# Gene-experience correlation during cognitive development: Evidence from the Adolescent Brain Cognitive Development (ABCD) Study[SM]

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**Running Title:** Gene-experience interplay in cognitive development

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Conflict of Interest Statement

Dr. Dale reports that he was a Founder of and holds equity in CorTechs Labs, Inc., and serves on its Scientific Advisory Board. He is a member of the Scientific Advisory Board of Human Longevity, Inc. He receives funding through research grants from GE Healthcare to UCSD. The terms of these arrangements have been reviewed by and approved by UCSD in accordance with its conflict of interest policies. All other authors have no conflicts to disclose.

Word count: 5,594

Abstract

**Background:** Findings in adults have shown more culturally sensitive 'crystallized' measures of intelligence have greater heritability, these results were not able to be shown in children.

**Methods:** With data from 8,518 participants, aged 9 to 11, from the Adolescent Brain Cognitive Development (ABCD) Study®, we used polygenic predictors of intelligence test performance (based on genome-wide association meta-analyses of data from 269,867 individuals) and of educational attainment (based on data from 1.1 million individuals), associating these predictors with neurocognitive performance. We then assessed the extent of mediation of these associations by a measure of recreational reading.

**Results:** more culturally sensitive 'crystallized' measures were more strongly associated with the polygenic predictors than were less culturally sensitive 'fluid' measures. This mirrored heritability differences reported previously in adults and suggests similar associations in children. Recreational reading more strongly statistically mediated the genetic associations with crystallized than those with fluid measures of cognition.

**Conclusion**: This is consistent with a prominent role of gene-environment correlation in cognitive development measured by "crystallized" intelligence tests. Such experiential mediators may represent malleable targets for improving cognitive outcomes.

## Introduction

Scores on cognitive tests in both children and adults have been linked to long term outcomes and to genetic variation(1–4). Some cognitive tests, e.g., those requiring literacy and mathematical skills, depend upon and are more sensitive to variability in cultural and socio-economic factors. These measures are often referred to as 'crystallized' intelligence measures. In contrast, other tests that tap the capacity to solve novel problems, or process novel information, often referred to as 'fluid' measures, are less culturally sensitive and are less strongly related to socio-economic variables(5,6). A recent review reported systematic differences in heritability (an estimate of trait variability attributable to genetic variation) of the traits measured by these different kinds of cognitive measures(7). Surprisingly, in studies of adult twins, more culturally sensitive tests exhibited higher, rather than lower, heritability; which runs counter to predictions from conventional models of intelligence. The authors described similar trends in the twin studies of children, but increased heritability of crystallized relative to fluid measures have not yet been established for children, in whom intellectual functions are continuing to mature.

The finding that the measures most strongly influenced by cultural factors exhibit higher heritability is perhaps counterintuitive; however previous authors have noted that genetic variation can be associated with environmental, cultural, or experiential (ECE) factors that themselves amplify effects of a genotype on the phenotype, a phenomenon often referred to as rGE (gene-environment correlation). These associations between genotypes and ECE factors could influence the development of cognitive and intellectual abilities in several ways. As an example, if others in the social environments of children recognize traits, e.g., precocious behavior, in those with a genetic propensity for a given cognitive ability, they may begin to treat

such individuals differently, rewarding them disproportionately for intellectual pursuits,

investing more in their instruction, and/or placing them in environments that drive learning more

effectively. Alternatively, the associations can be driven by the motivation of the children

themselves if for example they develop greater enthusiasm for intellectual activities for which

they have been more frequently rewarded, and which they then pursue more assiduously, thus

enjoying beneficial effects of the increased practice associated with these activities.  In either

case, the genetically advantaged abilities are disproportionately enhanced by these mediating

ECE factors.  Of course, individuals with less advantageous genotypes may experience the

converse of these social and motivational effects, resulting in languishing, or in the worst case

suppressed, intellectual development, even within similar environments. Such rGE effects can

increase variance in intellectual phenotypes and increase estimates of heritability using both

epidemiological and genomic methods(8).  The important implication is that a component of this

increased heritability requires the mediating ECE effects for its expression.  In essence, more

direct biological effects of the genotype *and* associated differences in the environments or

experiences of the child are both contributing causal factors influencing the mature phenotype,

but they act through dissociable mechanisms.

Heritability is a population statistic frequently measured using a twin design. For this

study, we used polygenic scores to examine variation in genetic and experiential factors and their

relationship to trait measures of cognitive function. Polygenic scores have the advantage that

they can be used to index relevant genetic factors in samples of unrelated individuals by

leveraging the statistical power of meta-analysis results from large Genome Wide Association

Studies (GWAS).  Using neurocognitive test scores, genomic data, and a measure of parent-

reported recreational reading assessed in a large sample of 8,618 children, aged 9 to11, from the

ABCD Study®, we used polygenic scores of intelligence test performance (based on GWAS of

269,867 individuals(9)) and educational attainment, sometimes considered a proxy for

intellectual ability (based on 1.1 million individuals(10)), to ask 3 questions:  First, do these

genomic predictors account for more of the variability in estimates of culturally sensitive

crystallized traits than fluid traits in children, as might be expected from reports of higher

heritability in adult twins? Second, does a parent-reported estimate of the time their children

spend reading for pleasure mediate the relationship between a genomic predictor and measures

of cognitive performance, consistent with a role of this experiential enhancer of performance in

increasing heritability?  Third, if mediation is observed, is this mediating effect larger for the

culturally sensitive crystallized than the fluid measures of cognitive performance, consistent with

a role for rGE in the higher heritability of these measures?

In additional analyses, we examined the degree to which the findings in the ethnically

diverse ABCD sample were similar between the subgroup of children with high genomic

European ancestry (EurA) and a remaining subgroup of children who were from diverse ancestry

groups (DivA). Finally, using simulations, we tested whether our observed findings may be due

to previously reported differences in test-retest reliabilities (for crystallized vs fluid measures).

## Materials and Methods

*2.1 Data available in the ABCD data release 2.0.1*

The ABCD study (http://abcdstudy.org) enrolled the families of 11,875 children aged 9 or

10 years at baseline(11). This longitudinal study follows the development of these children at 21

sites across the US for ten years. The cohort exhibits a large degree of sociodemographic

diversity. Exclusion criteria were limited to: 1) lack of English proficiency; 2) the presence of

severe sensory, neurological, medical or intellectual limitations that would inhibit the child's

ability to comply with the protocol; 3) an inability to complete an MRI scan at baseline. The

study protocols are approved by the University of California, San Diego Institutional Review

Board(12). Parent/caregiver permission and child assent from each participant were obtained.

Here, our data were drawn from the baseline assessments shared in ABCD release 2.0.1 (NDAR

DOI: 10.15154/1504041).

*2.1.1* Cognitive Measures

Seven of the 10 cognitive tasks were subtests from The NIH Toolbox Cognition Battery[®]

(NTCB) in the version recommended for ages 7+(http://www.nihtoolbox.org)(13). The average

time to complete this battery is approximately 35 minutes. The NTCB was administered in

English(14), using an iPad, with support from a research assistant when needed. The battery

yields individual test scores measuring specific constructs and composite scores that have been

shown to be highly correlated with 'gold standard' measures of intelligence in adults(15) and

children(5). Here, all 7 individual test scores and 2 composite scores were examined: the

Crystallized Cognition Composite Score (derived from scores on the Picture Vocabulary and

Oral Reading Recognition measures) and the Fluid Cognition Composite Score (derived from the

remaining measures). Additionally, three neurocognitive tasks were used that were not

components of the NTCB: Rey-Auditory Learning Task, Little Man Task and Matrix Reasoning.

Please see supplementary materials for a description of each task.

*2.1.2 Latent Neurocognitive Factors*

Thompson et al. derived a three factor, varimax rotated, solution for the latent structure

across the neurocognitive battery in ABCD using Bayesian Probabilistic PCA(16). The final

latent factor solution included the measures described above, except for the Matrix Reasoning

task which had very little effect on the solution. The factors will be referred to as Bayesian

Factors (BF) 1-3. Language tasks loaded most heavily on BF1, which was highly correlated with

the Crystallized Composite (r=0.93); executive functioning tasks loaded most heavily on BF2;

and learning/memory tasks loaded heavily on BF3.

*2.1.3 Recreational Reading*

Parents of ABCD participants were asked to complete a survey of their children's

activities. One question asked, "Does your child read for pleasure?"  The follow-up question

was, "About how many hours per week does your child read for pleasure?".  This estimate of

number of hours of recreational reading was log transformed due to skewness. To confine the

analyses of this variable to a homogenous group of children who read for pleasure, we included

only children whose parents answered 'yes' to the first question.

*2.1.4 Genetic Data and Computing Polygenic Scores*

Using genotype data we derived genetic ancestry using fastStructure(17) with four

ancestry groups. Genetic principal components were also calculated using PLINK. Variants were

imputed using the Michigan Imputation Server(18). Polygenic scores were computed using

PRSice(19). The Intelligence Polygenic Score (IPS) was trained on 269,867 individuals by

Savage et al.(9), and focused on neurocognitive tests considered to gauge fluid intelligence. The

Education Attainment Polygenic Score (EAPS) was generated from 1.1 million individuals,

predicting the phenotype of number of years of schooling completed. Please see supplementary

materials for further details on genetic data and analysis.

We were primarily focused on studying the IPS association with cognitive tests in

ABCD, due to it being trained on a more directly relevant phenotype. However, we additionally

examined EAPS as a secondary analysis for comparison as it has been previously used as a proxy

for cognitive ability and has a discovery sample size four times the size of the IPS.

*2.2 Analytic Methods*

*2.2.1 Ancestry Group Analyses*

Training and testing polygenic scores in different ancestry groups has been shown to

reduce predictive power(20–22). Given the ancestry differences between the polygenic score

discovery samples (predominantly European) and the ABCD study (multiple ancestry groups),

we wanted to confirm our main results in the full samples were not driven by population

structure. As such we additionally performed analyses in two subsamples: 1) children with

estimated proportion of European ancestry higher than 90% (EurA) and 2) a group of the

remaining children with diverse ancestry, which included those from other or mixed ancestry

(DivA).

*2.2.2 Statistical Model for Genomic Prediction of Behavioral Measures*

To assess the association between the polygenic scores and cognitive performance in

ABCD, we fit Generalized Linear Mixed-Effects Models (GLMMs) using the gamm4

package(23) in R. Each model predicted performance on a different cognitive measure.

Continuous variables were z-scored before model fitting to allow coefficients to be interpreted as

standardized effect sizes. To test if regression coefficients differed between regressions we

performed a z-test on the difference between coefficients, based on the propagated standard error

for the two regression coefficients as the sum of the error of variances for each measure. This test

assumes the standard errors are uncorrelated and so provides a conservative estimate of

significance. Please see supplementary materials for details and covariates used.

*2.2.3 Differential Mediation Analysis*

To assess whether recreational reading is a plausible ECE factor increasing heritability of

crystallized cognition, through rGE effects, we performed a mediation analysis.  Specifically, we

compared the statistical mediation effects of recreational reading experience on the associations

between the IPS and both the Crystallized Composite and Fluid Composite, respectively. We

achieved this by calculating the proportion of mediation of recreational reading on the i) IPS-

crystallized and the ii) IPS-fluid associations using an average causal mediation effect model(24)

across 10,000 bootstrap samples. With bootstrapped samples we tested if the mediation effect of

recreational reading on the IPS-crystallized association was greater than that of IPS-fluid , by

performing a Welch t-test on the samples. Mediation analysis was performed using general linear

models in the mediation package in R(25), see supplementary materials for details.

2.2.4 *Data and Code Availability*

The ABCD dataset is available approved researchers at https://nda.nih.gov/abcd. A

jupyter notebook of the analysis be found at

github.com/robloughnan/ABCD_Intelligence_Polygenic_Score.

## Results

*Demographics*

222       Figure *1* illustrates a flow-chart for sample selection. For the final analysis we have

223    8,518 individuals in the full sample, 4,885 in the EurA sample and 3,633 in the DivA sample.

*Behavioral Measures and Sociocultural Factors*

Mean performance, standard deviation (SD), median and estimates of variance explained

by age, sex, and the set of socio-cultural covariates (parental marital status, highest education

level of parent/caregiver, household income, ethnicity, genetic principal components) are given

for each behavioral measure examined in Table 2.  Consistent with previous reports, there are

substantial differences in the degree to which socio-cultural factors account for variability in

these measures.  The Crystallized Composite, its constituent Picture Vocabulary and Reading

Recognition measures, and BF1, on which these measures of language and literacy load heavily,

all exhibit higher levels of association with socio-cultural variables. This pattern persisted when

controlling for IPS (Sup. Table 2). Sex, age and socio-cultural factors explained little variability

in recreational reading. Partial correlations between the individual cognitive task measures

controlling for covariates (Figure 2), suggest that performance on the different tasks is modestly

correlated across children (rs=.08-.41) in this sample.  Correlations peak in the .3 range within

Fluid Composite measures, and the highest correlation is observed between the two Crystallized

Composite measures (Picture Vocabulary and Oral Reading r=.41).

*Genomic Prediction of Crystallized and Fluid Cognition Measures*

Table 3 summarizes the regression results for predicting the Crystallized and Fluid

Composites with IPS or EAPS in the full sample, and separately in the EurA and DivA

subsamples. The IPS was a significant predictor of both measures in all analyses. Importantly,

the standardized regression coefficient was significantly higher for the Crystallized than the

Fluid Composite regardless of ancestry group (full sample: z=4.8, p=$1.8\times10^{-6}$, EurA: z=4.6,

p=$5.1\times10^{-6}$ and DivA: z=2.5, p=$1.4\times10^{-2}$).

In no case did the EAPS, despite a much larger training sample size, appear to account

for more of the variance in the neurocognitive measures than did IPS.  However, across ancestry

groups and for both composite scores, combining both genomic predictors explained

significantly more variance in behavior than IPS alone (supplementary results). IPS + EAPS

explained 5.8% variance (p=$4.5\times10^{-64}$) in the Crystallized Composite for EurA (a 40% increase

compared to IPS alone). Supplementary Tables 3-8 show regression results for each behavior

using IPS, EAPS and IPS + EAPS within each ancestry group.

Fitting separate regression models for each individual task in the neurocognitive battery,

we found that the IPS was a significant predictor for each cognitive measure for the full sample

and the EurA subsample (all p values<$10^{-3}$), surviving the Bonferroni-corrected significance

threshold of 0.05/10=0.005. Within the DivA subsample only six of the ten tasks were

individually significantly predicted by the IPS (Sup. Table 8). Figure 3 shows the standardized

regression coefficients of IPS predicting performance on each task, as well as the Crystallized

and Fluid Composite measures from the NTCB and Bayesian Factors 1-3(16), in the full sample.

Individual cognitive measures included in the Crystallized Composite have consistently higher

IPS standardized regression weights than the measures included in the Fluid Composite. Other

neurocognitive tasks from the ABCD battery (shaded in gray) showed similar associations to the

Fluid Composite. The results for the Bayesian Factors mirrored these results: BF1, on which

'Crystallized" measures had the highest factor loadings (Sup. Figure 1)(16), displayed a stronger

association with IPS than BF2 and BF3 on which 'Fluid", executive function and memory

measures had higher loadings. The results in the subsamples (EurA and DivA) are provided in

supplemental material.

*Differential Mediation Results*

The mediation analysis showed that recreational reading partially mediated associations

between IPS and both composite measures, proportions of mediation: fluid 0.084 (95% CI:

0.047-0.14, $p \leq 10^{-4}$), crystallized 0.12 (95% CI: 0.88-0.16, $p \leq 10^{-4}$).  However, the differential

mediation analyses revealed a highly significant difference between the large degree of

attenuation of the association between IPS and the Crystallized Composite relative to that

between IPS and the Fluid Composite (Welch t-test: $t=125$, $df=19053$, $p<10^{-300}$), shown in Figure

*4.*

*Sensitivity Analyses to Address Test Reliability*

A previous study reported the test-retest reliability for the Fluid Composite from the

NTCB (.76) was somewhat lower than for the Crystallized Composite (.85)(5), raising questions

about whether differences in the strength of their associations with IPS could be attributed to

more noise in the Fluid Composite measure.  In supplementary sensitivity analyses we

demonstrate that our results are robust to the addition of simulated noise to the Crystallized

Composite that mimics this difference in test reliability. At this level of simulated noise we

estimated 1.0 power (alpha=0.05) to detect $Cryst_{noise}$ having a significantly greater IPS

standardized regression coefficient than the Fluid Composite. Moreover, additional sensitivity

analyses indicate that the observed differences in the mediation effects of recreational reading

are similarly robust against potential measurement error modelled as random noise.  These

analyses are described in detail in Supplemental Material.

## Discussion

We have shown that polygenic predictors of intelligence test performance and of

educational attainment are associated with neurocognitive performance in this large group of

children from diverse backgrounds. These results are consistent with previous findings

demonstrating that virtually all behavioral traits, including cognitive and intellectual phenotypes,

are heritable(26).  Moderate estimates of heritability of many behavioral phenotypes also

establish that a substantial portion of the variability is due to independent environmental

influences. Given that behavioral phenotypes emerge through interactions between children and

their physical, social, and cultural environments, much attention has been paid to how these

environmental factors modify the phenotypes, since they are presumably the malleable factors.

However, recently, more attention has been focused on the possible roles of mediating

nongenetic (ECE) factors that, through their statistical association with genetic variation (rGE),

may amplify heritability(7,8).

We found that a culturally dependent estimate of crystallized cognitive functions, the

Crystallized Composite measure from the NTCB, is more strongly associated with the best

available polygenic predictor of intelligence test performance than is the Fluid Composite

measure, consistent with earlier findings in adults of heritability differences(7) and polygenic

score performance(27) across similar measures.  This is despite the IPS being based on a large

meta-analysis of GWAS combining cognitive measures that were described by the authors as

primarily "fluid intelligence" measures(9). Indeed, the relative size of the IPS association across

the 15 measures examined here (Figure 2) closely mirrored the relative percent variance

explained in these measures by socio-cultural variables (Table 2), a pattern that persists after

accounting for IPS (Sup. Table2).  Moreover, for children who read for pleasure, the extent of

recreational reading was found to partially mediate the associations between IPS and both

Composite measures, but to a significantly greater degree for the crystallized than for the fluid

measure, consistent with a more prominent role of rGE in the development of abilities tapped by

measures that are both more heritable, and apparently more sensitive to socio-cultural variables.

In other words, even when controlling for *independent* contributions of more global sociocultural

variables, how often a child reads for pleasure more strongly mediates the association between

IPS and crystallized rather than in fluid performance.

It is perhaps unsurprising that recreational reading more strongly mediates the association

between IPS and culturally sensitive measures of intelligence since such measures are generally

sensitive to educational factors. Indeed, a measure of oral reading proficiency loads highly on

both measures of crystallized functions examined here, Crystallized Composite and BF1. One

can imagine that children with neurobehavioral phenotypes advantageous for learning to read

might be more likely to develop the habit of reading for pleasure than those with other

neurobehavioral phenotypes, for a variety of reasons. However, these results imply that choosing

to read for pleasure at 10 years of age is associated with having a genotype linked to intellectual

functions most dependent on reading, and that an estimate of the frequency of reading behavior

mediates that link. This is consistent with previous descriptions of  rGE effects, and with

analyses by Beam and Turkheimer(8), who showed that increasing rGE over time could explain

observed increases in the heritability of measures of cognitive function through development.

The ABCD study will provide an opportunity to measure changes in heritability at later time

points of this longitudinal study. Importantly, despite the lower test-retest reliability of the fluid

compared to the crystallized composite score from the NTCB(5), our supplementary analyses

show that this difference in test reliability is unlikely to explain our findings.

Though recreational reading would appear to be an enhancing mediator of intellectual

development, it is important to note that genotype-correlated ECE factors can also suppress

intellectual development.  As an example, early struggles to read by children with less

advantageous genotypes for reading may decrease the likelihood that these children will choose

reading activities, leading to slower progression of these faculties.  Worse, if children's early

reading attempts are experienced very negatively, these children may develop avoidant responses

to reading, which could result in active suppression of developing literacy.  Importantly, when

these kinds of differences originate with differences in children's genotypes, they can increase

heritability and exaggerate disparities.  Identifying ECE factors that contribute to heritability of

cognitive and intellectual phenotypes is important because it can point to practices that better

adapt to neurogenetic diversity among children.  Innovative pedagogical practices may lead to

approaches that increase "enhancing" ECE effects in the subset of children disadvantaged by

current practices and reduce ECE effects that suppress intellectual development and academic

achievement, which may lead to more equitable educational outcomes.

*Limitations and Caveats*

The proportion of variance in the cognitive measures accounted for by the genomic

predictors was larger in the EurA participants than in the DivA group(Sup. Tables 5 & 7), as

would be expected given the discovery samples were in individuals of European ancestry.

However, the patterns were generally similar in the DivA group.  This suggests similar genetic

architecture for these cognitive phenotypes across ancestry groups and supports the validity of

the results from the full sample.  Analyses in all three groups included as covariates the top ten

genetic principal components derived from the full sample.  Because of broad ancestral diversity

in the ABCD cohort, there is limited power for comparing the effects in different ancestry

groups.  As has been discussed in genetics generally(28,29), the lower predictive performance in

the DivA group once again underscores the importance of collecting genetic data from

ancestrally diverse populations and developing methods that can be used across ancestry groups.

One may have predicted that EAPS would have been a more powerful predictor of

cognitive measures in ABCD than IPS, due to it having over 4 times the discovery sample size.

However, we found generally the IPS had stronger associations (Sup. Tables 3-8), perhaps

because the phenotype is a better match between training and testing. This contrasts with results

of a previous study of adults, where EAPS explained 7-10% of the variance in cognition(10),

while IPS explained only 2-5%(9). This discrepancy may be due to methodological differences,

alternatively the young age of the cohort may be the key difference. Educational attainment,

while clearly related to scores on cognitive tests, may be influenced by other genetically

influenced traits (e.g. personality) that may contribute to greater persistence in formal education;

thus the EAPS is likely to reflect to a greater degree these traits. Such pleiotropic of EAPS

effects has been observed in adults(30).  When we include both EAPS and IPS in a single model,

together they explain 5.8% of the variability in the Crystallized Composite (EurA, Sup. Table 6),

substantially more than IPS alone explains (4.1%), indicating that these genomic predictors

capture unique sources of the relevant variance, and are likely measuring different (relevant)

constructs.

These results are consistent with previous evidence for a role of genetic variation in

developing cognitive functions, and they strengthen the evidence for rGE during cognitive

development.  However, it should be emphasized that the genomic predictors (together) account

for only 4.15% of cognitive performance variance in the full sample.  Furthermore, this was

observed for the Crystallized Composite measure, the culturally sensitive measure hypothesized

to exhibit increased genetic association as a result of rGE effects. The additive effects of

potentially confounding sociocultural covariates, even controlling for IPS, accounted for 13.2%

of the variability.  For the Fluid Composite the genomic predictors together accounted for only

1.1% of the variance, and sociocultural covariates accounted for almost 5%.  Of note, even with

the narrow 2 year age range in the cohort, age alone accounts for 10% of the variability in the

Crystallized Composite and 7% in the Fluid Composite. These effects may reveal clues about a

highly dynamic process of cognitive and intellectual development in children.

Finally, though the results of the mediation analysis focusing on recreational reading

strengthen the plausibility that such ECE mediators associate with genotypes and increase

genetic effects, these results do not prove a causal explanation, and none should be inferred.  In

the context of an observational study such as ABCD it is always possible that confounding

variables not accounted for in the analysis are responsible for the mediation effect we observed.

Funding

This research was in part supported by the    and National Institute of Mental Health under the

award number, R01MH122688.

ABCD Acknowledgement

Data used in the preparation of this article were obtained from the **Adolescent Brain Cognitive**

**Development□ Study** (**ABCD Study**®) (https://abcdstudy.org), held in the NIMH Data Archive

(NDA). This is a multisite, longitudinal study designed to recruit more than 10,000 children age

9-10 and follow them over 10 years into early adulthood. The ABCD Study is supported by the

National Institutes of Health and additional federal partners under award numbers:

U01DA041022, U01DA041028, U01DA041048, U01DA041089, U01DA041106,

U01DA041117, U01DA041120, U01DA041134, U01DA041148, U01DA041156,

U01DA041174, U24DA041123, and U24DA041147

A full list of supporters is available at https://abcdstudy.org/federal-partners/. A listing of

participating sites and a complete listing of the study investigators can be found at

https://abcdstudy.org/principal-investigators.html. ABCD Study consortium investigators

designed and implemented the study and/or provided data but did not necessarily participate in

analysis or writing of this report. This manuscript reflects the views of the authors and may not

reflect the opinions or views of the NIH or ABCD Study consortium investigators.

The ABCD data repository grows and changes over time. The ABCD data used in this came

from [NIMH Data Archive Digital Object Identifier (10.15154/1504041)].

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518     Figure 1 *Flow chart of sample selection and exclusion.*

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520 Figure 2 *Partial correlation matrix showing intercorrelations among individual task*
521 *performance measures (controlling for age, sex, parental marital status, parental education,*
522 *household income, principal components of genetic ancestry and Hispanic status) in the full*
523 *sample included in the present study of genomic predictors.*
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Figure 3 *Standardized regression coefficients of IPS for fitting linear mixed models to performance on Fluid and Crystallized Composites, each individual task from the NTCB, additional measures from the ABCD neurocognitive battery, and Bayesian (latent) Factors 1-3, in the full sample. Prediction of the Crystallized Composite is significantly stronger than for the Fluid Composite. Tasks included in the Fluid Composite (shaded in blue) have consistently lower regression coefficients than those included in the Crystallized Composite (shaded in red). Additional measures from the neurocognitive battery exhibit associations with IPS more similar to the Fluid Composite than to the Crystallized Composite, however Bayesian Factor 1, on which the verbal tasks load heavily, exhibits an association similar to the Crystallized Composite. Error bars show estimates of 95% confidence intervals as 1.96 × standard error.*

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543 Figure 4 *Differential mediation analysis in singletons (N= 5,210): histograms shows 10,000*
544 *bootstrap estimates for proportion of mediation of recreational reading on: i) IPS and*
545 *Crystallized Composite (red) and ii) IPS and Fluid Composite (blue). Recreational reading*
546 *attenuates the relationship between the IPS and the Crystallized Composite to a significantly*
547 *greater degree.*

|  | Full Sample | European Ancestry | Other Ancestry |
|---|---|---|---|
| Total N | 8518 | 4885 | 3633 |
| | **Mean (SD)** | | |
| Age - months | 119.05 (7.48) | 119.21 (7.49) | 118.85 (7.47) |
| | **N (%)** | | |
| Sex Male | 4438 (52.1) | 2576 (52.7) | 1862 (51.3) |
| Parent Married = Yes | 6024 (70.7) | 4066 (83.2) | 1958 (53.9) |
| **Parental Education** | | | |
|   < HS Diploma | 302 (3.5) | 21 (0.4) | 281 (7.7) |
|   HS Diploma/GED | 649 (7.6) | 138 (2.8) | 511 (14.1) |
|   Some College | 2149 (25.2) | 899 (18.4) | 1250 (34.4) |
|   Bachelor | 2318 (27.2) | 1548 (31.7) | 770 (21.2) |
|   Post Graduate Degree | 3100 (36.4) | 2279 (46.7) | 821 (22.6) |
| **Household Income** | | | |
|   [<50K] | 2353 (27.6) | 596 (12.2) | 1757 (48.4) |
|   [>=50K & <100K] | 2444 (28.7) | 1471 (30.1) | 973 (26.8) |
|   [>=100K] | 3721 (43.7) | 2818 (57.7) | 903 (24.9) |
| **Race** | | | |
|   White | 5715 (67.7) | 4750 (97.4) | 965 (27.1) |
|   Black | 1129 (13.4) | 1 (0.0) | 1128 (31.7) |
|   Asian | 199 (2.4) | 0 (0.0) | 199 (5.6) |
|   Other | 1397 (16.6) | 128 (2.6) | 1269 (35.6) |
| **Hispanic** | | | |
|   Hispanic | 1628 (19.1) | 131 (2.7) | 1497 (41.2) |

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574    *Table 1: Summary of demographics for individuals included in the full sample for the present*
575            *genomic prediction analyses, and for the genomic European Ancestry and genomic Other*
576            *Ancestry subgroups.*

| | Mean (SD) | Median | Sex | Age | Sociocultural |
|---|---|---|---|---|---|
| Crystallized Composite | 86.87 (6.93) | 87 | 0.01 | 9.51 | 21.57 |
| Fluid Composite | 92.18 (10.43) | 93 | 0.32 | 7.17 | 10.28 |
| Reading | 91.23 (6.73) | 91 | 0.01 | 5.97 | 13.18 |
| Picture Vocabulary | 85.04 (8.02) | 84 | 0.07 | 7.67 | 20.14 |
| Pattern | 88.29 (14.47) | 88 | 0.57 | 4.81 | 1.90 |
| List | 97.43 (11.81) | 97 | 0.13 | 2.04 | 9.47 |
| Picture | 103.33 (12.01) | 103 | 0.51 | 1.17 | 5.46 |
| Flanker | 94.42 (8.83) | 96 | 0.03 | 3.21 | 3.73 |
| Cardsort | 92.97 (9.26) | 94 | 0.48 | 3.76 | 5.22 |
| Rey Auditory Verbal | 43.78 (9.96) | 44 | 1.25 | 2.23 | 8.57 |
| Matrix Reasoning | 18.13 (3.74) | 18 | 0.34 | 2.74 | 9.15 |
| Little Man Task | 0.60 (0.17) | 0.56 | 0.48 | 5.13 | 6.25 |
| Bayesian Factor 1 | 0.05 (0.76) | 0.06 | 0.28 | 9.63 | 20.85 |
| Bayesian Factor 2 | 0.02 (0.76) | 0.06 | 0.22 | 5.49 | 2.65 |
| Bayesian Factor 3 | 0.04 (0.70) | 0.04 | 0.89 | 1.59 | 7.83 |
| Recreational Reading (hours) | 6.5 (10) | 4 | 0.45 | 0.18 | 0.86 |

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*Table 2 Mean (SD) and median for each behavioral measure in the full sample, estimated % variance explained by sex, age, and the set of socio-cultural covariates (parental marital status, parental education, household income, genetic ancestry PCs and Hispanic/non-Hispanic).*

| Sample | Fluid Composite | | | | Crystallized Composite | | | |
|---|---|---|---|---|---|---|---|---|
| | Stand. β | t | P value | % Var. Explained | Stand. β | t | P value | % Var Explained |
| | IPS | | | | | | | |
| Full Sample | 0.28 | 8.03 | 1.14E-15 | 0.75 | 0.50 | 15.82 | 1.31E-55 | 2.86 |
| EurA | 0.11 | 7.53 | 6.10E-14 | 1.15 | 0.21 | 14.48 | 1.44E-46 | 4.13 |
| DivA | 0.20 | 3.41 | 6.52E-04 | 0.32 | 0.40 | 7.34 | 2.68E-13 | 1.47 |
| | EAPS | | | | | | | |
| Full Sample | 0.11 | 7.23 | 5.26E-13 | 0.61 | 0.19 | 14.21 | 2.56E-45 | 2.32 |
| EurA | 0.09 | 6.60 | 4.66E-11 | 0.89 | 0.18 | 12.95 | 9.36E-38 | 3.34 |
| DivA | 0.08 | 3.38 | 7.28E-04 | 0.32 | 0.15 | 6.66 | 3.24E-11 | 1.21 |

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*Table 3 Regression results for GLMMs associating IPS (top) and EAPS (bottom) with Crystallized Composite and Fluid Composite of the NIH toolbox within full Sample and ancestry subgroups.*