# Efficacy of Mobile-Based Cognitive Training Program DoBrain in Preschool Children With or Without Developmental Disabilities: A Randomized, Single-Blind, Active-Controlled Trial

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**Objective** Mobile-based cognitive training programs can be a viable alternative to in-person interventions, but their efficacies have not been established yet. In this study, we examined the efficacy of DoBrain, a mobile-based cognitive training program designed for children with developmental disabilities (DDs), in comparison with general educational apps named Junior Naver and Kakao Kids.

**Methods** Children aged 34 to 77 months were recruited and randomized at a 1:1 ratio to use DoBrain or general educational apps. Each group used the assigned app on a daily basis at home for 30 minutes for 24 weeks. Parents were instructed to help the children with the app usage. A total of 166 children completed the post-test visit (DoBrain group, n=85,  $55.4\pm8.7$  months old; general educational app group, n=81,  $53.7\pm9.9$  months old). The primary outcome was cognitive development measured by Psychoeducational Profile-Revised (PEP-R), administered at baseline and at post-test.

**Results** DoBrain had no superior effect over general educational apps on the PEP-R Developmental Quotient. When the changes before and after app usage were compared, the DoBrain group and the general educational app group both showed declines in imitation (adjusted p=0.049 and 0.022), perception (adjusted p=0.004 and <0.001), and gross motor (adjusted p=0.003 and 0.002) domains of the PEP-R. Among the DoBrain group, children with DD showed a significantly greater gain in the eye-hand coordination domain of PEP-R compared with those without DD (adjusted p=0.047).

**Conclusion** DoBrain did not show a superior effect over general educational apps on overall cognitive development in preschool children, regardless of the presence of DD. Careful monitoring of the negative effect of mobile-based cognitive training programs is necessary. **Psychiatry Investig 2022;19(12):1000-1011** 

Keywords Cognitive training; Developmental disabilities; Randomized controlled trial; Cognition.

# **INTRODUCTION**

Children with developmental disabilities (DDs) are characterized by impairment in areas of physical, learning, language, or behavior. DD includes autism spectrum disorder (ASD), intellectual disability (ID), and language disorder (LD), all of which are described as neurodevelopmental disorders in the Diagnostic and Statistical Manual of Mental Disorders, 5th edition (DSM-5).<sup>1</sup> Generally, studies conducted in children with DD or their families suggest that intensive interventions from an early age are important.<sup>2,3</sup> However, most of the interventions require a high educator-to-child ratio because they are delivered to individuals or small groups of children. There are more barriers to general usage of intensive interventions, including high cost and uneven geographic accessibility.<sup>4</sup> As a result, only a limited proportion of children who are initially diagnosed with DD have the chance to receive appropriate individualized interventions.

During the last decade, mobile devices such as smartphones and tablets became widely available, and the potential of their use has also increased. Before the widespread availability of mobile devices, cognitive training programs were based on personal computers and their efficacy was studied in patients and healthy participants.<sup>5</sup> As mobile devices rapidly proliferated, mobile-based cognitive training programs gained popularity. Lumosity and BrainHQ are an example of mobile-

Received: May 17, 2022 Revised: August 24, 2022 Accepted: September 29, 2022

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based general educational apps that are developed to promote cognitive function,<sup>6,7</sup> and are marketed to be effective in improving cognitive outcomes in typically developing children. However, studies on those apps have methodological issues such as the absence of active controls and inclusion of insufficient number of participants.<sup>8</sup> Despite these issues, mobile-based apps are easy to access and mostly do not require trained educators, and thus can be an alternative to children with DD who cannot receive in-person interventions.

Recently, efforts have been made to develop cognitive training programs specifically for children with DD in addition to children with typical development.<sup>9,10</sup> DoBrain is a mobilebased cognitive training program designed for children with DD,<sup>11</sup> and targets areas of primary cognitive capacity (attention, orientation, memory), higher-level thinking abilities (problem-solving, reasoning, concept formation), and metaprocessing abilities (executive function, self-awareness). The program consists of multiple levels of games with different levels of difficulty, and only requires simple screen touches as an input. Although DoBrain is commercially available and actively used in countries including Korea and the United States, the efficacy of DoBrain has not been fully examined yet.

Therefore, in a community-based sample of preschool children, we compared the efficacy of DoBrain with Junior Naver and Kakao Kids (JNKK), which are the most popular general educational apps used in Korea for children with typical development. In addition, we examined whether the efficacy of DoBrain was different between children with DD and those without.

# **METHODS**

# Participants and procedure

A power analysis showed that with 100 children in each group, the study would have 80% power to detect a moderate effect (Cohen's d=0.4, p<0.05, two-tailed). A total of 200 children were enrolled upon consent from their parents (Figure 1). The children were enrolled at community-based daycare centers, kindergartens, and special education centers from May 1, 2020 to July 31, 2020. The enrolled children were between 34 and 77 months of age. Exclusion criteria included 1) history of neurologic diseases such as cerebral palsy, 2) any sensory disturbances (i.e., vision, hearing, taste, or smell), or 3) severe gross or fine motor problems that prevented them from participating in psychometric tests. Children who were receiving language or behavioral intervention were eligible if no changes in treatment were planned during the study duration of 24 weeks. Of the 200 children who were enrolled, 81 children were diagnosed with DDs and 119 children showed typical development. DDs included ASD, ID, and LD, and the diagnosis of each disorder was confirmed by board-certified child and adolescent psychiatrists based on DSM-5 and relevant psychometric tests including Psychoeducational Profile-Revised (PEP-R), Preschool Receptive-Expressive Language Scale (PRES) or Sequenced Language Scale for Infants (SEL-SI), and Korean version of the Childhood Autism Rating Scale (K-CARS). When a diagnosis of ID was confirmed, a diagnosis of LD was not applicable. If a child did not have any diagnosable DD, the child was classified as having normal development. The study was approved by the Institutional Review Board of Asan Medical Center (2020-0386). Trial registration:

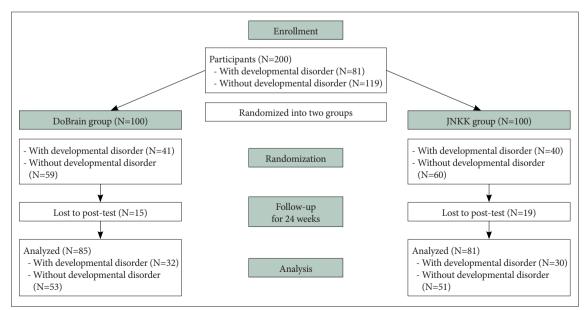


Figure 1. Study participant flowchart. JNKK, Junior Naver and Kakao Kids.

Clinical Research Information Service KCT0007096 (https:// cris.nih.go.kr/cris/index/index.do).

# Study design

DoBrain, which is a mobile-based cognitive training app designed for children with DD, was used in the DoBrain group (Supplementary Figure 1 in the online-only Data Supplement).<sup>12</sup> Junior Naver<sup>13</sup> and Kakao Kids,<sup>14</sup> which are general educational apps designed for typically developing children, were used in the JNKK group. The Junior Naver app includes educational videos, audiobooks, and games that are designed to stimulate imagination and promote development. The Kakao Kids app includes educational contents such as language games, mathematical quizzes, videos on social relationships, and fairy tales that are based on the national educational curriculum for preschool children. No difficulty adjustment was made for the general educational apps.

The present study was a single-blinded, randomized clinical trial. Children participating in the study were assigned to the DoBrain group or JNKK group in a 1:1 ratio using the block randomization method. The children and their parents were aware of the group assignment, but the investigators and evaluators were blinded to group assignment. Only the study coordinator was aware of the group assignment, who did not take part in evaluation or data analysis. To ensure the blinding, children and parents were instructed not to discuss the intervention method while meeting with the investigators and evaluators.

Children in the DoBrain group were instructed to use the DoBrain app for 30 minutes every day for 24 weeks. Children in the JNKK group were instructed to use the Junior Naver app for 30 minutes every day in the first 12 weeks and the Kakao Kids app for 30 minutes every day in the next 12 weeks. Parents were instructed to help the children with the app usage. App compliance was monitored via the central server system. When the expected time was not fulfilled, a text message encouraging additional usage was sent on a weekly basis. The app was delivered through smartphones or tablets.

#### Assessment and measures

The primary outcome of the present study was cognitive development, and the secondary outcomes were adaptive functioning, language skills, ASD symptoms, behavioral problems, and caregiver stress and depression. Relevant psychological tests and questionnaires were administered once at baseline after randomization and once more at post-test after the end of the 24-week study duration.

# **Cognitive development**

Cognitive development was assessed by the PEP-R<sup>15</sup> that

assesses seven domains of development: imitation, perception, fine motor, gross motor, eye-hand coordination, cognitive performance, and cognitive verbal performance. Previous studies reported that the PEP-R shows good internal consistency, test-retest reliability, and high concurrent validity with the Stanford-Binet Intelligence Scale and the Vineland Adaptive Behavior Scale.<sup>16,17</sup> The Developmental quotient (DQ) was calculated to assess the overall developmental level ([developmental age/chronological age]×100), where a lower DQ indicates more delay in cognitive development.

# Adaptive functioning

The Vineland Adaptive Behavior Scale, 2nd edition (VABS), which is a semi-structured interview, was administered to estimate adaptive functioning.<sup>18</sup> Overall standard Adaptive Behavior Composite Score (ABCS) and standardized scores for 4 domains were obtained. Standardized scores have a mean value of 100±15 and higher scores indicate a higher level of functioning. The VABS demonstrated good-to-excellent splithalf and test-retest reliability and modest concurrent validity.<sup>18</sup>

# Language skills

The PRES, which was standardized and validated in Korean children 2- to 6-years of age, was used as the primary measure for language skills.19 When PRES was not applicable due to inadequate language abilities or other child factors, the SELSI was administered.<sup>20</sup> SELSI is an indirect test for language skills where caregivers are questioned instead of children, which was standardized in 1,090 typically developing children. The Receptive and Expressive Vocabulary Test (REVT) was also administered whenever possible.<sup>21</sup> Speech therapists administered the PRES, SELSI, and REVT and the results were supervised by a senior speech therapist with more than 10 years of experience. A significant delay was defined as when the language age confirmed by PRES or SELSI was delayed by more than 12 months. Language quotient was calculated ([developmental age/chronological age]×100) to compare language development in children of different ages.

#### **ASD** symptoms

The K-CARS was administered by child psychologists to assess the ASD symptoms.<sup>22</sup> The K-CARS consists of 15 items rated on a 7-point scale with scores ranging from 1 to 4 (half points), in which higher total scores indicate higher severity. In addition, the Social Responsiveness Scale (SRS), a parent-completed 65-item questionnaire, was used to measure the frequency of ASD-related behaviors.<sup>23</sup>

# **Behavioral problems**

Behavioral and emotional problems were measured by the

Child Behavior Checklist for ages 1.5-5 (CBCL)<sup>24</sup> and the Aberrant Behavior Checklist (ABC).<sup>25</sup> CBCL is a 99-item scale rated by parents with scores ranging from 0 to 2 for each item. The scores for the following seven subscales were calculated: emotionally reactive, anxious/depressed, somatic complaints, withdrawn, sleep problems, attention problems, and aggressive behaviors. CBCL demonstrated good reliability and validity in clinical and non-clinical populations, and showed good cross-informant agreement.<sup>24</sup> ABC, which is a rating scale used to assess behavioral problems in individuals with DD, was also administered. The ABC consists of 58 items answered on a 4-point scale from 0 to 3 based on problem severity. The items are further categorized into the following five domains: irritability, lethargy/social withdrawal, stereotypic behavior, hyperactivity/noncompliance, and inappropriate speech. The psychometric properties of ABC have been assessed, and the subscales showed high internal consistency, adequate reliability, and established validity.25

# Caregiver stress and depression

Parenting stress and depression were measured using the Parenting Stress Index, 4th edition (PSI)<sup>26</sup> and the Center for Epidemiologic Studies Depression Scale-Revised (CESD-R),<sup>27</sup> respectively. The PSI is a 120-item measure with two domains: 1) the child characteristics domain and 2) the parent characteristics domain. Both domains showed excellent internal consistency, and the PSI displayed predictive validity in studies performed in other countries.<sup>26</sup> The CESD-R consists of 20 items related to depression rated on a 5-point scale based on symptom frequency. CESD-R showed good psychometric properties, including high internal consistency, and consistent convergent and divergent validity.<sup>27</sup>

#### Problematic smartphone use

The Smartphone Overuse Scale (SOS) was completed by parents to assess smartphone overuse in children. The scale was developed by the National Information Society Agency, and consists of 9-items that are rated in a 4-point scale.<sup>28</sup> The items are categorized into three subscales. The self-control failure subscale assesses whether controlling smartphone usage is possible; the salience subscale assesses whether smartphone use is the most important activity; and the problematic consequences assesses whether smartphone use is related to impairment in functioning. Psychometric properties of the SOS were explored in a preliminary survey sample of 201 children, and main survey sample of 2,348 children. Confirmatory factor analysis was conducted for SOS (Tucker-Lewis index= 0.91, comparative fit index=0.95, root-mean-square-error of approximation=0.06), and the reliability ranged from 0.58 to 0.75. The receiver operating characteristic analysis was conducted using the internet gaming disorder criteria in Section III of DSM-5 as a golden standard. As a result, a total score of  $\geq$ 24 in the SOS indicated potential risk (area under the curve [AUC]=0.668, sensitivity=70.4%, and specificity=63.3%), and  $\geq$ 28 indicated high risk (AUC=0.590, sensitivity=28.7%, and specificity=89.2%).

#### Statistical analyses

Descriptive statistics were calculated and the results were compared between the DoBrain group and the JNKK group. Psychological test and questionnaire outcomes were compared between the DoBrain group and the JNKK group. First, to estimate the change after app use in each group, paired ttest or Wilcoxon signed-rank test was conducted. Next, to compare the effect of the app between the DoBrain and the JNKK group, t-test or Wilcoxon rank-sum test was conducted. Test for normality was conducted using the Shapiro-Wilk test. Identical statistical analyses were performed to compare the effect of DoBrain in children with and without DD. Subscales were adjusted for multiple comparisons using Bonferroni's method. Statistical significance was set at p=0.05. All statistical analyses were performed using SAS software, version 9.4 (SAS Institute, Cary, NC, USA) and the R Statistical Software, version 4.0.2 (R Foundation for Statistical Computing, Vienna, Austria).

# RESULTS

Of the 200 children who were enrolled, 34 children were lost to post-test and 166 children were included in the final analysis of the study (Figure 1). There were no significant differences between the study children and those who were lost to post-test in terms of age and sex. However, the proportion of children diagnosed with ASD (final analysis group, 25.9% vs. lost-to-post-test group, 52.9%; p=0.002) or ID (final analysis group, 27.7% vs. lost-to-post-test group, 50.0%; p=0.011) was higher in the lost to post-test group. The reason for posttest loss included situations related to COVID-19 (n=17), unable to visit due to personal reasons (n=8), problems with using the app (n=3), concerns related to media exposure (n=1), and unknown (n=4).

The overall demographic characteristics of the study participants are summarized in Table 1. Children in the DoBrain group and the JNKK group did not show significant differences in the age at baseline, sex, diagnosis, number of siblings, birth order, paternal and maternal education, and family income.

Psychological test results were compared between the Do-Brain and JNKK groups (Table 2). Mean change from baseline ( $\Delta$ ) was calculated, and the between-group differences in  $\Delta$  were assessed. PEP-R domains including imitation (adjustEfficacy of Mobile-Based Cognitive Training

Table 1. Demographic characteristics of the study participants

Variable	DoBrain	JNKK	p-value
variable	(N=85)	(N=81)	p-value
Age at baseline, mean±SD (mon)	$55.4 \pm 8.7$	53.7±9.9	0.230
Male sex	50 (58.8)	57 (70.4)	0.120
Diagnosis			
ASD	24 (28.2)	19 (23.5)	0.482
ID	22 (25.9)	24 (29.6)	0.590
LD	6 (7.1)	5 (6.2)	0.819
Number of siblings			0.906
Only child	30 (35.3)	30 (37.0)	
Two	46 (54.1)	44 (54.3)	
Three or more	9 (10.6)	7 (8.6)	
Birth order			0.621
First	61 (71.8)	53 (65.4)	
Second	20 (23.5)	24 (29.6)	
Third or fourth	4 (4.7)	4 (4.9)	
Paternal education			0.156
Graduate school	17 (20.0)	20 (24.7)	
Bachelor's	55 (64.7)	56 (69.1)	
High school or less	13 (15.3)	5 (6.2)	
Maternal education			0.188
Graduate school	21 (24.7)	14 (17.3)	
Bachelor's	53 (62.4)	61 (75.3)	
High school or less	11 (12.9)	6 (7.4)	
Family income			0.985
High	18 (21.2)	18 (22.2)	
Moderate	54 (63.5)	51 (63.0)	
Low	13 (15.3)	12 (14.8)	

Values are N (%) unless specified otherwise. SD, standard deviation; ASD, autism spectrum disorder; ID, intellectual disability; JNKK, Junior Naver and Kakao Kids; LD, language disorder

ed p=0.049 and p=0.022), perception (adjusted p=0.004 and p<0.001), and gross motor (adjusted p=0.003 and p=0.002) showed significant declines from baseline in both the DoBrain and JNKK groups, but no significant difference in  $\Delta$  was observed between the two groups. The ABCS of the VABS showed improvement from baseline only in the DoBrain group (adjusted p=0.005), with no significant between-group difference in  $\Delta$ . The communication subscale of the VABS showed improvement from baseline in both the DoBrain and JNKK groups (adjusted p<0.001 and p=0.006), but no significant difference in  $\Delta$  was observed between the two groups. In the JNKK group, receptive language assessed by the SELSI showed a significant decline from baseline (adjusted p=0.046), whereas both receptive (adjusted p=0.022) and expressive (adjusted p=0.004) language assessed by the REVT showed improve-

ment from baseline, with no significant between-group differences in  $\Delta$ . The K-CARS score showed a significant difference in  $\Delta$  between the two groups (adjusted p=0.045).

Similarly, the questionnaire results were compared between the DoBrain and JNKK groups (Table 2). The SRS total score (adjusted p=0.040) and the aggressive behavior subscale score of the CBCL (adjusted p=0.013) improved from baseline only in the DoBrain group, and there were no significant differences in  $\Delta$  between the two groups. The irritability subscale of the ABC showed significant improvement from baseline in the DoBrain and JNKK groups (adjusted p=0.001 and p=0.007), with no significant between-group difference in  $\Delta$ . The lethargy/social withdrawal subscale (adjusted p=0.017) and the hyperactivity/noncompliance subscale (adjusted p=0.005) of the ABC showed significant improvement from baseline only in the DoBrain group, and there were no significant differences in  $\Delta$  between the two groups. The caregivers of the Do-Brain group reported significant improvement in the child characteristics subscale of PSI (adjusted p=0.012), whereas the caregivers of the JNKK group reported significant improvement of their CESD-R score (adjusted p=0.004), with no significant between-group differences in  $\Delta$ .

The DoBrain and JNKK groups were compared in a subset of children with DDs (Supplementary Table 1 in the onlineonly Data Supplement). In the JNKK group, the perception domain of PEP-R (adjusted p=0.008) and daily living skills of the VABS (adjusted p=0.020) showed significant decline from baseline. In addition, in the JNKK group, the K-CARS score showed significant aggravation from baseline (adjusted p= 0.022), with significant difference in  $\Delta$  between the two groups (adjusted p=0.005). In the DoBrain group, the irritability, lethargy/social withdrawal, and hyperactivity/noncompliance subscales (adjusted p=0.001, 0.017, and 0.017, respectively) of the ABC showed significant improvement. The caregivers of the JNKK group reported significant improvement of their CESD-R score (adjusted p=0.015).

In parallel, the DoBrain and JNKK groups were compared in a subset of children with normal development (Supplementary Table 2 in the online-only Data Supplement). PEP-R domains including imitation (adjusted p<0.001 and p=0.002), perception (adjusted p<0.001 and p<0.001), and gross motor (adjusted p<0.001 and p<0.001) showed significant declines from baseline in both the DoBrain and JNKK groups. The ABCS of the VABS showed improvement from baseline only in the DoBrain group (adjusted p=0.017), and the communication subscale of the VABS showed improvement from baseline in both the DoBrain and JNKK groups (adjusted p=0.002 and p=0.005). The caregivers of the DoBrain group reported significant improvement of their CESD-R score (adjusted p=0.043).

TZ-michle		DoBrain (N=85)			JNKK (N=81)		Group o	Group difference in $\Delta$
variable	Before	After	$\Delta^{\dagger}$	Before	After	$\Delta^{\dagger}$	p-value	Adjusted p-value <sup>†</sup>
PEP-R DQ	86.7±33.7	89.4±29.5	2.7±14.2	87.6±33.4	86.0±34.5	-1.6±13.7	0.054	0.432
Imitation	83.8±37.0	$81.2 \pm 30.9$	-2.7±20.7*	85.7±36.2	$80.1 \pm 34.7$	$-5.5\pm18.8^{*}$	0.932	>0.999
Perception	$88.4\pm 33.2$	83.4±27.4	$-5.0\pm18.8^{*}$	$94.0 \pm 34.1$	$84.1 \pm 34.1$	-9.8±20.8*	0.108	0.864
Fine motor	87.3±29.1	86.5±25.2	-0.8±17.9	$91.3 \pm 30.1$	87.5±28.7	-3.8±18.2	0.157	>0.999
Gross motor	$95.0\pm 25.3$	$91.7\pm 20.5$	-3.3±16.0*	99.4±29.4	91.1±29.6	-8.2±21.9*	0.105	0.840
Eye-hand coordination	92.3±25.9	94.8±21.9	$2.5\pm 15.7$	94.6±27.5	$91.0\pm 25.0$	-3.6±14.2	0.039	0.312
Cognitive performance	$91.3 \pm 35.4$	94.4±29.9	$3.0\pm17.9$	93.0±36.2	$90.3 \pm 37.1$	-2.7±19.1	0.038	0.304
Cognitive verbal	$85.7\pm 31.1$	87.9±29.8	$2.2\pm 13.9$	83.8±32.2	$84.5\pm 33.0$	$0.8 \pm 14.5$	0.660	>0.999
VABS ABCS	80.7±20.4	$83.9\pm 21.0$	3.3±8.7*	$80.4\pm 21.0$	$81.5 \pm 21.4$	$1.1 \pm 9.2$	0.124	0.620
Communication	$81.1 \pm 20.7$	86.2±23.0	$5.2\pm11.5^{*}$	$81.1\pm 21.5$	84.4±23.6	$3.4{\pm}11.1^{*}$	0.451	>0.999
Daily living skills	88.7±17.7	90.2±19.0	$1.6\pm 12.3$	90.3±17.3	$89.1 \pm 18.9$	$-1.2\pm10.3$	0.019	0.095
Socialization	$82.1\pm 23.6$	$84.1\pm 23.5$	$1.9\pm 13.5$	$81.6\pm 21.9$	$82.1\pm 23.1$	$0.5\pm 12.6$	0.400	>0.999
Motor skills	$85.5\pm16.8$	88.1±16.4	$2.6 \pm 13.4$	85.7±15.7	86.4±15.7	$0.6 \pm 9.3$	0.343	>0.999
Language								
PRES, receptive <sup>a</sup>	$93.1\pm 23.3$	95.7±22.0	$2.6\pm 15.7$	97.7±26.0	99.4±25.9	$1.8 \pm 15.1$	0.759	>0.999
PRES, expressive <sup>b</sup>	95.6±23.9	93.9±21.9	-1.7±14.5	$96.1\pm 25.0$	95.5±25.2	$-0.6\pm 14.3$	0.663	>0.999
SELSI, receptive <sup>c</sup>	$42.6\pm 12.9$	$41.5\pm 13.3$	-1.0±6.9	$41.3\pm 17.3$	$37.4\pm15.7$	-3.9±9.0*	0.159	0.317
SELSI, expressive <sup>d</sup>	$39.4\pm 14.0$	$38.1\pm13.4$	-1.3±4.8	$40.6 \pm 18.1$	$38.6\pm 16.4$	-2.0±9.4	0.748	>0.999
$\operatorname{REVT}$ , receptive <sup>e</sup>	$114.0\pm 31.9$	$119.1\pm 33.0$	$5.1 \pm 30.0$	$113.3\pm 26.7$	$124.4\pm 32.5$	$11.1\pm 28.8^{*}$	0.199	0.398
REVT, expressive <sup>f</sup>	$117.4\pm 27.8$	$119.6\pm 27.6$	$2.2\pm 23.7$	$118.6\pm 27.2$	$128.9\pm 27.2$	$10.4\pm 21.4^{*}$	0.070	0.140
K-CARS	$21.4\pm 8.5$	$20.9\pm 8.0$	-0.4±2.5	20.7±7.8	$21.1\pm 8.3$	$0.4\pm 2.4$	0.045	0.045
SRS total score	$53.6\pm 14.8$	52.0±12.2	-1.6±8.4*	$52.8\pm13.9$	$51.4\pm 12.0$	-1.4±6.1	0.745	0.745
CBCL								
Emotionally reactive	$53.4\pm5.0$	$53.0\pm 5.1$	$-0.4\pm4.3$	52.7±4.5	52.6±4.3	-0.2±4.6	0.471	>0.999
Anxious/depressed	53.0±5.2	52.7±5.5	$-0.3\pm3.3$	$51.9 \pm 4.1$	52.2±4.5	$0.3 \pm 5.7$	0.088	0.619
Somatic complaints	$53.6\pm 5.1$	52.9±4.7	-0.7±4.7	52.0±3.9	$52.6\pm 5.0$	$0.7 \pm 4.8$	0.190	>0.999
Withdrawn	$57.1\pm 8.2$	55.7±7.7	$-1.3\pm5.0$	56.9±9.9	$56.0 \pm 9.4$	-0.9±5.9	0.428	>0.999
Sleep problems	$54.2\pm 5.6$	$53.2\pm 5.1$	-0.9±4.9	$54.5\pm 6.9$	53.9±7.4	-0.6±6.1	0.623	>0.999
Attention problems	56.1±7.6	55.2±7.6	-0.9±4.9	55.4±7.0	55.3±7.4	$-0.1\pm 5.4$	0.468	>0.999
Aggressive behavior ABC	53.9±6.1	52.5±5.0	-1.4±4.8*	53.2±5.3	52.7±6.2	-0.5±5.6	0.189	>0.999
Irritability	$4.0 \pm 5.1$	2.6±3.8	$-1.4\pm3.8^{*}$	3.7±5.2	$2.9\pm 5.0$	-0.8±3.2*	0.342	>0.999
Lethargy/social withdrawal	$3.9\pm5.7$	2.9±4.7	$-1.0 \pm 4.3^{*}$	3.6±5.9	$3.2\pm6.0$	$-0.4\pm3.0$	0.133	0.666
Stereotypic behavior	$1.3\pm 2.9$	$1.0\pm 2.3$	-0.3±2.0	$0.9\pm 2.4$	$0.9\pm 2.3$	$0.0{\pm}1.0$	0.703	>0.999
Hyperactivity/noncompliance	$3.9 \pm 4.8$	$2.8 \pm 3.8$	$-1.1\pm3.2^{*}$	$3.4 \pm 4.1$	$2.9 \pm 4.5$	-0.5±3.2	0.482	>0.999
Inappropriate speech	$1.2 \pm 1.8$	$1.0\pm 1.5$	-0.2±1.5	$0.9 \pm 1.4$	$0.9\pm1.5$	-0.1±1.3	0.589	>0.999

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17		DoBrain (N=85)			JNKK (N=81)		Group	Group difference in $\Delta$
Variable	Before	After	$\Delta^{\dagger}$	Before	After	$\Delta^{\dagger}$	p-value	Adjusted p-value <sup>†</sup>
PSI								
Child characteristics	$51.5\pm11.0$	49.9±11.4	-1.6±5.3*	51.7±12.0	51.0±12.8	-0.6±6.7	0.302	0.604
Parent characteristics	50.9±9.4	$50.4 \pm 10.0$	-0.5±6.0	$51.2 \pm 10.4$	$51.5\pm 11.1$	$0.3\pm7.1$	0.580	>0.999
Caregiver CESD-R	$9.1 \pm 9.9$	7.8±8.8	-1.3±8.2	$9.8 \pm 11.2$	$8.0 \pm 11.4$	$-1.8\pm10.5^{*}$	0.321	0.321
SOS								
Self-control failure	$6.0\pm 2.1$	5.9±2.4	$-0.1\pm1.9$	$5.9\pm 2.0$	$6.1\pm 2.0$	$0.3\pm 1.9$	0.329	>0.999
Salience	6.7±3.0	6.9±2.6	$0.2\pm 2.1$	$5.9\pm 2.5$	6.7±2.6	$0.8\pm 2.2^{*}$	0.046	0.138
Problematic consequences	$4.9\pm 2.3$	$5.3\pm 2.1$	$0.4{\pm}1.6^{*}$	$4.7 \pm 2.0$	$5.2\pm 2.0$	$0.5\pm 1.9$	0.823	>0.999
Values are presented as mean±standard deviation. *adjusted p<0.05, significant difference observed between before and after values; <sup>†</sup> subscales were adjusted for multiple comparison using Bonferronis method. <sup>a</sup> DoBrain group (N=72), JNKK group (N=64); <sup>b</sup> DoBrain group (N=70), JNKK group (N=64); <sup>c</sup> DoBrain group (N=25), <sup>d</sup> DoBrain group (N=25), <sup>d</sup> DoBrain group (N=26); <sup>d</sup> DoBrain group (N=26); <sup>d</sup> DoBrain group (N=26); <sup>d</sup> DoBrain group (N=26), <sup>d</sup> DoBrain group (N=47), <sup>D</sup> EP-R DQ, Psychoeductional Profile-Revised Developmental Quotient, <sup>D</sup> DOBrain group (N=64); <sup>d</sup> DoBrain group (N=26); <sup>d</sup> DoBrain group (N=26); <sup>d</sup> DoBrain group (N=64); <sup>d</sup> Do	ard deviation. *adjus p (N=72), JNKK grc p (N=59), JNKK grc	sted p<0.05, signifi oup (N=64); <sup>b</sup> DoBr oup (N=47); <sup>f</sup> DoBr	cant difference obs ain group (N=70), ain group (N=56),	served between bef , JNKK group (N=4 JNKK group (N=4	ore and after value 54); 'DoBrain grou 7). PEP-R DQ Psy	s; <sup>†</sup> subscales were a p (N=24), JNKK gr cchoeducational Pro	djusted for mult roup (N=25); <sup>d</sup> D ofile-Revised De	p<0.05, significant difference observed between before and after values; <sup>†</sup> subscales were adjusted for multiple comparison using (N=64); <sup>b</sup> DoBrain group (N=70), JNKK group (N=64); <sup>c</sup> DoBrain group (N=25), <sup>d</sup> DoBrain group (N=25), <sup>d</sup> DoBrain group (N=75), <sup>f</sup> DoBrain group (N=77); <sup>f</sup> DoBrain group (N=56), JNKK group (N=47). PEP-R DQ, Psychoeducational Profile-Revised Developmental Quotient:

REVT, Receptive and Expressive Vocabulary Test, K-CARS, Korean version of the Childhood Autism Rating Scale; SRS, Social Responsiveness Scale; CBCL, Child Behavior Checklist; ABC, Aberrant Behavior Checklist, PSI, Parenting Stress Index; CESD-R, Center for Epidemiologic Studies Depression Scale-Revised; SOS, Smartphone Overuse Scale; JNKK, Junior Naver and

Kakao Kids

No significant side effects were reported in the study children. Problematic smartphone or tablet use was monitored by the SOS (Table 2). When the mean change from baseline ( $\Delta$ ) was calculated, the Problematic Consequences subscale showed a significant increase in the DoBrain group (adjusted p=0.012), and the salience subscale showed a significant increase in the JNKK group (adjusted p=0.006). However, when the DoBrain and JNKK groups were compared, no significant differences were observed in the subscales of the SOS.

A subset of children who used DoBrain were selected to compare the efficacy of DoBrain according to the presence of DD (Table 3). The results of the psychological tests and questionnaires were compared; the mean changes from baseline ( $\Delta$ ) were calculated, and differences in  $\Delta$  were assessed between children with DD and those without. The eve-hand coordination domain of the PEP-R showed significant differences in  $\Delta$  between children with DD and those without (adjusted p=0.047). Perception (adjusted p<0.001) and fine motor (adjusted p<0.001) domains of the PEP-R showed a significant decline from baseline in children without DD, but no significant difference in  $\Delta$  was observed between children with DD and those without. The SRS total score showed significant improvement from baseline in children with DD (adjusted p=0.001), and a significant difference in  $\Delta$  was observed between children with DD and those without (adjusted p=0.001). In children with DD, significant improvements from baseline were observed in irritability (adjusted p=0.001), lethargy/social withdrawal (adjusted p=0.017), and hyperactivity/noncompliance (adjusted p=0.017) subscales of the ABC. Significant differences in  $\Delta$  between children with DD and those without were observed in irritability (adjusted p=0.006), lethargy/social withdrawal (adjusted p=0.001), and stereotypic behavior (adjusted p=0.008) subscales of the ABC. The salience subscale of the SOS showed a significant increase from baseline in children with DD (adjusted p=0.030).

# DISCUSSION

In this study, we found that preschool children who used the DoBrain app for 24 weeks did not show significant differences in overall cognitive development compared with children who used general educational apps such as JNKK. Compared with the JNKK group, the DoBrain group showed significant improvements in the K-CARS assessment for ASD symptoms. When the efficacy of DoBrain was compared according to the presence of DD, children with DD showed greater gains in the eye-hand coordination domain of PEP-R, SRS total score, and irritability, lethargy/social withdrawal, and stereotypic behavior subscales of ABC than did those without DD. Compared to baseline, cognitive domains related to imitation, percep-

		With DD (N=32)		1	Without DD (N=53)		Group	Group difference in $\Delta$
Variable	Before	After	$\Delta^{\dagger}$	Before	After	$\Delta^{\dagger}$	p-value	Adjusted p-value <sup>†</sup>
PEP-R DQ	53.7±31.9	60.2±26.9	6.5±18.9	106.6±12.4	$107.0\pm11.6$	$0.3\pm9.9$	0.028	0.222
Imitation	$50.2 \pm 35.0$	52.5±28.8	2.3±26.8	$104.2\pm19.0$	98.5±15.2	$-5.6\pm15.5$	0.009	0.072
Perception	62.3±34.8	62.7±28.5	$0.4\pm 24.9$	$104.2\pm19.6$	95.9±17.4	-8.3±13.2*	0.086	0.685
Fine motor	$59.3\pm 26.3$	63.1±21.8	$3.9\pm 22.6$	$104.2\pm 13.6$	$100.6\pm 14.3$	$-3.5\pm13.7$	0.008	0.064
Gross motor	$78.8\pm 29.5$	79.7±21.6	$0.9\pm 23.1$	$104.8\pm 15.9$	$99.0 \pm 16.1$	-5.8±8.9*	0.078	0.627
Eye-hand coordination	$69.1\pm 24.6$	76.4±21.2	7.3±17.6	$106.4\pm 13.5$	$105.9\pm 13.1$	$-0.5\pm13.7$	0.006	0.047
Cognitive performance	$58.1\pm36.5$	66.4±28.3	$8.3\pm 24.8$	$111.4\pm 11.6$	$111.3\pm 14.2$	$-0.2\pm11.1$	0.076	0.609
Cognitive verbal	56.7±28.4	59.7±27.2	$3.0 \pm 16.5$	$103.1\pm 16.1$	$104.8\pm 14.5$	$1.7\pm 12.2$	0.679	>0.999
VABS ABCS	61.5±13.9	63.6±14.8	$2.1 \pm 7.4$	92.2±13.8	96.2±13.1	$4.0 \pm 9.4^{*}$	0.341	>0.999
Communication	$62.8 \pm 15.2$	66.0±19.4	$3.3 \pm 10.5$	92.1±14.9	98.4±15.0	$6.3\pm 12.0^{*}$	0.234	>0.999
Daily living skills	$74.8\pm 13.2$	$74.5\pm17.4$	$-0.4\pm12.8$	$97.0\pm 14.6$	99.8±12.5	$2.8 \pm 12.0$	0.473	>0.999
Socialization	$60.6 \pm 16.4$	$60.1\pm17.3$	-0.6±9.9	$95.1\pm16.9$	98.5±12.1	$3.4{\pm}15.2$	0.149	0.747
Motor skills	$75.1\pm 14.3$	$75.8\pm 15.4$	$0.7 \pm 11.8$	$91.8\pm 15.1$	$95.5\pm 12.0$	$3.7 \pm 14.3$	0.802	>0.999
Language								
PRES, receptive <sup>a</sup>	$65.5\pm 21.3$	69.0±22.8	$3.5\pm 17.1$	$103.0\pm 14.2$	$105.3\pm 11.2$	$2.3\pm 15.3$	0.776	>0.999
PRES, expressive <sup>b</sup>	$64.8\pm 20.8$	$65.0\pm 20.0$	$0.2\pm 17.1$	$105.5\pm 14.7$	$103.1\pm 12.3$	$-2.3\pm13.7$	0.531	>0.999
SELSI, receptive <sup>c</sup>	$41.3\pm 12.0$	$40.1 \pm 11.9$	$-1.1 \pm 7.1$	56.9±19.3	$57.0\pm 24.5$	$0.1 \pm 5.2$	0.794	>0.999
SELSI, expressive <sup>d</sup>	$38.1\pm13.5$	$36.5\pm11.9$	$-1.6\pm4.6$	$54.7\pm 14.1$	$56.5\pm 21.7$	$1.7 \pm 7.6$	0.515	>0.999
REVT, receptive <sup>e</sup>	76.6±30.5	$84.2 \pm 34.0$	$7.6\pm10.8$	$119.9\pm 28.1$	$124.6\pm 29.6$	$4.7 \pm 32.0$	0.499	0.998
REVT, expressive <sup>f</sup>	$80.8 \pm 28.7$	$82.6 \pm 31.6$	$1.8 \pm 9.6$	$124.3\pm 21.6$	$126.6\pm 20.4$	$2.3\pm 25.6$	0.738	>0.999
K-CARS	30.7±7.3	29.6±7.1	$-1.1 \pm 3.5$	$15.9\pm 1.4$	$15.9\pm 1.8$	$0.0{\pm}1.5$	0.153	0.153
SRS total score	$66.1\pm15.1$	61.5±12.2	$-4.6\pm8.6^{*}$	46.0±7.9	46.3±7.8	$0.2 \pm 7.9$	0.001	0.001
CBCL								
Emotionally reactive	$54.8\pm 6.2$	54.2±4.6	$-0.6\pm 5.0$	$52.6 \pm 4.0$	$52.3\pm5.3$	-0.3±3.9	0.540	>0.999
Anxious/depressed	$53.8 \pm 4.9$	$53.0\pm 5.2$	$-0.8 \pm 4.1$	52.5±5.3	$52.4\pm 5.7$	-0.1±2.7	0.111	0.775
Somatic complaints	$54.8\pm 5.9$	$54.3\pm 5.6$	$-0.5\pm5.0$	$52.8 \pm 4.5$	$52.1\pm 3.8$	-0.7±4.5	0.889	>0.999
Withdrawn	64.3±6.6	$61.8 \pm 7.2$	-2.5±6.3	$52.6\pm 5.4$	$52.1\pm 5.2$	-0.6±3.8	0.029	0.203
Sleep problems	$56.3\pm6.4$	53.7±6.0	-2.6±5.7	52.9±4.6	52.9±4.5	$0.0 \pm 4.0$	0.035	0.246
Attention problems	$62.4\pm 8.0$	$60.3 \pm 9.1$	-2.1±6.8	$52.3\pm 3.8$	$52.1 \pm 4.3$	-0.2±3.3	0.171	>0.999
Aggressive behavior ABC	57.1±7.9	54.6±6.2	-2.4±6.3	52.0±3.7	$51.2 \pm 3.5$	-0.8±3.5	0.059	0.410
Irritability	7.3±6.6	$4.3\pm 5.0$	-3.0±4.5*	$1.9\pm 2.3$	$1.5\pm 2.3$	-0.5±2.9	0.001	0.006
Lethargy/social withdrawal	$8.8 \pm 6.1$	$6.2\pm5.1$	-2.7±4.7*	$0.9\pm 2.7$	$0.9\pm 2.9$	$0.0 \pm 3.8$	0.000	0.001
Stereotypic behavior	$3.2 \pm 4.0$	$2.1 \pm 3.0$	$-1.1\pm2.7$	$0.1 \pm 0.4$	$0.3 \pm 1.3$	$0.2 \pm 1.1$	0.002	0.008
Hyperactivity/noncompliance	$8.1 \pm 5.4$	5.7±4.3	-2.4±4.3*	$1.4 \pm 1.6$	$1.1 \pm 2.0$	-0.3±2.0	0.025	0.124
Inappropriate speech	$2.6\pm 2.2$	$1.9\pm 2.0$	-0.6±2.1	$0.3 \pm 0.6$	$0.3 \pm 0.7$	$0.0 \pm 0.9$	0.041	0.204

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111		With DD (N=32)		F	Without DD (N=53)	3)	Group	Group difference in $\Delta$
variable	Before	After	$\Delta^{\dagger}$	Before	After	$\Delta^{\dagger}$	p-value	Adjusted p-value <sup>†</sup>
ISd								
Child characteristics	59.8±7.5	58.3±7.9	-1.6±5.2	46.4±9.6	$44.8 \pm 10.2$	-1.7±5.3	0.934	>0.999
Parent characteristics	$54.1\pm 9.2$	54.1±7.8	$0.1\pm 5.3$	49.0±9.2	$48.2 \pm 10.6$	-0.8±6.4	0.545	>0.999
Caregiver CESD-R	$11.0\pm 12.6$	$10.1 \pm 11.1$	-0.9±9.9	7.9±7.8	6.4±6.9	-1.5±7.1*	0.307	0.307
SOS								
Self-control Failure	6.7±2.4	7.1±2.7	$0.4\pm 2.0$	$5.5 \pm 1.8$	$5.2 \pm 1.8$	$-0.3\pm1.8$	0.290	0.869
Salience	7.4±2.9	7.8±2.5	$0.4\pm2.4^{*}$	6.3±2.9	6.3±2.5	$0.1\pm 2.0$	0.710	>0.999
Problematic consequences	5.2±2.7	5.6±2.2	$0.4\pm 2.1$	$4.8 \pm 2.0$	$5.2\pm 2.0$	$0.4\pm 1.3$	0.513	>0.999
Values are presented as mean±standard deviation. *adjusted p<0.05, significant difference observed between before and after values, <sup>†</sup> subscales were adjusted for multiple comparison using Bonferronis method. *with DD group (N=19), without DD group (N=53); <sup>b</sup> with DD group (N=51); <sup>w</sup> with DD group (N=53); <sup>w</sup> with DD group (N=22), without DD group (N=22), without DD group (N=22), without DD group (N=22), without DD group (N=21); <sup>w</sup> with DD group (N=22), <sup>w</sup> with DD group (N=22), <sup>w</sup> with DD group (N=21); <sup>w</sup> with DD gro	ard deviation. *adju p (N=19), without L V=2); *with DD grou	sted p<0.05, signifi DD group (N=53); <sup>b</sup> up (N=8), without	cant difference obs with DD group (N DD group (N=51);	erved between bef =17), without DD <sup>f</sup> with DD group (	fore and after value group (N=53); <sup>w</sup> ii N=9), without DD	s; <sup>†</sup> subscales were th DD group (N=2 group (N=47). DI	adjusted for mul (2), without DD (2), developmenta	tiple comparison using group (N=2); <sup>d</sup> with DD I disability; PEP-R DQ

anguage Scale; SELSI, Sequenced Language Scale for Infants; REVT, Receptive and Expressive Vocabulary Test; K-CARS, Korean version of the Childhood Autism Rating Scale; CBCL, Child Behavior Checklist, SRS, Social Responsiveness Scale; ABC, Aberrant Behavior Checklist, PSI, Parenting Stress Index; CESD-R, Center for Epidemiologic Studies Depression Scale-Re-

vised; SOS, Smartphone Overuse Scale

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tion, and gross motor declined in both the DoBrain group and the JNKK group, while SRS total score and behavioral problems as measured by the aggressive behavior subscale of CBCL, lethargy/social withdrawal and hyperactivity/noncompliance domains of ABC, and child characteristics domain of PSI improved only in the DoBrain group.

To our concern, we found that cognitive domains related to imitation, perception, and gross motor declined in the Do-Brain group and the JNKK group alike. Increased screen time may be one possible explanation for this observation. During the study period, smartphones and tablets were regularly used, and problematic screen use assessed by SOS was increased. Excessive screen time negatively affects social interaction,<sup>29</sup> physical activity,<sup>30</sup> and perception,<sup>31</sup> which can result in the decline of the imitation, gross motor, and perception domains of the PEP-R. More screen time can lead to less time engaged in other activities that promote normal development. As such, mobile-based cognitive training is not free from the negative effects of screen use. In the same context, a recent study that used computer-assisted intervention reported that increased program use was associated with worse receptive language outcomes.32 Further research is necessary to evaluate the detrimental effect of mobile-based cognitive training.

Contrary to our expectations, the effect of DoBrain on overall cognitive development was not superior to general educational apps. This result is in line with other studies that have questioned the effect of cognitive training in children.<sup>33,34</sup> While some studies proposed that cognitive training leads to cognitive improvement,<sup>35,36</sup> methodological issues should be taken into account, including the absence of an active control group and appropriate blinding. The strength of this study lies in the use of comparison with general educational apps, Junior Naver and Kakao Kids, which served as an active control. In addition, although double-blinding was not possible due to the nature of this study, the single-blinded design of the study would have minimized the assessor bias. In contrast to previous studies that included older children or adolescents,35-37 our sample included children with a mean age younger than 5 years, and may better represent the efficacy of mobile-based cognitive training in preschool children.

ASD symptoms measured by K-CARS showed a significant improvement in the DoBrain group than in the JNKK group. This difference between two groups was observed in a subset of children with DD, but no difference was observed in children with normal development. Furthermore, ASD symptoms measured by SRS showed a significant decline from baseline in the DoBrain group but not in the JNKK group. When a subgroup of children using DoBrain were assessed separately, the SRS total score showed a significant improvement in children with DD compared with those without. Existing evidence suggests that cognitive training may improve cognitive function, and thus may have beneficial effects on ASD symptoms.<sup>9</sup> However, in our study, the improvement of ASD symptoms was not accompanied by the improvement of cognitive function. Therefore, the improvements in ASD symptom observed in our study require careful interpretation.

Language skills were assessed by PRES, SELSI, and PRES, which yielded heterogeneous results. Compared with baseline, no significant differences were observed in the DoBrain group, whereas the JNKK group showed a significant decline in the SELSI receptive language domain and improvements in REVT receptive and expressive language domains. This inconsistent finding can be explained in part by the different dimensions of language that the two tests assess. SELSI assesses overall language abilities including semantics, phonology, syntax, and pragmatics through parental report, while REVT only assesses vocabulary abilities.<sup>20,21</sup> In this study, receptive and expressive vocabulary improved in the JNKK group, which is consistent with the characteristics of the JNKK app that present diverse vocabularies for practice. However, this improvement was confined to vocabulary. In contrast to the JNKK group, the DoBrain group did not show significant changes in language skills; the DoBrain app is more focused on problemsolving and reasoning, and thus minimal language stimuli are provided, which can be the reason for the overall null effect. In line with previous studies,<sup>38,39</sup> our results suggest that a welldesigned software that uses multimodal stimuli may be helpful in improving vocabulary skills.

Children in the DoBrain group showed significant improvements in behavioral problems as measured by the aggressive behavior subscale of CBCL, irritability, lethargy/social withdrawal, and hyperactivity/noncompliance domains of ABC, and child characteristics domain of PSI. In particular, irritability, lethargy/social withdrawal, and hyperactivity/ noncompliance domains of ABC showed significant improvement in children with DD, but not in children with normal development. This finding is in line with a recent meta-analysis in which behavioral measures showed significant improvements after using cognitive training programs.<sup>36</sup> Behavioral problems may have improved as a consequence of using DoBrain, considering that neural and behavioral plasticity is greater in younger children.<sup>40</sup> However, it should be taken into account that CBCL, ABC, and PSI are completed by caregivers of the children, and that placebo effects may also have played a part. In addition, children who spent more time using DoBrain may simply have had less time to engage in problematic behaviors.

We used the SOS to monitor problematic smartphone or tablet use, which is a possible side effect of a mobile-based cognitive training program. Compared with baseline, the problematic consequences subscale increased in the DoBrain group, and the salience subscale increased in the JNKK group. Although the total score did not exceed the cutoff point and the increase in the total score was less than one point, a significant increase suggests that monitoring the side effects of screen use is necessary. To the best of our knowledge, no other studies assessed the impact of mobile-based cognitive training programs on problematic screen use. Further studies should be performed while accounting for the changes in total screen time.

Several limitations of this study warrant discussion. First, the participants of the study were informed of which group they were allocated to. Parents, who completed the questionnaires, were aware of the group status, and this may have resulted in bias. In particular, questionnaires such as SRS, CBCL, ABC, PSI, and caregiver CESD-R rely on parental evaluation, and parents in the DoBrain group may report more positive outcomes. However, the JNKK group were active controls for this study, which may result in less placebo effect when compared with passive controls. Second, children with DD were diagnosed with heterogeneous disorders including ASD, ID, and LD, which exhibit distinct clinical characteristics. Therefore, the effect observed in children with DD as a whole may not be applicable to children with each disorder. Third, among the children who were lost to post-test, there were significantly more children who were diagnosed with ASD or ID. Although the majority of the reasons for follow-up loss were related to COVID-19, some of the children had difficulties in continuing the app. Therefore, there may be a bias in the sample included in the final analysis. Fourth, parents were instructed to help the children with the app usage, but the exact extent of parental help was not assessed, which may have an impact on study outcomes. Fifth, the study population was not sufficiently large to represent the general population. Sixth, for assessments that are not standardized for age, the effect of natural developmental trajectory on improvement must be considered. Finally, no passive control group was included in the study.

Despite these limitations, the strength of our study comes from a relatively large sample of children who were included in a randomized trial. The need for mobile-based cognitive training for children with or without DD is rapidly growing, with diverse apps being readily available. Our current results indicate that DoBrain has no superior effect over general educational apps on overall cognitive development. In the domains of cognitive development that are frequently practiced inside the DoBrain app including eye-hand coordination, possible improvement was implicated in a subset of children with DD, which requires careful interpretation. DoBrain was more effective in reducing ASD symptoms compared with the general educational apps, but the effect size was small. Further study with a better-designed, personalized mobilebased cognitive training app is required to verify its effect on cognitive development. In addition, careful monitoring of the detrimental effect of mobile-based cognitive training on specific domains of development is necessary.

#### **Supplementary Materials**

The online-only Data Supplement is available with this article at https://doi.org/10.30773/pi.2022.0136.

# Availability of Data and Material

The datasets generated or analyzed during the study are available from the corresponding author on reasonable request.

## **Conflicts of Interest**

The authors have no potential conflicts of interest to disclose.

#### **Author Contributions**

Conceptualization: Hyo-Won Kim. Data curation: Taeyeop Lee, Jichul Kim, Kee Jeong Park. Formal analysis: Seonok Kim, Taeyeop Lee. Funding acquisition: Hyo-Won Kim. Methodology: Taeyeop Lee, Seonok Kim. Project administration: Hyo-Won Kim. Resources: Hyo-Won Kim. Software: Taeyeop Lee, Seonok Kim. Supervision: Hyo-Won Kim. Validation: Hyo-Won Kim. Visualization: Taeyeop Lee, Seonok Kim. Writing—original draft: Taeyeop Lee. Writing—review & editing: Hyo-Won Kim.

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#### **Funding Statement**

This work was supported by the National Research Foundation of Korea (NRF) grant funded by the Ministry of Science and ICT (NRF-2020R1A5A8017671).

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Mariahla	D	oBrain (N=3	32)	J	INKK (N=30	))	Group	difference in $\Delta$
Variable	Before	After	$\Delta^{\dagger}$	Before	After	$\Delta^{\dagger}$	p-value	Adjusted p-value <sup>†</sup>
PEP-R DQ	53.7±31.9	60.2±26.9	6.5±18.9	50.2±25.7	50.1±28.9	-0.1±15.7	0.024	0.194
Imitation	50.2±35.0	52.5±28.8	2.3±26.8	46.4±26.1	45.4±29.3	-1.0±19.0	0.593	>0.999
Perception	62.3±34.8	62.7±28.5	0.4±24.9	61.9±31.8	52.3±31.7	-9.7±27.4*	0.058	0.460
Fine motor	59.3±26.3	63.1±21.8	3.9±22.6	62.7±30.5	61.2±27.0	-1.5±19.3	0.330	>0.999
Gross motor	78.8±29.5	79.7±21.6	$0.9 \pm 23.1$	73.2±29.8	69.7±30.1	-3.5±28.0	0.914	>0.999
Eye-hand coordination	69.1±24.6	76.4±21.2	7.3±17.6	68.9±26.8	66.4±22.8	-2.5±15.9	0.026	0.205
Cognitive performance	58.1±36.5	66.4±28.3	8.3±24.8	52.3±27.0	52.5±31.8	0.2±18.9	0.063	0.508
Cognitive verbal	56.7±28.4	59.7±27.2	3.0±16.5	48.4±21.7	50.6±26.1	2.3±14.2	0.778	>0.999
VABS ABCS	61.5±13.9	63.6±14.8	2.1±7.4	59.4±14.4	59.4±15.5	0.1±6.4	0.253	>0.999
Communication	62.8±15.2	66.0±19.4	3.3±10.5	60.7±17.8	60.9±17.2	0.2±11.4	0.480	>0.999
Daily living skills	74.8±13.2	74.5±17.4	-0.4±12.8	75.1±15.9	71.3±16.7	-3.8±6.7*	0.026	0.131
Socialization	60.6±16.4	60.1±17.3	-0.6±9.9	60.8±15.5	59.2±17.6	-1.7±8.5	0.413	>0.999
Motor skills	75.1±14.3	75.8±15.4	0.7±11.8	72.0±12.6	73.6±12.6	1.6±8.4	0.826	>0.999
K-CARS	30.7±7.3	29.6±7.1	-1.1±3.5	28.2±8.4	29.6±8.1	1.4±3.2*	0.005	0.005
SRS total score	66.1±15.1	61.5±12.2	-4.6±8.6*	64.6±14.3	61.5±12.5	-3.1±7.4*	0.352	0.352
CBCL								
Emotionally reactive	54.8±6.2	54.2±4.6	-0.6±5.0	53.9±5.3	53.3±4.6	-0.6±4.5	0.749	>0.999
Anxious/depressed	53.8±4.9	53.0±5.2	-0.8±4.1	52.8±5.1	53.2±5.0	0.4±6.4	0.048	0.339
Somatic complaints	54.8±5.9	54.3±5.6	-0.5±5.0	52.9±4.7	53.4±5.3	0.5±4.3	0.575	>0.999
Withdrawn	64.3±6.6	61.8±7.2	-2.5±6.3	65.4±11.0	63.2±11.3	-2.2±7.1	0.782	>0.999
Sleep problems	56.3±6.4	53.7±6.0	-2.6±5.7	54.6±6.6	55.0±7.9	0.4±6.9	0.050	0.352
Attention problems	62.4±8.0	60.3±9.1	-2.1±6.8	60.1±8.0	60.0±8.3	-0.1±6.8	0.238	>0.999
Aggressive behavior	57.1±7.9	54.6±6.2	-2.4±6.3	55.7±6.5	53.7±6.4	-2.1±6.7	0.738	>0.999
ABC								
Irritability	7.3±6.6	4.3±5.0	-3.0±4.5*	6.5±6.8	5.0±6.5	-1.6±3.8	0.130	0.651
Lethargy/social withdrawal	8.8±6.1	6.2±5.1	-2.7±4.7*	8.4±7.2	7.5±7.9	-1.0±4.2	0.060	0.302
Sterotypic behavior	3.2±4.0	2.1±3.0	-1.1±2.7	2.2±3.6	2.2±3.4	0.0±1.3	0.071	0.354
Hyperactivity/noncompliance	8.1±5.4	5.7±4.3	-2.4±4.3*	6.1±4.8	5.2±5.2	-0.9±4.2	0.152	0.761
Inappropriate speech	2.6±2.2	$1.9{\pm}2.0$	-0.6±2.1	1.7±1.8	$1.8 \pm 1.9$	0.0±1.9	0.270	>0.999
PSI								
Child characteristics	59.8±7.5	58.3±7.9	-1.6±5.2	59.8±11.3	58.3±12.3	-1.5±6.4	0.979	>0.999
Parent characteristics	54.1±9.2	54.1±7.8	0.1±5.3	56.5±9.5	55.3±11.9	-1.2±6.1	0.411	>0.999
Caregiver CESD-R	11.0±12.6	10.1±11.1	-0.9±9.9	14.7±13.5	10.5±14.0	-4.1±11.6*	0.074	0.074
SOS								
Self-control failure	6.7±2.4	7.1±2.7	$0.4 \pm 2.0$	6.6±1.9	7.1±2.3	0.4±2.0	0.908	>0.999
Salience	7.4±2.9	7.8±2.5	0.4±2.4*	6.9±3.0	8.1±2.8	1.2±2.4*	0.177	0.530
Problematic consequences	5.2±2.7	5.6±2.2	$0.4{\pm}2.1$	5.5±2.5	6.0±2.3	0.5±1.9	0.885	>0.999

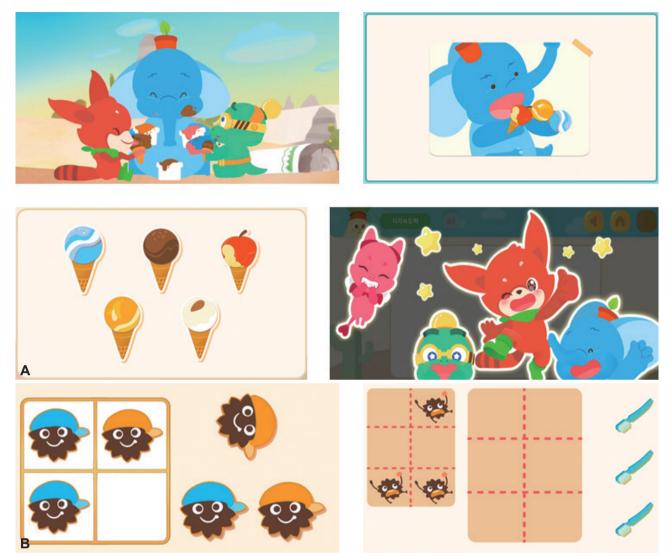
Supplementary Table 1. Comparison of efficacy of DoBrain and general educational apps in children with developmental disabilities

Values are presented as mean±standard deviation. \*adjusted p<0.05, significant difference observed between before and after values; <sup>†</sup>subscales were adjusted for multiple comparison using Bonferroni's method. PEP-R DQ, Psychoeducational Profile-Revised Developmental Quotient; VABS ABCS, Vineland Adaptive Behavior Scale Adaptive Behavior Composite Score; K-CARS, Korean version of the Childhood Autism Rating Scale; SRS, Social Responsiveness Scale; CBCL, Child Behavior Checklist; ABC, Aberrant Behavior Checklist; PSI, Parenting Stress Index; CESD-R, Center for Epidemiologic Studies Depression Scale-Revised; SOS, Smartphone Overuse Scale; JNKK, Junior Naver and Kakao Kids

Supplementary Table 2. Comparison of et	fficacy of DoBrain and	general educational apps in childre	n with normal development
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Variable	Γ	0oBrain (N=5	(3)		JNKK (N=51	)	Group	difference in $\Delta$
variable	Before	After	$\Delta^{\dagger}$	Before	After	$\Delta^{\dagger}$	p-value	Adjusted p-value <sup>†</sup>
PEP-R DQ	106.6±12.4	107.0±11.6	0.3±9.9	108.9±10.8	106.5±15.6	-2.4±12.5	0.725	>0.999
Imitation	104.2±19.0	98.5±15.2	-5.6±15.5*	108.0±16.9	99.9±18.1	-8.1±18.4*	0.495	>0.999
Perception	104.2±19.6	95.9±17.4	-8.3±13.2*	112.2±18.2	102.3±18.8	-9.9±16.2*	0.578	>0.999
Fine motor	104.2±13.6	100.6±14.3	-3.5±13.7	107.5±13.0	102.4±16.2	-5.1±17.6	0.304	>0.999
Gross motor	104.8±15.9	99.0±16.1	-5.8±8.9*	114.2±15.8	103.3±21.5	-10.9±17.3*	0.023	0.186
Eye-hand coordination	106.4±13.5	105.9±13.1	-0.5±13.7	109.3±13.8	105.1±12.0	-4.2±13.4	0.304	>0.999
Cognitive performance	111.4±11.6	111.3±14.2	-0.2±11.1	116.1±12.4	111.7±17.6	-4.4±19.1	0.282	>0.999
Cognitive verbal	103.1±16.1	104.8±14.5	1.7±12.2	103.9±15.3	103.8±17.2	-0.1±14.7	0.738	>0.999
VABS ABCS	92.2±13.8	96.2±13.1	4.0±9.4*	92.8±12.7	94.5±11.5	1.7±10.6	0.256	>0.999
Communication	92.1±14.9	98.4±15.0	6.3±12.0*	93.1±12.7	98.3±13.8	5.2±10.6*	0.613	>0.999
Daily living skills	97.0±14.6	99.8±12.5	2.8±12.0	99.2±10.7	99.5±10.5	0.3±11.7	0.210	>0.999
Socialization	95.1±16.9	98.5±12.1	3.4±15.2	93.9±14.6	95.6±13.0	1.7±14.4	0.562	>0.999
Motor skills	91.8±15.1	95.5±12.0	3.7±14.3	93.8±11.0	93.9±12.0	0.1±9.9	0.275	>0.999
K-CARS	15.9±1.4	15.9±1.8	0.0±1.5	16.3±1.5	16.1±1.6	-0.2±1.6	0.949	0.949
SRS total score	46.0±7.9	46.3±7.8	0.2±7.9	45.9±7.7	45.5±6.6	-0.5±5.0	0.590	0.590
CBCL								
Emotionally reactive	52.6±4.0	52.3±5.3	-0.3±3.9	52.1±3.8	52.1±4.1	0.1±4.7	0.225	>0.999
Anxious/depressed	52.5±5.3	52.4±5.7	-0.1±2.7	51.4±3.4	51.6±4.2	0.2±5.2	0.702	>0.999
Somatic complaints	52.8±4.5	52.1±3.8	-0.7±4.5	51.4±3.2	52.2±4.8	0.8±5.1	0.191	>0.999
Withdrawn	52.6±5.4	52.1±5.2	-0.6±3.8	$51.8 \pm 4.1$	51.8±4.2	0.0±4.9	0.209	>0.999
Sleep problems	52.9±4.6	52.9±4.5	0.0±5.0	54.4±7.2	53.2±7.0	-1.2±5.6	0.303	>0.999
Attention problems	52.3±3.8	52.1±4.3	-0.2±3.3	52.7±4.5	52.6±5.1	-0.1±4.5	0.847	>0.999
Aggressive behavior	52.0±3.7	51.2±3.5	-0.8±3.5	51.7±3.7	52.2±6.1	0.5±4.6	0.162	>0.999
ABC								
Irritability	1.9±2.3	1.5±2.3	-0.5±2.9	2.0±3.1	1.7±3.3	-0.4±2.6	0.858	>0.999
Lethargy/social withdrawal	0.9±2.7	0.9±2.9	0.0±3.8	0.8±1.9	$0.7 \pm 1.8$	-0.1±1.9	0.921	>0.999
Sterotypic behavior	$0.1 \pm 0.4$	0.3±1.3	0.2±1.1	0.2±0.5	0.1±0.5	-0.1±0.7	0.088	0.439
Hyperactivity/noncompliance	1.4±1.6	1.1±2.0	-0.3±2.0	$1.8 \pm 2.7$	1.6±3.4	-0.2±2.4	0.964	>0.999
Inappropriate speech	0.3±0.6	0.3±0.7	0.0±0.9	0.5±0.9	$0.4{\pm}0.8$	-0.1±0.8	0.595	>0.999
PSI								
Child characteristics	46.4±9.6	44.8±10.2	-1.7±5.3	46.8±9.7	46.7±11.1	-0.1±6.9	0.210	0.421
Parent characteristics	49.0±9.2	48.2±10.6	-0.8±6.4	48.1±9.7	49.2±10.1	1.1±7.5	0.259	0.518
Caregiver CESD-R	7.9±7.8	6.4±6.9	-1.5±7.1*	6.9±8.5	6.4±9.3	-0.5±9.7	0.799	0.799
SOS								
Self-control failure	5.5±1.8	5.2±1.8	-0.3±1.8	5.4±1.9	5.5±1.5	0.1±1.9	0.237	0.711
Salience	6.3±2.9	6.3±2.5	0.1±2.0	5.4±2.0	5.9±2.0	0.6±2.0	0.136	0.409
Problematic consequences	4.8±2.0	5.2±2.0	0.4±1.3	4.3±1.6	4.8±1.6	0.5±1.9	0.646	1.939

Values are presented as mean±standard deviation. \*adjusted p<0.05, significant difference observed between before and after values; †subscales were adjusted for multiple comparison using Bonferroni's method. PEP-R DQ, Psychoeducational Profile-Revised Developmental Quotient; VABS ABCS, Vineland Adaptive Behavior Scale Adaptive Behavior Composite Score; K-CARS, Korean version of the Childhood Autism Rating Scale; SRS, Social Responsiveness Scale; CBCL, Child Behavior Checklist; ABC, Aberrant Behavior Checklist; PSI, Parenting Stress Index; CESD-R, Center for Epidemiologic Studies Depression Scale-Revised; SOS, Smartphone Overuse Scale; JNKK, Junior Naver and Kakao Kids



Supplementary Figure 1. Screenshot of the DoBrain app.