Impact of the COVID-19 Pandemic on Early Child Cognitive Development: Initial Findings in a Longitudinal Observational Study of Child Health

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ABSTRACT

Since the first reports of novel coronavirus in the 2020, public health organizations have advocated preventative policies to limit virus, including stay-at-home orders that closed businesses, daycares, schools, playgrounds, and limited child learning and typical activities. Fear of infection and possible employment loss has placed stress on parents; while parents who could work from home faced challenges in both working and providing full-time attentive childcare. For pregnant individuals, fear of attending prenatal visits also increased maternal stress, anxiety, and depression. Not surprising, there has been concern over how these factors, as well as missed educational opportunities and reduced interaction, stimulation, and creative play with other children might impact child neurodevelopment. Leveraging a large on-going longitudinal study of child neurodevelopment, we examined general childhood cognitive scores in 2020 and 2021 vs. the preceding decade, 2011-2019. We find that children born during the pandemic have significantly reduced verbal, motor, and overall cognitive performance compared to children born pre-pandemic. Moreover, we find that males and children in lower socioeconomic families have been most affected. Results highlight that even in the absence of direct SARS-CoV-2 infection and COVID-19 illness, the environmental changes associated COVID-19 pandemic is significantly and negatively affecting infant and child development.

INTRODUCTION

Since the beginning in March of 2020, the outbreak of the SARS-CoV-2 (COVID-19) pandemic in the USA, and the accompanying economic shut-down has brought about significant upheavals to the social, economic, and public health environments in which children live, grow and play. While children, and those under age 5, have largely been spared from the severe health and mortality complications associated with SARS-CoV-2 infection [1, 2], they have not been immune to the impact of the stay-athome, masking, and social distancing policies. These policies, meant to limit spread of the SARS-CoV-2 virus, have closed daycares, schools, parks, and playgrounds [3, 4], and have disrupted children's educational opportunities [5], limited explorative play and interaction with other children [6], and reduced physical activity levels [7]. From the beginning of the pandemic, there has been concern that these public-health policies would adversely impact infant and early child development and mental health. While there is no past analogue or example of non-conflict related wide-spread and prolonged lock-downs from which to draw information from, concern for child development stemmed principally from the known impact that family and home stress, parent and child anxiety, lack of stimulating environments, and other economic and environmental adversities can have on the developing infant and child brain [8, 9].

Like many other states and regions in the United States, elementary, secondary, and post-secondary schools were closed throughout the state of Rhode Island (RI) to in-person instruction from March 16, 2020 through to the beginning of the 2020/2021 school year. However, with the surge of virus infections in the fall of 2020, most schools in RI continued to operate with fully remote or with hybrid in-person/on-line learning until January 2021. For younger children, daycare centers were also closed in March of 2020, but were allowed to reopen with reduced capacity in June, 2020. Restrictions on daycare capacity remained in effect until June 2021. Broader state-wide travel restrictions and stay-athome orders were enforced from March to May, 2020, with many businesses operating with reduced on-site work forces and/or work-from-home options until mid-2021. Indoor and outdoor mask policies were also in place throughout 2020 and 2021 following CDC guidance. Despite being one of the smaller US states with a population of just over 1 million, RI has suffered high numbers of SARS-CoV-2 infections, with approximately 154,000 cases of COVID-19 illness and almost 3,000 deaths. RI has mirrored national trends with respect to disproportionate infections and deaths in Hispanic or Latino and Black or African American communities [10, 11] as well as lower income families [12].

From 2009, Brown University and the Warren Alpert Medical School at Brown University has been home to a longitudinal study of child health and neurodevelopment, termed the RESONANCE study. Now part of the NIH Environmental influences on Child Health Outcomes (ECHO) program, the RES-ONANCE cohort consists of approximately 1600 caregiver-child dyads, who have been continuously

enrolled between 0 and 5 years of age since 2009 and have been followed through infancy, childhood, and early adolescence. This cohort, therefore, offers a unique opportunity to explore the impact of the COVID-19 pandemic on child health trends in RI, which may reflect broader trends in the US.

In the absence of direct SARS-CoV-2 infection, environmental exposures associated with COVID-19 pandemic can affect the developing infant and child through multiple pathways. The human brain is unique in its prolonged developmental timeline [13, 14]. Infants are born with relatively immature brains that, like them, are simultaneously competent and vulnerable. Infants are inherently competent in their ability to initiate relationships, explore, seek meaning, and learn; but are vulnerable and depend entirely on caregivers for their survival, emotional security, modeling of behaviors, and the nature and rules of the physical and socio-cultural world that they inhabit [15]. The infant brain is likewise born with immense capacity to learn, remodel, and adapt, but is sensitive and vulnerable to neglect and environmental exposures that begin even before birth [16-18]. Optimal brain development depends on secure and trusting relationships with knowledgeable caregivers who are responsive to the infant's needs and interests. Neurodevelopmental processes, including myelination and synaptogenesis, for example, are stimulated by external cues and experiences like maternal interaction, and physical skinto-skin "kangaroo" care, touch, and warmth [19-22]. The brain's adaptive plasticity, however, is a double-edged sword. While positive and enriching environments can promote healthy brain development [23-27], neglect insecurity, stress, and lack of stimulation can impair maturing brain systems and disrupt cognitive and behavioral outcomes [28-30].

Maternal stress, anxiety, and depression in pregnancy can impact the developing fetal and infant brain structure and connectivity, leading to potential delays in motor, cognitive, and behavioral development [31, 32]. It is believed that alterations in fetal exposure to stress-related hormones, including cortisol, affect theses changes in brain structure and function [33-35]. Past analysis has revealed strong associations between maternal prenatal stress and anxiety related to maternal or paternal displacement and job loss and infant health (birth weight and gestation duration), mortality, temperament, and cognitive development [36]. Throughout the COVID-19 pandemic, maternal and paternal job-loss, employment furloughs, or increased food / housing insecurity have been experienced by many families. Survey results at the beginning of the pandemic showed significantly increased rates of clinically relevant symptoms of maternal depression and anxiety [37].

Given these changes in children's home, education, and social environments, it is not surprising that cross-sectional and longitudinal studies of child and adolescent mental health throughout the current pandemic have revealed increased stress, anxiety, and depression [38]. Studies of child learning further show reduced academic growth in math and language arts in elementary and high school children [4]. The impact on infant and child cognitive development, however, remains less clear. Despite the

known impact of parent and family stress, economic adversity, reduced parent and peer interaction and other stimulating environments, on child brain development [9], direct observation of reduced development as a result of the COVID-19 pandemic remains elusive, despite early findings of altered temperament in 3month old infants [39].

In this work, therefore, we specifically sought to explore individual and population-level trends in infant and early child neurodevelopment. Analyses of cognitive development, assessed using Mullen Scales of Early Learning [40], a population normed and clinically administered tool that assesses function across the five primary domains of fine and gross motor control, visual reception, and expressive and respective language via direct observation and performance, provides some of the first direct evidence of the developmental impact of the COVID-19 pandemic. Comparing yearly mean scores since 2011, controlling for age, gender, demographic, and socioeconomic indicators, we find striking evidence of declining overall cognitive functioning in children beginning in 2020 and continuing through 2021. We find that males appear significantly more impacted than females, and that higher socioeconomic status (SES, as measured by maternal education [41]) helps buffer against this negative impact. On a more individual level, we examined longitudinal pre and during-pandemic trends in the same children from 2018 to 2021, again finding declines in ability in 2020 and 2021.

Finally, to examine the impact of the pandemic during pregnancy on subsequent child development, we compared developmental scores in children up to 1 year of age born prior to 2019 and after July 2020 (i.e., those born prior to the pandemic and those whose *in utero* development may have been impacted by the COVID-19 environment for at least the last trimester of pregnancy). Again, we find significantly reduced scores in children born since the start of the pandemic, with lower SES individuals and males more significantly affected. Throughout all analyses, however, we found that the perceived stress of mothers in our cohort was unchanged throughout the pandemic, and was not significantly associated with developmental scores.

These results provide compelling early evidence of the impact of the COVID-19 pandemic, even in the absence of direct SARS-CoV-2 infection, on infant and early child neurodevelopment.

METHODS

All data were acquired in accordance with ethical approval and oversight by the Rhode Island Hospital institutional review board, with informed consent obtained from all parents or legal guardians.

Since 2011, 1224 cognitive assessments were collected from 672 healthy, full-term, and neurotypically-developing children between 3 months to 3 years of age. Repeated measures were separated by at least 1 year (mean = 384 +/- 41 days). A general overview of all child assessment timings are shown in **Figure 1a**. This dataset included 1070 assessments (from 605 children) prior to March 2020; 154 assessments (from 118 children) between March 2020 and June 2021; with 39 children who were born just prior to the pandemic in 2018 and 2019, and following during the pandemic to 2021.

To start, we grouped measures by testing year from 2011 to 2021 and performed an analysis of covariance to compare the three primary composite measures: the early learning composite (ELC), verbal development quotient (VDQ) and non-verbal development quotient (NVDQ) between each pre and during pandemic year-pair; with child age, and maternal education (as a proxy for socioeconomic status, SES) as covariates.

To investigate the cognitive trends more rigorously, we next constructed a series of general linear mixed-effects models that modeled the cognitive measures while taking a step-wise approach to include additional predictive factors, beginning with

$$CM_{i,j} = \beta_{0,j} + \beta_{1,j} \times age_{i,j} + \beta_{2,j}COVID_{model}$$
^[1]

Where CM is the cognitive measure of interest (ELC, VDQ, or NVDQ) of child *j* at time-point *i*. $\beta_{0,j}$ is the intercept and $\beta_{1,j}, \dots, \beta_{n,j}$, are the regression coefficients. $\beta_{0,j}, \beta_{1,j}$ combine a sample fixed effect and a subject-specific random effect (u_j), e.g.,

$$\beta_{0,j} = \beta_0 + \mu_{0,j},$$
 [1b]

and

$$\beta_{1,j} = \beta_1 + \mu_{1,j}.$$
 [1c]

allow individual differences in mean cognitive function and change with age. The $COVID_{model}$ term is a binary factor that is 0 for any testing date prior to March 2020, and 1 for testing dates thereafter. Equation [1] was fit to the complete cohort dataset using the fitlme function in Matlab (MathWorks, Cam-

bridge, MA v2019b). The COVID_{model} term allowed us to test the hypothesis that pandemic-related environmental changes are associate with a significant negative change in cognitive functioning.

Building from our simple function, we used a step-wise regression approach to systematically include additional factors and interactive terms in our general model, i.e.,:

$$CM_{i,j} = \beta_{0,j} + \beta_{1,j} \times age_{i,j} + \beta_{2,j}COVID_{model} + \beta_{3,j}SES,$$
[2]

$$CM_{i,j} = \beta_{0,j} + \beta_{1,j} \times age_{i,j} + \beta_{2,j}COVID_{model} + \beta_{3,j}SES + \beta_{4,j}Gender_j$$
^[3]

to control for potential differences in gender and socioeconomic factors in the population. Given past findings associating COVID-19 stay-at-home orders and prematurity or potential low birth weight [42], we further included these birth outcomes as additional predictors,

$$CM_{i,j} = \beta_{0,j} + \beta_{1,j} \times age_{i,j} + \beta_{2,j}COVID_{model} + \beta_{3,j}SES + \beta_{4,j}Gender_j + \beta_{5,j}Birthweight + \beta_{6,j}Gestation$$
[4]

Finally, we also tested the interaction between the COVIDmodel and SES and Gender terms to test whether these factors had additive or buffering effects,

$$CM_{i,j} = \beta_{0,j} + \beta_{1,j} \times age_{i,j} + \beta_{2,j}COVID_{model} + \beta_{3,j}SES \times COVID_{model} + \beta_{4,j}Gender_j \times COVID_{model}$$

[5]

At each stage of analysis, we examined the parsimonious fit of the model using the Bayesian Information Criterion (BIC) [43], stopping when additional factors no longer improved the model fit. In the final model, we then examined the significance level of each model parameter, specifically the COVID_{model} term, which denotes a potential significant difference in pre- and during pandemic scores.

Recognizing that the analysis from 2011 to 2021 contained differing sets of children we next reduced our dataset to 39 children who were enrolled up to a year prior to the pandemic and who had at least 1 visit before and 1 visit during the pandemic (**Table 1, Figure 1b**). To this reduced set of data, we performed analogous mixed-effects modeling using the series of general linear models constructed above (Eqns. 1 to 5).

Given the potential impact of the pandemic on maternal stress and, consequently, the developing fetal brain, we next compared the cognitive measures in 291 children under 1 year of age who were born

before January 2019 (i.e., their prenatal and first postnatal year of development occurred before the pandemic) to 118 children born after July 2020 (i.e., at least one term of pregnancy and the first postnatal year occurred during the pandemic) (**Table 1, Figure 1c**). As above, we used a mixed-effects approach to model our series of general linear models (Eqns. 1 to 5).

To test whether maternal stress was a potential causative factor, we replaced the Model term in our general linear models with the total score from the perceived stress scale (PSS) [44], a 10-item self-report that provides a continuous scale of perceived and experienced stress due to life situations.

$$CM_{i,j} = \beta_{0,j} + \beta_{1,j} \times age_{i,j} + \beta_{2,j}PSS + \beta_{3,j}SES \times PSS + \beta_{4,j}Gender_j \times PSS$$
[6]

PSS scores were obtained from each pregnant mother prior to birth and at each child visit starting in 2017. We performed this analysis twice. In the first set, we used the post-natal scores collected at the same time as the child cognitive measures, in the second, we used the prenatal PSS scores obtained from the mother during pregnancy.

RESULTS

Demographic summaries of the children included in each set of analysis (complete set; children followed pre- and during pandemic; and children under 1 year of age born before and during the pandemic) are shown in **Table 1**.

Results from our ANCOVA comparing each year pre- and during-pandemic year pair is shown in **Table 2**. In general, we find mean ELC values from 2011 to 2019 ranging from 98.5 to 107.3, with standard deviations of 15.2 to 19.7 (**Fig. 2**), in general agreement with the expected mean of 100 and standard deviation of 15. Means and standard deviations for 2020 (March to December) and 2021 (January through Aug) were: 86.3+/-17.9 and 78.9+/-21.6, respectively. Controlling for differences in age and maternal education, we find inconsistent differences in mean ELC between 2011-2019 and 2020, but consistent and significant reductions between 2011-2019 and 2021 (p<0.001). Results are similar for the verbal and non-verbal composite measures (**Fig. 2, Table 2**). In all cases maternal education was a significant and positive factor on ELC, VDQ, and NVDQ measures.

Results from our mixed model analysis using our complete data set are summarized in **Table 3** and reinforce the results obtained from our initial ANCOVA. For each composite score, the model including child age, maternal education, child gender, birth weight, gestation duration, and test timing (pre vs. during the pandemic)), as well as interaction terms of timing, gender, and maternal education was the most parsimonious model (lowest BIC). Across all measures, we found cognitive scores were significantly reduced during the pandemic by 27 to 37 points (or almost two full standard deviations), p values < 0.01, with higher maternal education, increased birth weight and increased gestation duration being protective; while males were more heavily affected. We did not find significant differences in birth weight or gestation duration overall in the pre vs during pandemic children (p > 0.3), **Fig. 3**.

Repeating this set of analysis in the subset of 39 children who were born immediately before the beginning of the pandemic and whom we have followed up over the past 18 months (**Table 4**) presents contrasting results. Longitudinal plots of individual cognitive measures are shown in **Fig. 4**. Across all composite cognitive measures, we find that the inclusion of a term that distinguishes between pre vs. during the COVID-19 pandemic does not improve the model fit, and no significant decrease in cognitive measures in these children. This result suggests that much of the reductions observed in **Figure 2**, and **Tables 2** and **3** may be driven by the infants born during the pandemic. Results from our last set of analysis (**Table 5**), restricted to newborns and infants under 1 year of age born before or during the pandemic, support this hypothesis. Here we find a significant decrease in child cognitive scores in children born during the pandemic with males affected more than females, and higher maternal education being a protective factor.

The strong effect of the pandemic on early neurodevelopment suggests that maternal stress, already shown to be increased in mothers at the start of the pandemic may be an important factor. However, including maternal stress, recorded both prenatally and following delivery into our model, in place of the Model term was not significant (**Table 6**). Indeed, when examining perceived stress in mothers by year we find (**Fig. 5**) no significant increase or decrease during the pandemic compared to prepandemic years.

DISCUSSION

Children are inherently shaped by their environment. Across the fetal, infant, and early childhood lifestages, a child's brain undergoes immense structural and functional growth that is driven by an integrative mixture of genetic and environmental factors. The outbreak of the COVID-19 pandemic, and the associated economic shut-down, school disruptions, and social distancing, stay-at-home, and mask policies have fundamentally altered the environment in which children and pregnant individuals have lived, over the past 18 months. Despite much speculation, the short and long-term impacts of the COVID-19 pandemic on fetal and child health and neurodevelopment in the absence of direct infection are not yet known [45]. This gap makes it difficult to develop evidence-based guidelines of care for expectant mothers and individuals, design effective strategies for follow-up care of sensitive infants; or provide informed guidance for school and daycare reopening and in-person vs. online learning.

Leveraging data collected continuously over the past decade in Providence, RI and surrounding areas, we sought to investigate how the pandemic has impacted cognitive development and function in newborns and young children. Included pregnant individuals and children reported no symptoms of SARS-CoV-2 infection or had evidence of positive antibody or RT-PCR testing. Families also reported having adhered to stay-at-home and on-going mask and social distancing policies, suggesting observed effects are environmentally driven rather than due to potential direct effects of infection. However, we did not perform antibody testing to confirm past infection status.

Overall, we find that measured verbal, non-verbal, and overall cognitive scores are significantly lower since the beginning of the pandemic. Looking further, we find that children born before the pandemic and followed through the initial stages do not show a reduction in skills or performance, but rather that young infants born since the beginning of the pandemic show significantly lower performance than infants born before January 2019. Thus, our results seem to suggest that early development is impaired by the environmental conditions brought on by the pandemic.

In contrast to other on-going studies through the pandemic [37, 39], we did not find an increase in general maternal stress and, thus, this was not a significant predictor factor in our analysis. This may reflect a general insensitivity of the PSS tool used here to pandemic-related stress; the potential selection bias in the families included in our study; or strong family and social support networks available to the pregnant individuals. The PSS is a standardized ten-item questionnaire that asks about general life stressors and how stressful individuals find their lives but does not include specific questions related to health or wellbeing. In contrast, the MOM-COPE study utilized retrospective data collection using an ad-hoc developed questionnaire focused on worry and anxiety of COVID-19 infection, pregnancy risk, and their own and their infant's health [39]. The survey-based study by Lebel and colleagues [37] also used a specially developed questionnaire to gauge maternal concern of the pandemic and its im-

pact on their own and their infant's health. Additional measures of depression and anxiety were assessed using the standardized Edinburgh Depression Scale (EDS [46]) and PROMIS Anxiety Adult 7item short form [47] instruments.

With respect to included participants, families in our study were drawn from the local Providence and surrounding communities with focus on typical child development. Exclusion criteria for enrollment has been consistent across the duration of the study and includes: twin or multiple pregnancy; preterm birth prior to 37 weeks; small for gestational age and/or birthweight less and 1500g; 5min APGAR scores <= 8; major psychiatric illness in the mother, including depression requiring medication within 6 months of pregnancy; use of alcohol, tobacco, or illicit substances during pregnancy; diagnosed neurological disorder (e.g., epilepsy) in the child. During the pandemic, we did not target individuals with COVID-19 exposure or illness. However, since all study visits take place in a clinical setting, parents less concerned about the pandemic, and those with strong social support networks, may have been more likely to participate than those with greater concerns. Thus, our observation that maternal stress (PSS) did not significantly increase may simply reflect the reality that we only tested less stressed and anxious mothers. These parents may also have greater financial security or other socioeconomic characteristics. However, in ad hoc analysis of maternal education between families tested before or during the pandemic (**Fig. 5b**). This, however, is an area requiring further investigation.

Family and social support is also an important contributor to maternal health and wellbeing [48] that can also affect infant temperament, behavior, and cognitive development [49]. Building from our discussion of maternal stress, we unfortunately did not collect additional measures related to daycare or working from home status of our parents. It is possible that parents of lower socioeconomic means may have been less able to afford daycare or prolonged parental leave. These are additional areas of focus, which could provide important guides for public health policies.

The first 1,000 days of a child's life are commonly cited as an important and sensitive period of child development. Environmental factors, including maternal mental and physical health, nutrition, stimulation, and supportive caregiving can individually and in combination affect the developing fetal and infant brain throughout this early life period [19-22]. Many of these factors have been substantively impacted by public health policies enacted in response to the SARS-CoV-2 outbreak. Work-from-home and shelter-in-place orders, for example, along with closed daycares, nurseries, and preschools may have dramatically changed the quantity and quality of parent, caregiver, and teach-child interaction and stimulation. Studies in older children and adolescents over the past year have found reduced social interaction, increased media consumption, and reduced physical activity [50-52]. It is likely these same trends are true for younger children and infants as well with the closure or reduced capacity of

daycares and preschools, and may be associated with impaired motor development, motor coordination and visual processing, language development, and socioemotional processing. In addition, masks worn in public settings and in school or daycare settings may impact a range of early developing skills, such as attachment, facial processing, and socioemotional processing.

Unfortunately, we do not have direct or parent-reported measures indicative of parent or caregiverchild interaction, early media exposure, or physical activity to investigate the potential causative role of these factors.

One aspect also not investigated here is the impact of mask-wearing by the study staff during child visits and assessments [53]. The inability of infants to see full facial expressions may have eliminated non-verbal cues, muffled instructions, or otherwise altered the understanding of the test questions and instructions.

For parents who were able to work from home, and did not face furlough or employment loss, the dual role of childcare and work has increased the strain on parents, in particular mothers, resulting in increased parental stress and anxiety. For families who have suffered job loss, they may experience greater stress, depression, anxiety, as well as food and housing insecurity. Although we found no significant changes in the degree of maternal perceived stress, and we did, however, find that maternal education, often used as a marker of surrogate of socioeconomic status, was generally associated with improved cognitive function and, as an interactive term, had a buffering effect against the impact of the pandemic. This is particularly salient given the disproportionate effect the pandemic has had on lower income families [54], who have not only dealt with job loss and financial insecurity, but are also over-represented in front line and essential service employment with increased risk of SARS-CoV-2 infection and COVID-19 illness [55].

What is unclear from our data, however, is if observed declines or impairments are temporary and will normalize as employment and school closures are lifted and children return to pre-pandemic levels of play and interaction, and family financial insecurity and mental health challenges subside. Unfortunately, when that will occur is also unknown given the ongoing surge of infections associated with new virus variants. It is clear, however, that young infants and children are developing differently than prepandemic, and that addressing this now while their brain is at its most plastic and responsive, is imperative. Programs such as unemployment insurance, Supplemental Nutrition Assistance Program (SNAP), the Special Supplemental Nutrition Program for Women, Infants, and Children (WIC), and housing assistance, may help minimize the impact of the pandemic on the most sensitive of children. In addition, further research directly exploring aspects of parent-child attachment, interaction, nutrition,

food security, and environmental stimulation is needed to understand the primary driving factors underlying the trends presented here.

CONCLUSION

The COVID-19 pandemic has fundamentally altered the child health landscape, with pregnant mothers and individuals, and children living in a strikingly different economic, psychosocial, and educational environment than what was present just 18 months ago. Against this environmental backdrop, unanswered questions remain regarding the impact of the work-from-home, shelter-in-place, and other public health policies that have limited social interaction and typical childhood experiences on early child neurodevelopment. In this work, we provide early evidence suggestive of significant reductions in attained cognitive function and performance in children born over the past 18 months during the pandemic. While socioeconomic factors appear to mitigate against the negative consequences of the pandemic, the primary factors underlying our observed trends remain unknown. Understanding these factors are critical to helping ensure affected children rebound as the pandemic winds down and they re-enter daycares and schools; as well as implementing additional public health and educational policies that address the most affected of children, particularity those in lower income families.

DATA SHARING

All data acquired and presented here is freely available upon request to the authors.

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CONTRIBUTOR ROLES

All listed authors were involved in the study design, data acquisition and analysis, drafting and revising this manuscript, and provide final and accountable approval for its contents. SCD and DV verify the underlying data.

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The authors report no significant financial conflicts of interest with respect to the subject matter of this manuscript.

Tables & Figures

Table 1. Group demographics for each child cohort, including the complete set of data tested over the past decade; children born just before the pandemic and followed during the pandemic; and children under 1 year of age born before January 2019 or after July 2020.

		Complete Set	Matched Pre & During	Under 1 Year
Biological Sex	Male	292	17	186
biological Sex	Female	381	22	223
Dana	Hispanic	112	9	70
Race	Not Hispanic	561	30	339
	Caucasian / White	320	29	248
	African American / Black	43	2	28
Ethnicity	Asian	8	0	6
	Mixed Race	111	5	76
	Unknown / Declined	191	3	51
	Some Primary School	0	0	0
	Primary School Graduate	0	0	0
	Some High School	20	2	10
Maternal Educa- tion	High School Graduate	91	7	66
	Some College	157	13	104
	College Graduate	143	8	82
	Graduate	262	9	147
Birth Outcomes	Birth Weight (lbs)	7.3 (1.3)	7.36 (1)	7.3 (1.3)
Birth Outcomes	Gestation (weeks)	38 (3.7)	38 (1.7)	39 (2)

Table 2. Year-pair comparisons of mean ELC, VDQ, and NVDW composite scores measured before and during the COVID-19 pandemic. While we see inconsistent differences between each pre-pandemic year and 2020, we find consistent statistically significant reductions between each pre-pandemic year and 2021, controlling for child age and maternal education.

	ELC			VDQ		NVDQ			
	2020	2021		2020	2021		2020	2021	
2011	0.023	<0.0001	2011	0.045	< 0.0001	2011	0.029	< 0.0001	
2012	0.79	<0.0001	2012	0.97	< 0.0001	2012	0.92	< 0.0001	
2013	< 0.0001	<0.0001	2013	<0.0001	< 0.0001	2013	0.007	< 0.0001	
2014	< 0.0001	<0.0001	2014	<0.0001	< 0.0001	2014	<0.0001	< 0.0001	
2015	< 0.0001	<0.0001	2015	0.0004	< 0.0001	2015	0.0006	< 0.0001	
2016	0.0001	<0.0001	2016	0.14	< 0.0001	2016	<0.0001	< 0.0001	
2017	0.015	<0.0001	2017	0.042	< 0.0001	2017	0.16	< 0.0001	
2018	0.12	<0.0001	2018	0.11	< 0.0001	2018	0.35	< 0.0001	
2019	0.002	<0.0001	2019	0.008	< 0.0001	2019	0.04	< 0.0001	

Table 3. Results of our sequential or step-wise general linear models investigating differences in pre vs. during-pandemic composite ELC, VDQ, and NVDQ measures. The COVID-model term is 0 for all test dates prior to March 2020, and 1 for all dates following, allowing us to test for a signifiant change in pre vs. during-pandemic scores. Across all models, we find this term is significant (p < 0.01). We also note that males appear more heavily affected that females, but that higher maternal education appears to be protective against the effects of the pandemic.

ELC	Mod	lel 1	Mod	lel 2	Mod	lel 3	Mod	lel 4	Mod	lel 5
BIC	10288.0628	346	9321.27150	05	9305.27828	35	8469.12013	36	8461.15408	35
Term	Estimate	pValue								
Intercept	95.27373 2	0.000000	77.15468 5	0.000000	79.75920 5	0.000000	5.697486	0.635849	7.342544	0.540383
Child Age	0.004234	0.002910	0.004015	0.005853	0.004240	0.003387	0.004745	0.001152	0.004722	0.001149
COVID-Model	- 12.10102 5	0.000000	- 11.99400 3	0.000000	- 12.03450 1	0.000000	- 12.56996 4	0.000000	- 31.70724 2	0.000091
Maternal Ed.			3.375311	0.000000	3.376083	0.000000	3.062195	0.000000	2.553374	0.000000
Male					-4.795305	0.000020	-4.218355	0.000122	-3.629912	0.001307
Gestation							1.644557	0.000005	1.670624	0.000004
Birth Weight							1.618325	0.003349	1.608542	0.003496
Male x Model									-5.210031	0.118383
Maternal Ed. x Model									4.089579	0.003280
VDQ	Mod	lel 1	Mod	lel 2	Mod	lel 3	Mod	lel 4	Mod	lel 5
BIC	112	252.715679	101	190 924489	101	179.861822	92	299 916193	92	299.341288
Term	Estimate	pValue								
Intercept	93.53301 0	0.000000	66.89222 9	0.000000	69.93796 3	0.000000	3.376040	0.844570	4.581158	0.790121
Child Age	0.007594	0.000044	0.005348	0.004688	0.005635	0.002886	0.005894	0.002397	0.005890	0.002391
COVID-Model	20.12408 6	0.000000	19.70636 5	0.000000	19.74255 3	0.000000	19.45135 4	0.000000	37 34855 2	0.001706
Maternal Ed.			5.109479	0.000000	5.105734	0.000000	4.503137	0.000000	4.038469	0.000000
Male					-5.633027	0.000276	-5.549127	0.000455	-4.986241	0.002484
Gestation							1.408398	0.005929	1.440316	0.005010
Birth Weight							2.083655	0.008176	2.072341	0.008587
Male x Model									-4.726815	0.338660

ELC	Model 1		Model 2		Mod	Model 3		lel 4	Model 5	
BIC	10288.0628	346	9321.27150)5	9305.278285		8469.12013	36	8461.154085	
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Maternal Ed. x Model									3.775018	0.065723

NVDQ	Mod	el 1	Mod	el 2	Mod	lel 3	Mod	lel 4	Mod	lel 5
BIC	11011.785624		9960.852998		9950.889997		9037.219460		9036.929494	
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Intercept	108.5042 97	0.000000	81.26229 8	0.000000	84.14567 1	0.000000	8.099286	0.600946	9.080857	0.558075
Child Age	-0.004783	0.005940	-0.007278	0.000047	-0.007061	0.000072	-0.008527	0.000002	-0.008541	0.000002
COVID-Model	15 30586 5	0.000000	- 14.71208 9	0.000000	14.70860 6	0.000000	14.67272 4	0.000000	27.07196 1	0.009252
Maternal Ed.			5.149235	0.000000	5.117169	0.000000	4.791398	0.000000	4.425289	0.000000
Male					-4.982514	0.000539	-4.711620	0.000963	-4.120723	0.005689
Gestation							1.743029	0.000149	1.764651	0.000133
Birth Weight							1.568929	0.026826	1.566894	0.027418
Male x Model									-5.459957	0.207356
Maternal Ed. x Model									2.836307	0.116332

Table 4. Results of our step-wise general linear models investigating the impact of the pandemic on longitudinal cognitive development in children born up to a year prior to the pandemic. Overall, we note no significant reductions in development trends. However, given the same sample size (n=39) care should be taken when interpreting these results.

ELC	Mod	lel 1	Mod	lel 2	Mod	lel 3	Model 4		Mod	lel 5
BIC	1	1102.12489	11	101.898327	11	01.612437	ç	951.013815	ļ g	953.514413
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Intercept	100.2663 60	0.000000	97.40534 0	0.000000	100.4188 25	0.000000	2.425663	0.963978	5.410311	0.920276
Child Age	-0.019395	0.027243	-0.019942	0.022518	-0.018752	0.028939	-0.014696	0.106447	-0.014752	0.101032
COVID-Model	-2.047941	0.599668	-1.582132	0.684468	-2.147518	0.577349	-2.401197	0.551321	-4.655415	0.579965
Maternal Ed.			0.684191	0.175319	0.624238	0.203671	1.235866	0.099346	0.960662	0.239703
Male					-5.524167	0.110745	-4.711777	0.195625	-4.057912	0.280674
Gestation							1.846022	0.244516	1.776850	0.264256
Birth Weight							2.767081	0.234533	2.879715	0.217461
Male x Model									-4.696733	0.439359
Maternal Ed. x Model									0.979381	0.441595
VDQ	Mod	lel 1	Mod	lel 2	Mod	lel 3	Mod	lel 4	Mod	el 5
BIC	1:	226.191366	1:	226.774931	1:	226.374071	1	064.219276	1	067.622234
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Intercept	101.3552 48	0.000000	97.85392 6	0.000000	102.4927 23	0.000000	9.015940	0.919235	5.232606	0.952853
Child Age	-0.023287	0.097212	-0.025017	0.075669	-0.022803	0.098320	-0.019001	0.222603	-0.019044	0.218556
COVID-Model	-4.743447	0.416743	-3.787672	0.518215	-4.945343	0.394431	-5.742650	0.348743	1.291075	0.926868
Maternal Ed.			0.900950	0.232498	0.769328	0.297368	1.885437	0.104399	2.121262	0.143755
Male					-8.271020	0.111431	-7.058668	0.235855	-4.752222	0.472669
Gestation							1.668816	0.518236	1.689203	0.509866
Birth Weight							2.607091	0.477348	2.691657	0.460487
Male x Model									-7.675059	0.445066
Maternal Ed. x Model									-0.650951	0.763981

NVDQ	Mod	lel 1	Model 2		Mod	Model 3		lel 4	Model 5	
BIC	1	1179.560518		1181.306747		1182.293166		028.841428	1032.726942	
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Intercept	116.3016 31	0.000000	115.0556 96	0.000000	117.8856 55	0.000000	61.13313 6	0.429975	62.08237 9	0.427188
Child Age	-0.028338	0.004479	-0.029330	0.003731	-0.028594	0.004333	-0.025756	0.019887	-0.025880	0.019502
COVID-Model	-4.038943	0.429808	-3.518395	0.498170	-3.961846	0.441532	-3.823532	0.505631	-4.243381	0.715800
Maternal Ed.			0.346362	0.610793	0.252290	0.709778	0.435734	0.675374	0.283903	0.829291
Male					-4.714430	0.305498	-4.454052	0.390088	-3.770467	0.526075
Gestation							0.437372	0.845883	0.418870	0.852975
Birth Weight							4.950345	0.127611	4.979334	0.126569
Male x Model									-2.011659	0.807478
Maternal Ed. x Model									0.306913	0.861942

Table 5. Results of our step-wise general linear models investigating the impact of the pandemic on newborns and young infants under 1 year of age born pre-pandemic (before Jan 2019) and during the pandemic (after July 2020). Like results in **Table 3**, we not again that child born during the pandemic exhibit significantly lower cognitive performance than children born before the pandemic (p<0.001) but that maternal education is a protective factor.

ELC	Mod	lel 1	Mod	lel 2	Mod	lel 3	Mod	lel 4	Mod	lel 5
BIC	45	574.859772	41	149.364403	41	144.183573	35	562.855010	35	551.746593
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Intercept	98.86778 5	0.000000	89.28601 5	0.000000	91.23906 0	0.000000	-4.010653	0.784098	-1.668205	0.907809
Child Age	-0.007007	0.213556	-0.010064	0.080259	-0.009064	0.112001	-0.008868	0.127818	-0.009452	0.100363
COVID-Model	10.32953 9	0.000000	10.54104 0	0.000000	10.59565 1	0.000000	11.641199	0.00000	49.88214 4	0.000070
Maternal Ed.			1.955167	0.000549	1.922774	0.000617	1.567565	0.003179	0.990045	0.066967
Male					-3.540089	0.006954	-2.253041	0.072222	-1.719909	0.176304
Gestation							2.280328	0.000000	2.286943	0.000000
Birth Weight							1.168585	0.071640	1.233054	0.054573
Male x Model									-3.132217	0.504954
Maternal Ed. x Model									7.239489	0.000396
VDQ	Mod	lel 1	Mod	lel 2	Mod	Iodel 3 Model 4		Mod	lel 5	
BIC	52	201.271496	47	723.637711	47	723.500494	41	100.649566	40	95.222650
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Intercept	104.1753 52	0.000000	93.96528 6	0.000000	95.98510 7	0.00000	25.91846 6	0.320788	23.80413 1	0.356684
Child Age	-0.038547	0.000114	-0.046407	0.000007	-0.045482	0.000010	-0.046321	0.000018	-0.047249	0.000011
COVID-Model	- 18.52041 7	0.000000	18.81946 1	0.000000	18.84788 7	0.000000	17.28608 0	0.000016	76.61423 7	0.000738
Maternal Ed.			2.346477	0.012257	2.267559	0.015649	1.435786	0.123670	0.639073	0.503533
Male					-3.241651	0.138492	-0.972784	0.658666	-0.352928	0.875775
Gestation							2.974616	0.000141	3.023880	0.000096
Birth Weight							1.445501	0.205085	1.491077	0.188468
Male x Model									-0.876683	0.915634

ELC	Model 1		Model 2		Model 3		Model 4		Model 5	
BIC	4!	574.859772	4:	149.364403	4403 414		3562.855010		3	551.746593
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Maternal Ed. x Model									10.73971 4	0.003465

NVDQ	Mod	el 1	Mod	el 2	Mod	lel 3	Mod	lel 4	Model 5	
BIC	52	210.416306	47	711.799610	47	05.756790	40	030.416662	40)18.537463
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue	Estimate	pValue
Intercept	108.7211 97	0.000000	86.78188 7	0.000000	90.97108 4	0.000000	47.58487 1	0.058604	44 25826 8	0.074154
Child Age	0.010480	0.248158	0.003817	0.677479	0.005243	0.566518	-0.000470	0.958481	-0.001571	0.859054
COVID-Model	- 18.31754 3	0.000000	17.46835 5	0.000000	- 17.44950 6	0.000000	16.60589 7	0.000004	82 32681 4	0.000112
Maternal Ed.			4.357814	0.000009	4.183439	0.000016	3.910971	0.000017	2.950514	0.001422
Male					-6.370348	0.004507	-3.660405	0.083655	-2.640348	0.221525
Gestation							3.425030	0.000005	3.447173	0.000003
Birth Weight							1.066360	0.327691	1.193121	0.268472
Male x Model									-4.956133	0.503528
Maternal Ed. x Model									12.32378 7	0.000384

Table 6. To determine the influence of maternal pre and post-natal perceived stress, we replaced the COVID-model term in our general linear model with perceived stress scale (PSS) scores self-reported by the child's mother. Overall, we found that this term was not significant and did not contribute to the model fit, suggesting that while stress may play an important biological role in fetal and infant brain development, it does not appear to be solely explain the reduced cognitive scores presented in this work.

Postnatal Stress	EI	C	VI	DQ	NVDQ		
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	
PSS	0.002642	0.412242	0.003645	0.462384	-0.475995	0.613823	
Male x PSS	0.133976	0.683750	0.206981	0.683043	0.363321	0.406477	
Maternal Ed. x PSS	-0.042414	0.732502	-0.273539	0.155058	0.010763	0.948116	
Prenatal Stress	EI	.C	VI	DO	NVDQ		
Term	Estimate	pValue	Estimate	pValue	Estimate	pValue	
PSS	1.045007	0.157838	1.832849	0.177509	1.712337	0.226585	
Male x PSS	-0.216887	0.555744	-0.277688	0.676534	-1.050890	0.128812	
Maternal Ed. x PSS	-0.095475	0.496830	-0.215181	0.389910	-0.067473	0.798480	

Figure 1. Visual overview of all child study visits used in each set of analysis. (a) All children between 0 and 3 years of age; (b) Children with at least one assessment before and one during the pandemic; and (c) Children under 1 year of age born prior to 2019 or after July 2020.

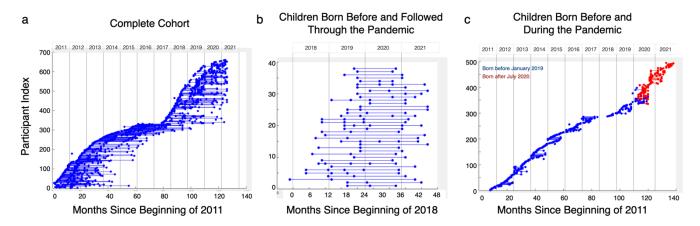


Figure 2. Visual comparison and trends of yearly ELC, VDQ, and NVDQ composite scores of the Mullen Scales of Early Learning. In each panel, the black line and gray bar represent the expect mean (100) and standard deviation (15). Overall, we note a consistent trend of measures from 2011 to 2019, and then a significant decline in 2020 and 2021, corresponding to the COVID-19 pandemic. Significance of these reductions, controlling for important confounders, is presented in Table 3.

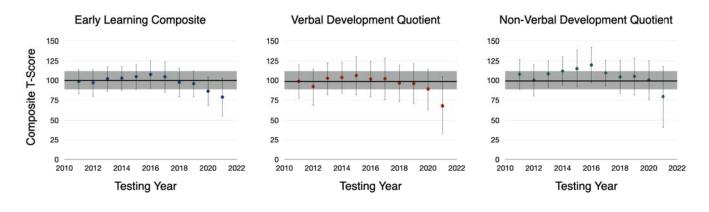


Figure 3. Comparison of yearly gestation duration and child birth weights for children tested in our study. Overall, we find no significant change between pre- or during the pandemic.

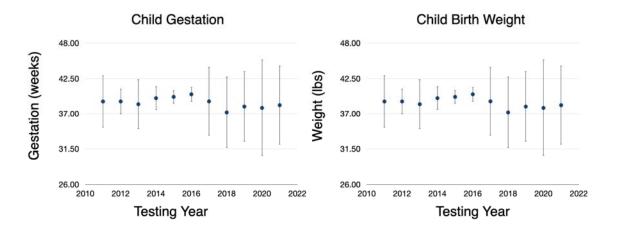


Figure 4. Plots of individual longitudinal ELC, VDQ, and NVDQ measures in the children born before the pandemic (2018) with at least 1 pre- and during pandemic measure. We do not observe any clear increasing or decrease trends through the pandemic.

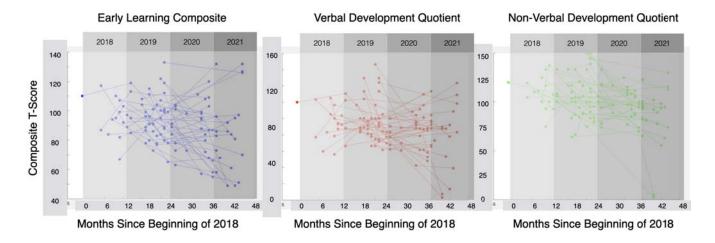
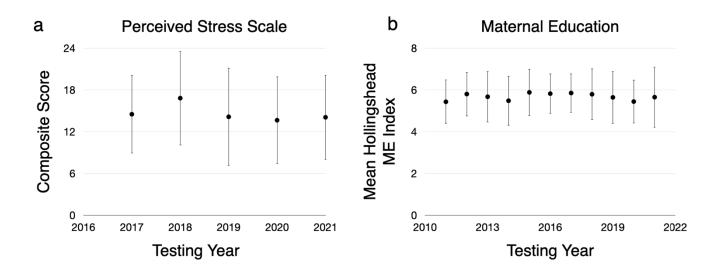


Figure 5. (a) Yearly maternal PSS scores, obtained during pregnancy and following child birth. In contrast to other studies reported at the beginning of the pandemic, we do not observe a significant increase in maternal stress in 2020 or 2021. We also examined maternal education as a marker of socioeconomic status of families tested in each year (b). Like PSS, we found no significant increase or decrease in families tested during the COVID-19 pandemic.



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