

would have lower scores on this measure than children exposed for shorter periods.

Methods

Data were drawn from the Quebec Longitudinal Study of Child Development, whose protocol was approved by the Quebec Institute of Statistics and the Sainte-Justine Hospital Research Centre ethics committees. Participants were recruited via the Quebec Birth Registry using a stratified procedure based on living area and birth rate. The initial sample included $n = 2120$ infants born in Quebec in 1997-1998. Our analysis sample included $n = 1073$ mother-child pairs for whom data was available for MDS at 2 or more time points from 5 months to 5 years, verbal abilities for at least 1 time point from 5 to 10 years, and all covariates. At each data collection, informed written consent was obtained from all participants.

MDS were assessed at 5 months, 1.5, 3.5, and 5 years using a short version (5-12 questions) of the Center for Epidemiologic Studies Depression Scale (CES-D).¹² Responses were standardized to a score between 0 and 10. This version of the CES-D is highly correlated with the original.¹³ The CES-D does not provide a clinical diagnosis of depression, but instead captures MDS. It is a valid and reliable measure of MDS assessing the occurrence and severity of symptoms during the previous week. Responses ranged between 0 (none) to 3 (all the time); all scores were significantly correlated with each other ($r = 0.36-0.44$; $P < .0001$). A threshold of 2.67 (out of 10) was used to approximate the conventional cut-off (16/60) for elevated MDS from the original CES-D.^{12,14,15} We created a single variable for MDS with 4 categories of interest including mothers who (1) never met criteria for elevated MDS ("no exposure"; 63.8%, $n = 684$), (2) met criteria for "early exposure" (at either 5 months and/or 1.5 years; 18.1%, $n = 194$), (3) met criteria for "late exposure" (at either 3.5 and/or 5 years; 16.8%, $n = 181$), and (4) mothers who met criteria for chronic exposure (at 5 months, 1.5, 3.5, and 5 years; 1.3%, $n = 14$).

Verbal abilities were assessed at ages 5, 6, and 10 years using the Peabody Picture Vocabulary Test-Revised (PPVT-R)¹⁶; the child was presented with pictures and had to identify the picture that matched the word read out by the interviewer. Test scores were age-corrected and used in linear regression models. Preliminary analyses showed that the pattern of associations between the exposure and outcome was largely similar across ages. Longitudinal research shows that development of verbal abilities in young children remains relatively stable over time.^{17,18} PPVT-R scores at ages 5, 6, and 10 years were strongly correlated with each other ($r = 0.54-0.64$; $P < .0001$); therefore, outcomes between 5 and 10 years were combined into a single mean score ($n = 480$ had a PPVT-R score at all 3 ages, $n = 380$ had at least 2 and $n = 213$ had at least 1 score). Tests were administered in either French¹⁹ or English; therefore, we adjusted for the language of test administration in analyses. To facilitate interpretation, our global verbal abilities score was converted to a standardized score (mean = 100, SD = 15).

Maternal education, verbal IQ, maternal language spoken at home (dichotomized as English and/or French and/or

another language or neither French nor English), age at birth of target child (dichotomized as ≤ 21 years or < 21 years),²⁰ and birth order of target child were assessed at baseline and considered as potential controls.^{21,22} To isolate the role of MDS in children's verbal abilities, we adjusted for 2 main types of maternal psychopathology symptoms that may co-occur with MDS. General, trait-like maternal anxiety was assessed when the target child was 4.5 years using validated items inspired by *Diagnostic and Statistical Manual of Mental Disorders, 4th Edition* criteria.²³ Antisocial behavior in adolescence was assessed by asking mothers whether they had exhibited 5 different conduct problems.²⁴ The scale ranged from 0 to 5. Mother-child interactions at baseline were assessed by an observer using the Home Observation Measurement of the Environment.²⁵ Maternal parenting practices and family functioning at baseline were self-reported using the Parental Cognitions and Conduct toward the Infant Scale²⁶ and the family dysfunction scale,²⁷ respectively. Socioeconomic status (SES) of the family at baseline was derived from 5 variables including maternal education (years of schooling), spouse's education and occupational status, maternal occupational status, and household income. The final SES composite was standardized for all families. Further information on the questionnaires and methods of data collection can be found online at <http://www.jesuisjeserai.stat.gouv.qc.ca>.

Statistical Analyses

Data analyses included 3 steps. First, we selected control variables on the basis of previous literature indicating an association between a given variable and/or MDS and children's verbal abilities, and epidemiologic guidelines for modeling longitudinal data, whereby potential confounders are selected at baseline and not at subsequent time points^{28,29} and, bivariate association at $P < .05$ between a control variable and either MDS or verbal abilities. Next, linear regression models were used to examine the association between the timing and chronicity of MDS and children's verbal abilities. Finally, to adjust for attrition, we identified variables which differed significantly between the initial and analysis samples and created inverse probability weights based on these variables.

In post hoc regression analyses, we tested the association between the number of times a mother had elevated MDS and children's verbal abilities using the same covariates. We created a categorical variable with 5 categories of mothers who (1) never had elevated MDS (63.8%, $n = 684$), (2) had elevated MDS once (21.4%, $n = 230$), (3) twice (9.2%, $n = 99$), (4) 3 times (4.3%, $n = 46$), and (5) 4 times (1.3%, $n = 14$). All statistical analyses were conducted in SAS v 9.4 (SAS Institute, Cary, North Carolina).

Results

Our analysis sample ($n = 1073$) significantly differed from our initial cohort sample ($n = 2120$). Mothers included in our analysis sample were less likely to have male children (50% in analysis sample vs 57% in initial sample, $\chi^2 = 9.31$, $P = .002$), be unemployed (45% vs 55%, $\chi^2 = 22.39$, $P < .0001$), have a high

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