Clarifying the role of maternal childhood trauma on offspring behavioral outcomes: A systematic review of genetic and environmental contributions to transgenerational associations

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Trans generational mechanisms linking maternal childhood trauma (MCT) to offspring behavioral outcome (OBO)



FIGURE 1. Causal influences on the association between maternal childhood trauma and offspring behavior

Dashed arrows: Environmental pathway Solid arrows: Genetic pathway Variables measured or indexed in color, unmeasured causal influences in gray

Transmission operating through environmental pathways alone:



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Layering in consideration of genetic influences



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Research Question

What is the state of the literature on the role of genetic and environmental factors in the transgenerational association between maternal childhood trauma (MCT) and offspring behavioral outcome (OBO)?

Method

- Systematic review: goal is to synthesize findings of all relevant studies on a certain topic/issue in order to make findings more digestible / accessible for other researchers or practitioners
- Followed the Preferred reporting items for systematic review and metaanalysis protocols (PRISMA-P) reporting guidelines (2015)
- Registered the review with the International Prospective Register of Systematic Reviews (PROSPERO) in October 2019

Eligibility criteria

- (1) published in a peer-reviewed journal in English
- (2) included measurement of maternal childhood trauma
- (3) included measurement of child variation in emotional and behavioral outcomes from six months to six years of age, and
- (4) inclusion of control for a genetic contribution to the association between (2) and (3).

Search Strategy & Screening Process



Author (Year)	Study Design	Sample Characteristics	Measure of MCT	Measure of OBO	Genetic	Analytic Methods	Baseline Association	Genetic
(real)					contribution		between wich and obo	Attendation
Bouxette, Turcot AA et al. (2015)	 Longitudinal, elinisally_ ascertained- sample (Maternal Adversity, Vulnerability, and Neurodevelopment Study, MAVAN) 	 Sample size: n=154 mothers-infant dyads, recruited 13-20 weeks gestation to offspring age 36 months Cohort Demographics: 88.7% European/Caucasian, 8.1% African descent/African American, 3.2% Hespanic/Latino 	 Measures used: Childhood Trauma Questionnaire (CTQ; Bernstein et al. 1994); Parental Bonding Instrument (PBI, Parker et al. 1979) Used previously validated principal component analysis to reduce measures from the two scales 	Measures used: Early Childhood Behaxiaua Questionnaire (ECBQ; Putnam et al. 2006); Strengths and Difficulties Questionnaire (SDQ; Goodman, 1997) at 60 months Principal component analysis done for ECBQ results to derive infant negative emotionality/behavioral dysregulation (NE/BR) at 18 and 36 months	Measurers used: Center for Epidemiologic Studies Depression Scale (CES- D, BadJoff, 1997) at 6 and 36 months postpartum • Genetic data? Y/N • Genetic data? Infant buccal swabs obtained at 36 months; Maternal buccal swabs obtained at 36 months	Statistical method: Multiple linear hierarchical regression Genetic analysis: DNA extraction and 5- HTTLPR genotyping; Compared La/La homozygote infants to S/Le allele carriers Other covariates: demographic data	Effect of maternal adversity on child NE/BR (r = .20, p < .01)	Significant nteraction effect of maternal adversity and offspring genotype on child NE/BR (B=1.03, p<.05)
Bouxette Turcot AA et al. (2019)	• Longitudinal, elinically_ accortained sample (MAVAN)	 Sample size: n=239 mothers-infant dyads, recruited 13-20 weeks gestation to offspring age 36 months Cohort Demographics: 88.7% European/Caucasian, 8.1% African descent/African American, 3.2% Hepanic/Latino 	 Measures used: Childhood Trauma Questionnaire (CTQ; Bernstein et al. 1994); Parental Bonding Instrument (PBI, Parker et al. 1979) Used previously validated principal 	Measures used: ECBQ at 36 months Principal component analysis done for ECBQ results to derive infant negative emotionality/behavioral dysregulation (NE/BR)	Measurers used: Edinburgh Postnatal Depression Scale (EPDS, Cox, Holden, & Sage, Sky, 1987) at 6 months • Genetic data? Y/N	Statistical method: Mediation analyses (direct and indirect effects with bootstrapped confidence intervals) Other covariates: maternal sensitivity (29- minute nonfeeding mother-infant	Significant total effect of maternal adversity on Court SEBR It \$3, p	Indirect effect for maternal adversity on child NE/BR via maternal depression: B = .033, p < .05

Finding 1: Baseline association between MCT and OBO

- 10 studies reported significant baseline associations between MCT and OBO
- 3 articles did not report or find a significant baseline association
 - Racine et al. (2018) infant outcomes subdivided into five different domains of the Ages and Stages Questionnaire
 - Madigan et al. (2017) ascertainment of infant outcomes based on scales that rated frequency of 8 behaviors on a 3-point scale (never, sometimes, often) with only adequate internal consistency
 - Hipwell et al. (2019) infant temperament measured on 6 items of fussy/difficult subscale with only adequate convergent validity; did find an association with infant negative reactivity during Still Face Paradigm

Finding 2: Genetic Attenuation of the Association between MCT and OBO

- Of the 10 articles that reported a significant baseline association between MCT and OBO, 7 articles reported attenuated effects when taking genetic influences into account
- 2 additional studies (Madigan et al. 2017, Racine et al) reported that the association operated exclusively through the indirect genetic pathway
- 3 studies did not find improved association between MCT and OBO
 - Choi et al. (2017): maternal depression attenuated relationship between MCT and bonding, but not offspring behavior specifically
 - McDonnell (2016): mediation model was not significant
 - Hipwell (2019): also did not find a baseline association

Discussion / Conclusion

- Observed associations between MCT and OBO may be at least partially mediated by genetic factors, in addition to environmental factors
- Current state of the literature demonstrates there is still a bit of a schism between environmental research in the social sciences vs genetic research
- In both cases, I believe articles would be enriched if they borrowed from one another increasing environmental considerations in genomics research and vice versa
- Next steps: use some of the methodological limitations of the papers included in this review to apply this question in a new secondary data analysis asking virtually the same question



Thank You & Questions