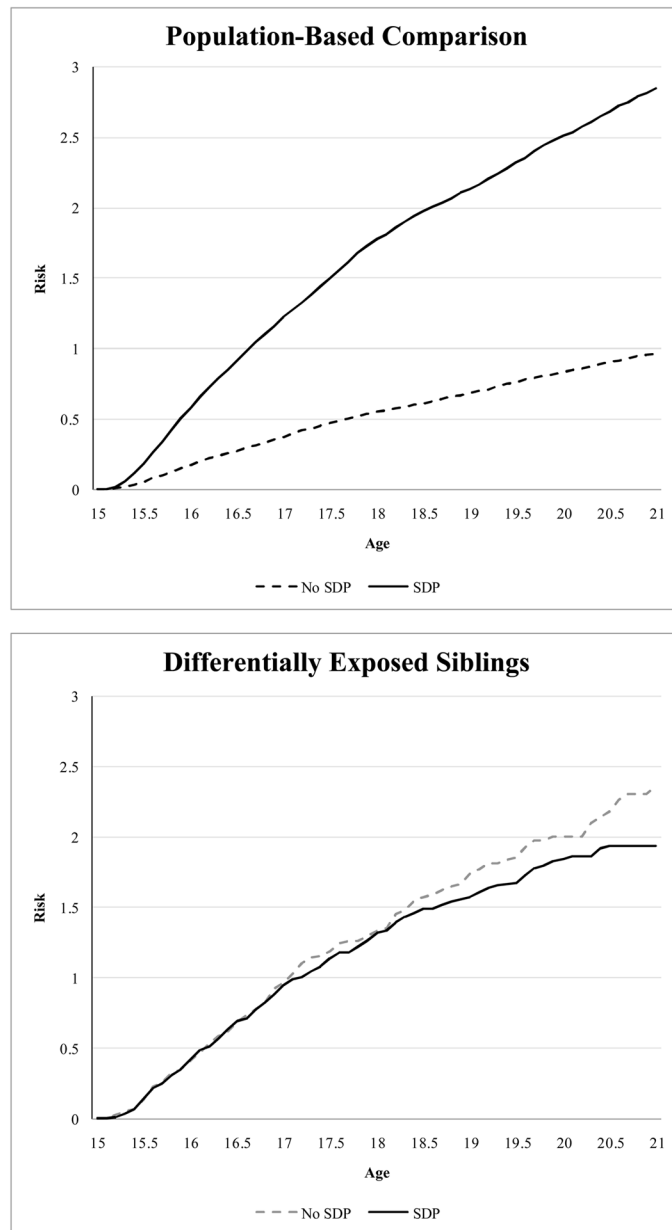
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## References

1. Wakschlag LS, Hans SL. Maternal smoking during pregnancy and conduct problems in high-risk youth: A developmental framework. *Development and Psychopathology*. 2002; 14:351–369. [PubMed: 12030696]
2. Cnattingius S. The epidemiology of smoking during pregnancy: Smoking prevalence, maternal characteristics, and pregnancy outcomes. *Nicotine and Tobacco Research*. 2004; 6:S125–S140. [PubMed: 15203816]
3. Huizink AC, Mulder EJH. Maternal smoking, drinking or cannabis use during pregnancy and neurobehavioral and cognitive functioning in human offspring. *Neuroscience and Biobehavioral Reviews*. 2006; 30:24–41. [PubMed: 16095697]
4. Knopik VS. Maternal smoking during pregnancy and child outcomes: Real or spurious effect? *Developmental Neuropsychology*. 2009; 34:1–36. [PubMed: 19142764]
5. D'Onofrio BM, Van Hulle CA, Waldman ID, et al. Smoking during pregnancy and offspring externalizing problems: An exploration of genetic and environmental confounds. *Development and Psychopathology*. 2008; 20:139–164. [PubMed: 18211732]
6. Gilman SE, Gardener H, Buka SL. Maternal smoking during pregnancy and children's cognitive and physical development: A causal risk factor? *American Journal of Epidemiology*. 2008; 168:522–531. [PubMed: 18653646]
7. Wakschlag LS, Keenan K. Clinical significance and correlates of disruptive behavior symptoms in environmentally at-risk preschoolers. *Journal of Clinical Child Psychology*. 2001; 30:262–275. [PubMed: 11393926]
8. Fergusson DM, Woodward LJ, Horwood LJ. Maternal smoking during pregnancy and psychiatric adjustment in late adolescence. *Archives of General Psychiatry*. 1998; 55:721–727. [PubMed: 9707383]
9. Wakschlag LS, Lahey BB, Loeber R, Green SM, Gordon RA, Leventhal BL. Maternal smoking during pregnancy and the risk of conduct disorder in boys. *Archives of General Psychiatry*. 1997; 54:670–676. [PubMed: 9236551]
10. Weissman MM, Warner V, Wickramaratne PJ, Kandel DB. Maternal smoking during pregnancy and psychopathology in offspring followed to adulthood. *Journal of the American Academy of Child and Adolescent Psychiatry*. 1999; 38:892–899. [PubMed: 10405508]
11. Brennan PA, Grekin ER, Mednick SA. Maternal smoking during pregnancy and adult male criminal outcomes. *Archives of General Psychiatry*. 1999; 56:215–219. [PubMed: 10078497]
12. Rasanen P, Hakko H, Isohanni M, Hodgins S, Jarvelin J, Tiihonen J. Maternal smoking during pregnancy and risk of criminal behavior among adult male offspring in the Northern Finland 1966 birth cohort. *American Journal of Psychiatry*. 1999; 156:857–862. [PubMed: 10360123]
13. Wakschlag LS, Pickett KE, Cook E, Benowitz NL, Leventhal BL. Maternal smoking during pregnancy and severe antisocial behavior in offspring: A review. *American Journal of Public Health*. 2002; 92:966–974. [PubMed: 12036791]
14. Fergusson DM. Prenatal smoking and antisocial behavior. *Archives of General Psychiatry*. 1999; 56:223–224. [PubMed: 10078498]
15. Cnattingius S, Reilly M, Pawitan Y, Lichtenstein P. Maternal and fetal genetic factors account for most of familial aggregation of preeclampsia: a population-based Swedish cohort study. *Am J Med Genet A*. Nov 1; 2004 130(4):365–371. [PubMed: 15384082]
16. Matthews TJ. Smoking during pregnancy in the 1990s. *National Vital Statistics Reports*. 2001; 49(7)
17. Zimmer MH, Zimmer M. Socioeconomic determinants of smoking behavior during pregnancy. *The Social Science Journal*. 1998; 35:133–142.

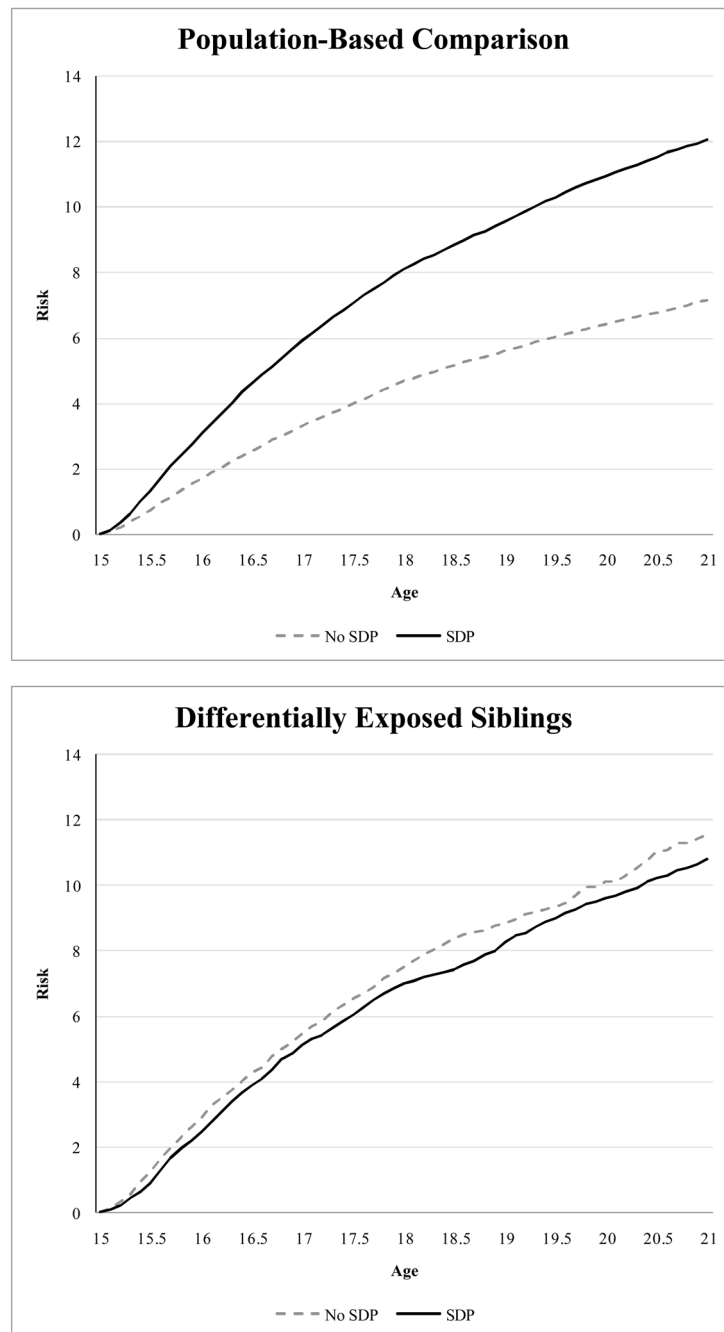
18. Maughan B, Taylor A, Caspi A, Moffitt TE. Prenatal smoking and early childhood conduct problems. *Archives of General Psychiatry*. 2004; 61:836–843. [PubMed: 15289282]
19. Brook JS, Brook DW, Whiteman M. The influence of maternal smoking during pregnancy on the toddler's negativity. *Archives of Pediatric and Adolescent Medicine*. 2000; 154:381–385.
20. D'Onofrio BM, Turkheimer E, Eaves LJ, et al. The role of the Children of Twins design in elucidating causal relations between parent characteristics and child outcomes. *Journal of Child Psychology and Psychiatry*. 2003; 44:1130–1144. [PubMed: 14626455]
21. Silberg JL, Parr T, Neale MC, Rutter M, Angold A, Eaves LJ. Maternal smoking during pregnancy and risk to boys' conduct disturbance: An examination of the causal hypothesis. *Biological Psychiatry*. 2003; 53:130–135. [PubMed: 12547468]
22. Moffitt TE. The new look of behavioral genetics in developmental psychopathology: Gene-environment interplay in antisocial behaviors. *Psychological Bulletin*. 2005; 131:533–554. [PubMed: 16060801]
23. D'Onofrio BM, Singh AL, Iliadou A, et al. A quasi-experimental study of maternal smoking during pregnancy and offspring academic achievement. *Child Development*. in press.
24. Agrawal A, Knopik VS, Pergadia ML, et al. Correlates of cigarette smoking during pregnancy and its genetic and environmental overlap with nicotine dependence. *Nicotine and Tobacco Research*. 2008; 10:567–578. [PubMed: 18418779]
25. Scarr S, McCartney K. How people make their own environments: A theory of genotype->environment effects. *Child Development*. 1983; 54:424–435. [PubMed: 6683622]
26. Lambe M, Hultman C, Torrang A, MacCabe J, Cnattingius S. Maternal smoking during pregnancy and school performance at age 15. *Epidemiology*. 2006; 17:524–530. [PubMed: 16878043]
27. Lundberg F, Cnattingius S, D'Onofrio B, et al. Maternal smoking during pregnancy and intellectual performance in young adult Swedish male offspring. Submitted.
28. Knopik VS, Sparrow EP, Madden PAF, et al. Contributions of parental alcoholism, prenatal substance exposure, and genetic transmission to child ADHD risk: A female twin study. *Psychological Medicine*. 2005; 35:625–635. [PubMed: 15918339]
29. Rice, F.; Harold, GT.; Boivin, J.; Hay, DF.; Van den Bree, M.; Thapar, A. Disentangling prenatal and inherited influences in humans with an experimental design. *Proceedings of the National Academy of Sciences of the United States of America*; 2009. Early Edition Online
30. Thapar A, Rice F, Hay D, et al. Prenatal smoking may not cause ADHD. Evidence from a novel design. *Biological Psychiatry*. in press.
31. Rutter M. Proceeding from observed correlation to causal inference: The use of natural experiments. *Perspectives on Psychological Science*. 2007; 2:377–395.
32. Rutter M, Pickles A, Murray R, Eaves LJ. Testing hypotheses on specific environmental causal effects on behavior. *Psychological Bulletin*. 2001; 127:291–324. [PubMed: 11393298]
33. Rodgers JL, Cleveland H, van den Oord E, Rowe D. Resolving the debate over birth order, family size, and intelligence. *American Psychologist*. 2000; 55:599–612. [PubMed: 10892201]
34. Lahey BB, D'Onofrio BM, Waldman ID. Using epidemiologic methods to test hypotheses regarding causal influences on child and adolescent mental disorders. *Journal of Child Psychology and Psychiatry*. in press
35. Swedish\_Centre\_for\_Epidemiology. Welfare NBoHa. The Swedish Medical Birth Registry: A summary of content and quality. 2001.
36. Magnusson PKE, Gunnell D, Tynelius P, Smith GD, Rasmussen F. Strong inverse association between height and suicide in a large cohort of Swedish men: Evidence of early life origins of suicidal behavior? *American Journal of Psychiatry*. 2005; 162:1373. [PubMed: 15994722]
37. Statistics\_Sweden. Multi-Generation Register 2002, A description of contents and quality. Örebro; Sweden: 2003.
38. Fazel S, Grann M, Carlström E, Lichtenstein P, Långström N. Risk factors for violent crime in schizophrenia: a national cohort study of 13,806 patients. *Journal of Clinical Psychiatry*. in press.
39. Långström N, Grann M, Ruchkin V, Sjöstedt G, Fazel S. Violent offending in autism spectrum disorder: National study of risk factors among hospitalised individuals. *Journal of Interpersonal Violence*. 2008

40. Otto U. Function of male youths during military service: A follow-up study of youth clientele. *Acta Psychiatrica Scandinavica*. 1980; 282:5–60.
41. Lindqvist R, Lendahls L, Tollbom O, Aberg H, Hakansson A. Smoking during pregnancy: Comparison of self-reports and cotinine levels in 496 women. *Acta Obstetrica et Gynecologica Scandinavica*. 2002; 81:240–244. [PubMed: 11966481]
42. Hatziaandreu EJ, Pierce JP, Fiore MC, Grise V, Novotny TE, Davis RM. The reliability of self-reported cigarette consumption in the United States. *American Journal of Public Health*. 1989; 79:1020–1023. [PubMed: 2751017]
43. Cnattingius S, Haglund B. Decreasing smoking prevalence during pregnancy in Sweden: the effect on small-for-gestational-age births. *American Journal of Public Health*. 1997; 87:410–413. [PubMed: 9096542]
44. Statistics\_Sweden. Swedish socioeconomic classification: Reports on statistical coordination. Orebro; Sweden: 1982.
45. Lichtenstein P, Pedersen NL, McClearn GE. The origins of individual differences in occupational status and educational level. *Acta Sociologica*. 1992; 35:13–31.
46. Greene, WH. *Econometric Analysis*. 5. Upper Saddle River, NJ: Prentice-Hall; 2003.
47. Allison, PD. *Fixed Effects Regression Models*. Washington DC: Sage; 2009.
48. Neuhaus JM, McCulloch CE. Separating between- and within-cluster covariate effects by using conditional and partitioning methods. *Journal of the Royal Statistical Society*. 2006; 68:859–872.
49. Schafer JL, Graham JW. Missing data: Our view of the state of the art. *Psychological Methods*. 2002; 7:147–177. [PubMed: 12090408]
50. Sexton M, Hebel JR. A Clinical Trial of Change in Maternal Smoking and Its Effect on Birth Weight. *JAMA*. 1984; 251:911–915. [PubMed: 6363731]
51. Pickett KE, Wood C, Adamson J, Desouza L, Wakschlag LS. Meaningful differences in maternal smoking behaviour during pregnancy: implications for infant behavioural vulnerability. *J Epidemiol Community Health*. 2008; 62:318–324. [PubMed: 18339824]
52. Meyer KA, Williams P, Hernandez-Diaz S, Cnattingius S. Smoking and the risk of oral clefts: Exploring the impact of study designs. *Epidemiology*. 2004; 15:671–678. [PubMed: 15475715]
53. Neiderhiser JM, Leve LD, Ge X, et al. The impact of prenatal drug exposure on toddler behavior: Distinguishing genetic effects from exposure using an adoption design. *Behavior Genetics*. 2007; 37:780.
54. Knopik VS, Heath AC, Jacob T, et al. Maternal alcoholism and offspring ADHD: Disentangling genetic and environmental effects using a children-of-twins design. *Psychological Medicine*. 2006; 2006:1461–1471. [PubMed: 16734942]
55. Arseneault L, Moffitt TE, Caspi A, Taylor PJ, Silva PA. Mental disorders and violence in a total birth cohort: results from the Dunedin Study. *Archives of General Psychiatry*. 2000; 57:979–986. [PubMed: 11015816]
56. Glantz MY, Campbell Chompers J. Prenatal drug exposure effects on subsequent vulnerability to drug abuse. *Development and Psychopathology*. 2006; 18:893–922. [PubMed: 17152406]
57. Winzer-Serhan UH. Long-term consequences of maternal smoking and developmental chronic nicotine exposure. *Front Biosci*. 2008; 13:636–649. [PubMed: 17981576]
58. Sexton M, Hebel RH. A clinical trial of change in maternal smoking and its effect on birth weight. *JAMA*. 1984; 251:911–915. [PubMed: 6363731]
59. Johansson ALV, Dickman PW, Kramer MS, Cnattingius S. Maternal smoking and infant mortality: Does quitting smoking reduce the risk of infant death? *Epidemiology*. 2009; 20:1–8. [PubMed: 19234393]



**Figure 1.** Association between Maternal Smoking during Pregnancy and Offspring Convictions for Violent Offenses  
 Note. SDP=Smoking during pregnancy. The first panel presents the risk across the age range for the entire population. The second panel presents the risk in the sample of differentially exposed siblings. Estimates of risk are based on Kaplan Meier Estimates controlling for offspring sex and birth order.





**Figure 2.** Association between Maternal Smoking during Pregnancy and Offspring Convictions for Nonviolent Offenses  
 Note. SDP=Smoking during pregnancy. The first panel presents the risk across the age range for the entire population. The second panel presents the risk in the sample of differentially exposed siblings. Estimates of risk are based on Kaplan Meier Estimates controlling for offspring sex and birth order.

**Table 1**

## Demographic Characteristics

Variable	N	Mean (SD)/%
Maternal age at childbirth (years)	609372	29.9 (5.1)
Paternal age at childbirth (years)	606951	31.4 (5.9)
Gestational age (weeks)	608895	39.8 (1.7)
Offspring Sex	609372	48.6 % (Female)
Birth order	609372	
First Born		41.6%
Second Born		35.5%
Third Born		16.4%
Fourth + Born		6.4%
Parents not cohabiting at childbirth	599110	2.7%
Maternal Education	601271	
Maternal less than 9 years of education		2.4%
9 years of education		11.2%
1–2 years upper secondary education		38.5%
3 years upper secondary education		13.3%
Less than 3 years post-secondary education		16.8%
3+ years post-secondary education		17.4%
Postgraduate education		0.5%
Maternal Occupation	608551	
Maternal unskilled blue collar		28.6%
Skilled blue collar		11.2%
Low-level white collar		13.9%
Intermediate-level white collar		16.9%
High-level white collar		6.0%
Self-employed		2.6%
Employed, uncategorized		6.2%
Not working/Missing		14.6%
Maternal History of criminal conviction	609372	11.9%
Paternal Education	583771	
Paternal less than 9 years of education		5.6%
9 years of education		16.3%
1–2 years upper secondary education		37.5%
3 years upper secondary education		12.3%
Less than 3 years post-secondary education		12.8%
3+ years post-secondary education		14.0%
Postgraduate education		1.4%
Paternal Occupation	603283	
Paternal unskilled blue collar		21.1%
Skilled blue collar		22.1%

<b>Variable</b>	<b>N</b>	<b>Mean (SD)/%</b>
Low-level white collar		8.3%
Intermediate-level white collar		15.6%
High-level white collar		13.3%
Self-employed		7.5%
Employed, uncategorized		4.8%
Not working/Missing		7.3%
Paternal History of criminal conviction		40.7%

Table 2

## Survival Models Predicting Violent Crimes

Risk Factor	Model 1 <sup>a</sup>		Model 2 <sup>b</sup>		Model 3 <sup>c</sup>	
	b	SE	b	SE	b	SE
No SDP	1.00 - Ref		1.00 - Ref		1.00 - Ref	
Moderate SDP (1-9 cigarettes/day)	0.90*	0.03	0.45*	0.03	0.02	0.12
High SDP (10+ cigarettes/day)	1.23*	0.03	0.62*	0.03	0.03	0.14
Offspring Sex	-1.54*	0.03	-1.57*	0.03	-1.40*	0.07
Birth order	0.10*	0.01	0.22*	0.01	0.01	0.03
Maternal age at childbirth (years)			-0.06*	0.00		
Paternal age at childbirth (years)			-0.01*	0.00		
Gestational age (weeks)			0.01*	0.00		
Parents not cohabiting at childbirth			0.46*	0.05		
Maternal less than 9 years of education			-	-		
9 years of education			-0.13*	0.06		
1-2 years upper secondary education			-0.25*	0.06		
3 years upper secondary education			-0.46*	0.08		
Less than 3 years post-secondary education			-0.45*	0.08		
3+ years post-secondary education			-0.59*	0.08		
Postgraduate education			-1.25*	0.51		
Maternal unskilled blue collar			-	-		
Skilled blue collar			-0.09*	0.04		
Low-level white collar			-0.12*	0.04		
Intermediate-level white collar			-0.14*	0.06		
High-level white collar			-0.24*	0.10		
Self-employed			-0.18*	0.09		
Employed, uncategorized			0.10*	0.05		
Not working/Missing			0.10*	0.03		

Risk Factor	Model 1 <sup>a</sup>		Model 2 <sup>b</sup>		Model 3 <sup>c</sup>	
	b	SE	b	SE	b	SE
Maternal History of criminal conviction	0.57*	0.03	-	-	-	-
Paternal less than 9 years of education	-0.11*	0.05	-0.12*	0.04	-0.22*	0.06
9 years of education	-0.12*	0.04	-0.22*	0.06	-0.40*	0.07
1-2 years upper secondary education	-0.22*	0.06	-0.40*	0.07	-0.50*	0.08
3 years upper secondary education	-0.40*	0.07	-0.50*	0.08	-0.57*	0.23
Less than 3 years post-secondary education	-0.50*	0.08	-0.57*	0.23	-	-
3+ years post-secondary education	-0.57*	0.23	-	-	-0.12*	0.03
Postgraduate education	-	-	-0.19*	0.06	-0.30*	0.05
Paternal unskilled blue collar	-	-	-0.33*	0.07	-0.26*	0.05
Skilled blue collar	-0.12*	0.03	-0.26*	0.05	0.20*	0.05
Low-level white collar	-0.19*	0.06	0.20*	0.05	0.29*	0.04
Intermediate-level white collar	-0.30*	0.05	0.29*	0.04	0.73*	0.03
High-level white collar	-0.33*	0.07	0.73*	0.03	-	-
Self-employed	-0.26*	0.05	-	-	-	-
Employed, uncategorized	0.20*	0.05	-	-	-	-
Not working/Missing	0.29*	0.04	-	-	-	-
Paternal History of criminal conviction	0.73*	0.03	-	-	-	-

Note.

<sup>a</sup>Based on the total sample (N=609,372).

<sup>b</sup>Based on data with no missing data (N=566,183).

<sup>c</sup>Based on fixed effects model that examined differentially exposed siblings (N= SDP=50,339). SDP=Smoking during pregnancy. Parameters are distributed as logits.

\*  $P < 0.05$ .



Table 3

## Survival Models Predicting Nonviolent Crimes

Risk Factor	Model 1 <sup>a</sup>		Model 2 <sup>b</sup>		Model 3 <sup>c</sup>	
	b	SE	b	SE	b	SE
No SDP	1.00 - Ref		1.00 - Ref		1.00 - Ref	
Moderate SDP (1–9 cigarettes/day)	0.48*	0.01	0.27*	0.01	-0.12	0.07
High SDP (10+ cigarettes/day)	0.63*	0.01	0.32*	0.02	-0.12	0.07
Offspring Sex	-0.71*	0.01	-0.72*	0.01	-0.68*	0.011
Birth order	0.02*	0.00	0.07*	0.01	0.08*	0.02
Maternal age at childbirth (years)			-0.02*	0.00		
Paternal age at childbirth (years)			0.00	0.00		
Gestational age (weeks)			0.02*	0.00		
Parents not cohabiting at childbirth			0.25*	0.03		
Maternal less than 9 years of education			-	-		
9 years of education			-0.01	0.04		
1–2 years upper secondary education			-0.03	0.03		
3 years upper secondary education			-0.09*	0.04		
Less than 3 years post-secondary education			-0.15*	0.04		
3+ years post-secondary education			-0.10*	0.04		
Postgraduate education			0.00	0.10		
Maternal unskilled blue collar			-	-		
Skilled blue collar			-0.03	0.02		
Low-level white collar			-0.07*	0.02		
Intermediate-level white collar			-0.07*	0.02		
High-level white collar			-0.08*	0.03		
Self-employed			0.01	0.04		
Employed, uncategorized			0.12*	0.02		
Not working/Missing			0.04*	0.02		
Maternal History of criminal conviction			0.37*	0.01		

Risk Factor	Model 1 <sup>a</sup>		Model 2 <sup>b</sup>		Model 3 <sup>c</sup>	
	b	SE	b	SE	b	SE
Paternal less than 9 years of education	-	-	-	-	-	-
9 years of education	0.03	0.03	0.03	0.03	0.03	0.03
1–2 years upper secondary education	0.01	0.02	0.01	0.02	0.01	0.02
3 years upper secondary education	-0.04	0.03	-0.04	0.03	-0.04	0.03
Less than 3 years post-secondary education	-0.10*	0.03	-0.10*	0.03	-0.10*	0.03
3+ years post-secondary education	-0.08*	0.03	-0.08*	0.03	-0.08*	0.03
Postgraduate education	-0.02	0.06	-0.02	0.06	-0.02	0.06
Paternal unskilled blue collar	-	-	-	-	-	-
Skilled blue collar	-0.01	0.02	-0.01	0.02	-0.01	0.02
Low-level white collar	-0.06*	0.03	-0.06*	0.03	-0.06*	0.03
Intermediate-level white collar	-0.10*	0.02	-0.10*	0.02	-0.10*	0.02
High-level white collar	-0.08*	0.03	-0.08*	0.03	-0.08*	0.03
Self-employed	-0.02	0.02	-0.02	0.02	-0.02	0.02
Employed, uncategorized	0.18*	0.03	0.18*	0.03	0.18*	0.03
Not working/Missing	0.22*	0.03	0.22*	0.03	0.22*	0.03
Paternal History of criminal conviction	0.43*	0.01	0.43*	0.01	0.43*	0.01

Note.

<sup>a</sup>Based on the total sample (N=609,372).

<sup>b</sup>Based on data with no missing data (N=566,183).

<sup>c</sup>Based on fixed effects model that examined differentially exposed siblings (N= SDP=50,339). SDP=Smoking during pregnancy. Parameters are distributed as logits.

\*  $p < 0.05$ .