Attention-Deficit/Hyperactivity Disorder in Young Children: Predictors of Diagnostic Stability

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KEY WORDS
attention-deficit/hyperactivity disorder, cohort, diagnostic stability, income-to-needs ratio, predictors, preschool, socioeconomic status

ABBREVIATIONS
ADHD—attention-deficit/hyperactivity disorder
ASD—autism spectrum disorder
DISC-IV—Diagnostic Interview Schedule for Children—Version IV
OR—odds ratio
PGA—parent global assessment
SES—socioeconomic status

Dr Law conceptualized and designed the study, coordinated and supervised data collection, conducted the initial analyses, drafted the initial manuscript, and reviewed and revised the manuscript; Dr Sideridis critically reviewed the initial analyses, conducted all remaining analyses, and reviewed and revised the manuscript; Dr Albers Prock supervised data collection and reviewed and revised the manuscript; Dr Sheridan conceptualized and designed the study with Dr Law, supervised data collection, reviewed all analyses completed by Drs Law and Sideridis, and critically reviewed and revised the manuscript; and all authors approved the final manuscript as submitted.

WHAT’S KNOWN ON THIS SUBJECT: Approximately 50% of children diagnosed with attention-deficit/hyperactivity disorder (ADHD) at <7 years of age in the community do not meet criteria for ADHD over time. There is a need to examine predictors of diagnostic stability in young children with ADHD.

WHAT THIS STUDY ADDS: Predictors of diagnostic stability from early to middle childhood include child’s baseline externalizing and internalizing symptoms, parental history of psychopathology, and socioeconomic status. These predictors may guide treatment planning at the time of ADHD diagnosis.

OBJECTIVES: The goals of this study were (1) to provide estimates of diagnostic stability for a sample of young children diagnosed with attention-deficit/hyperactivity disorder (ADHD) after undergoing comprehensive multidisciplinary assessments and (2) to identify baseline child and family characteristics that predict diagnostic stability over time.

METHODS: Children aged 3 to 6 years, 11 months consecutively diagnosed with ADHD after multidisciplinary consultations at a tertiary care clinic between 2003 and 2008 were recontacted in 2012 and 2013 (N = 120). At follow-up, the primary outcome was the proportion of children who continued to meet diagnostic criteria for ADHD. To identify predictors of diagnostic stability, logistic regression models were used. In addition, a latent class model was used to independently classify subjects into distinct clusters.

RESULTS: In this cohort, 70.4% of the children contacted at follow-up continued to meet diagnostic criteria for ADHD. Predictors of diagnostic stability included externalizing and internalizing symptoms at baseline, parental history of psychopathology, and family socioeconomic status. The latent class model independently identified 3 distinct profiles: (1) children who no longer met ADHD criteria; (2) children with persistent ADHD and high parental psychopathology; and (3) children with persistent ADHD and low family socioeconomic status.

CONCLUSIONS: Young children who underwent comprehensive developmental and psychological assessments before receiving an ADHD diagnosis, had higher rates of diagnostic stability than in previous studies of community samples. Child and family factors that predict diagnostic stability have the potential to guide treatment planning for children diagnosed with ADHD before 7 years of age. Pediatrics 2014;133:659–667
Predictors of ADHD stability among children diagnosed during preschool or kindergarten are poorly understood. In the studies available, children with persistent ADHD have higher psychiatric comorbidities, family history of psychopathology, and exposure to childhood adversity than children who no longer meet criteria for ADHD at follow-up.14,15 However, in these studies, initial ADHD diagnoses have been assessed by using parent report of symptoms without in-depth evaluation of medical or developmental diagnoses that may mimic ADHD symptoms. To the best of our knowledge, the present study is the first study to identify predictors of diagnostic stability in children aged <7 years who received an ADHD diagnosis after medical, developmental, and psychological assessments.

METHODS
Subjects
Participants were 120 children ages 3 to 6 years, 11 months consecutively diagnosed with ADHD at the Developmental Medicine Center of Boston Children’s Hospital between January 1, 2003, and December 31, 2008. These children were prospectively recontacted at a mean interval of 7 years (range: 5.6–9.5 years) after their initial diagnosis. Children who were adopted or had autism spectrum disorder (ASD), genetic syndromes, or brain injuries at the time of initial ADHD diagnosis were excluded. The Boston Children’s Hospital institutional review board approved all study procedures.

Identification of ADHD in Young Children at Time 1
At time 1, each child completed a multidisciplinary assessment in which developmental–behavioral pediatricians obtained a detailed medical and social history, and completed a physical examination on each child; pediatric psychologists assessed each child’s developmental skills by using cognitive, language, and early academic standardized tests, reviewed day care and school intake questionnaires, and obtained ratings of child behavior by using the Behavior Assessment System for Children.31 The team of clinicians met subsequent to the assessment to review the child’s diagnostic criteria for ADHD and to discuss alternative diagnoses and potential comorbid conditions; the clinicians then systematically recorded all findings in a formal report.

Determination of Psychiatric Disorders at Time 2
After families were recruited for the follow-up time 2 study, they were scheduled for a visit in which parents completed the Diagnostic Interview Schedule for Children–Version IV (DISC-IV), Vanderbilt ADHD Diagnostic Parent Rating Scales, the parent global assessment (PGA), and the Children with Special Health Care Needs CSHCN Screener.32–40 The PGA is a 7-point scale ranging from 1 (no impairments) to 7 (needs 24-hour supervision due to severe impairments). For children who were being treated with stimulant medications, parents were asked to complete rating scales describing their child’s behavior on and off medication (e.g., before the medication took effect in the morning). Parents also completed the Conflict Tactic Scales, Revised, which provides information about psychological and physical aggression in the family,41 and the Social Communication Questionnaire–Lifetime Form,42 which screens for difficulties in social and communication functioning. Parents were also asked about duration and intensity of behavioral therapy, school services, and medication use since time 1. Lastly, the Family Interview for Genetic Studies43 and demographic information not uniformly available from records (including time 1 parental education and income) were obtained. Teachers were also asked to complete the
ADHD Case Definition at Time 2
Children who did not meet DSM-IV-TR criteria on both the DISC-IV and the Vanderbilt ADHD Diagnostic Parent Rating Scale had a PGA score <3 (i.e., minimal impairments), and <6 symptoms on the Vanderbilt ADHD Diagnostic Teacher Rating Scale were classified as no longer meeting criteria for ADHD at time 2. Nine children received an intermediate diagnosis on the DISC-IV, which indicated that symptoms and impairments were present but did not meet full diagnostic criteria for ADHD. Review of these 9 children showed that all of them had at least 4 symptoms in either the inattentive and/or hyperactive/impulsive criteria on the DISC-IV and the Vanderbilt ADHD Diagnostic Parent Rating Scale, impairments in multiple settings on the DISC-IV, a PGA score >3 (i.e., moderate to severe impairments), and >6 symptoms on the Vanderbilt ADHD Diagnostic Teacher Rating Scale. For the purpose of this research study, we classified these intermediate cases as ADHD at time 2.

Covariates of Interest
All covariates were child and family characteristics at time 1 when each child initially received his or her ADHD diagnosis.

Child-level covariates included: (1) age at initial diagnosis; (2) gender; (3) presence of comorbid developmental, behavioral, or mental health diagnoses; (4) race/ethnicity; (5) composite nonverbal cognitive score on one of the following measures: the Wechsler Intelligence Scale for Children, the Wechsler Preschool and Primary Scale of Intelligence, the Differential Ability Scale, and the Stanford-Binet Intelligence Scales; (6) severity of externalizing symptoms (i.e., attention problems, hyperactivity, aggression); and (7) severity of internalizing symptoms (i.e., anxiety, depression, somatization). The severity of externalizing and internalizing symptoms was taken from the Behavior Assessment System for Children and stratified into a T score ≥70 and <70.

Family-level covariates included: (1) parental history of psychopathology; (2) parental education level; (3) income-to-needs ratio (a proxy for family economic resources); and (4) percentage of residents living below the poverty level based on the child’s address. Parental education level was a categorical variable divided into high school or less, some college, and at least a college degree. Income-to-needs ratio was calculated by dividing the subject’s household income by the US Census Bureau poverty threshold income for a family of that size.

Data Analysis
Statistical analyses were conducted by using SPSS version 21.0 (IBM SPSS Statistics, IBM Corporation, Armonk, NY [2012]) and Latent GOLD 4.5 (Statistical Innovations Inc, Belmont, MA [2008]). To compare means of continuous variables, t tests were used; χ² statistics were used to compare proportions of categorical variables.

Logistic Regression
A multivariate logistic regression model was implemented to predict the stability of ADHD diagnosis from a combination of child-level and family-level factors. These factors were chosen on the basis of information from previous literature and were initially included in separate logistic regression models. A best synthesis model followed and included predictors with coefficients significant at P < .10.

Latent Class Model
Data were analyzed by means of latent class mixture modeling in an effort to identify the presence of subgroups that shared similar levels across a combination of independent variables. The latent class models were developed initially for dichotomous variables and were extended to include nominal and, subsequently, continuous indicators. Because the likelihood ratio χ² statistic had been shown to be sensitive to sparse data, bootstrapping P values by using 500 replicated data sets was used to test the improvement in fit between 2 models in this study. The magnitude of R² values, correct classifications based on group membership (stable versus unstable diagnosis), significance of independent variables in defining cluster group membership, and a significant reduction in the likelihood ratio test (L²) were all considered when comparing nested models.

RESULTS
Study Participants
At time 1, a total of 330 children aged <7 years received an ADHD diagnosis in the Developmental Medicine Center of Boston Children’s Hospital. Of these, 120 met inclusion criteria and were seen by the multidisciplinary teams described earlier (Fig 1). Longitudinal follow-up was possible for 88 children (73.3% of the cohort). Table 1 presents baseline characteristics of the cohort. Characteristics of children who were and were not followed up at time 2 were compared: there was no significant difference between the 2 groups in age of initial ADHD diagnosis, current age, gender, nonverbal cognitive score, and per cent living below the poverty level.

Behavioral, Developmental, and Psychiatric Profiles of the Cohort
The cohort as a whole was found to have high rates of comorbid developmental disorders after multidisciplinary assessment as well as high rates of comorbid psychiatric disorders at follow-up (Table 2). After a mean interval of 7 years, 62 children (70.4% of...
the cohort) continued to meet diagnostic criteria for ADHD.

**Prediction of Stable ADHD Diagnosis by Using Logistic Regression Analysis**

Within the child-level model, significant predictors of ADHD stability included elevated externalizing symptoms ($B = 2.21, P < .01; \text{OR}: 9.0$), elevated internalizing symptoms ($B = 1.53, P < .05; \text{OR}: 4.6$), and lower nonverbal composite score ($B = -0.05, P < .05; \text{OR}: 0.94$) (Table 3). Family-level predictors were positive parental history of psychopathology ($B = 2.04, P < .05; \text{OR}: 7.6$) and lower income-to-needs ratio ($B = -0.57, P < .001; \text{OR}: 0.56$). In the combined model, the significant predictors discussed earlier remained significant except for the child’s nonverbal cognitive score. All results were in the same direction and were more pronounced.

**Exploring the Presence of Subgroups Based on ADHD Diagnostic Stability By Using Latent Class Modeling**

A set of ancillary analyses was conducted to evaluate whether specific clusters of individuals and profiles would explain the stability of ADHD
diagnosis (Table 4). A 1-class, baseline model was used as a reference point, and this model was compared with a 2-class model. The 2-class model provided a good fit to the data and suggested that the group of children with stable ADHD diagnosis had baseline characteristics different from the group of children with unstable ADHD diagnosis ($R^2 = 0.256$). Although the 2-class model provided a good fit to the data, examination of the bivariate residuals suggested that some were above the expected value of 1.0, and the probability of misfit was equal to 3.4%. Thus, on the basis of findings from previous research, the model was fit with the addition of a 3-class model, in an attempt to minimize those residual covariations and to improve model fit. The difference between a 2-class and a 3-class model was evaluated by using the log-likelihood –2LL statistic based on the bootstrap distribution and 500 replications (Table 5), suggesting that a 3-class model provided significantly improved fit ($-2\text{LL}_{\text{Diff}} = 165.76, P < .001$).

In the 3-class model, there were 2 classes of children that belonged to the stable ADHD group and 1 class of children that fit into the unstable ADHD group (Fig 2). Only 8.14% were considered misclassified cases. Class 1 represented children with a 99.5% probability of having a stable ADHD diagnosis. These were children with elevated internalizing and externalizing symptoms, had parents with a history of psychopathology and low educational level, and also came from families with low family economic resources. Class 2 described a cluster of children with an 82% probability of having a stable ADHD diagnosis. The major difference, however, was that children in class 2 tended to have parents with more psychopathology but a higher probability of attending college and higher income-to-needs ratio compared with class 1 children. Last, class 3 described children who no longer met criteria for ADHD. These children were characterized by relatively low levels of internalizing and externalizing symptoms during early childhood, came from families with higher economic resources, and had parents who were more likely to attend college.

Profiles of Children With Remitted ADHD

Of the 26 children who did not meet diagnostic criteria for ADHD at follow-up,
new diagnoses emerged in 15 children. Specifically, 2 children met criteria for anxiety disorders, 10 had new diagnoses of ASD, and another 3 had new learning disorders. The remaining children did not meet diagnostic criteria for any developmental, learning, or behavioral disorder, including ADHD, and did not shift from one externalizing disorder to another, which suggests remittance of ADHD symptoms at time 2. For children who were reported to have new ASD by parents, psychological reports were available for the majority of these children. Their Social Communication Questionnaire total scores (median: 17.40 points) were significantly higher compared with those without a new ASD diagnosis (median: 5.93 points; \( P < .001 \)).

**DISCUSSION**

Several important findings emerged from the present cohort study. First, ADHD stability over a 7-year period was 70% when young children underwent comprehensive developmental and psychological evaluations. Although diagnostic stability in this cohort was higher overall than previous studies of community samples\(^9\)–\(^16\), this study found that diagnosing ADHD in young children is not an easy task, even for professionals with expertise in the pediatric ADHD population. More than one-half of children who no longer met criteria for ADHD were later found to have alternative developmental diagnoses, including anxiety, ASD, and learning disorders. However, there was a group of children who no longer met criteria for ADHD and did not have any new onset of psychiatric disorders, suggesting ADHD remittance over time. Clinicians caring for young children with ADHD symptoms, therefore, should be equipped to monitor learning, behavioral, and mood difficulties over time. Importantly, children with an early ADHD diagnosis should have continuity of care with clinicians who have expertise in

**TABLE 4** Percentage of Sample Belonging to Each Class and Significance of Predictor

<table>
<thead>
<tr>
<th>Fixed Effect</th>
<th>Class 1 Stable Diagnosis</th>
<th>Class 2 Stable Diagnosis</th>
<th>Class 3 Unstable Diagnosis</th>
<th>Z Test</th>
<th>( P )</th>
<th>( R^2 )</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cluster size, ( n,(%) )</td>
<td>33 (38.4)</td>
<td>29 (33.7)</td>
<td>2 (27.9)</td>
<td>3.009</td>
<td>.0047**</td>
<td>65.8%</td>
</tr>
<tr>
<td>Stability of ADHD diagnosis Stable</td>
<td>99.5%</td>
<td>82.06%</td>
<td>8.71%</td>
<td>2.223</td>
<td>.08</td>
<td>8.92%</td>
</tr>
<tr>
<td>Parental history of psychopathology Present</td>
<td>91.08%</td>
<td>93.86%</td>
<td>69.86%</td>
<td>2.360</td>
<td>.0096**</td>
<td>14.19%</td>
</tr>
<tr>
<td>Externalizing symptoms T score &gt;70 in ( \geq1 ) subscale</td>
<td>53.63%</td>
<td>42.82%</td>
<td>27.34%</td>
<td>2.108</td>
<td>.11</td>
<td>9.30%</td>
</tr>
<tr>
<td>Internalizing symptoms T score &gt;70 in ( \geq1 ) subscale</td>
<td>22.52%</td>
<td>23.82%</td>
<td>5.63%</td>
<td>1.894</td>
<td>.011*</td>
<td>60.98%</td>
</tr>
<tr>
<td>Parents going to college Yes</td>
<td>4.65%</td>
<td>88.94%</td>
<td>78.39%</td>
<td>4.810</td>
<td>&lt;.001***</td>
<td>55.83%</td>
</tr>
<tr>
<td>Income-to-needs ratio Income greater than expected for a family of that size</td>
<td>6.44%</td>
<td>20.89%</td>
<td>41.92%</td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

\( * P < .05, ** P < .01, *** P < .001. \)

\( a \) These values indicate amount of variance of each variable explained by using the 3-class solution.

**TABLE 5** Nested Latent Class Models Suggesting the Superiority of a 3-Class Solution Over Competing 1- and 2-Class Models for Explaining Stability in ADHD Diagnosis

<table>
<thead>
<tr>
<th>Model Tested</th>
<th>BICLL(^a)</th>
<th>LL</th>
<th>Class Error(^b), ( P )</th>
</tr>
</thead>
<tbody>
<tr>
<td>1-class independence model</td>
<td>1278.330</td>
<td>−610.212</td>
<td>&lt;.0001***</td>
</tr>
<tr>
<td>2-class</td>
<td>1199.742</td>
<td>−553.100</td>
<td>.039*</td>
</tr>
<tr>
<td>3-class</td>
<td>1183.840</td>
<td>−527.332</td>
<td>.732</td>
</tr>
</tbody>
</table>

\( a \) BICLL, Bayesian Information Criterion based on log-likelihood.

\( b \) A nonsignificant model indicates no difference between the hypothetical and obtained model.

**FIGURE 2**

Latent class model displaying the presence of 2 stable diagnosis classes and 1 unstable diagnosis class.
ADHD or should receive a comprehensive reevaluation during school age. Although it is not surprising that elevated externalizing symptoms are associated with stability in ADHD diagnosis, we additionally observed that children with a greater risk of ADHD persistence had higher internalizing symptoms at a young age. This finding is consistent with previous literature on older children with ADHD,52–54 and it points to the importance of screening for internalizing symptoms at initial diagnostic visits, even when parental concerns are about disruptions due to externalizing symptoms.

Finally, we observed that family factors, including parental history of psychopathology and family economic resources, are predictive of ADHD diagnostic stability. The idea that family contexts contribute to the maintenance of preschool-aged problem behaviors and long-term impairments is not new. Previous research studies have shown that variables of maternal psychopathology, family climate, and SES are predictive of ADHD diagnostic stability. The idea that family contexts contribute to the maintenance of preschool-aged problem behaviors and long-term impairments is not new. Previous research studies have shown that variables of maternal psychopathology, family climate, and SES are predictive of ADHD diagnostic stability.52–54 Given the convergence of these findings in these previous samples and our own research, it seems that factors related to the family should be explored more explicitly in clinical settings. Furthermore, these findings indicate that parent-training programs may have a potentially important role in ADHD treatment in early childhood and may enhance child-specific treatments in this young age range. Evidence-based parent management training has been shown to help parents actively acquire parenting skills that target relationship building and improve parent–child interactions, as well as lower children’s externalizing behaviors.55,56 The American Academy of Pediatrics therefore recommends that primary care physicians prescribe such evidence-based therapy as a first line of treatment for preschool-aged children.57

Interestingly, parental education emerged in the latent class model as the most discriminating variable in the prediction of ADHD stability ($R^2 = 60.98\%$). One possible explanation is that parents with different levels of education provide a different day-to-day environment for children and that these environmental factors mediate changes in the prefrontal cortex, decreasing ADHD symptoms over time. This speculation is bolstered by recent data pointing to the impact of early environments, particularly those linking the effects of SES on prefrontal cortex development and neurocognitive measures associated with ADHD symptoms.23,24,58 Potential mediators of the relationships between low parental education and high ADHD persistence include differences in the quality of the home environment (eg, use of complex language, enrichment activities), quality of parent–child interactions, and degree of family conflicts. The influences of these variables on diagnostic stability in ADHD are beyond the scope of the present study but are of critical importance to explore in future studies.

Several limitations should be noted when interpreting our findings. One limitation is that this cohort consisted of children referred to a single tertiary care clinic. However, these children were not a convenience sample or recruited from advertisement. Our cohort consisted of every child aged <7 years diagnosed with ADHD in our specialty clinic in 2003 to 2008; almost all of them were referred to this clinic because pediatricians would not otherwise give a diagnosis of ADHD or start treatment at such a young age. Another limitation is that we were not able to recontact every child diagnosed at time 1. However, a comparison of demographic data indicated that subjects lost to follow-up did not differ from subjects with full participation. Lastly, families in this cohort had a higher mean income compared with the mean income of the US population, even when family size was considered. Use of a cohort with higher SES may have biased the study toward less severe cases and, consequently, may have produced more conservative results.

CONCLUSIONS

Although ADHD symptoms may improve in a subset of children, for many, ADHD symptoms are chronic and associated with many psychiatric comorbid conditions. Those with higher risk for long-term ADHD have more externalizing symptoms and internalizing symptoms at baseline, more parental psychopathology, and fewer family economic resources. This study adds to the growing literature demonstrating that: (1) child-specific therapies may not suffice as the only armamentarium for practitioners treating children with ADHD; and (2) the preschool years may serve as an important, sensitive period for intervention.

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