Preterm children’s developmental coordination disorder, cognition and quality of life: a prospective cohort study

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ABSTRACT
Objective To evaluate the rate of developmental coordination disorder (DCD) and its correlation to cognition and self-experienced health-related quality of life (HRQoL) in children born very preterm.

Design Prospective follow-up study.

Setting Regional population of children born very preterm in Turku University Hospital, Finland, in 2001–2006.

Patients A total of 170 children born very preterm were followed up until 11 years of age.

Main outcome measures Motor and cognitive outcomes were evaluated using the Movement Assessment Battery for Children - Second Edition (Movement ABC-2) and the Wechsler Intelligence Scale for Children - Fourth Edition, respectively, and HRQoL using the 17-Dimensional Illustrated Questionnaire (17D). The Touwen neurological examination was performed to exclude other neurological conditions affecting the motor outcome.

Results Eighteen children born very preterm (17 boys) (11.3%) had DCD, defined as Movement ABC-2 total test score ≤5th percentile. A positive correlation between motor and cognitive outcome (r=0.22, p=0.006) was found. Children born very preterm with DCD had lower cognitive scores than those without DCD (Full-Scale IQ mean 76.8 vs 91.6, p=0.001). Moreover, children born very preterm with DCD reported lower HRQoL than children born very preterm without motor impairment (17D mean 0.93 vs 0.96, p=0.03). However, HRQoL was higher in this group of children born very preterm compared with population-based normative test results (p=0.001).

Conclusions DCD was still common at 11 years of age in children born very preterm in 2000s. DCD associated with adverse cognitive development and lower self-experienced HRQoL. However, this group of children born very preterm reported better HRQoL in comparison with Finnish norms.

INTRODUCTION
The incidence of cerebral palsy (CP) has decreased among children born very preterm.1–6 However, the rate of non-CP motor impairments such as developmental coordination disorder (DCD) has not decreased,7 and children born preterm are still at increased risk for cognitive impairment compared with term peers.8–11

DCD is defined as motor problems interfering with academic achievement or activities of daily living which cannot be explained by medical, neurological or cognitive impairment.12 The aetiology of DCD is multifactorial, and neuroimaging studies have shown alterations in the brain development and functioning in children with DCD.13–16 The prevalence of DCD has been shown to vary from 5% to 6% in school-aged children and from 8% to 51% in those born preterm.8 12 17–19 DCD has been shown to co-occur with developmental disorders such as social, behavioural and attention problems, and learning difficulties.12 17 20–22 Nevertheless, data on the relationship between DCD and cognitive development in early adolescence are limited.22
Severe neurodevelopmental impairments such as CP, cognitive impairment, and hearing and visual impairment have been reported to associate with poorer self-experienced health-related quality of life (HRQoL) in school-aged children born preterm, while those without these morbidities have reported HRQoL equal to peers.23 24 The effect of preterm birth on HRQoL seems to be most significant in younger years and seems to decrease over time.25 The impact of motor impairments such as DCD on HRQoL is not well known. Dewey and Volkovinskaia26 have found no differences in total HRQoL scores between adolescents with DCD and/or attention deficit hyperactivity disorder (ADHD) and typically developing adolescents. However, they found that adolescents with DCD and ADHD had lower HRQoL on the mood and emotions subscale and school environment subscale. Their additional comparisons indicated that on both these subscales adolescents with DCD and ADHD had significantly lower scores than adolescents with DCD only. Karras et al27 have found that children with DCD reported significantly lower scores in 4 out of 10 HRQoL subscales: psychological well-being, mood and emotions, parent relations and home life, and school environment.

The aims of this study were to evaluate the rate of DCD and to study the correlation between motor and cognitive development at 11 years of age in children born very preterm and/or with very low birth weight in 2000s, and to study the effect of DCD on self-experienced HRQoL. We hypothesised that DCD is still common in children born very preterm and/or with very low birth weight, that poorer motor outcome correlates with adverse cognitive performance, and that DCD correlates with lower perceived HRQoL as compared with children born very preterm and/or with very low birth weight without motor impairment.

METHODS
Participants
This prospective study is part of the PIPARI (Piennapaijoisten raskilasten käyttäytyminen ja toimintakyky imevääsiästä kouluikään; The Development and Functioning of Very Low Birth Weight Infants from Infancy to School Age) study of infants born very preterm.28 29 The participants were born to Finnish-speaking or Swedish-speaking families from January 2001 to December 2006 in Turku University Hospital, Finland, which is one of the five level III hospitals in Finland. From 2001 to 2003 the inclusion criteria were birth weight ≤1500 g and prematurity (<37 gestational weeks). From 2004, the inclusion criteria were broadened to all infants born <32 weeks of gestational age irrespective of birth weight. The exclusion criteria were severe congenital anomalies or diagnosed syndrome affecting cognitive development. The flow chart of the participants is shown in figure 1. Written informed consent for this follow-up study was provided by parents and children.

Patient involvement
This study was done without patient and public involvement.

Motor outcome
The diagnosis of CP was confirmed based on the classification proposed by Himmelmann et al30 after a systematic clinical follow-up by 2 years of corrected age by an experienced child neurologist. The motor outcome of the children born very preterm without CP was evaluated at 11 years of age by one of the three physicians using the Movement Assessment Battery for Children - Second Edition (Movement ABC-2).31 32 The three physicians performing the motor and neurological assessments were PhD students of the PIPARI study group. The raw scores were converted into total standard scores and percentile scores according to the test manual, using age band 3 (11–16 years) and the norms for 11-year-old children. A total test score >15th percentile indicated no movement difficulty, >5th to 15th percentile indicated risk
of movement difficulties, and ≤5th percentile indicated DCD. The Touwen neurological examination was used to confirm that there were no other neurological conditions such as muscle diseases affecting the motor development. All the Movement ABC-2 assessments and Touwen neurological examinations were video-recorded. In case of any hesitation regarding the assessments, the videos were reassessed by one experienced child neurologist (LH).

Cognitive outcome
The cognitive development of children born very preterm at 11 years of age was assessed with the Wechsler Intelligence Scale for Children - Fourth Edition (WISC-IV), Finnish translation. The assessments were performed either in Finnish or Swedish according to the child’s native language. Finnish assessments were performed by one of the two psychologists, who were PhD students of the PIPARI study group. Swedish-speaking children were assessed by a native Swedish-speaking psychologist. General intelligence was measured with Full-Scale IQ, which consisted of the Verbal Comprehension Index, the Perceptual Reasoning Index, the Working Memory Index and the Processing Speed Index. The classification was based on the test manual. The scores were classified as severe cognitive impairment.

Health-related quality of life
The self-experienced HRQoL of children born very preterm at 11 years of age was evaluated using a generic self-assessment measure, the 17-Dimensional Illustrated Questionnaire (17D). It consisted of 17 multiple-choice questions of health and function. The domains were mobility, vision, hearing, breathing, sleeping, eating, speech, excretion, school and hobbies, learning and memory, discomfort and symptoms, depression, distress, vitality, appearance, friends, and concentration. Each domain had a five-level tick box functioning scale alternating from a perfect level to a severe dysfunction. The children completed the questionnaire by themselves before the motor assessment, except for one child who was not able to read and was interviewed by the physician before the assessment. The relative weights of each dimension were defined in the instrument’s home page. The overall HRQoL was calculated from the health state descriptive system using population-based preference or utility weights for 11-year-old healthy Finnish school children. The HRQoL score varied from 0 (worst score, equal to death) to 1 (best score, equal to complete health).

Statistical analysis
Differences in continuous background characteristics between the study children born very preterm and the children who withdrew were studied using a two-sample t-test or a Wilcoxon two-sample test. For categorical background characteristics, χ² test or Fisher’s exact test was used. Correlations between the percentiles for the total scores of the Movement ABC-2 and Full-Scale IQ, and between the percentiles for the total scores of the Movement ABC-2 and WISC-IV indexes, were calculated using Pearson correlations. Associations between DCD and background characteristics were studied using logistic regression analysis. Differences in the Full-Scale IQ and indexes between children born very preterm with and without DCD were studied using two-sample t-test. The associations between motor outcome (children born very preterm with and without DCD), cognitive outcome (Full-Scale IQ and indexes) and background characteristics (birth weight, gestational age, and mother’s and father’s education) were used linear multiple regression model. The background characteristics were chosen a priori.

If up to three dimensions were missing from the 17D, multiple imputation was used to replace missing values with one value, as suggested by the instrument’s home page (http://www.15d-instrument.net/15d/replacing-missing-data/), in order to calculate the 17D total score. If more than three dimensions were missing, the questionnaire was not used in the analyses. Differences in the 17D scores between the groups of (1) children with and without DCD, (2) with and without CP, and (3) with and without severe cognitive impairment were studied using Mann-Whitney U test. The correlations between the Movement ABC-2 and the 17D as well as the Full-Scale IQ and the 17D were studied using Spearman’s correlation. The differences in 17D scores in the study children born very preterm compared with the Finnish population-based normative results were studied using Mann-Whitney U test. Statistical analyses were carried out using SAS V.9.4 for Windows. P values <0.05 were considered statistically significant.

RESULTS
A total of 170 children born very preterm were followed until 11 years of age. The follow-up rate was 77.6% (out of 219 participants). The background characteristics of the study children and the children who withdrew are shown in table 1. The rate of CP did not differ between the study children and the children who withdrew (p=0.5). The mothers of the children who withdrew had lower educational level compared with the mothers of the study children (53% vs 36% with ≤12 years of education; p=0.04). No other differences in background characteristics shown in table 1 were found between the study children and the children who withdrew.

Motor development
All children born very preterm, including those with Full-Scale IQ <70, were able to follow the given instructions without any adaptations of test items and completed the Movement ABC-2. Accordingly, children born very preterm with Full-Scale IQ <70 were included in the analyses regarding DCD, as suggested by the recent European
Table 1  Background characteristics of the 11-year-old children (n=170) born at very low gestational age (<32 weeks) or with a very low birth weight (≤1500 g)

| Characteristics                                      | Study children, n=170 | Children who withdrew, n=49 | P value |
|------------------------------------------------------|-----------------------|----------------------------|---------|
| Gestational age, mean (SD) (minimum, maximum), weeks | 29.1 (2.7) (23.0, 35.9) | 29.0 (2.7) (23.7, 34.1)    | 0.9     |
| Birth weight, mean (SD) (minimum, maximum), g        | 1134.4 (315.3) (400.0, 2120.0) | 1184.6 (374.5) (565.0, 1970.0) | 0.3     |
| Small for gestational age (<−2 SD), n (%)            | 56 (32.9)             | 11 (22.5)                  | 0.2     |
| Male, n (%)                                          | 94 (55.3)             | 30 (61.2)                  | 0.5     |
| Caesarean section, n (%)                             | 101 (59.4)            | 32 (65.3)                  | 0.5     |
| Bronchopulmonary dysplasia, n (%)                    | 22 (12.9)             | 7 (14.3)                   | 0.8     |
| Operated necrotising enterocolitis, n (%)            | 7 (4.2)               | 3/48 (6.3)                 | 0.7     |
| Sepsis, n (%)                                         | 30 (17.7)             | 7 (14.3)                   | 0.6     |
| Laser-treated retinopathy of prematurity, n (%)      | 4 (2.4)               | 3/47 (6.4)                 | 0.2     |
| Brain MRI at term age, n (%)*                        |                       |                            | 0.9     |
| Normal findings                                      | 96/165 (58.2)         | 29/48 (60.4)               |         |
| Minor pathologies                                    | 27/165 (16.4)         | 7/48 (14.6)                |         |
| Major pathologies                                    | 42/165 (25.5)         | 12/48 (25.0)               |         |
| Mother's education, n (%)                            |                       |                            | 0.04    |
| ≤12 years                                            | 61/168 (36.3)         | 24/45 (53.3)               |         |
| >12 years                                            | 107/168 (63.7)        | 21/45 (46.7)               |         |
| Father's education, n (%)                            |                       |                            | 0.4     |
| ≤12 years                                            | 110/166 (66.3)        | 32/44 (72.7)               |         |
| >12 years                                            | 56/166 (33.7)         | 12/44 (27.3)               |         |

*The specific MRI protocol and details about the classification of the findings have been previously described by Setänen et al.14

Academy of Childhood Disability recommendations.12 There were nine (5.3%) children born very preterm with CP who were assessed at 11 years of age; they were excluded from the analyses regarding the Movement ABC-2.

Of all the 161 children born very preterm without CP, one did not complete the Movement ABC-2. A total of 142 (88.8%) had a total test score >5th percentile. Of these children born very preterm, 12 (8.5%) had their score between the 5th and 15th percentile in the Movement ABC-2, indicating a risk for motor problems. There were 18 children born very preterm (11.3%) with a total test score ≤5th percentile in the Movement ABC-2; these children were denoted to have DCD after confirming with the Touwen neurological examination that they did not have such neurological findings or other neurological disorders which could explain their poor performance. Twelve of the children born very preterm with DCD were born extremely preterm (<28 gestational weeks) and/or with extremely low birth weight (<1000 g), representing 18.2% of all (n=66) extremely preterm and/or extremely low birthweight children. All but one of the children born very preterm with DCD were boys. Of the other background characteristics shown in table 1, lower gestational age (p=0.04), bronchopulmonary dysplasia (p=0.04), sepsis (p=0.04) and major brain pathologies on MRI at term age (p=0.02) were associated with DCD.

Cognitive development
The mean value (SD (minimum, maximum)) of the Full-Scale IQ for the whole very preterm study cohort (n=170) was 88.5 (17.0 (40.0, 131.0)). The mean value for the verbal comprehension was 90.3 (14.8 (46.0, 122.0)), for the perceptual reasoning 92.0 (17.1 (40.0, 122.0)), for the working memory 92.6 (16.3 (46.0, 133.0)) and for the processing speed 93.9 (17.4 (47.0, 153.0)). Of all the 161 children born very preterm without CP, 89 (55.3%) performed within the average range (Full-Scale IQ ≥90), 34 (21.1%) had low average performance (Full-Scale IQ ≥80–89), 25 (15.5%) had borderline cognitive development (Full-Scale IQ ≥70–79) and 13 (8.1%) had severe cognitive impairment (Full-Scale IQ <70). The mean values of the Full-Scale IQ and its four indexes are shown by categories of motor outcome in table 2. The Movement ABC-2 scores of children born very preterm without CP correlated positively with Full-Scale IQ (r=0.2, p=0.006), working memory (r=0.3, p<0.001), processing speed (r=0.2, p=0.03) and perceptual reasoning (r=0.2, p=0.03). The scatter plot of the Full-Scale IQ and the Movement ABC-2 is shown in figure 2.
Table 2  Cognitive outcome shown in 11-year-old children born very preterm with CP and according to the performance in the Movement Assessment Battery for Children in children without CP

|                        | CP, n=9          | DCD, ≤5th percentile, n=18 | Children without motor impairment, >5th percentile, n=142 | P value |
|------------------------|------------------|-----------------------------|----------------------------------------------------------|---------|
| Full-Scale IQ          | 62.4 (22.8)      | 76.8 (18.2)                 | 91.6 (14.3)                                              | <0.001* |
|                        | (40.0, 97.0)     | (40.0, 100.0)               | (52.0, 131.0)                                           |         |
| Verbal comprehension   | 75.1 (21.1)      | 83.8 (16.3)                 | 92.1 (13.4)                                              | 0.02*   |
|                        | (46.0, 98.0)     | (46.0, 108.0)               | (60.0, 122.0)                                           |         |
| Perceptual reasoning   | 64.8 (23.1)      | 85.7 (17.9)                 | 94.8 (14.6)                                              | 0.02*   |
|                        | (40.0, 100.0)    | (51.0, 109.0)               | (62.0, 122.0)                                           |         |
| Working memory         | 77.0 (20.5)      | 79.2 (13.9)                 | 95.4 (15.0)                                              | <0.001* |
|                        | (46.0, 109.0)    | (46.0, 97.0)                | (55.0, 133.0)                                           |         |
| Processing speed       | 72.4 (21.4)      | 83.3 (19.4)                 | 96.5 (15.5)                                              | 0.001*  |
|                        | (47.0, 106.0)    | (47.0, 118.0)               | (56.0, 153.0)                                           |         |

The mean values (SD) (minimum, maximum) of Full-Scale IQ and its four indexes are shown. The outcomes are compared between children with DCD and children without motor impairment (two-sample t-test). *The results remained statistically significant after adjusting for birth weight, gestational age, and mother’s and father’s education.

CP, cerebral palsy; DCD, developmental coordination disorder.

Children born very preterm with DCD had lower Full-Scale IQ than children born very preterm without motor impairment (mean 76.8 vs 91.6) (p<0.001). Similarly, children born very preterm with DCD scored lower than children born very preterm without motor impairment in all indexes, as shown in table 2. The results remained statistically significant after adjusting for birth weight, gestational age, and mother’s and father’s education.

**Health-related quality of life**

A total of 167 (98.2%) of the followed children born very preterm completed the 17D questionnaire. There were no statistically significant correlations between the Movement ABC-2 and the 17D questionnaire (r=0.1, p=0.06), nor between the Full-Scale IQ and the 17D questionnaire (r=0.07, p=0.4). However, children born very preterm with DCD had lower self-experienced HRQoL compared with children born very preterm without DCD (0.93 vs 0.96, p=0.03). Children born very preterm with DCD showed more problems than children born very preterm without DCD on the dimensions considering vision (0.96 vs 0.99, p=0.008), hearing (0.92 vs 0.98, p=0.01) and speech (0.96 vs 0.99, p=0.007). The HRQoL of children born very preterm with CP did not differ from the HRQoL of children born very preterm without CP (0.94 vs 0.96, p=0.6), nor did the HRQoL in children born very preterm with severe cognitive impairment (Full-Scale IQ <70) from the children born very preterm without cognitive impairment (0.93 vs 0.96, p=0.2). This cohort of children born very preterm reported better self-experienced HRQoL compared with Finnish population-based normative results (p<0.001). Children born very preterm showed less problems than the normative population on the dimensions considering sleeping (p=0.02), discomfort and symptoms (p<0.001), depression (p<0.001), vitality (p=0.02), appearance (p=0.03), friends (p=0.01), and concentration (p=0.001).

DISCUSSION

This study showed that DCD is still common in 11-year-old children born very preterm in 2000s. Children born very preterm with DCD had worse cognitive development.
than children born very preterm without motor impairment. Moreover, children born very preterm with DCD reported lower HRQoL than children born very preterm without DCD. However, the HRQoL was higher in this study cohort of children born very preterm than in Finnish norm population.

The finding of a high rate of DCD in children born very preterm in early adolescence is parallel to the recently reported rising trend of non-CP motor impairments in children born extremely preterm at the age of 6.5 and 8 years.21 The rate of DCD in children born extremely preterm of this PIPARI study cohort was 18%, while two recent studies from Sweden and Australia have reported non-CP motor impairment rates of 26%–37% in extremely preterm populations.21 The Swedish study reported that the rate of non-CP motor impairment was 37% when they used a cut-off based on their control group, but if the normative cut-offs had been used the rate would have been 12.5%. Some studies have reported the prevalence of non-CP motor impairment as being higher in boys,21 12 while others have shown no significant difference in the prevalence in boys and girls.21 In the present study, all but one of the children with DCD were boys. However, the small number of children with DCD did not enable reliable statistical analysis regarding sex.

A positive correlation between motor outcome and cognitive development in children born very preterm was found even if the correlations were not strong in magnitude. Children born very preterm with DCD had lower mean scores in the Full-Scale IQ and in all indexes compared with children born very preterm without motor impairment. The differences were clinically significant in magnitude, that is, 15 points for Full-Scale IQ, 8 points for verbal comprehension, 9 for perceptual reasoning, 16 for working memory and 13 for processing speed, all in favour of children without DCD. This is in line with previous studies that have reported lower Full-Scale IQ and processing speed in children born very preterm with DCD at 5 years of age22 and lower perceptual reasoning and processing speed in children born extremely preterm with DCD at 6.5 years of age.21 However, as WISC-IV has some items in the processing speed and the perceptual reasoning index subtests requiring fine motor control (eg, holding a pen, drawing in a small space and manipulating blocks), it is possible that motor impairment may have an effect on the child’s performance in these subtests. According to our results, DCD might also indicate problems in cognitive development at 11 years of age in children born very preterm. Lower motor scores accumulated among boys born very preterm in the present study. Future research may expand current findings about possible mechanisms leading to vulnerability according to sex.

This study showed lower self-experienced HRQoL in children born very preterm with DCD compared with children born very preterm without motor impairment in early adolescence. The affected domains were vision, hearing and speech. The absolute differences in HRQoL results between the groups were minor since the scoring system ranges from 0 to 1. Whether these statistically significant differences have clinical importance is not definite. A previous review using various instruments suggested difficulties in fine motor skills (and causing difficulty, eg, with brushing teeth, washing hair, dressing up and using knife and fork) and in social skills (causing, eg, loneliness and spending more time alone).39 Nevertheless, comparing different instruments should be treated with caution. Self-experienced HRQoL at 11 years of age was better in our study cohort of children born very preterm compared with the test normative at the same age in the Finnish population. This is an unexpected finding as children born very preterm have many impairments potentially lowering their HRQoL. The mothers of the children who withdrew from the study had lower educational level compared with the mothers of the study children. This may have influenced the results and may offer one explanation why the study cohort of children born very preterm reported their HRQoL better compared with the norms. However, we are not aware of differences in general health outcomes in the 1990s and 2000s. In any case, good HRQoL in children born very preterm at 11 years of age is a reassuring information for families with a preterm infant.

The strength of this study was its relatively high follow-up rate (78%) from birth to 11 years of age. The examinations were performed with the latest version of the Movement ABC-2, and a thorough Touwen neurological examination was used to support the definition of DCD. A possible limitation was that the motor assessments were not done repeatedly as suggested by the latest European Academy of Childhood Disability recommendations.12 However, these new guidelines were not available during the data collection. We also chose to use the strict cut-off of fifth percentile to define clinically significant non-CP motor impairment. There was no possibility to compare the rate of DCD with peers born at term due to lack of a control group. To assess cognitive development we used WISC-IV, which is a validated and widely used tool in Finland, and the national cut-offs are precise and up-to-date. Regarding HRQoL, the results the Finnish normative of the same age population were available, although these were based on data collection before 1996. Although the sample size of the whole study group was satisfactory, the total number of children born very preterm with DCD and CP was small, which restricts the power of the statistical analysis concerning these groups and the generalisability of the results.

**CONCLUSIONS**

This study supports previous findings that, even though more preterm born infants survive without CP, they still have an increased risk for DCD. Children born very preterm with DCD showed lower cognitive performance than children born very preterm without DCD. It is important to recognise motor problems early to provide
interventions and support services needed and to provide cognitive assessments with a low threshold. The HRQoL of children born very preterm was to a large extent good, but did differ between children born very preterm with DCD and those without motor impairment.

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Contributors KU collected the data, drafted the initial manuscript and revised the manuscript. LH and LL conceptualised and designed the study, designed the data collection instrument, supervised the data collection, revised the manuscript, and critically reviewed the study for intellectual content. AN collected the data and revised the manuscript. MH, LR and PTR revised the manuscript and critically reviewed the study for intellectual content. AN collected the data and revised the manuscript. LH and LL conceptualised and designed the study, designed the data collection instrument, supervised the data collection, revised the manuscript, and critically reviewed the study for intellectual content. AM, PC and LA designed the study and supervised the project and critically reviewed the study for intellectual content. All authors approved the final manuscript as submitted and agree to be accountable for all aspects of the work.

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