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Association between small for gestational age and motor coordination difficulties in children aged 5-6 years: Insights from the Hokkaido Study on Environment and Children's Health

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(ii) Structured abstract (up to approximately 250 words) and 3-9 keywords;

Abstract

Background: Developmental coordination disorder (DCD) presents motor skill delays in early childhood and has been associated with later maladaptation, necessitating early intervention. However, research on the potential risk factors, particularly in preschool-aged children, remains scarce.

Aims: We aimed to explore the association between small for gestational age (SGA) and other factors and motor coordination problems in 5–6-year-olds from the Hokkaido Study on Environment and Children’s Health Cohort.

Study Design: Prospective

Subjects: We analyzed data from >3,500 participants from the Hokkaido study, a prospective birth cohort, and assessed children aged 5–6 years.

Outcome measures: Participants underwent assessment with the Developmental Coordination Disorder Questionnaire Japanese version (DCDQ-J). We conducted linear regression analyses adjusted for variables such as the child’s sex, maternal age, and maternal smoking history during pregnancy, while also examining the independent associations of each risk factor.

Results: Among the 3,883 children analyzed for SGA, children with SGA exhibited significantly lower DCDQ-J total scores than non-SGA children (mean difference: -2.25, 95% confidence

interval [-4.19, -0.30], $p=0.02$). On the subscales, children with SGA demonstrated significantly lower “Control During Movement” scores than non-SGA children (mean difference: -0.96, 95% confidence interval [-1.78, -0.13], $p=0.02$). Furthermore, the child’s sex, maternal smoking, maternal age, and preterm birth were independently associated with DCD.

Conclusions: SGA was shown to be one of the risk factors for the manifestation of motor coordination difficulties in 5–6-years old children. In combination with other factors, screening for motor coordination difficulties in SGA children will be an important means of initiating appropriate interventions.

Keywords:

Small for gestational age, Developmental coordination disorder, Motor coordination difficulties, Preschool age, Cohort study, Japan

(iii) Text and acknowledgments;

Introduction

Developmental coordination disorder (DCD) requires early detection and intervention; therefore, the establishment of the potential risk factors associated with DCD is crucial[1-3]. DCD is classified within the realm of neurodevelopmental disorders and is diagnosed based on the manifestation of difficulties such as delays in acquiring motor skills and clumsiness [4, 5]. The lack of motor skills hinders age-appropriate daily activities, has long-term effects on productivity in academic performance and employment, and influences play and social activities [5, 6]. Moreover, DCD has been shown to be associated with both externalizing and internalizing problem behaviors [7, 8], and it has been suggested that these associations may be secondary impairments mediated by difficulties in participating in school activities and peer relationships [9]. Despite such a significant need for understanding and support, there is a lack of awareness of DCD within educational institutions [10]. Many parents of children with DCD have serious concerns about their children's current situations and future prospects [11].

According to the Diagnostic and Statistical Manual of Mental Disorders Text Revision published by the American Psychiatric Association (DSM-5TR), the reported prevalence of DCD in children aged 5–11 years is 5–8%. Additionally, prevalence rates of 7–8% have been reported in Canada, Sweden, and Taiwan. However, a 2022 survey in Japan [12], which randomly selected

2000 childcare facilities nationwide, revealed that only 0.2% of children were diagnosed with DCD, while 4.8% showed signs of motor coordination difficulties without diagnosis. However, it has also been reported that there is no significant difference between the worldwide prevalence of attention-deficit hyperactivity disorder, known for its high comorbidity with DCD [13], and its prevalence in Japan [14]. These findings indicate the potential under-recognition of DCD in Japan.

In children with DCD, 50–70% continue to experience coordination difficulties into adolescence[4]. Due to significant individual differences in early childhood motor skill acquisition, diagnosing DCD before the age of 5 years is atypical [4, 15]. However, considering the potential benefits of early intervention for improving developmental trajectories and preventing secondary consequences in at-risk children, diagnosing DCD or at least identifying motor coordination difficulties before a child enters elementary school is encouraged [2, 3].

Extensive research has been conducted on the risk factors associated with the development of DCD in children. Low birth weight and preterm birth have been identified as significant early risk factors for the development of DCD[1, 4, 16, 17]. Recently, small for gestational age (SGA), which has some overlap with low birth weight and preterm birth, has attracted attention. SGA infants can be identified at birth, and this early detection can have a positive impact on later neurodevelopment[18-22]. Faebo Larsen et al. investigated early risk factors for DCD in a study

1 involving 33,354 mothers and their children from the Danish National Birth Cohort. In this study,
2 SGA was defined as birth weight below -2 standard deviations (SD) relative to gestational age^{[23,}
3 ^{24]}. The results indicated that male sex, preterm birth, and SGA were strong predictors of DCD at
4 the age of 7 years[23]. In a cohort study conducted in three regions of Italy, risk factors for DCD
5 were investigated in 803 children aged 8–11 years, revealing that SGA is a risk factor for DCD[25].
6 In this study, SGA was defined as a birth weight below the 10th percentile for gestational age.
7 Both studies suggested that SGA is an independent risk factor for later DCD, distinct from low
8 birth weight and preterm birth. However, neither study reported findings concerning the risk of
9 DCD in preschool-aged children born as SGA. In Japan, Suyama et al. conducted a study on birth-
10 related risk factors for DCD and reported that a child's sex and maternal smoking during
11 pregnancy are associated with a higher likelihood of DCD in preschool-aged children [26].
12 However, the association between SGA and DCD has not been reported in Japan to date.

13 Head circumference (HC) serves as a metric for assessing central nervous system
14 development and identifying neonates vulnerable to neurodevelopmental issues [27]. Due to
15 the link between DCD and neurological development, a small HC at birth is deemed a
16 potential risk factor. Prior research has indicated that infants with a small HC may experience
17 negative impacts on later neurodevelopment, affecting motor skills and cognitive abilities
18 [28] up to age 4.5 years and intellectual capacities [29] up to age 8. However, evidence

regarding coordination difficulties in preschool-aged SGA children or those with a small HC is lacking.

Therefore, this study aimed to investigate the relationship between SGA and small HC and motor coordination difficulties in preschool-aged children (aged 5–6 years) in a Japanese birth cohort. One of the significant aspects of this study is that it was conducted using large-scale cohort data from Japan.

Materials and methods

Study participants

The Hokkaido Study on Environment and Children's Health is an ongoing prospective cohort study that was initiated in 2002. The details of this study have been described previously [30] [31]. The study population comprised women in their first trimester of pregnancy who visited one of the 37 affiliated hospitals or clinics in the study area (Hokkaido, Japan) and were enrolled between February 2003 and March 2012. Demographic characteristics, medical history, and obstetric history were obtained from the mothers at enrollment (first trimester of pregnancy) using a baseline questionnaire. Information on smoking during pregnancy was obtained using an additional questionnaire after childbirth. The initial cohort comprised 20,926 pregnant women, and the total number of eligible children was 20,945. We sent follow-up questionnaires, including the Japanese version of the Developmental Coordination Disorder Questionnaire (DCDQ-J), to

children born between April 2008 and November 2011 during the academic year when they turned 5 years old, and obtained responses from 4,851 participants.

Figure 1 depicts the participant selection process from the Hokkaido Study on Environment and Children's Health, initially involving 20,945 participants. Those lacking essential information to determine SGA and small HC (birth weight, height, HC, gestational age, and sex of the infant) were excluded. Additionally, participants who were born as multiples or had Down's syndrome were excluded. Individuals without parity information were considered primiparous and were included. Furthermore, considering that the target age for DCDQ-J is ≥ 5 years, along with the purpose of focusing on preschool-aged children, individuals who were aged less than 60–85 months during the DCDQ-J assessment were excluded. Following these criteria, 3,883 and 3,848 children were included in the SGA and small HC analyses, respectively.

Definition of SGA and small HC

In this study, SGA was defined as having a birth weight and height below the 10th percentile relative to parity, infant sex, and gestational age[32]. The definition of SGA varies, and no single consensus exists[33]. Criteria such as birth weight less than the 10th percentile [25], below the 5th percentile[18], or below -2SD of weight [18, 23] are utilized. Some criteria consider both birth weight and height less than the 10th percentile [34] or less than -2SDs for both weight and height [35]. In this study, we refined the definition of SGA from a previous study in this cohort [36],

1 which was based solely on gestational age, sex, and weight. By integrating birth height into our
2 criteria [37], we aimed to identify factors of growth restriction with greater precision and capture
3 more severe cases through a more stringent definition [33]. The Hokkaido Study was equipped to
4 incorporate and validate definitions of SGA that included birth height, owing to its access to
5 professionally measured birth height data from medical institutions. This was facilitated using a
6 database furnished by the Japanese Paediatric Society[38]. Small HC was defined as a birth HC
7 less than the 10th percentile of the reference according to gestational age and infant sex, following
8 previous research guidelines [28].

9 ***Measurement tool: the Japanese version of the DCDQ-J***

10 To assess motor coordination in preschool-aged children, we used the DCDQ-J[39]. The
11 Japanese version was adapted for the Japanese population using the DCDQ 2007 and the
12 recommended international guidelines[40] for measuring children's motor coordination
13 impairments. The questionnaire comprises 15 items, and parents evaluate their child using the
14 following 5-point scale: (1) "Not at all like your child;" (2) "A bit like your child;" (3)
15 "Moderately like your child;" (4) "Quite a bit like your child;" and (5) "Extremely like your child."
16 The total score ranges from 15 to 75 points, with lower scores indicating potential issues with
17 motor coordination. While standardized cut-off scores for the total score are established in the
18 DCDQ 2007[41], they have not been set for the DCDQ-J. Therefore, we conducted analyses using

continuous variables. Additionally, as the interpretive guideline, we used the cut-off in the DCDQ 2007, which is for children aged 5–7 years; a total score of 46 or lower suggests or raises suspicion of DCD[41]. The DCDQ-J comprises three subscales. The first, "Control During Movement," includes six items pertaining to motor control, for example, "Your child throws a ball with control and accuracy." The "Fine Motor/Handwriting" subscale encompasses four items that gauge fine motor skills, illustrated by "Your child's writing or drawing keeps pace with peers." The "General Coordination" subscale comprises five items assessing overall movement coordination, such as "Your child enjoys and engages in sports or active games that demand proficient motor skills." To comprehensively assess the link between SGA and DCD, we analyzed each subscale in conjunction with the aggregate DCDQ-J score.

Covariates

Covariates in our statistical analysis included, based on prior research, the child's sex [4, 42], maternal smoking during pregnancy [26, 36], and maternal age at childbirth [25, 43]. Previous research from the ongoing Hokkaido Study identified male sex and maternal smoking during pregnancy as significant risk factors in 2020 [26]. Additionally, a 2021 study demonstrated that smoking during pregnancy mediated socioeconomic determinants, contributing to increased SGA incidence[36]. Smoking during pregnancy status is based on postpartum questionnaire responses, categorizing participants as "Smoked during pregnancy" or "Did not smoke during pregnancy".

In this study, the information of gestational age was used to identify SGA and small HC; therefore, gestational age itself was not included as a covariate to avoid multicollinearity. However, as previous research has indicated that preterm birth is a risk factor for DCD, we conducted a univariate analysis on the association between gestational age (≥ 37 weeks or below) and DCD, providing this as supplementary information in Supplemental Table 1. [1, 4, 16, 17].

Statistical analysis

The baseline research participants' characteristics were evaluated using descriptive analysis. As of 2024, given that a standard cut-off value for the Japanese version of the DCDQ[41] remains unestablished and focuses on observing differences in motor coordination characteristics rather than diagnosis in preschool-aged children, we have recorded the results as continuous variables. Consequently, we performed a linear regression analysis on the individual scores for "Control During Movement," "Fine motor/handwriting," and "General Coordination" in the DCDQ-J, in addition to the total score. We set the presence of SGA or small HC as the exposure variable. After conducting the univariate regression analysis, we implemented multivariate regressions adjusted for the child's sex, maternal smoking during pregnancy, and maternal age at childbirth. Referring to previous research and considering multicollinearity, the associations between SGA and DCDQ-J scores, as well as small HC and DCDQ-J scores, were analyzed separately. Statistical analyses were performed using R version 4.3.2 (R Development Core Team, Vienna, Austria;

1 <https://www.R-project.org>), and a *p*-value less than 0.05 indicated statistical significance.

3 **Results**

4 The present study encompassed data on 3,883 participants for SGA in relation to motor
5 coordination issues and on 3,848 participants concerning small HC and its association with motor
6 coordination challenges. Figure 1 illustrates a flowchart of the participants included in this study's
7 analyses. Table 1 shows the characteristics of SGA and non-SGA children, as well as those with
8 small HC and non-small HC. The proportion of SGA children according to weight and height was
9 2.9% of the total sample (N=3,883). The proportion of children with a small HC was 9.1% of the
10 total sample (N=3,848). Infants born SGA and those born with weight and height in the
11 appropriate range for gestational age, as well as those with a small HC and those with an HC in
12 the appropriate range for gestational age, shared a median gestational age of 39 weeks. The
13 median birth weight of SGA infants was 2,400 g, whereas that of non-SGA infants was 3,050 g.
14 Regarding small HC, the median birth weight for infants with small HC was 2,760 g, whereas in
15 infants who were non-small HC, the median birth weight was 3,060 g. Among SGA children,
16 9.7% (N=11) were born at or before 36 weeks of gestation, compared to 3.3% (N=125) of children
17 not born SGA. Similarly, 3.4% (N=12) of children with small HC were born at or before 36 weeks
18 of gestation, compared to 3.5% (N=122) of those without small HC.

1 The association between child's sex and the occurrence of SGA was examined using the Chi-
2 square test. The results indicated that the association between child's sex and SGA was not
3 statistically significant ($p = 0.37$). Similarly, the association between child's sex and the
4 occurrence of small HC was also examined using the Chi-square test. The results showed that the
5 association between child's sex and small HC was statistically significant ($p < 0.01$). These
6 findings suggest that there may be no association between child's sex and SGA, but there may be
7 an association between child's sex and small HC.

8 Table 2 presents the results of the univariate linear regression analysis of changes in DCDQ-
9 J scores based on the presence or absence of SGA or small HC. Univariate linear regression
10 analysis demonstrated that the total DCDQ-J score was 2.25 points lower in SGA children than
11 in non-SGA children ($p=0.02$). Regarding the subscales, "Control During Movement" was 0.96
12 points lower in SGA children than in non-SGA children ($p=0.02$). However, no consistent trend
13 was observed for "Fine Motor/Handwriting" and "General Coordination" scores based on the
14 presence or absence of SGA. Regarding the presence or absence of small HC, no consistent trend
15 was observed in either the total score or in each subscale of the DCDQ-J. For the effect sizes, the
16 R^2 values for SGA and small HC in relation to the total DCDQ-J score and all subscales were
17 below 0.01, indicating that the impact of these variables on DCDQ-J scores is relatively minimal.
18 While SGA and small HC have a statistically significant effect on motor coordination issues, the

1 magnitude of this effect is considered to be limited.

2 The results of the multivariate linear regression analysis are shown in Table 3. Factors
3 considered, apart from SGA and small HC, included the child's sex, maternal smoking during
4 pregnancy, and the mother's age at childbirth. Figure 2A and Figure 2B present the associations
5 between SGA and small HC with DCDQ-J scores, adjusted for the covariates, based on the results
6 of the multivariate linear regression analysis shown in Table 3. Estimated differences are
7 displayed along with their 95% confidence intervals. The results revealed that even when
8 accounting for the aforementioned factors, the total scores on the DCDQ-J were 2.27 points lower
9 in SGA children than in non-SGA children ($p=0.02$). Regarding the subscales of DCDQ-J, the
10 "Control During Movement" scores were 0.9 points lower in SGA children than in non-SGA
11 children ($p=0.03$). Additionally, in SGA children, the score for "Fine Motor" was reduced by 0.67
12 points ($p=0.049$, listed as $p=0.05$ in Table 3) compared to those without SGA. In terms of small
13 HC, children with small HC had total DCDQ-J scores 1.05 points lower than those in children
14 without small HC. However, the 95% confidence interval ranged from -2.19 to 0.09, indicating
15 no statistically significant difference. Additionally, consistent with the findings of the univariate
16 linear regression analysis, the multivariate linear regression analysis revealed no consistent trends
17 in DCDQ-J subscales based on the presence or absence of small HC. The results of the
18 multivariate regression analysis, considering the child's sex, maternal smoking during pregnancy,

1 and maternal age at childbirth, showed that even after adjusting for SGA and other variables, boys
2 had DCDQ-J total scores that were 3.03 points lower ($p<0.01$). Additionally, children whose
3 mothers did not smoke during pregnancy had DCDQ-J total scores that were 1.37 points higher
4 ($p<0.01$), and for each additional year of mother's age at childbirth, DCDQ-J total scores
5 decreased by 0.11 points ($p<0.01$). Similarly, after adjusting for small HC and other variables,
6 boys had DCDQ-J total scores that were 3.09 points lower ($p<0.01$), children whose mothers did
7 not smoke during pregnancy had DCDQ-J total scores that were 1.46 points higher ($p<0.01$), and
8 each additional year of mother's age at childbirth was associated with a 0.11-point decrease in
9 DCDQ-J total scores ($p<0.01$). These results suggest that, in addition to SGA, factors such as sex,
10 maternal smoking during pregnancy, and maternal age significantly influence children's motor
11 coordination.

12 However, considering the effect sizes, the R^2 values for SGA and small HC in relation to the
13 total DCDQ-J score and all subscales were below 0.1, indicating that even for variables showing
14 statistically significant differences, the magnitude of their impact should be considered limited.

15 The t -test results for the DCDQ-J scores based on the presence or absence of SGA and small
16 HC are presented in Supplemental Table 2. Significant differences were observed for total DCDQ-
17 J scores based on the presence or absence of SGA ($p=0.02$). Specifically, we found significant
18 differences in the scores for "Control During Movement," indicating body coordination in motor

activities ($p=0.02$). However, no significant differences were observed in the scores for “Fine Motor/Handwriting” and “General Coordination.” Additionally, no significant differences were observed in the DCDQ-J total score or any subscale score based on the presence of a small HC.

In this study, based on previous research, we used the child’s sex, maternal smoking during pregnancy, and maternal age at childbirth as covariates in the multivariate analysis. The results of the univariate regression analysis examining the independent associations between each variable and DCDQ-J scores are presented in Supplemental Table 1. Additionally, we assessed the independent association between preterm birth, which was excluded from the multivariate analysis due to concerns about multicollinearity with SGA, and DCDQ-J scores. The results are shown in Supplemental Table 1.

First, boys had DCDQ-J total scores that were 2.99 points lower than those of girls ($p<0.01$), fine motor scores that were 2.18 points lower ($p<0.01$), and general coordination scores that were 0.92 points lower ($p<0.01$). Next, children whose mothers smoked during pregnancy had DCDQ-J total scores that were 1.37 points lower than those of children whose mothers did not smoke ($p<0.01$), fine motor scores that were 0.62 points lower ($p<0.01$), and general coordination scores that were 0.44 points lower ($p=0.02$). Regarding the mother’s age, for each 1-year increase in maternal age, the DCDQ-J total score decreased by 0.09 points ($p=0.01$), the fine motor score decreased by 0.04 points ($p<0.01$), and the general coordination score decreased by 0.02 points

($p<0.01$). For preterm birth, when comparing children born at 36 weeks or earlier with those born at 37 weeks or later, the group born at 37 weeks or later had DCDQ-J total scores that were 2.71 points lower ($p<0.01$), control during movement scores that were 1.10 points lower ($p<0.01$), and fine motor scores that were 0.93 points lower ($p<0.01$).

Discussion

This study demonstrated the association between SGA and motor coordination difficulties in Japanese preschool-aged children from a prospective birth cohort using *t*-tests and regression analyses, suggesting that SGA is a risk factor for motor coordination difficulties during early childhood. We observed no statistically significant association between small HC and motor coordination difficulties in preschool-aged children. Therefore, our results suggest the importance of evaluating motor coordination difficulties at an earlier age in children born SGA. Additionally, other risk factors besides SGA were examined, and it was suggested that being male, maternal smoking during pregnancy, and maternal age at birth are associated with DCD in preschool-aged children.

One key finding was that SGA children had significantly lower mean DCDQ-J scores than non-SGA children, even after adjusting for variables such as the child's sex, maternal smoking during pregnancy, and maternal age at childbirth. Children born SGA particularly struggled in the "Control During Movement" subscale, indicating having difficulties in activities requiring body

control, such as walking, running, jumping over obstacles, stacking objects, and throwing. This underscores the importance of early identification and intervention for motor coordination difficulties. Additionally, it was indicated that SGA children might also exhibit difficulties in "fine motor" tasks. This shows that they may struggle with tasks requiring hand movements and finger manipulation. It is considered advantageous to initiate training and support before the commencement of elementary school, where tasks such as writing and personal grooming become more frequent. Task-oriented approaches have the highest level of evidence and are recommended as methods to improve coordinated motor skills in children with DCD [15]. However, the association between SGA and fine motor skills is statistically weak, considering the large sample size of this study; thus, the necessity for intervention should be evaluated cautiously. It is advantageous to assess individual characteristics with subscale metrics such as the DCDQ and to formulate personalized support strategies crucial for mastering skills required in everyday life. Another key observation was that when compared to the cut-off value of 46 points used as a reference for the DCDQ 2007[41, 26], even SGA children scored 6.2 points higher than the cut-off value. This could be attributed to significant individual differences in the acquisition of motor skills among preschool-aged children, making it challenging to distinguish between developmental variations and actual coordination issues. Additionally, Asian individuals frequently exhibit greater dexterity than their European counterparts, who were the original focus

of the questionnaire. This suggests the potential need for a higher cut-off value [44]. Independent of a diagnosis of DCD, it is advisable to conduct regular assessments and observations of motor coordination skills in SGA children to support optimal developmental outcomes.

This study aimed to investigate the association between SGA and small HC, and developmental coordination difficulties in early childhood. Our findings indicate that SGA is associated with DCD even after adjusting for covariates. However, it is important to recognize the presence of other risk factors when interpreting these results. Consistent with previous studies, this study also demonstrated independent associations between factors such as child's sex, maternal smoking during pregnancy, and maternal age at childbirth, and preterm birth with scores indicating coordination difficulties. These results can be interpreted as follows: First, regarding sex, boys were found to have lower DCDQ-J scores than girls, indicating more significant motor coordination problems. This trend has been consistently reported in previous studies, with boys being known to be at higher risk for DCD [4, 25]. Next, regarding maternal smoking during pregnancy, children whose mothers smoked were found to have lower DCDQ-J scores. It is widely recognized that smoking during pregnancy negatively affects fetal development, which may also lead to motor coordination difficulties [26, 36]. Therefore, smoking cessation counseling during pregnancy should be emphasized as a critical intervention to support healthy child development. Furthermore, it was found that DCDQ-J scores slightly decreased as maternal age

1 increased. The impact of advanced maternal age on a child's neurodevelopment has been the
2 subject of various debates; however, higher maternal age may lead to relatively reduced nutrient
3 and oxygen supply to the fetus, potentially affecting motor coordination [25]. Finally, preterm
4 birth was also identified as a risk factor for DCD. Preterm birth is particularly associated with
5 immature neurological development, which is known to influence motor coordination [1] [16]. In
6 addition to SGA, continuous developmental monitoring and early intervention for preterm
7 children are considered especially important. These factors may independently influence DCD,
8 and it is also possible that multiple factors interact to amplify their effects. Therefore, it is essential
9 to consider these factors comprehensively when assessing the risk of DCD.

10 Our findings are consistent with those of several prior studies. Research conducted by Faebo
11 Larsen et al. on school-aged children (aged 7 years) found that being SGA is a risk factor for later
12 DCD, aligning with our findings[23]. Similarly, studies on low birth weight, which frequently
13 overlap with SGA, such as Deshmukh et al.'s study on school-aged children up to 10 years old,
14 support our findings[17]. Scammon's growth curves [45] evidently show that motor neuron
15 development, crucial for motor coordination, reaches approximately 80% and 100% by ages 5
16 and 12, respectively. Therefore, it is considered adequate to conduct screening for coordination
17 difficulties around preschool age. For children born SGA, it will be desirable to conduct regular
18 observations from preschool age (5–6 years) through the end of elementary school (approximately

1 12 years old).

2 This study boasts several strengths and expected external validity. First, the Hokkaido Study
3 stands as Japan's inaugural prospective birth cohort study, boasting a robust sample size and
4 comprehensive collection of children's growth and birth-related data. The cohort's profile has been
5 previously outlined by Kishi et al. [30] [30, 31]. Second, the assessment team comprised child
6 psychologists and experts in the DCDQ, as demonstrated in a prior study by Suyama et al. [26].
7 Furthermore, we utilized the validated Japanese version of the DCDQ 2007, as confirmed by
8 Nakai et al. [39].

9 The delineation of SGA using both height and weight as opposed to weight alone warrants
10 careful consideration across different studies. In one investigation, predictions of height at 18
11 years did not vary, whether based solely on weight or both weight and height [46]. Nonetheless,
12 within neurodevelopmental research, it is acknowledged that such differences in SGA criteria
13 could influence the prognosis of neurodevelopmental outcomes. In this study, we defined SGA
14 using weight, height, gestational age, sex, and birth order (parity), focusing on DCD risk
15 assessment. Additionally, supplementary data on the risk of DCD in children classified as below
16 the 10th percentile for weight are presented in Supplemental Tables 3–6. Supplemental Table 3
17 shows the characteristics of SGA and non-SGA children defined by weight. Supplemental Table
18 4 presents the results of a univariate regression analysis of SGA and DCDQ-J scores in preschool

1 children, while Supplemental Table 5 provides the results of a multivariable linear regression
2 analysis adjusted for covariates. Supplemental Table 6 shows the results of t-tests for SGA and
3 DCDQ-J scores. In all analyses, no significant differences were observed between SGA defined
4 by weight and DCDQ-J scores at ages 5–6. While children classified as SGA showed a tendency
5 toward motor coordination difficulties, motor coordination difficulties, statistical significance
6 was not confirmed. This underscores the importance of vigilant monitoring of
7 neurodevelopmental progress in children small for both height and weight at birth relative to their
8 gestational age. Such insights bear crucial clinical implications for enhancing long-term
9 neurodevelopmental outcomes in these children through early identification and intervention.

10 As a limitation of this study, although a statistically significant association was observed
11 between SGA and motor coordination problems in early childhood, the effect size was small. This
12 suggests that the impact of SGA and other variables examined in this study on motor coordination
13 is limited. Therefore, when considering interventions and support, it is important to fully account
14 for the possibility that other risk factors may interact and contribute to these issues. Further
15 research is needed to comprehensively understand the risk factors for DCD.

16 In conclusion, our research findings are considered to have both clinical and practical
17 implications. Particularly, the importance of early screening for motor coordination issues is
18 emphasized for children born SGA, especially when both height and weight are below the

1 expected levels at birth. This is mainly crucial for boys, as well as for those whose mothers
2 smoked during pregnancy or were of advanced maternal age during childbirth. Future studies
3 should explore whether the associations between SGA, other risk factors, and motor coordination
4 difficulties are consistent across different large-scale populations and cohorts. Additionally,
5 through longitudinal data collection and analysis, a deeper understanding of the link between
6 these risk factors and motor coordination during early childhood can be achieved. This will
7 enhance our comprehension of the neurodevelopmental patterns in children born smaller than
8 expected for their gestational age and contribute to the development of tailored screening and
9 intervention protocols.

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12 study, as well as the hospitals and clinics that cooperated with us. We also extend our
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15
16 (iv) Author contributions;

17 **Author contributions:** Maki TOJO: Conceptualization; Data curation; Formal analysis;
18 Investigation; Methodology; Project administration; Software; Validation; Visualization;

1 Writing – original draft; Writing – review & editing. Hiroyoshi IWATA:
2 Conceptualization; Data curation; Formal analysis; Investigation; Methodology; Project
3 administration; Supervision; Writing – original draft; Writing – review & editing. Naomi
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5 Methodology; Project Administration; Resources; Supervision; Writing – original draft;
6 Writing – review & editing.

9 ***Ethics***

10 Written informed consent was obtained from all participants(the guardians of the
11 children). The Institutional Ethics Board for Epidemiological Studies at Hokkaido
12 University Center for Environmental and Health Sciences and Hokkaido University
13 Graduate School of Medicine approved the study protocol (approval number: 14-27, on
14 June 26th, 2014).

16 **Data availability**

17 The data used in this study are not suitable for deposition in public databases due to ethical
18 constraints and specific legal frameworks in Japan. The Ethical Guidelines for

Epidemiology issued by the Japan Ministry of Education, Culture, Sports, Science, and Technology and the Ministry of Health, Labor, and Welfare restrict the public sharing of epidemiological data. Additionally, the Act on the Protection of Personal Information prohibits the public deposition of personal data. Therefore, any inquiries regarding access to data should be directed to rkishi@med.hokudai.ac.jp. This email address is managed by Professor Reiko Kishi, the Principal Investigator of the Hokkaido Study on Environment and Children's Health at the Center for Environmental and Health Sciences, Hokkaido University.

(v) Declaration of competing interest;

Declaration of competing interest

The authors declare that they have no competing financial interests or personal relationships that could have influenced the work reported in this study.

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(vii) Figure legends*;

Figure 1. Flow diagram of the participants included in the statistical analysis selected from the Hokkaido Study on Environment and Children’s Health.

SGA, small for gestational age; HC, head circumference; DCDQ-J, Developmental Coordination Disorder Questionnaire Japanese version.

Figure 2. Estimated differences in DCDQ-J scores associated with the presence of SGA (Figure 2A) or small HC (Figure 2B).

CI, confidence interval; SGA, small for gestational age; HC, head circumference.

Tables with their legends

Table 1. Characteristics of children and mothers according to the presence of SGA or small HC from the Hokkaido Study on Environment and Children’s Health

	SGA			small HC		
	SGA	Non-SGA	Overall	Small HC	Non-small HC	Overall
	(N=113)	(N=3770)	(N=3883)	(N=350)	(N=3498)	(N=3848)
Mother:						
Age at birth						
Median [IQR]	32.0 [29.0, 34.0]	32.0 [28.0, 35.0]	32.0 [28.0, 35.0]	31.0 [28.0, 34.0]	32.0 [28.0, 35.0]	32.0 [28.0, 35.0]
Smoked during pregnancy						
Smoked during pregnancy	29 (25.7%)	549 (14.6%)	578 (14.9%)	64 (18.3%)	504 (14.4%)	568 (14.8%)
Did not smoke during pregnancy	83 (73.5%)	3193 (84.7%)	3276 (84.4%)	283 (80.9%)	2968 (84.8%)	3251 (84.5%)

Child:

Weeks at birth

Median [IQR]	39 [38, 40]	39 [38, 40]	39 [38, 40]	39 [38, 40]	39 [38, 40]	39 [38, 40]
≤ 36	11 (9.7%)	125 (3.3%)	136 (3.5%)	12 (3.4%)	122 (3.5%)	134 (3.5%)
37 ≤	102 (90.3%)	3645 (96.7%)	3747 (96.5%)	338 (96.6%)	3376 (96.5%)	3714 (96.5%)

Sex

Female	62 (54.9%)	1890 (50.1%)	1952 (50.3%)	215 (61.4%)	1722 (49.2%)	1937 (50.3%)
Male	51 (45.1%)	1880 (49.9%)	1931 (49.7%)	135 (38.6%)	1776 (50.8%)	1911 (49.7%)

Weight at birth

Median [IQR]	2400 [2210, 2580]	3050 [2820, 3300]	3040 [2800, 3290]	2760 [2530, 2980]	3060 [2840, 3320]	3030 [2800, 3290]
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Age (months) at answering DCDQ-J

Median[IQR]	61 [60, 65]	61 [60, 65]	61 [60, 65]	61 [60, 65]	61 [60, 65]	61 [60, 65]
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DCDQ-J Total

Median[IQR]	53.0 [44.0, 61.0]	55.0 [48.0, 62.0]	55.0 [48.0, 62.0]	54.0 [47.0, 61.8]	55.0 [48.0, 62.0]	55.0 [48.0, 62.0]
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DCDQ-J Control during movement

Median[IQR]	21.0 [17.0, 23.0]	21.0 [18.0, 24.0]	21.0 [18.0, 24.0]	21.0 [18.0, 24.0]	21.0 [18.0, 24.0]	21.0 [18.0, 24.0]
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DCDQ-J Fine motor

Median [IQR]	15.0 [12.0, 19.0]	16.0 [13.0, 18.0]	16.0 [13.0, 18.0]	16.0 [12.0, 18.0]	16.0 [13.0, 18.0]	16.0 [13.0, 18.0]
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DCDQ-J General coordination

Median [IQR]	18.0 [15.0, 21.0]	18.0 [15.0, 21.0]	18.0 [15.0, 21.0]	18.0 [15.0, 21.0]	18.0 [15.0, 21.0]	18.0 [15.0, 21.0]
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IQR, inter-quartile range in the 75th and 25th percentile

SGA, Small for Gestational Age; HC, Head Circumference; IQR, Interquartile Range; DCDQ-J, Developmental Coordination Disorder Questionnaire - Japanese version

Data set: Hokkaido Study on Environment and Children’s Health Cohort

Table 2. Univariate linear regression analysis results for DCDQ-J in SGA patients and those with small HC

DCDQ-J	SGA							small HC						
	β	95% CI Lower	95% CI Upper	Std. Error	<i>t</i>	<i>Pr(> t)</i>	Effect Size (R_squared)	β	95% CI Lower	95% CI Upper	Std. Error	<i>t</i>	<i>Pr(> t)</i>	Effect Size (R_squared)
Total	-				-			-				-		
		-4.19	-0.30	0.99		0.02	*	<0.01		-1.86	0.43	0.58		<0.01
	2.25				2.26			0.72				1.23		
Control	-				-			-				-		
During Movement		-1.78	-0.13	0.42		0.02	*	<0.01		-0.87	0.10	0.25		<0.01
	0.96				2.27			0.38				1.55		
Fine Motor	-				-			-				-		
		-1.30	0.08	0.35		0.09	.	<0.01		-0.54	0.27	0.21		<0.01
	0.61				1.72			0.14				0.66		
General	-				-			-				-		
Coordinator		-1.45	0.09	0.39		0.08	.	<0.01		-0.65	0.25	0.23		<0.01
	0.68				1.73			0.20				0.86		

β, Estimated score difference due to the presence of SGA or small HC; DCDQ-J, Developmental Coordination Disorder Questionnaire - Japanese version; SGA, Small for Gestational Age; HC, Head Circumference; CI, Confidence Interval;

Std. Error, Standard Error.

*P< 0.05, P< 0.1 as revealed using univariable linear regression analysis

Data set: Hokkaido Study on Environment and Children’s Health Cohort

Table 3. Multivariate linear regression analysis results for DCDQ-J in SGA patients or those with small HC

Outcome	Predictors	SGA						Predictors	small HC					
		β	95% CI Lower	95% CI Upper	Std. Error	$Pr(> t)$	Effect Size (R_squared)		β	95% CI Lower	95% CI Upper	Std. Error	$Pr(> t)$	Effect Size (R_squared)
Total	SGA	-2.27	-4.20	-0.33	0.42	0.02 *	0.03	Small HC	-1.05	-2.19	0.09	0.58	0.07 .	0.03
	Child sex	-3.03	-3.68	-2.38	0.14	<0.01 ***		Child sex	-3.09	-3.75	-2.44	0.33	<0.01 ***	
	Did not smoke during pregnancy	1.37	0.46	2.29	0.20	<0.01 **		Did not smoke during pregnancy	1.46	0.54	2.38	0.47	<0.01 **	
	Mother's age at birth	-0.11	-0.17	-0.04	0.02	<0.01 **		Mother's age at birth	-0.11	-0.18	-0.04	0.04	<0.01 **	
Control During Movement	SGA	-0.90	-1.73	-0.07	0.42	0.03 *	<0.01	Small HC	-0.38	-0.87	0.11	0.25	0.13	<0.01
	Child sex	0.09	-0.18	0.37	0.14	0.51		Child sex	0.07	-0.35	0.21	0.14	0.64	
	Did not smoke during pregnancy	0.34	-0.05	0.73	0.14	0.09 .		Did not smoke during pregnancy	0.37	-0.47	0.07	0.20	0.07 .	
	Mother's age at birth	-0.04	-0.07	-0.01	0.02	0.00 **		Mother's age at birth	-0.05	-0.07	-0.01	0.02	<0.01 **	
Fine Motor	SGA	-0.67	-1.33	0.00	0.34	0.05 *	0.09	Small HC	-0.38	-0.76	0.02	0.20	0.06 .	0.09
	Child sex	-2.20	-2.42	-1.97	0.11	<0.01 ***		Child sex	-2.21	1.98	2.43	0.11	<0.01 ***	
	Did not smoke during pregnancy	0.61	0.30	0.92	0.16	<0.01 ***		Did not smoke during pregnancy	0.65	-0.70	-0.26	0.16	<0.01 ***	
	Mother's age at birth	-0.04	-0.06	-0.02	0.01	<0.01 ***		Mother's age at birth	-0.04	-0.06	-0.02	0.01	<0.01 ***	
General Coordination	SGA	-0.70	-1.43	0.11	0.39	0.08 .	0.02	Small HC	-0.29	-0.74	0.16	0.23	0.21	0.02
	Child sex	-0.93	0.67	1.19	0.13	<0.01 ***		Child sex	-0.95	-1.21	-0.69	0.13	<0.01 ***	
	Did not smoke during pregnancy	0.43	-0.59	-0.09	0.19	0.02 *		Did not smoke during pregnancy	0.45	0.08	0.82	0.19	0.02 *	

Mother's age at birth	-0.02	-0.05	0.01	0.01	0.14	Mother's age at birth	-0.02	-0.05	0.01	0.01	0.13

β, Estimated score difference due to the presence of SGA or small HC, adjusted for child sex (male), maternal smoking during pregnancy, and maternal age at child's birth

CI, Confidence Interval; SGA: Small for Gestational Age; HC: Head Circumference.

***P< 0.001, **P< 0.01, *P< 0.05, P< 0.1 as revealed using multivariable linear regression analysis

Data set: Hokkaido Study on Environment and Children's Health Cohort

Abbreviations

ADHD- Attention Deficit Hyperactivity Disorder

ASD - Autism Spectrum Disorder

DCD - Developmental Coordination Disorder

DCDQ - Developmental Coordination Disorder Questionnaire

DCDQ-J - Japanese version of the Developmental Coordination Disorder Questionnaire

DSM-5TR - Diagnostic and Statistical Manual of Mental Disorders Text Revision

HC - Head Circumference

SGA - Small for Gestational Age

SD - Standard Deviations

Total number of participating children in the Hokkaido Study ($n = 20,945$)

SGA

- No data on SGA criteria ($n = 7,450$)
Exclude: no data (birth weight, height, sex of the infant, and gestational age)

($n = 13,495$)

- No data on DCDQ-J for 4-7 years old ($n=9,543$)

($n = 3,952$)

- Multiple births ($n = 63$)

($n = 3,889$)

- Chromosome abnormality ($n = 3$)

($n = 3,886$)

- Less than or more than 5 or 6 years old ($n = 3$)

Participants included for statistical analysis ($n = 3,883$)

Small HC

- No data on small HC criteria ($n = 7,579$)
Exclude: no data (birth head circumference, sex of the infant, and gestational age)

($n = 13,366$)

- No data on DCDQ-J for 4-7 years old ($n=9,449$)

($n = 3,917$)

- Multiple births ($n = 63$)

($n = 3,854$)

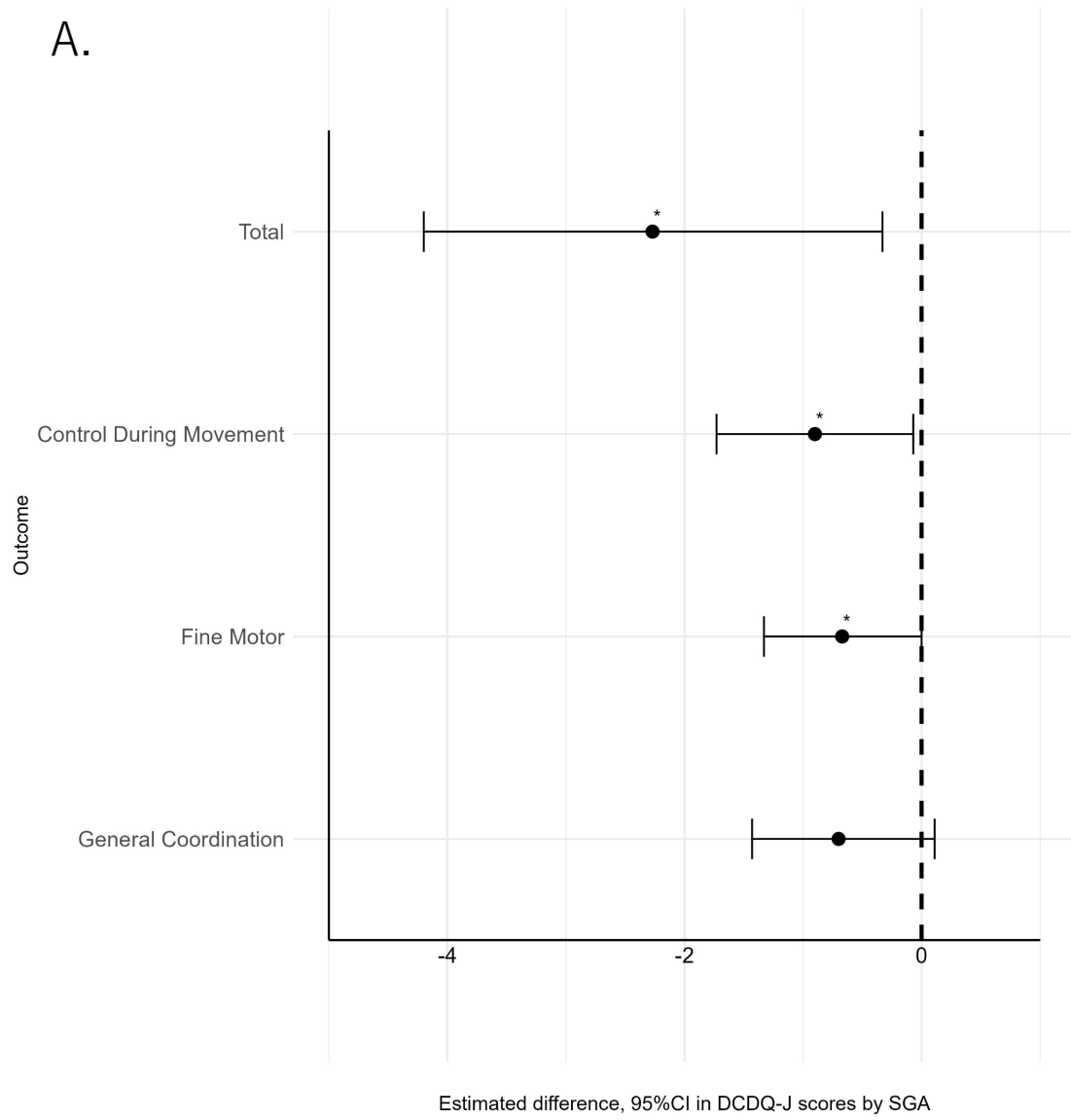
- Chromosome abnormality ($n = 3$)

($n = 3,851$)

- Less than or more than 5 or 6 years old ($n = 3$)

Participants included for statistical analysis ($n = 3,848$)

A.



B.

