

Child maltreatment, adaptive functioning, and polygenic risk: A structural equation mixture model

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Abstract

This study used a structural equation mixture model to examine associations between child maltreatment, polygenic risk, and indices of adaptive functioning. Children aged 6 to 13 years ($N = 1,004$), half maltreated, half nonmaltreated, were recruited to attend a research day camp. Multi-informant indicators of prosocial behavior, antisocial behavior, withdrawn behavior, and depression were collected and used in a latent class analysis. Four classes emerged, characterizing “well-adjusted,” “externalizing,” “internalizing,” and “socially dominant” groups. Twelve genetic variants, previously reported in the Gene \times Environment literature, were modeled as one weighted polygenic risk score. Large main effects between maltreatment and adaptive functioning were observed (Wald = 35.3, $df = 3$, $p < .0001$), along with evidence of a small Gene \times Environment effect (Wald = 13.5, $df = 3$, $p = .004$), adjusting for sex, age, and covariate interaction effects.

In the year 2015, Child Protective Services in the United States received child maltreatment referrals involving approximately 7.2 million children (US Department of Health and Human Services, Administration for Children and Families, Administration on Children, Youth and Families, Children’s Bureau, 2017). Children with a history of abuse or neglect are often deprived of the intellectual stimulation, guaranteed safety, nutrition, and emotional support necessary for healthy development. Experiences of maltreatment place children at heightened risk of developing diverse forms of psychopathology, as well difficulties in academics, interpersonal relationships, and cognitive functioning (Cicchetti, 2016). The specific effects of particular subtypes of maltreatment are largely undetermined and extremely difficult to study as most maltreated children experience multiple subtypes, or *maltreatment multiplicity* (Cicchetti & Rogosch, 2012). Several studies demonstrate that with every additional form of caregiver victimization, maladaptive outcomes across multiple psychological domains increase in severity (Cicchetti & Rogosch, 2012; Edwards, Probst, Rodenhizer-Stämpfli, Gidycz, & Tansill, 2014). There is some evidence of subtype specificity such as the association between sexual abuse

and substance use in women, as well the role of physical abuse in the development of antisocial behaviors (Jaffee, Caspi, Moffitt, & Taylor, 2004; Wilsnack, Vogeltanz, Klassen, & Harris, 1997). However, in general, research shows that child maltreatment influences psychological functioning in a nonspecific (affecting multiple domains) and equivalent manner (no one type of maltreatment is particularly less damaging; Vachon, Krueger, Rogosch, & Cicchetti, 2015). Therefore, the focus of the current study is on impact of maltreatment multiplicity on children’s development.

Resilience and Child Maltreatment

Consistent with the developmental psychopathology framework of multifinality, influences across multiple systems, from genes to society, give rise to divergent trajectories of development for maltreated children (Cicchetti & Rogosch, 1996). That is, despite the trauma of abuse and neglect, not all maltreated children follow along paths of maladaptation. Instead, children have a remarkable capacity to demonstrate resilient functioning (Cicchetti, 2013). Resilience refers to the complex and dynamic capacity of individuals to recover or withstand adversities that have the potential to significantly undermine development (Masten, 2015; Sapienza & Masten, 2011). Rather than simply referring to an absence of psychopathology, resilience signifies that children have obtained competence in multiple domains of psychological health, or *adaptive functioning*, despite experiencing adversity. This categorical notion of manifesting resilience (having both experienced adversity and exhibiting adaptive functioning) leads many researchers to study and characterize resilience using *person-centered* statistical approaches. Person-centered

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statistical approaches attempt to characterize similarities and differences among individuals based on the assumption that the population is heterogeneous in terms of the joint distribution of the observable indicator variables (Masyn, 2013). However, modeling the dynamic processes involved in obtaining adaptive functioning in contexts of adversity, such as interactions between risk and protective factors, is generally best achieved using *variable-centered* approaches. Variable-centered, as opposed to person-centered, approaches tend to analyze the relations among variables. Hybrid models have grown in popularity because of their ability to incorporate both person-centered and variable-centered approaches to understanding resilience (Masten, 2015). One type of hybrid model used in this study is referred to as a structural equation mixture model (SEMM). An SEMM involves testing a structural model with one or more categorical latent variables along with one or more continuous factors.

Protective and vulnerability factors

A key step in resilience research is the identification of protective and vulnerability factors that either enhance or decrease, respectively, the likelihood of healthy development for children faced with hardship. Greater attention to these factors in child maltreatment research will likely contribute to more effective means of identifying children at highest risk of maladaptation as well as improve prevention/intervention strategies (Luthar & Cicchetti, 2000). Examples of factors that have been established as protective for maltreated children include a child's connection to safe neighborhoods and quality education as well as a child's self-regulatory and self-esteem capacities (Cicchetti, 2013; Haskett, Nears, Ward, & McPherson, 2006). Identified vulnerability factors include low self-efficacy and exposure to high-crime neighborhoods (Jaffee, Caspi, Moffitt, Polo-Tomás, & Taylor, 2007; Kim & Cicchetti, 2003).

Beyond environmental influences, there has been growing emphasis, in the child maltreatment and resilience literature, on identifying genetic vulnerability and protective factors. By and large, these types of studies test for candidate-variant, *Gene × Environment interaction* ($G \times E$), whereby associations between environmental exposures and psychological phenotypes depend on genotype (e.g., Caspi et al., 2002). The $G \times E$ literature has been highly contentious due to replication failures, reported effect sizes at odds with hypothesis-free/genome-wide studies of behavior, improper statistical techniques, and inconsistent findings with respect to direction of effects (see Border & Keller, 2017; Duncan & Keller, 2011; Keller, 2014). Many of the reported candidate genetic moderators of childhood adversity, with roles in multiple biological systems, have been described as conferring individual differences in *environmental sensitivity* (ES; Pluess, 2015). ES refers broadly to differences in the degree to which individuals perceive, process, and respond to environmental influences. At its core, ES is a concept based on the hypothesis of neurosensitivity; namely, more sensitive ner-

vous systems lead to heightened awareness and deeper processing of environmental stimuli (i.e., high ES; Pluess, 2015). Though still in its infancy as a concept, the notion is that high levels of ES may potentiate the harm inflicted in children by maltreatment. Genetic variants relating to less efficient dopamine functioning or dysregulated hypothalamic–pituitary–adrenal axis activity have been hypothesized to increase ES (Bakermans-Kranenburg & van IJzendoorn, 2011; Canli & Lesch, 2007; Hostinar, Cicchetti, & Rogosch, 2014). Commonly studied genes in this arena the *serotonin transporter* (*5-HTT*) and the *dopamine receptor D4* (*DRD4*), which have been reported as moderators of maltreatment outcomes, albeit with varying success in replication (e.g., Byrd & Manuck, 2014; Cutuli, Raby, Cicchetti, Englund, & Egeland, 2013; Risch et al., 2009).

Often limited by the scope of a study's initial genotyping efforts, the vast majority of work on the genetics of psychological well-being in children faced with adversity has been focused on the moderating effects of a dozen or so genetic variants. These studies either examine variants individually or summed as a single polygenic risk score/s (PGRS). Often, candidate PGRS involve a count of the number of so-called *sensitivity* alleles that a given person carries (Beaver, 2008; Cicchetti & Rogosch, 2012). In general, these studies have reported that the higher the sensitivity allele count, the more susceptible individuals are to the effects of maltreatment, in line with the ES concept. Cicchetti and Rogosch (2012), for example, found that maltreated children with multiple sensitivity genetic variants scored lowest on a composite index of adaptive functioning as compared to maltreated children carrying less.

At least within the context of candidate $G \times E$ research on childhood adversity, most PGRS are unweighted and rely on a priori assumptions regarding which specific alleles of a given genetic variant confer sensitivity and which alleles confer protection. Such approaches are completely reliant on the robustness of previous literature, which has been strongly called into question (see Duncan & Keller, 2011). Moreover, unweighted PGRS assume that each genetic variant contributes equally to the overall main or interactive effect. When phenotypes and genotypes have been assessed on a large number of people, PGRS have the benefit of being weighted based on genome-wide association studies (Dudbridge, 2013). These PGRS can be considered a type of *fixed-weight composite* (Grace & Bollen, 2008). However, at least to date, there are no genome-wide $G \times E$ studies of child maltreatment to pull data from. Likely, these studies do not exist because the immense sample sizes required are currently prohibitive. If the previous literature's robustness is questionable, candidate $G \times E$ PGRS may only be able to be reasonably weighted by a study's own sample, otherwise referred to as *unknown weights composites* (Grace & Bollen, 2008).

In addition to considering effect size weighting, the coding of genetic alleles in candidate PGRS, in terms of designating which alleles confer sensitivity versus protection, is important. If researchers hypothesize that ES increases as the num-

ber of sensitivity variants carried increases, then it is imperative that there be either strong rational based on previous literature for selecting particular variants or a flexible statistical method that does not require such a priori assumptions. Unfortunately, there exists inconsistent evidence that any particular allele is robustly associated with any broad sensitivity to the environment phenotype. For example, some studies report the *long* allele, rather than the traditionally assumed *short* allele, of the usual suspect 5-HTTLPR variant is associated with greater reaction to environmental conditions (e.g., Uher & McGuffin, 2010).

In an attempt to overcome both issues of weighting and assumptions regarding allelic sensitivity, the current study utilized an unknown weights composite referred to as a *formative factor*. This approach has never been used in $G \times E$ research to date. Unlike a *reflective factor* from a confirmatory factor analysis, a formative factor is conceptualized as being formed by its indicators, requiring no correlation between indicators. Formative indicators' contribution to a formative factor are weighted as a function of what the formative factor is predicting, such that the indicator weights represent the indirect effect of each indicator on an outcome via the factor (Coltman, Devinney, Midgley, & Venaik, 2008). Moreover, formative factors allow for indicator weights to be estimated freely as either positive or negative. This means that the coding designation of sensitivity or protective alleles is not required a priori; only the selection of variants themselves is needed in order to construct a genetic formative factor.

The Current Study

The goal of the current study was to examine if the relation between maltreatment multiplicity and adaptive functioning, in children, depends on variation in candidate variants commonly studied in the ES, $G \times E$ literature. As a means of quantifying adaptive functioning in maltreated and nonmaltreated children, a latent class analysis (LCA) was performed on four commonly studied domains of psychological health: prosocial behavior, antisocial behavior, withdrawn behavior, and depression (Cicchetti & Rogosch, 2012). LCA accounts for measurement error and classification accuracy, does not treat all indicators the same, and avoids relying on researcher cut-offs (i.e., naïve profiling) making it an ideal person-centered modeling technique (Masyn, 2013). Next, 12 genetic variants were selected for use in a polygenic formative factor, based on a review of $G \times E$ findings in the maltreatment literature (see online-only Supplemental Material). Finally, the latent class variable representing adaptive functioning was embedded in a variable-centered, structural model wherein class membership was predicted by maltreatment multiplicity, the polygenic formative factor, and their interaction ($G \times E$). Our study used an all African American sample of children. This was done for three primary reasons. First, research with African American children is severely underrepresented in the current behavioral science literature, especially the genetic literature (Lewis, 2003). Second, genetic analyses have

greater statistical power when only one ancestral group is studied at a time because genetic variation covaries with ancestral origin. Third, the vast majority of the original sample of maltreated and nonmaltreated children is African American.

This study has two major aims:

1. Describe heterogeneity in multidomain, adaptive functioning in African American maltreated and nonmaltreated children of low socioeconomic background. That is, in an exploratory manner, elucidate the number and nature of a finite number of “adaptive functioning” groups of children.
2. Predict membership in “adaptive functioning” classes based on maltreatment multiplicity, a polygenic formative factor composed of 12 ES genetic variants, and the interaction between them ($G \times E$).

Method

Participants

Children aged 6 to 13 years ($N = 1,004$; M age = 10.09, $SD = 1.60$) were recruited in cohorts across 19 years of data collection to participate in a research-based, summer camp developed for low-income youth. Nonmaltreated comparison children ($n = 512$) and maltreated children ($n = 492$) encompassed the complete sample of participants. Among the participants, 495 were girls and 509 were boys. A single nucleotide polymorphism panel of 106 ancestral informative genetic markers was utilized to classify individuals into African, European, and Native American descent (Yaeger et al., 2008). This sample had a mean proportion of African American ancestry of .93, validating genetic homogeneity with self-reported ethnicity.

Recruitment procedures. Maltreated children were identified by the county Department of Human Services (DHS) as having experienced child maltreatment and were representative of youth receiving DHS services. Nonmaltreated children from sociodemographically comparable backgrounds were recruited from families receiving Temporary Assistance for Needy Families. Informed consent was obtained from parents of all participants. Furthermore, consent was given for examination of DHS records pertaining to the recruited families. See the online-only Supplemental Materials for more details on recruitment procedures.

Procedure

Maltreated and nonmaltreated children attended weeklong day camps and participated in research assessments. The camp lasted 7 hr/day for 5 days, providing 35 hr of child–counselor and peer–peer interactions. Trained research assistants, blind to study hypotheses and maltreatment status, conducted individual research assessments with children.

Clinical consultation/intervention was provided if any concerns over danger to self or others surfaced during the camp week.

Measures

Measures of adaptive functioning. We utilized multi-informant (peer, camp counselor, and self) measures of prosocial behavior, antisocial behavior, withdrawn behavior, and depressive symptomatology. All four adaptive functioning domains, except depression, included multiple measures. Measures within those three respective domains were averaged together and formed into composite parcels (representing each domain). Specifically, facet-representative parcels were used (Little, Cunningham, Shahar, & Widaman, 2002). Three confirmatory factor analyses were conducted using the respective indicators for each of the three parcels. Fit indices from all three confirmatory factor analysis models indicated good fit to the data, providing support for the cohesion of the indicators for use in parcels. The prosocial, antisocial, and withdrawn parcels along with the measure of depression served as the four indicators in the LCA. Because of the nature of the limited response noninterval scale of these four variables, we discretized them into six-category polytomous variables. See online-only Supplemental Material for more details.

Peer measure: Peer behavior ratings. After interacting with their peers during the week of summer camp, children evaluated the characteristics of their camp group peers via a sociometric peer ratings method on the last day of camp (Coie & Dodge, 1983). For each peer in the camp group, children were given six behavioral descriptors characterizing different types of social behavior. Children were asked to rate each peer on how characteristic the behavioral descriptor was for that peer on a 3-point scale. In the current study, ratings from peers for cooperative behavior, disruptive behavior, shyness, and fighting behavior were used. All ratings from peers on each child for each of the social behavioral descriptors were averaged.

Counselor measures.

Pupil evaluation inventory. At the end of each camp week the Pupil Evaluation Inventory (Pekarik, Prinz, Liebert, Weintraub, & Neale, 1976) was completed by camp counselors for children in their respective groups. The Pupil Evaluation Inventory consists of 35 items yielding three homogeneous and stable social behavior factors, including likeability, aggression, and withdrawn behavior. Interrater reliabilities based on intraclass correlations across the years of camp ranged from 0.72 to 0.85 ($M = 0.78$) for likeability, 0.85 to 0.90 ($M = 0.88$) for aggression, and 0.72 to 0.84 ($M = 0.78$) for withdrawal.

Counselor behavior ratings. Camp counselors completed 7-point ratings of children's behavior each day during three

separate, 45-min, observations during camp (see Cicchetti & Rogosch, 2012; Wright, 1983). Counselors rated children on 9 items tapping three domains of interpersonal functioning, including aggressive behavior, socially withdrawn behavior, and prosocial behavior. Individual counselor assessments for each of the three scales across the three assessment occasions were averaged to generate individual child scores. Interrater reliabilities based on average intraclass correlations among pairs of raters across the years of assessment ranged from 0.68 to 0.80 ($M = 0.76$) for prosocial, 0.70 to 0.84 ($M = 0.77$) for aggression, and 0.61 to 0.77 ($M = 0.71$) for withdrawn behavior.

Teacher Report Form. Counselor-rated behaviors were evaluated at the end of each week by counselors' completion of the Teacher Report Form (TRF; Achenbach, 1991). The TRF is a validated, reliable, and widely used assessment of behavioral functioning from the perspective of teachers. This measure was used in the present study because camp counselors are able to observe children in a similar manner as teachers. The TRF contains 118 items rated for frequency and assesses multiple dimensions of child behavioral symptomatology. In the present study, we examined the rule breaking, aggressive problems, and withdrawn subscales.

Self-reported measure: Children's Depression Inventory. The Children's Depression Inventory (Kovacs, 1992/1982) is a widely used, valid, and reliable, self-report questionnaire to assess depressive behaviors in school-aged children. For each item, children choose from among three option statements, depicting increasing levels of depressive symptoms, in order to characterize their experiences in the past 2 weeks. Internal consistency for the total scale has ranged from 0.71 to 0.89.

Predictors of adaptive functioning latent class variable.

Maltreatment classification. The Maltreatment Classification System (MCS; Barnett, Manly, & Cicchetti, 1993) was used to index maltreatment. The MCS codes all available information from DHS records, making independent determinations of maltreatment experiences rather than relying on case dispositions and official descriptions. For more information on the MCS, see the online-only Supplemental Materials. *Maltreatment multiplicity*, or the number of maltreatment subtypes experienced, ranging from 0 (*nonmaltreated*) to 4 (*having documented experience of all forms of child maltreatment at least once in their childhood*) was used as the primary maltreatment variable.

ES genetic variables. Based on a literature review, 12 genetic variants, thought to confer ES as well as having shown moderation effects on child maltreatment outcomes, were chosen for inclusion in the polygenic formative factor. See the online-only Supplementary Material for the basis of selecting these variants as well as genotyping procedures.

While the formative factor approach does not require the coding of variants in any particular order, we nevertheless designated one allele from each variant as the sensitivity allele. By doing so, the direction and magnitude of the factor loadings will shed some light on the accuracy and/or application of prior research to an all African American sample. These genes and alleles include the following (a) the 7-repeat allele of *DRD4*-VNTR, (b) the C allele of *DRD4*-rs1800955, (c) the val/G allele of *COMT* Val¹⁵⁸Met (rs4680), (d) the A allele of *DRD2*-rs1800497, (e) the del allele of *DRD2*-rs1799732, (f) the 10-repeat allele of *DAT1*-VNTR, (g) the T allele of *DAT1*-rs40184, (h) the T allele of *DAT1*-rs27072, (i) the short allele of 5-HTTLPR, (j) the T allele of *CRHR1*-rs110402, (k) the G allele of *OXTR*-rs53576, and (l) the T allele of *FKBP5*-rs1360780. No variants significantly deviated from Hardy–Weinberg equilibrium, except for the *DRD4*-VNTR variant, $\chi^2(44, N = 995) = 255.28, p = .005$. A departure from Hardy–Weinberg equilibrium is not unusual for the *DRD4*-VNTR nor is it likely to impact the results given strict quality control to prevent genotyping error (see DeYoung et al., 2011). This variant was therefore not excluded as an indicator of the polygenic factor (see Sensitivity Analyses where this variant was dropped). It was unclear which genetic model (additive, dominant, or recessive) was most appropriate for each variant based on the extant G \times E literature and within the context of an unordered categorical dependent variable. Model comparisons were conducted using each type of coding scheme and ultimately an additive coding model was selected (see online-only Supplementary Material).

Validation of latent class labeling: California Child Q-Set (CCQ). Validation of the LCA “class labeling” was carried out using a card sorting task called the CCQ (Block, Block, & Keyes, 1988). The CCQ was completed by camp counselors and involves sorting cards, with psychological descriptions written on them, into categories based on how characteristic each card’s description is to a given child’s observed behavior. The CCQ is widely used for the various psychological scales and criteria developed from it. This measure was not used as a latent class indicator, affording the possibility to independently validate the construct of each class label.

Data analytic approach

LCA. Using *Mplus* version 7.4 data analysis software (Muthén & Muthén, 1998–2012), a LCA was conducted. Class-specific item probabilities and class probabilities are the two primary sets of measurement and structural parameters derived from an LCA, respectively. Item probabilities refer to the probability of endorsing a particular item (or category of an indicator) conditional on class membership. Class probabilities correspond to the distribution of the categorical latent variable, reflecting the proportion of the population predicted to belong in each class (Masyn, 2013).

Model building process. Because there were no a priori assumptions about the number or nature of latent classes in this sample, an exploratory process of determining the best fitting unconditional LCA was undertaken. Following the use of best practices outlined by Masyn (2013) and Nylund, Asparouhov, and Muthén (2007), a series of LCA models was fit in an iterative fashion. The LCA procedure involves fitting a series of *K*-class models beginning with a one-class model and stopping at a *K*-class model that becomes not well identified (K_{\max}). Models were deemed not well identified if one or more of the following criteria were met: lack of best log likelihood value replication across a set of random start values, lack of model convergence, or an extraction of a class with a small estimated class size, which may indicate overextraction.

Model estimation. To estimate latent class analysis parameters, *Mplus* uses the expectation-maximization algorithm for full information maximum-likelihood estimation from incomplete data assuming data is missing at random. Random start values were utilized in an attempt to replicate a global maximum of the likelihood function rather than a local solution.

Evaluating model fit. A combination of statistical indicators and substantive interpretation were used to determine the best number of latent classes based on use of best practices by Masyn (2013). The goal is to extract a well-fitting, parsimonious, yet meaningful measurement model. These indices include the likelihood ratio test (LRT) statistic for nested models, the adjusted Lo–Mendell–Rubin likelihood ratio test (adjusted LMR-LRT), the parametric bootstrapped LRT, the Akaike information criterion (AIC), the Bayesian information criterion (BIC), the consistent Akaike information criterion (CAIC), the approximate weight of evidence (AWE), the approximate Bayes Factor (*BF*), and the correct model probability (*cmP*). Classification accuracy and class homogeneity were evaluated using relative entropy (E_K), average posterior class probability (*AvePP*), and the odds of correct classification ratio for a given class (OCC_k). The *K*-class model deemed to have the best combination of absolute fit, relative fit, classification accuracy, and model usefulness was chosen as the final model to be included in the structural component of the SEMM.

SEMM. The final unconditional LCA model chosen above was embedded into a larger SEMM. To do so, an alternative three-step method was utilized (Asparouhov & Muthén, 2014). The first step involves selecting the best fitting unconditional LCA model. In Step 2, the modal class assignments and classification error rates are saved. Finally, in Step 3, the structural model (with predictors added) is run with a nominal modal class assignment variable used as a single indicator of a latent class variable with the measurement error rates prefixed. A latent class variable is an unordered categorical variable; therefore, the relation between class membership and

predictors is parameterized as a multinomial logistic regression.

To test for main effects of maltreatment, predictors included maltreatment multiplicity along with sex and age as control variables. To test for $G \times E$, predictors included maltreatment multiplicity as well as the polygenic formative factor, and a Polygenic Formative Factor \times Maltreatment interaction term. The control predictors of age, sex, and any interaction terms with covariates that associated with either maltreatment multiplicity or the polygenic formative factor were also added to complete the final structural model (Keller, 2014).

Results

Selecting an optimal latent class solution

The latent class enumeration results are provided in Table 1. A solution of seven classes was not well identified, and thus consideration was only given for the 1- through 6-class solutions. The LMR-LRT indicated no significant improvement in model fit beyond a 3-class solution. The Bayesian LRT was significant for all model comparisons, providing no helpful information for the selection of an optimal class, and was therefore not included in Table 1. The lowest BIC and CAIC values were for the 3-class solution. The most parsimonious K -class model with a high BF ratio, along with the K -class solution with a cmP near 1 was also the 3-class solution. The lowest AWE was for the 2-class solution. The smallest K -class solution with a log-likelihood (LL) value lower than that of a 1-class model with all indicators allowed to freely covary with all other indicators within its class was the 4-class solution. That is, the 4-class solution was the most parsimonious solution to fit the data better (i.e., lower LL value) than a fully saturated mean and variance/covariance model that is an exact fit to the data (in terms of the first and second order moments of the data). Finally, examination of an AIC “elbow” plot indicated a bend at the 4-class solution.

Because of the ambiguity provided by the results of the fit indices, the 3- through 5-class solutions were further evaluated in terms of classification precession, class separation, and class homogeneity. In all three k -class solutions, the primary driver of class separation appeared to be variation in prosocial and antisocial behavior. No one k -class solution seemed to stand out as having the best class separation. In all three k -class solutions, the prosocial and antisocial behavior indicators had the best homogeneity. Again, no one k -class solution seemed to have the best looking class homogeneity with respect to individual indicators. The 3-class solution masked too much population heterogeneity and the 5-class solution overextracted classes, arbitrarily dividing a homogenous subgroup. Ultimately, we choose the 4-class solution as the final unconditional model given its good absolute fit, relative fit, classification precision, and substantive meaningfulness. As seen in Table 2, the 4-class solution

Table 1. Fit indices for LCA of adaptive functioning

Model	LL χ^2 test			Ajd. LMR-LRT				$cmP(K)$				
	LL	$npar$	χ^2	df	p	BIC	CAIC		AWE	$\chi^2 (df = 6)$	p	$BF (K, K+1)$
1-Class	-6269.6	20	1667.5	1275	<.001	12677	12697	12875	542.835	<.001	3.9×10^{-87}	0
2-Class	-5998.2	41	1103.5	1252	1.00	12279	12320	12685	183.39	<.001	4.4×10^{-9}	0
3-Class	-5906.5	62	921.2	1231	1.00	12241	12303	12855	108.97	1	6.4×10^7	1
4-Class	-5852.0	83	811.6	1210	1.00	12277	12360	13099	58.79	0.374	5×10^{18}	0
5-Class	-5822.6	104	749.8	1188	1.00	12363	12467	13393	52.36	0.897	1.3×10^{20}	0
6-Class	-5796.4	125	721.2	1170	1.00	12455	12580	13693	—	—	—	0
Saturated	-5854.5	26	798.2	1267	1.00	11888	11914	12146	—	—	—	0

Note: LL , model log likelihood value. $npar$, number of free parameters. $LL \chi^2$ test, log likelihood ratio model chi-square goodness-of-fit test. BIC, Bayesian information criterion. CAIC, consistent Akaike information criterion. AWE, approximate weight of evidence criterion. Ajd, LMR-LRT, adjusted Lo-Mendell-Rubin likelihood ratio test. p , p value corresponding to the adjusted LMR-LRT χ^2 statistic ($df = 6$) comparing $H_0: K$ classes vs. $H_1: K + 1$ classes. Bootstrapped LRT p value. $BF (K, K+1)$, Bayes factor ratio of Models $K, K+1$. $cmP(K)$, correct model probability. Bolded values correspond to “best” model according to the fit index—columns with no bolded values indicate that the best value for that index was not reached prior to the maximum class extraction supported by the data.

Table 2. Model classification diagnostics for the four-class solutions

K-class solution	k-class	Estimated k-class proportion	90% CI ^a	<i>mcaP_k</i>	<i>AvePP_k</i>	<i>OCC_k</i>	Entropy
4-Class	Class 1	.33	[.26, .39]	.318	.913	21.82	.785
	Class 2	.15	[.09, .22]	.119	.853	32.62	
	Class 3	.24	[.19, .28]	.239	.917	35.65	
	Class 4	.29	[.24, .34]	.324	.814	10.84	

Note: ^aBias-corrected bootstrapped 95% confidence intervals.

had good classification accuracy as indexed by high AvePP, *OCC_k*, and entropy values. The 3- and 5-class solutions had comparable degrees of classification accuracy.

Well-adjusted class

The largest model-estimated class (32.5%) from the 4-class solution was given the label of “well-adjusted” and can be visualized in Figure 1a. This class is characterized by high probabilities of exhibiting high levels of prosocial behavior (i.e., 66% chance of endorsing levels of prosocial behavior

0.5 SD above the mean or greater, or 95% chance of exhibiting levels of prosocial behavior greater than the sample mean). The well-adjusted class is also characterized by a high probability of low antisocial behavior. Finally, children in this class have high probabilities of endorsing low levels of depression and low levels of withdrawn behavior. This class was given the label of “well-adjusted” because the profile of competence characterized by this class is similar to profiles considered developmentally adaptive and predictive of positive life success (Masten et al., 1999). Because all children in the sample were from high-risk contexts (i.e., either poverty

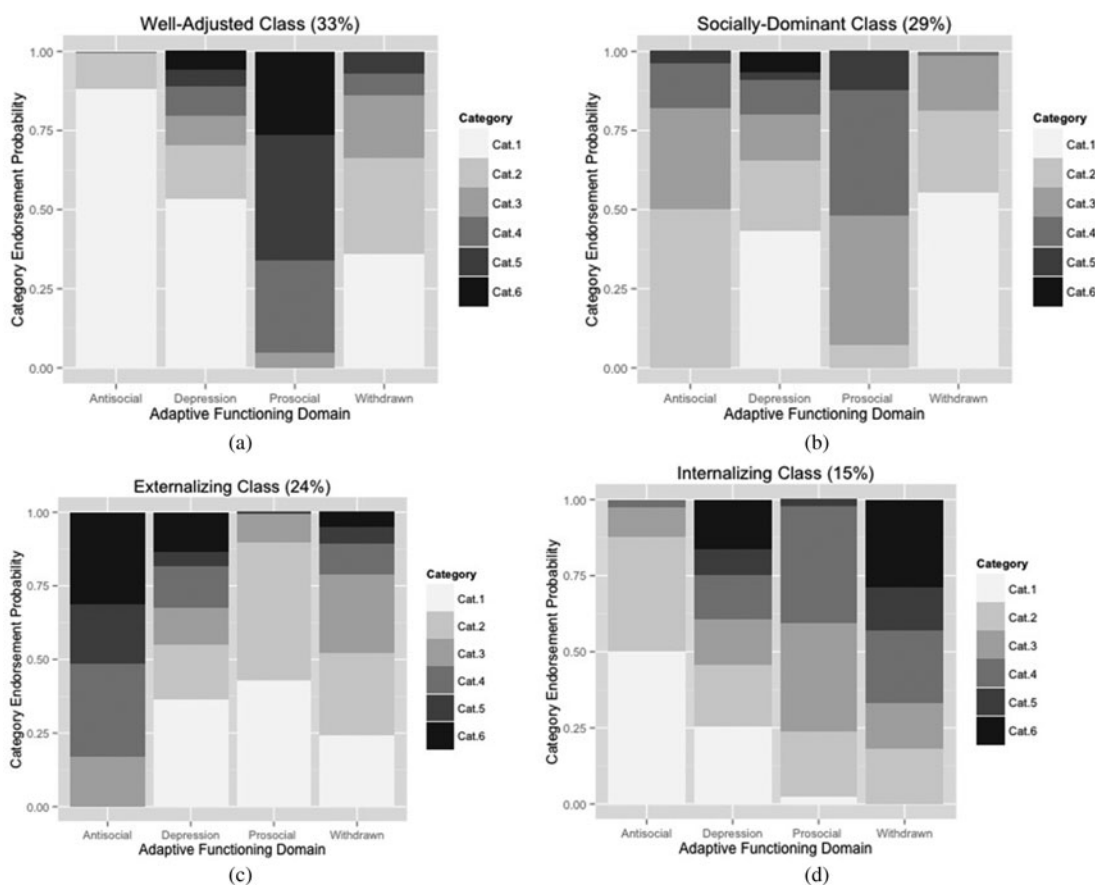


Figure 1. Profiles of adaptive functioning: (a) the well-adjusted class, (b) the socially dominant class, (c) the externalizing class, and (d) the internalizing class. The z score range cutoffs for each category of the antisocial, withdrawn, and depression variables is Cat1 = <-0.5 SD; Cat2 = >0.5 to 0; Cat3 = >0 to 0.5 SD; Cat4 = >0.5 to 1 SD; Cat5 = >1 to 1.5 SD; Cat6 = >1.5 SD. The z score range cutoffs for each category of prosocial variable is Cat1 = <-1 SD; Cat2 = >-1 to -0.5 SD; Cat3 = >-0.5 to 0 SD; Cat4 = >0 to 0.5 SD; Cat5 = >0.5 to 1 SD; Cat6 = >1 SD.

or poverty with maltreatment), children in this class may be considered to be demonstrating *resilient* functioning.

Socially dominant class

The next largest model-estimated class (28.8%) was labeled “socially dominant” and can be visualized in [Figure 1b](#). This class had the lowest levels of withdrawn behavior compared to all other classes. This class is characterized by medium levels of prosocial behavior and medium to medium-high levels of antisocial behavior. Finally, this class had a high probability of exhibiting low levels of depression. This class was labeled “socially dominant” because the profile it characterizes matches the literature-defined expression of social dominance, including high social presence (low withdrawn behavior) as well as use of both cooperation (prosocial behavior) and aggression (antisocial behavior; Hawley, 1999; Teisl, Rogosch, Oshri, & Cicchetti, 2012). Social dominance refers to a pattern of resource acquisition and social control whereby both cooperation and coercion are used. Socially dominant children are often admired and socially influential though not necessarily without the use of antisocial behaviors such as bullying to gain such social centrality (Hawley, 1999).

Externalizing class

The third largest class that emerged from the 4-class LCA was termed the “externalizing class” and is plotted in [Figure 1c](#). This class’s most prominent feature was the high probability of exhibiting extremely low levels of prosocial behavior and the high probability of high levels of antisocial behavior. Moreover, this class was characterized by relatively low withdrawn behavior. The somewhat mixed endorsement probabilities for depression, in this class, is perhaps to be expected because some children exhibit impulsivity and even agitation as the result of depressive feelings (Ryan et al., 1987). The relatively low levels of withdrawn behavior in this class is consistent with the notion that children characterized by externalizing behaviors are not generally inhibited or reserved, socially. However, this class still has an estimated 21% of children exhibiting withdrawn behavior greater than 0.5 *SD* above the sample mean. This is also not necessarily an unexpected finding nor does it negate the rationale for labeling this class as externalizing. A subset of children are both aggressive and withdrawn; in these cases, antisocial behavior is thought to contribute to social rejection and victimization, which may in turn contribute to lonely and reserved behaviors (Ladd & Burgess, 1999). In summary, based on the profile of category endorsement probabilities and consistency with definitions in the literature, this class is considered to represent children who exhibit primarily externalizing symptoms.

Internalizing class

The smallest estimated class (15.5%) is considered the “internalizing class” and is plotted in [Figure 1d](#). This class is char-

acterized by low levels of antisocial behaviors and medium levels of prosocial behavior. This class had relatively high levels of withdrawn behavior. Finally, category endorsement probabilities for depression were mixed. This class had the highest probability (16%) of endorsing the highest category of depressed behavior out of all the classes. Moreover, individuals in this class are estimated to be more likely to have depression scores above the mean than below the mean. Nevertheless, 25% of individuals in this class are expected to have depression scores below 0.5 *SD* below the mean. These mixed endorsement probabilities for difference levels of depression suggest poor item homogeneity. Though, because of low base rates, uncovering a class of children homogeneous for very high levels of depression is unlikely, even in a class characterized by high withdrawn and low antisocial behavior.

Construct validity of latent class labels

The ego-resiliency and social competence *q*-sorts of the CCQ were used to validate the well-adjusted class. To validate the externalizing, internalizing, and socially dominant classes, the internalizing *q*-sort, externalizing *q*-sort, and social dominance *q*-scale, respectively, were utilized (Block et al., 1988; Teisl et al., 2012). Each CCQ measure associated positively ($p < .001$) with membership in the respective class being validated when compared to membership in each other class. These results suggest good construct validity and provide a degree of assurance in the labels used for each class.

SEMM results

Global maltreatment multiplicity results. Because the latent class variable has three unordered categories, regression results are based on a multinomial logistic regression, in which simultaneous pairs of logistic regressions are tested (for a given predictor, the effect on the odds of being in one class versus a reference class). To test for global effects, Wald statistics were estimated. Maltreatment multiplicity was significantly associated with latent class membership (Wald = 35.3, $df = 3$, $p < .0001$). To visualize this global effect, an overall probability plot is depicted in [Figure 2](#). As the number of maltreatment subtypes increases, the probability of being in the externalizing class increases and the probability of being in the well-adjusted class decreases. There are slight declines in the probability of being in both the socially dominant and internalizing classes as maltreatment increases.

Multinomial logistic regression results. To further interrogate the global maltreatment multiplicity effects on class membership, we examined all pairwise logistic regressions. When compared to the well-adjusted class, maltreatment multiplicity is associated with membership in the externalizing (odds ratio [*OR*] = 1.8, $p < .001$) and socially dominant classes ($OR = 1.28$, $p = .019$) but not the internalizing class ($OR = 1.24$, $p = .14$). When compared to the socially dom-

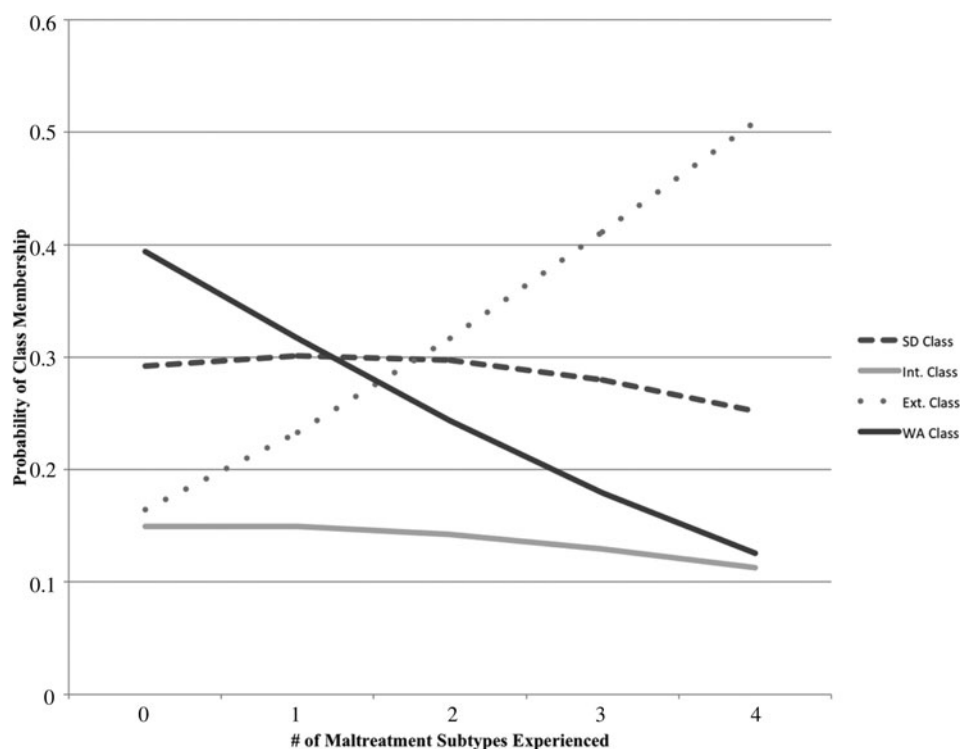


Figure 2. Main effects of child maltreatment probability curve plot. SD, socially dominant. Int., internalizing. Ext., externalizing; WA, well-adjusted. Controlling for sex and age.

inant class, maltreatment multiplicity is associated with membership in the externalizing ($OR = 1.38, p = .002$) but not the internalizing class ($OR = 0.97, p = .81$). Finally, maltreatment multiplicity was associated with membership in the externalizing class when compared to the internalizing class ($OR = 1.42, p = .007$).

Selecting a formative factor model. The “additive” model was ultimately selected for use in the final SEMM (see online-only Supplementary Materials for more details). Initially, each genetic variant was scored in terms of the number of putative sensitivity alleles. However, not all of the genetic variable indicator weights loaded onto the factor model in the same direction. That is, not all alleles contributed to the overall $G \times E$ effect in the same manner. Some variants, assumed to have a vulnerability-like effect exhibited a protective effect and vice versa (see Supplemental Table S.2). To simplify the interpretation of the formative factor, variants were reverse-coded in the final SEMM so that all variants contributed positively to the factor. This did not change the size of the Wald statistic for the global $G \times E$ effect.

Final SEMM with control variables. After testing all covariate interaction effects, there were only significant associations between the polygenic factor and age ($\beta = 0.182, p < .001$) and sex and maltreatment ($OR = 0.93, p = .035$). The association between polygenic factor and age is unusual, and may be spurious. Nevertheless, this association may still bias the

$G \times E$ effects. Because of these associations, it was necessary to add Sex \times Factor and Age \times Maltreatment interaction terms as control variables, in addition to sex and age as main effects (Keller, 2014). The final model regressed the latent class variable onto maltreatment multiplicity, the polygenic factor, the interaction between maltreatment and the polygenic factor ($G \times E$), age, the interaction between age and maltreatment, sex, and the interaction between sex and the polygenic factor.

Global $G \times E$ results. A Wald test for a global $G \times E$ effect was significant controlling for all other variables (Wald = 13.5, $df = 3, p = .004$). That is, the effect of maltreatment on class membership depends on the level of genetic variation. There was no significant association between the polygenic factor and maltreatment, ruling out gene–environment correlation (rGE). The results of the polygenic factor indicator weights are presented in Supplemental Table S.2. All but one genetic variant (*DAT1-VNTR*) loaded significantly to the formative factor. Five genetic variants loaded negatively onto the factor, meaning for these variants, the alleles assumed to confer “sensitivity” contributed protection. These variants included (a) the 7-repeat allele of *DRD4*, (b) The del allele of *DRD2*-rs1799732, (c) the short allele of 5-HTTLPR, (d) the T allele of *CRHR1*-rs110402, and (e) the T allele of *FKBP5*-rs1360780. With genetic variants loading in both positive and negative directions, factor scores can be difficult to interpret, and therefore we reverse-coded those variants. As can be seen from the *well-adjusted* $G \times E$ plot, higher factor scores

decrease the probability of class membership as maltreatment multiplicity increases. Based on these vulnerability relations, we considered higher factors scores as potentially representing higher levels of *ES genetic load*, bearing in mind that the reverse-coded five variants contributed to this vulnerability effect in an unexpected manner.

Multinomial logistic regression results. To further interrogate the global $G \times E$ effect, the results of the six pairwise logistic regressions are presented in Supplemental Table S.3. Only two significant pairwise $G \times E$ effects emerged, controlling for all other variables. One $G \times E$ effect corresponds to the comparison between the well-adjusted class and the externalizing class. Specifically, the effect of an increasing maltreatment multiplicity on the odds of being in the externalizing class versus the well-adjusted class increases as *ES genetic load* increases ($\Delta \log \text{odds} = 0.17, p = .001$). The other $G \times E$ effect corresponds to the comparison between the externalizing and socially dominant classes. Specifically, the effect of an increasing maltreatment multiplicity on the odds of being in the externalizing class versus the socially dominant class increases as *ES genetic load* increases class ($\Delta \log \text{odds} = 0.18, p = .001$). Only one pairwise comparison had a significant maltreatment main effect that was not in the context of a $G \times E$. Maltreated children were slightly more likely to be in the socially dominant class versus the well-adjusted class ($OR = 1.3, p = .04$).

Sensitivity analyses. A variety of sensitivity analyses were conducted to insure the robustness of this model. See the online-only Supplementary Materials for more details. In summary, our sensitivity analyses tested and confirmed four assumptions: (a) maltreatment multiplicity could be treated as a continuous variable, (b) there was no differential item functioning with respect to maltreatment status, (c) the formative factor results were not dependent on which genetic variant was used as the anchor, and (d) maltreatment multiplicity was not associated with study cohort, and finally that an unweighted polygenic index also demonstrated $G \times E$ effects.

Discussion

Study overview

This study had two primary aims. The first was to characterize adaptive functioning in maltreated and nonmaltreated, African American children. The second aim was to predict variation in adaptive functioning based on maltreatment multiplicity, *ES genetic variation*, and their interaction ($G \times E$). Child maltreatment was highly associated with adaptive functioning, in particular increasing the odds of membership in maladapted classes such as the externalizing group. Moreover, an interaction between child maltreatment and a polygenic formative factor composed of putative sensitivity variants significantly predicted latent class membership. This study was the

first of its kind to combine latent class analysis with the use of a formative factor to test for $G \times E$.

Comment on the LCA

A 4-class solution was chosen as the best overall unconditional latent class model. The four classes were labeled as “well-adjusted”, “externalizing,” “internalizing,” and “socially dominant.” Approximately 31% of maltreated children were modally assigned to the well-adjusted class (33% of nonmaltreated children were assigned to well-adjusted class) and would therefore be considered *resilient*. These proportions closely resemble those from similar resilience studies on child maltreatment (e.g., Cicchetti & Rogosch, 2012). In general, profiles of externalizing and internalizing behaviors are considered maladaptive for children’s development. For example, children with high levels of these broadband behaviors tend to have more difficulty maintaining friendships, are at heightened risk for abusing substances in adolescence and adulthood, tend to display difficulties in school performance, and tend to be victimized more by their peers (e.g., Colder et al., 2013; Masten et al., 2005). In order to climb the social ladder, enhance likability, and control resources, some children engage in highly competitive dominance strategies marked by both cooperative and coercive behaviors. Such profiles of social dominance, as seen in this study, can increase the odds of obtaining social influence and resource control especially in the early years of life (Hawley, 1999; Teisl et al., 2012). As children age, however, patterns of behavior that are predominately prosocial in nature (such as the well-adjusted class in this study) are more consistently linked with positive peer preferences. Competitive social behaviors in middle childhood, even those that include the use of cooperative strategies, that are marked by hostility and aggression (such as the socially dominant class in this study) can lead to peer rejection and victimization (Coie & Dodge, 1983; Teisl et al., 2012).

Comparing the results of this measurement model with other studies is difficult. For one reason, a LCA with similar indicators of adaptive functioning has not been used with maltreatment data to our knowledge. Moreover, the majority of mixture modeling techniques used in resilience research employ growth mixture modeling as opposed to cross-sectional LCA (Bonanno et al., 2012). Yates and Grey (2012) conducted an LCA on a similar sample of high-risk children (former foster care youth) using indicators of adaptive functioning. A four-class solution was extracted with subgroups labeled as *resilient*, *maladapted*, *internally resilient*, and *externally resilient*. While some competence indicators used by Yates and Grey were similar to those used in the present study (relational competence and depression), other indicators were not (educational competence, occupational competence, civic engagement, and self-esteem), making a direct comparison of models problematic. Nevertheless, the profiles of behavior observed in this study are commonly reported in both typical and high-risk child populations (e.g.,

Bornstein, Hahn, & Haynes, 2010; Mendez, Fantuzzo, & Cicchetti, 2002).

Maltreatment multiplicity and $G \times E$ predictors of latent class membership

Beyond simply describing heterogeneity in adaptive functioning, via an LCA, this study aimed to understand how maltreatment experiences and ES genetic variation relate to adaptive functioning. The association between maltreatment multiplicity and adaptive functioning class membership was first tested outside the context of a $G \times E$. In terms of overall probabilities, as maltreatment multiplicity increased, children became more likely to exhibit profiles consistent with externalizing behaviors and less likely to exhibit profiles of well adjustment. The association between abusive histories and behaviors such as aggression, impulsivity, and delinquency are some of the most robust findings in all of psychology (Jaffee et al., 2004). The chaotic, hostile, and dysfunctional parent-child relationships evident in cases of maltreatment can disrupt the healthy development of children's ability to adaptively regulate emotions and actions. These regulatory deficits compounded by problems in social information processing, known to result from maltreatment, greatly increase children's antisocial tendencies (Cicchetti, Rogosch, & Thibodeau, 2012; Thibodeau et al., 2015). Such antisocial behaviors, as characterized in the externalizing class in this study, are associated with negative outcomes, including risk for substance use and academic problems (Cicchetti et al., 2012).

While maltreated children had a significantly higher likelihood of being in the socially dominant class than the well-adjusted class, this effect was small and reflected the rather slight increase in the overall probability of socially dominant class membership as a function of maltreatment multiplicity. These results are largely in line with the findings from Teisl et al. (2012). That research group also performed mixture modeling, identifying three groups of children characterized by nondominant behaviors, primarily coercive dominant behaviors, and primarily competent/cooperative dominant strategies. Maltreated children, regardless of the type of maltreatment, were more likely to be in the primarily coercive dominant and nondominant classes as compared to the competent class ($OR = 3.07$ and $OR = 2.23$, respectively). It is unclear why Teisl et al. (2012) found larger effects relating to the effects of abuse on maladaptive social dominance behaviors. One explanation was that we only identified one *bistrategic* socially dominant class rather than subclasses characterized by primarily coercive or competent dominating strategies. Had we identified these subclasses, the maltreatment associations may have been stronger. That is, the small maltreatment associations with social dominance in our study may be muddied by the presence of distinct dominance groups within the bistrategic class. Moreover, it is possible that the inclusion of measures of internalizing behaviors may have created difficulty in extracting more fine-grained

profiles of social dominance. In any case, in our sample, maltreated children were more likely to be characterized by dominating behaviors. Social learning perspectives have suggested that exposure to adult models of instrumental aggression and dominance common to abusive parenting may lead children to overemphasize the need to use socially dominant behaviors (Teisl et al., 2012).

The lack of findings suggesting increased odds of being in the internalizing class as a function of maltreatment multiplicity is inconsistent with a large body of literature suggesting otherwise (i.e., Cicchetti, Rogosch, & Oshri, 2011; Teisl et al., 2012). While maltreatment multiplicity was associated directly with the individual, ordinal measure of depression (ordinal $OR = 1.17$, $p = .006$), there was no association with the individual measure of withdrawn behavior. These somewhat discrepant findings could explain the overall lack of association with class membership. Moreover, the estimated class size was relatively small for the internalizing class (15.5%). Therefore, low power in this study could have contributed to Type II error. Finally, some children with otherwise internalizing struggles may exhibit irritability and even aggression, making a distinction between purely internalizing versus externalizing behaviors somewhat difficult to make (Ryan et al., 1987). There was some nonnegligible probability of exhibiting moderate levels of withdrawn and depression behavior in the externalizing class.

A formative factor composed of 12 genetic variants, shown to exhibit $G \times E$ effects in previous studies, was constructed and included as a moderator of the association between maltreatment multiplicity and adaptive functioning class membership. There was an overall significant $G \times E$ effect, such that as genetic load increased, maltreated children became increasingly more likely to be members in the externalizing class and less likely to be in the well-adjusted or socially dominant classes. An advantage of using a formative factor to construct polygenic indices is that the contribution of each variant to the factor can load either in a negative or a positive direction. Five of these variants loaded onto the formative factor negatively, thereby exhibiting protective rather than the expected sensitivity effects. These findings highlight the complexity of $G \times E$ interplay as well as further add to the mixed-findings commonly reported in the candidate $G \times E$ research community. It is not uncommon for studies to report discrepant findings for these particular genetic variants. A number of reports, for example, have found the short allele as opposed to the long allele of *5-HTTLPR* confers protective effects in contexts of abuse (e.g., Cicchetti et al., 2011; Sharpley, Palanisamy, Dillingham, & Agnew, 2014). Contradictory vulnerability/protection effects with regards to *FKBP5*-rs1360780 and *CRHRI*-rs110402 have also been reported (Roy, Gorodetsky, Yuan, Goldman, & Enoch, 2010; Sumner, McLaughlin, Walsh, Sheridan, & Koenen, 2014). These results add to a growing number of studies demonstrating polygenic moderation of child maltreatment outcomes, and the relative, but modest role that some candidate genes have in predicting risk and resilience (i.e., Thibodeau et al., 2015).

This study built upon earlier work by Cicchetti and Rogosch (2012) sharing some similarities and differences. In general, Cicchetti and Rogosch's findings indicated that ES gene variation was differentiating, in terms of adaptive functioning, for both nonmaltreated children and maltreated children more in line with differential susceptibility theory (Belsky & Pluess, 2009). However, by "eye-balling" the plots in Figure 3, our results appear to display interactions more consistent with diathesis stress (Monroe & Simons, 1991). There are statistical tools to decipher the Person \times Environment interaction effects of vantage sensitivity, differential susceptibility theory, and diathesis stress (i.e., Roisman et al., 2012). These tools are most effective when a study's environmental range stretches from positive to negative. The current study has a restricted environmental range with only children from low-income backgrounds sampled. Thus, these tools were not employed. Moreover, there is no clear way of identifying interaction patterns such as those listed when the outcome variable is unordered and categorical. One potential reason for why our results are slightly different than Cicchetti and Rogosch (2012) is that they used a continuous variable of adaptive functioning and did not employ a weighted composite for their polygenic index.

In summary, we replicate and extend very well known findings in the child maltreatment literature, namely, that child maltreatment is strongly associated with negative outcomes and that common genetic variation does not play a major role in differentiating children in terms of adaptive func-

tioning. We demonstrate that candidate $G \times E$ polygenic risk scores can be sample weighted and that a priori assumptions about allelic risk/protection is not required. While we report significant $G \times E$ findings, they should be treated with great caution, and replication attempts should be made. The effects were small, and roughly half of the genetic variants had the opposite $G \times E$ trend as commonly reported in previous work in the field of childhood adversity.

With any study of parent behavior–child behavior associations, the directionality of causation can be difficult to infer. Because this study did not employ a twin or adoption design, nor utilize longitudinal data, definitive conclusions regarding the causality of the impacts of parental abuse/neglect cannot be made. Although statistical $G \times E$ interactions were detected, these interactions say little about how the interactions operate biologically at a mechanistic level. Furthermore, not all unobserved contextual factors common to both caregiver and child were controlled for, which can increase Type 1 errors, though in practice adding all such factors in a model is extremely difficult. Finally, not all commonly utilized indicators of adaptive functioning were measured, such as academic success (Masten, 2015). This study had a number of notable strengths. First, maltreatment was measured objectively and prospectively. Second, this study examined adaptive functioning in four major domains of competence based on measures collected by multiple informants (peers, counselors, and self) via an LCA. Third, rather than focusing on a single candidate gene variant, 12 variants were examined across 8

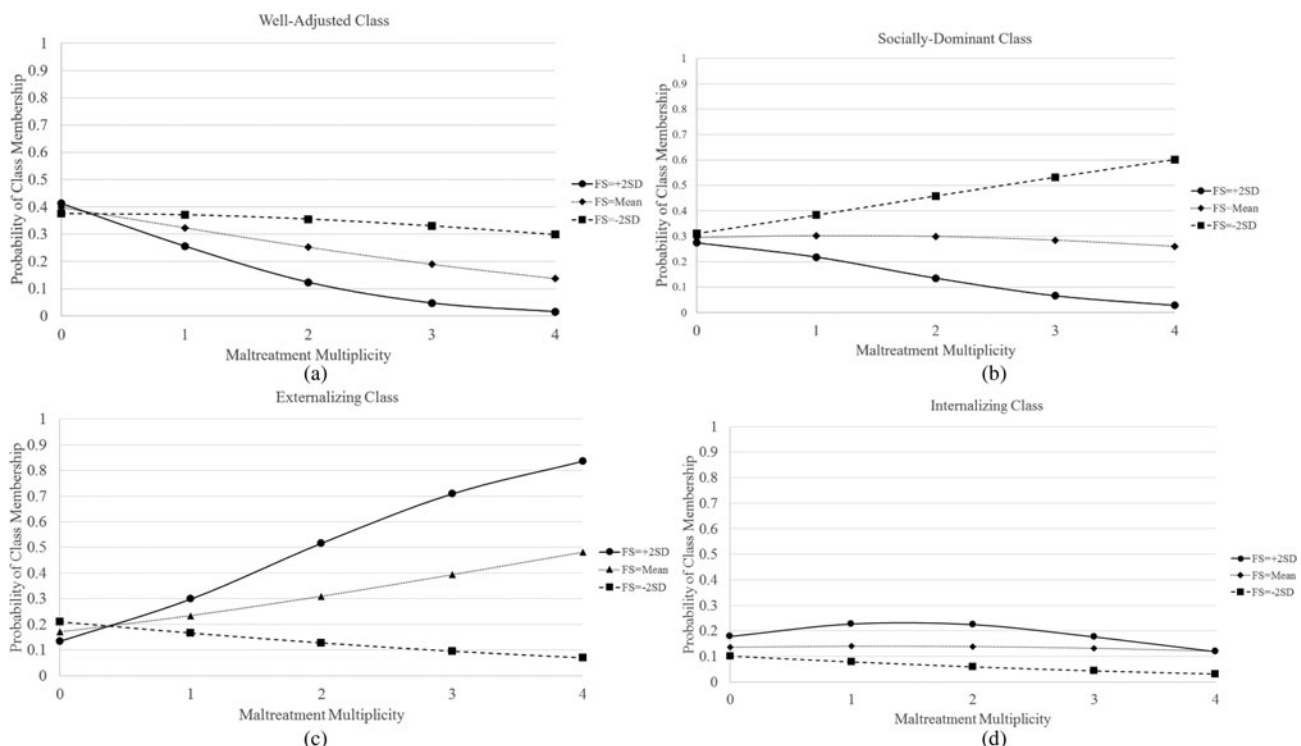


Figure 3. $G \times E$ probability curve plots for (a) the well-adjusted class, (b) the socially dominant class, (c) the externalizing class, and (d) the internalizing class.

genes collectively via formative factor. The formative factor not only provides a weighted composite but also allows variants to load either negatively or positively. Because of this, interaction effects were not dependent on the a priori assumptions made about plasticity effects. To the best of our knowledge, this is the first time a formative factor has been used to weight genetic effects; it is also the first time a formative factor has been used in an interaction and the first time that a formative factor has predicted a latent class variable. Finally, $G \times E$ research is fraught with underpowered studies. This study used a relatively large, ancestrally homogeneous sample. In accordance with recommendations made by Keller (2014), covariate interaction terms were added, ensuring proper use of control variables.

Concluding remarks

Given the amount of novel statistical techniques used in this study, caution is warranted regarding the $G \times E$ effects. $G \times E$ studies are notorious for nonreplication (Duncan & Keller,

2011). Despite proper use of control variables and large sample sizes, definitive conclusions regarding the impact of these 12 ES gene variants on maltreatment multifinality awaits repeated replication. Researchers are encouraged to test the utility of formative factors when conducting $G \times E$ research and/or mixture modeling research. In no way do these results suggest that any child is “immune” to the detrimental effects of maltreatment. Child abuse affects all children regardless of genotype; every effort should be made to prevent maltreatment from occurring. If anything, these results suggest that common variation in just a few genes has a small (yet statistically significant) impact on explaining different behaviors observed in maltreated children of African American ancestry. At any point in a child’s life, environmental protective factors such as supportive caregivers, strong peer groups, or an influential teacher can dramatically promote resilient functioning. Child maltreatment is one of the most detrimental environmental exposures that any individual can experience; continued efforts to study psychological health outcomes of maltreated children will likely help develop more effective prevention and treatment tools.

References

- Achenbach, T. M. (1991). *Manual for the Child Behavior Checklist/4-18 and 1991 Profile*. Burlington, VT: University of Vermont, Department of Psychiatry.
- Asparouhov, T., & Muthén, B. (2014). Auxiliary variables in mixture modeling: Three-step approaches using M plus. *Structural Equation Modeling, 21*, 329–341.
- Bakermans-Kranenburg, M. J., & van IJzendoorn, M. H. (2011). Differential susceptibility to rearing environment depending on dopamine-related genes: New evidence and a meta-analysis. *Development and Psychopathology, 23*, 39–52.
- Barnett, D., Manly, J. T., & Cicchetti, D. (1993). Defining child maltreatment: The interface between policy and research. In D. Cicchetti, & S. L. Toth (Eds.), *Child abuse, child development, and social policy* (pp. 7–73). Norwood, NJ: Ablex.
- Beaver, K. M. (2008). The interaction between genetic risk and childhood sexual abuse in the prediction of adolescent violent behavior. *Sexual Abuse: a Journal of Research and Treatment, 20*, 426–443.
- Belsky, J., & Pluess, M. (2009). Beyond diathesis stress: Differential susceptibility to environmental influences. *Psychological Bulletin, 135*, 885.
- Block, J. H., & Block, J. H., & Keyes, S. (1988). Longitudinally foretelling drug usage in adolescence: Early childhood personality and environmental precursors. *Child Development, 59*, 336–355.
- Bonanno, G. A., Mancini, A. D., Horton, J. L., Powell, T. M., LeardMann, C. A., Boyko, E. J., . . . Smith, T. C. (2012). Trajectories of trauma symptoms and resilience in deployed US military service members: Prospective cohort study. *British Journal of Psychiatry, 200*, 317–323.
- Border, R., & Keller, M. C. (2017). Commentary: Fundamental problems with candidate gene-by-environment interaction studies—Reflections on Moore and Thoenes (2016). *Journal of Child Psychology and Psychiatry, 58*, 328–330.
- Bornstein, M. H., Hahn, C. S., & Haynes, O. M. (2010). Social competence, externalizing, and internalizing behavioral adjustment from early childhood through early adolescence: Developmental cascades. *Development and Psychopathology, 22*, 717–735.
- Byrd, A. L., & Manuck, S. B. (2014). MAOA, childhood maltreatment, and antisocial behavior: Meta-analysis of a gene-environment interaction. *Biological Psychiatry, 75*, 9–17.
- Canli, T., & Lesch, K. P. (2007). Long story short: The serotonin transporter in emotion regulation and social cognition. *Nature Neuroscience, 10*, 1103–1109.
- Caspi, A., McClay, J., Moffitt, T. E., Mill, J., Martin, J., Craig, I. W., . . . Poulton, R. (2002). Role of genotype in the cycle of violence in maltreated children. *Science, 297*, 851–854.
- Cicchetti, D. (2013). Annual research review: Resilient functioning in maltreated children—Past, present, and future perspectives. *Journal of Child Psychology and Psychiatry, 54*, 402–422.
- Cicchetti, D. (2016). Socioemotional, personality, and biological development: Illustrations from a multilevel developmental psychopathology perspective on child maltreatment. *Annual Review of Psychology, 67*, 187–211.
- Cicchetti, D., & Rogosch, F. A. (1996). Equifinality and multifinality in developmental psychopathology. *Development and Psychopathology, 8*, 597–600.
- Cicchetti, D., & Rogosch, F. A. (2012). Gene \times Environment interaction and resilience: Effects of child maltreatment and serotonin, corticotropin releasing hormone, dopamine, and oxytocin genes. *Development and Psychopathology, 24*, 411–427.
- Cicchetti, D., Rogosch, F. A., & Oshri, A. (2011). Interactive effects of corticotropin releasing hormone receptor 1, serotonin transporter linked polymorphic region, and child maltreatment on diurnal cortisol regulation and internalizing symptomatology. *Development and Psychopathology, 23*, 1125–1138.
- Cicchetti, D., Rogosch, F. A., & Thibodeau, E. L. (2012). The effects of child maltreatment on early signs of antisocial behavior: Genetic moderation by tryptophan hydroxylase, serotonin transporter, and monoamine oxidase A genes. *Development and Psychopathology, 24*, 907–928.
- Coie, J. D., & Dodge, K. A. (1983). Continuities and changes in children’s social status: A five-year longitudinal study. *Merrill-Palmer Quarterly, 29*, 261–282.
- Colder, C. R., Scalco, M., Trucco, E. M., Read, J. P., Lengua, L. J., Wieczorek, W. F., & Hawk, L. W., Jr. (2013). Prospective associations of internalizing and externalizing problems and their co-occurrence with early adolescent substance use. *Journal of Abnormal Child Psychology, 41*, 667–677.
- Coltman, T., Devinney, T. M., Midgley, D. F., & Venaik, S. (2008). Formative versus reflective measurement models: Two applications of formative measurement. *Journal of Business Research, 61*, 1250–1262.
- Cutuli, J. J., Raby, K. L., Cicchetti, D., Englund, M. M., & Egeland, B. (2013). Contributions of maltreatment and serotonin transporter genotype to depression in childhood, adolescence, and early adulthood. *Journal of Affective Disorders, 149*, 30–37.
- DeYoung, C., Cicchetti, D., Rogosch, F. A., Gray, J., Eastman, M., & Grigorenko, E. (2011). Sources of cognitive exploration: Genetic variation in the prefrontal dopamine system predicts Openness/Intellect. *Journal of Research in Personality, 45*, 364–371.
- Dudbridge, F. (2013). Power and predictive accuracy of polygenic risk scores. *PLOS Genetics, 9*, e1003348.

- Duncan, L. E., & Keller, M. C. (2011). A critical review of the first 10 years of candidate gene-by-environment interaction research in psychiatry. *American Journal of Psychiatry*, *168*, 1041–1049.
- Edwards, K. M., Probst, D. R., Rodenhizer-Stämpfli, K. A., Gidycz, C. A., & Tansill, E. C. (2014). Multiplicity of child maltreatment and biopsychosocial outcomes in young adulthood: The moderating role of resiliency characteristics among female survivors. *Child Maltreatment*, *19*, 188–198.
- Grace, J. B., & Bollen, K. A. (2008). Representing general theoretical concepts in structural equation models: The role of composite variables. *Environmental and Ecological Statistics*, *15*, 191–213.
- Haskett, M. E., Nears, K., Ward, C., & McPherson, A. (2006). Diversity in adjustment of maltreated children: Predictors of resilient functioning. *Clinical Psychology Review*, *26*, 796–812.
- Hawley, P. H. (1999). The ontogenesis of social dominance: A strategy-based evolutionary perspective. *Developmental Review*, *19*, 97–132.
- Hostinar, C. E., Cicchetti, D., & Rogosch, F. A. (2014). Oxytocin receptor gene polymorphism, perceived social support, and psychological symptoms in maltreated adolescents. *Development and Psychopathology*, *26*, 465–477.
- Jaffee, S. R., Caspi, A., Moffitt, T. E., Polo-Tomás, M., & Taylor, A. (2007). Individual, family, and neighborhood factors distinguish resilient from non-resilient maltreated children: A cumulative stressors model. *Child Abuse & Neglect*, *31*, 231–253.
- Jaffee, S. R., Caspi, A., Moffitt, T. E., & Taylor, A. (2004). Physical maltreatment victim to antisocial child: Evidence of an environmentally mediated process. *Journal of Abnormal Psychology*, *113*, 44.
- Keller, M. C. (2014). Gene \times environment interaction studies have not properly controlled for potential confounders: The problem and the (simple) solution. *Biological Psychiatry*, *75*, 18–24.
- Kim, J., & Cicchetti, D. (2003). Social self-efficacy and behavior problems in maltreated and nonmaltreated children. *Journal of Clinical Child and Adolescent Psychology*, *32*, 106–117.
- Kovacs, M. (1992). *The children's depression inventory: A self-rated depression scale for school-aged youngsters*. Pittsburgh, PA: University of Pittsburgh Press. (Original work published 1982)
- Ladd, G. W., & Burgess, K. B. (1999). Charting the relationship trajectories of aggressive, withdrawn, and aggressive/withdrawn children during early grade school. *Child Development*, *70*, 910–929.
- Lewis, B. F. (2003). A critique of literature on the underrepresentation of African Americans in science: Directions for future research. *Journal of Women and Minorities in Science and Engineering*, *9*, 361–374.
- Little, T. D., Cunningham, W. A., Shahar, G., & Widaman, K. F. (2002). To parcel or not to parcel: Exploring the question, weighing the merits. *Structural Equation Modeling*, *9*, 151–173.
- Luthar, S. S., & Cicchetti, D. (2000). The construct of resilience: Implications for interventions and social policies. *Development and Psychopathology*, *12*, 857–885.
- Masten, A. S. (2015). *Ordinary magic: Resilience in development*. New York: Guilford Press.
- Masten, A. S., Hubbard, J. J., Gest, S. D., Tellegen, A., Garmezy, N., & Ramirez, M. (1999). Competence in the context of adversity: Pathways to resilience and maladaptation from childhood to late adolescence. *Development and Psychopathology*, *11*, 143–169.
- Masten, A. S., Roisman, G. I., Long, J. D., Burt, K. B., Obradović, J., Riley, J. R., . . . Tellegen, A. (2005). Developmental cascades: Linking academic achievement and externalizing and internalizing symptoms over 20 years. *Developmental Psychology*, *41*, 733.
- Masyn, K. E. (2013). Latent class analysis and finite mixture modeling. In T. D. Little (Ed.), *The Oxford handbook of quantitative methods in psychology* (Vol. 2, pp. 551–611). Oxford: Oxford University Press.
- Mendez, J. L., Fantuzzo, J., & Cicchetti, D. (2002). Profiles of social competence among low-income African American preschool children. *Child Development*, *73*, 1085–1100.
- Monroe, S. M., & Simons, A. D. (1991). Diathesis-stress theories in the context of life stress research: Implications for the depressive disorders. *Psychological Bulletin*, *110*, 406.
- Muthén, L. K., & Muthén, B. O. (1998–2012). *Mplus user's guide* (7th ed.). Los Angeles: Author.
- Nylund, K. L., Asparouhov, T., & Muthén, B. O. (2007). Deciding on the number of classes in latent class analysis and growth mixture modeling: A Monte Carlo simulation study. *Structural Equation Modeling*, *14*, 535–569.
- Pekarik, E. G., Prinz, R. J., Liebert, D. E., Weintraub, S., & Neale, J. M. (1976). The Pupil Evaluation Inventory: A sociometric technique for assessing children's social behavior. *Journal of Abnormal Child Psychology*, *4*, 83–97.
- Pluess, M. (2015). Individual differences in environmental sensitivity. *Child Development Perspectives*, *9*, 138–143.
- Risch, N., Herrell, R., Lehner, T., Liang, K. Y., Eaves, L., Hoh, J., . . . Merikangas, K. R. (2009). Interaction between the serotonin transporter gene (5-HTTLPR), stressful life events, and risk of depression: A meta-analysis. *Journal of the American Medical Association*, *301*, 2462–2471.
- Roisman, G. I., Newman, D. A., Fraley, R. C., Haltigan, J. D., Groh, A. M., & Haydon, K. C. (2012). Distinguishing differential susceptibility from diathesis-stress: Recommendations for evaluating interaction effects. *Development and Psychopathology*, *24*, 389–409.
- Roy, A., Gorodetsky, E., Yuan, Q., Goldman, D., & Enoch, M. A. (2010). Interaction of FKBP5, a stress-related gene, with childhood trauma increases the risk for attempting suicide. *Neuropsychopharmacology*, *35*, 1674–1683.
- Ryan, N. D., Puig-Antich, J., Ambrosini, P., Rabinovich, H., Robinson, D., Nelson, B., . . . Twomey, J. (1987). The clinical picture of major depression in children and adolescents. *Archives of General Psychiatry*, *44*, 854–861.
- Sapientza, J. K., & Masten, A. S. (2011). Understanding and promoting resilience in children and youth. *Current Opinion in Psychiatry*, *24*, 267–273.
- Sharpley, C. F., Palanisamy, S. K., Glyde, N. S., Dillingham, P. W., & Agnew, L. L. (2014). An update on the interaction between the serotonin transporter promoter variant (5-HTTLPR), stress and depression, plus an exploration of non-confirming findings. *Behavioural Brain Research*, *273*, 89–105.
- Sumner, J. A., McLaughlin, K. A., Walsh, K., Sheridan, M. A., & Koenen, K. C. (2014). CRHR1 genotype and history of maltreatment predict cortisol reactivity to stress in adolescents. *Psychoneuroendocrinology*, *43*, 71–80.
- Teisl, M., Rogosch, F. A., Oshri, A., & Cicchetti, D. (2012). Differential expression of social dominance as a function of age and maltreatment experience. *Developmental Psychology*, *48*, 575.
- Thibodeau, E. L., Cicchetti, D., & Rogosch, F. A. (2015). Child maltreatment, impulsivity, and antisocial behavior in African American children: Moderation effects from a cumulative dopaminergic gene index. *Development and Psychopathology*, *27*(4, Pt. 2), 1621–1636.
- Uher, R., & McGuffin, P. (2010). The moderation by the serotonin transporter gene of environmental adversity in the etiology of depression: 2009 update. *Molecular Psychiatry*, *15*, 18–22.
- US Department of Health and Human Services, Administration for Children and Families, Administration on Children, Youth and Families, Children's Bureau. (2017). *Child Maltreatment 2015*. Available from <http://www.acf.hhs.gov/programs/cb/research-data-technology/statistics-research/child-maltreatment>
- Vachon, D. D., Krueger, R. F., Rogosch, F. A., & Cicchetti, D. (2015). Assessment of the harmful psychiatric and behavioral effects of different forms of child maltreatment. *JAMA Psychiatry*, *72*, 1135–1142.
- Wilsnack, S. C., Vogeltanz, N. D., Klassen, A. D., & Harris, T. R. (1997). Childhood sexual abuse and women's substance abuse: National survey findings. *Journal of Studies on Alcohol*, *58*, 264–271.
- Wright, J. (1983). *The structure and perception of behavioral consistency*. (Unpublished doctoral dissertation, Stanford University).
- Yaeger, R., Avila-Bront, A., Abdul, K., Nolan, P. C., Grann, V. R., Birchette, M. G., . . . Ziv, E. (2008). Comparing genetic ancestry and self-described race in African Americans born in the United States and in Africa. *Cancer Epidemiology Biomarkers & Prevention*, *17*, 1329–1338.
- Yates, T. M., & Grey, I. K. (2012). Adapting to aging out: Profiles of risk and resilience among emancipated foster youth. *Development and Psychopathology*, *24*, 475–492.