Cortical Thickness in Children Receiving Intensive Therapy for Idiopathic Apraxia of Speech

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Abstract Children with idiopathic apraxia experience difficulties planning the movements necessary for intelligible speech. There is increasing evidence that targeted early interventions, such as Prompts for Restructuring Oral Muscular Phonetic Targets (PROMPT), can be effective in treating these disorders. In this study, we investigate possible cortical thickness correlates of idiopathic apraxia of speech in childhood, and changes associated with participation in an 8-week block of PROMPT therapy. We found that children with idiopathic apraxia \((n = 11)\), aged 3–6 years, had significantly thicker left supramarginal gyri than a group of typically-developing age-matched controls \((n = 11)\), \(t(20) = 2.84, p \leq 0.05\). Over the course of therapy, the children with apraxia \((n = 9)\) experienced significant thinning of the left posterior superior temporal gyrus (canonical Wernicke’s area), \(t(8) = 2.42, p \leq 0.05\). This is the first study to demonstrate experience-dependent structural plasticity in children receiving therapy for speech sound disorders.

Keywords Motor speech disorder · Childhood apraxia of speech (CAS) · Plasticity · Supramarginal · Wernicke · MRI

Idiopathic apraxia, a form of Speech Sound Disorder (SSD), is broadly characterized as deficits in the planning of movements necessary for intelligible speech. Children with apraxia of speech have difficulty producing target phonemes, either in isolation or succession, and typically present with inaccurate word production and dysfluency. Childhood apraxias emerge in the first years of life, and have the potential to negatively impact later language, academic, and social skills development. Unlike dysarthria, which is generally
considered a neuromuscular disorder, apraxia may or may not be associated with neurological insult (see, American Speech-Language-Hearing Association 2007a, b, for a discussion on childhood apraxia of speech, CAS, and possible presentations and diagnostic challenges). In their recent literature review (Lieggeois and Morgan 2012) found no evidence for unilateral lesions leading to apraxia in children; in the few cases of neurological insult associated with CAS, bilateral abnormalities in brain regions known to support language processing (i.e., perisylvian cortex) and motor control (basal ganglia, Rolandic cortex) were documented. Idiopathic apraxia of speech is among the most common SSDs (Tyler2008; see also, Gierut 1998; Law et al. 2004). One approach, known as Prompts for Restructuring Oral Muscular Phonetic Targets (PROMPT), has been shown to be effective in treating childhood SSDs (Tyler 2008; see also, Gierut 1998; Law et al. 2004). One approach, known as Prompts for Restructuring Oral Muscular Phonetic Targets (PROMPT), has been shown to be effective in treating CAS and other motor speech disorders (Chumpelik 1984; Grigos et al. 2010; Hayden 2006; Namasivayam et al. 2013). During PROMPT sessions, therapists provide direct tactile and kinesthetic cues, in addition to auditory and visual cues, to promote correct speech production. Clients practice requisite oral-motor positions and trajectories in words and phrases, gaining experience and familiarity with target sound production. Although researchers have begun to document the efficacy of targeted interventions such as PROMPT (e.g., Chumpelik 1984; Grigos et al. 2010; Hayden 2006; Namasivayam et al. 2013), the mechanisms for change and the neural correlates of participation in therapy remain unknown.

Recent advances in MRI technology and analyses permit quantitative and objective study of gross brain structure, which can be used to compare groups or to characterize neuroanatomical change over time. In their seminal study, (Maguire et al. 2000) found that London taxi drivers had larger posterior and smaller anterior hippocampi than age-matched controls, a difference thought to reflect the relative navigational experience of the two groups. Indeed, regional differences were associated with time spent driving a taxi, providing compelling evidence for experience-dependent structural plasticity in the mature hippocampus. Others have shown that experience-dependent structural changes are not limited to deep structures. Using voxel-based morphometry (VBM), (Draganski et al. 2004) documented increased grey matter density in posterior temporal and inferior parietal regions in a group of adults learning to juggle. Cortical growth, although transient, occurred in regions known to support visual processing of moving objects. Using similar approaches, others have documented visual spatial or navigational training-induced effects on cortical structure (Ilg et al. 2008; Wenger et al. 2012).

Increasingly, researchers are documenting training-induced structural changes in pediatric populations. Hyde et al. (2009) used deformation-based morphometry (DBM) to compare the brains of healthy children, 6 years of age, who received instrumental musical training with a cohort who did not. At baseline (prior to first music lessons), no structural differences were observed. Over the course of 15 months, however, both cortical and subcortical effects emerged: the group receiving musical training showed morphometric differences in several areas, including increased volume in the right precentral gyrus, right Heschl’s gyrus, and a mid region of the corpus callosum, consistent with previous reports of structural differences in musicians and non-musicians. Changes in cortical regions were correlated with performance on motor and melodic/rhythmic discrimination tasks, supporting the argument that musical training can drive structural change. Others have documented cortical volume changes in children receiving behavioral interventions for ADHD (Hoekzema et al. 2011), and dyslexia (Krafnik et al. 2011; see also Gebauer et al. 2012; Keller and Just 2009). Quantitative neuroanatomical investigations suggest a massive potential for structural plasticity in pediatric therapeutic contexts.

The goals of this study are to (1) assess cortical thickness correlates of idiopathic verbal apraxia in childhood, and (2) characterize changes in cortical thickness associated with participation in PROMPT therapy. We chose vertex-based thickness analyses over conventional VBM and DBM, as the approach shows relative sensitivity to subtle cortical effects (see Hutton et al. 2009; Scanlon et al. 2011). To assess structural correlates of the disorder and intervention, we collected high-resolution MR images from 14 young children referred for treatment of verbal apraxia, and a group of 14 typically developing controls. The apraxia group then received 8 weeks of PROMPT therapy, before returning for follow-up scans. The children with apraxia had presumably idiopathic disorders; however, in the absence of obvious structural abnormalities or lesions, we expect subtle cortical differences in regions known to support speech and language (see Liegeois and Morgan 2012). In children receiving therapy, we expect gains to be reflected in cortical thickness change in canonical language areas and sensory-motor regions (i.e., perisylvian and Rolandic cortices).

Method

Participants

Fourteen children with idiopathic apraxia of speech, and 14 typically developing children (Controls) participated
in this study. Children with apraxia were recruited from a large pool of children referred to The Speech and Stuttering Institute (Toronto, ON, Canada) for treatment of speech sound disorders. The apraxia group consisted of 9 males and 5 females, aged 3.9–6.6 years (M = 4.5, SD = 0.8) with confirmed moderate to severe speech difficulties, characterized by focal motor planning deficits (apraxia), in the absence of known neurological disorders, neuromuscular deficits (dysarthria), or hearing problems. Children in the apraxia group completed baseline, intervention, and follow-up components of the study over a 10-week period. Baseline testing involved brief cognitive assessment, comprehensive speech language assessment, and neuroimaging. Within 1 week of completing baseline assessments, the apraxia group began an 8-week block (16 sessions total) of PROMPT therapy (in all cases, therapy was provided without charge to children and their families). Within 1 week of completing PROMPT, participants with apraxia completed a repeat speech language assessment and neuroimaging.

The Control group consisted of 8 males and 6 females, 4.1–6.3 years of age (M = 4.1, SD = 0.7). Controls were recruited from the community, and were negative for history of developmental delay, neurological disorder, hearing problems, and speech language impairment. Controls underwent baseline cognitive assessment and neuroimaging, but did not receive speech assessments or any speech training. All Controls were invited to return for repeat neuroimaging after 10 weeks; however, only a small subset (n = 4) were available at the required interval. Demographic information for both groups is presented in Table 1.

Speech assessments and therapy were conducted at The Speech and Stuttering Institute; cognitive assessments and neuroimaging were carried out at the Hospital for Sick Children (Toronto, ON). The study was approved by the Hospital’s Research Ethics Board.

### Procedure

#### Cognitive Assessment

To estimate gross cognitive functioning, both groups underwent brief baseline assessment with the Peabody Picture Vocabulary Test (PPVT; Dunn and Dunn 2007), Expressive Vocabulary Test (EVT; Williams 2007), and the Wechsler Nonverbal Scale of Ability (WNV; Wechsler and Naglieri 2006). These are standardized tests of receptive language, expressive language, and nonverbal functioning, respectively. Both groups scored within normal limits across all measures (see Table 1); for the apraxia group, intact gross language functioning suggests a focality of speech-motor deficit.

### Table 1 Demographic and neuropsychological profile of participants

|                        | Apraxia mean (SD) | Control mean (SD) | Between-groups t-test |
|------------------------|-------------------|-------------------|-----------------------|
| Age in years           | 4.54 (0.83)       | 4.95 (0.72)       | t(26) = 1.4, p > 0.05  |
| Handedness             |                   |                   |                       |
| EHI                    | 52.53 (69.95)     | 60.43 (42.44)     | t(26) = 0.4, p > 0.05  |
| Receptive language     |                   |                   |                       |
| PPVT—z score           | 0.27 (0.94)       | 1.16 (0.91)       | t(26) = 2.5, p ≤ 0.05  |
| Expressive Language    |                   |                   |                       |
| EVT—z score            | 0.56 (0.78)       | 1.14 (0.88)       | t(25) = 1.8, p > 0.05  |
| Nonverbal Functioning  |                   |                   |                       |
| WNV, 2-subtest FSIQ estimate, z score | −0.10 (1.17) | 0.12 (1.09) | t(26) = 1.7, p > 0.05  |

EHIT Edinburgh handedness inventory, PPVT Peabody picture vocabulary test, EVT expressive vocabulary test, WNV Wechsler nonverbal scale of intelligence
the clinician would model production of the word “pop”, and then guide the child’s mandibular movement using their thumb and index finger while the child repeatedly attempts “pop” production. In this study, PROMPT was provided by the same therapist (author DG); although participants present with variable speech production errors, therapy followed a consistent routine (Namasivayam et al. 2013), with the common goal of promoting development of new and stable motor programs. Between PROMPT sessions, the children worked with their parents to practice prescribed daily home-based activities for approximately 5–10 min per day (e.g., parent reminds child to keep mouth opening small and jaw in midline while producing the word “pop”, and then plays game involving blowing and catching bubbles and saying “pop”).

Speech Assessment

Children in the apraxia group underwent comprehensive speech assessments before and after speech therapy. We report on performance on three widely-used standardized devices (see below). Assessments were videotaped and speech samples collected (16-bit 44.1 kHz recordings) for analyses. Performance of individuals contributing both pre- and post-intervention scans is presented in Supplementary Table 1.

Goldman-Fristoe Test of Articulation 2 (GFTA-2; Goldman and Fristoe 2000)

The GFTA-2 is used to assess articulation in English speakers between the ages of 2 and 21 years. Word production is analyzed in initial, medial, and final positions, along with consonant blends in the initial position. For this study, we report on performance on the Sounds-in-Words subtest; raw scores are analyzed for therapy-induced change.

Hodson Computerized Analysis of Phonological Patterns (HCAPP; Hodson 2003)

The HCAPP is used to systematically measure phonological deviations in speech. Using 50 target words, errors are quantified along several dimensions. We report on Total Occurrence of Major Phonological Deviations.

Verbal Motor Production Assessment for Children (VMPAC; Hayden and Square 1999)

The VMPAC tests neuromotor integrity for speech in children ages 3–12 years. Scores on the Focal Oromotor Control and Sequencing subtests are reported as percentage correct values.

MRI Acquisition and Processing

T1-weighted magnetization prepared rapid gradient echo (MPRAGE) images (sagittal, 3D; TR/TE/TI = 2300/2.96/900 ms, respectively; flip angle = 9°) were acquired at 1.0 × 1.0 × 1.0 mm resolution using a Siemens Trio 3T scanner (Siemens Aktiengesellschaft, Munich, Germany). Images were submitted to CIVET version 1.1.10, an automated processing pipeline, involving non-uniformity correction (Sled et al. 1998), stereotactic registration (Collins et al. 1994), skull stripping (Smith 2002), and tissue classification (Tohka et al. 2004). Inner and outer cortical surfaces were extracted for vertex-based analyses (Kim et al. 2005; Lerch and Evans 2005). Cortical thickness was calculated as the difference between linked inner and outer surface vertices (40,962 pairs per hemisphere), smoothed using a 20 mm surface-based kernel (Lerch and Evans 2005).

Regions-of-Interest (ROIs)

To minimize the number of comparisons, we confined our analyses to areas known to support language, speech, and voluntary oral-motor control. We developed discreet ROIs for canonical Broca’s area (left pars opercularis) and Wernicke’s area (the superior temporal gyrus posterior to Heschl’s convolutions) and their right hemisphere homologues, neighboring gyri of the frontal and temporal lobes, and inferior aspects of the pre- and post-central gyri. Because the temporal-parietal junction is frequently included as an extracanonical region contributing to Wernicke’s area (see Bogen and Bogen 1976), we also developed bilateral ROIs for the posterior half of the supramarginal gyrus, inferior to the ascending ramus of the Sylvian fissure. See Supplementary Fig. 1 online. ROIs were drawn, extending to the depths of sulci, on an average surface of all subjects’ scans after non-linear surface-based registration (Lyttelton et al. 2007). Cortical thickness maps, computed in native space for each scan, were then brought into average space using the same surface deformation field. The transformed thickness maps were then used to compute mean thickness within each ROI for each subject, for each scan visit. In order to minimize variability and focus on local effects, cortical thickness maps were scaled by dividing the thickness at each vertex pair by the mean thickness for the respective hemisphere; mean scaled thickness for each ROI was used in both between-groups and within-subjects analyses.

Statistical Analyses

To assess correlates of the disorder, we compared the apraxia and Control groups for baseline scaled cortical
thickening at each ROI using independent samples t-tests. For any region showing significant group differences, we compared baseline speech performance (GFTA-2, HCAPP, and VMPAC scales) to scaled cortical thickness in the apraxia group using bivariate Pearson correlations. To assess correlates of therapy, we analyzed pre- to post-intervention thickness changes occurring at each ROI in the clinical group using within-subjects t-tests. For any region showing significant within-subjects effects, we compared changes in speech performance (GFTA-2, HCAPP, and VMPAC scales) to changes in scaled cortical thickness using bivariate Pearson correlations, and describe changes occurring in the small subset of Controls who were scanned serially. Due to the limited sample and exploratory nature of this study, we did not correct for multiple comparisons across ROIs.

Results
Efficacy of PROMPT Therapy
In all cases, parents of children with idiopathic verbal apraxia reported that PROMPT therapy was beneficial for their children. Two children in the apraxia group did not return for follow-up speech assessment or MRI, due to anxiety experienced during baseline assessment; we report on the remaining 12 children with apraxia, who showed significant (p < 0.05) gains on all speech measures. Change scores and within-subjects statistics are presented in Table 2.

Cortical Thickness Correlates of Apraxia
Of the 28 children in this study, 11 children with apraxia (8 males, mean age 4.7 years) and 11 Controls (5 males, mean age 4.8 years) had baseline MRIs suitable for analyses (i.e., with cortical morphology retained after segmentation; rejected scans showed obvious movement-related artifact). Suitability was determined by agreement of two independent raters, blind to group membership.

No significant between-groups differences were observed for overall cortical thickness or mean thickness within each hemisphere (t-tests, p > 0.05). Only one significant between-groups difference in ROI analyses was observed: children with idiopathic apraxia had significantly thicker left posterior supramarginal gyrus than Controls, t(20) = 2.84, p ≤ 0.05. See Fig. 1. Left posterior supramarginal gyrus thickness did not correlate with any of the baseline measures of speech performance in the clinical group, p > 0.05.

Cortical Thickness Changes Following PROMPT
Nine children (6 male; mean age 4.5 years) with mild apraxia had pre- and post-intervention MRIs that were suitable for cortical thickness analyses. Of the four Controls studied at follow-up, three (2 male; mean age 5.5) had suitable baseline and follow-up scans; due to the small number with repeat neuroimaging, we excluded Controls from formal longitudinal analyses, and provide only a description of change for informal comparison with the group receiving intervention.

In children receiving therapy for apraxia, ROI analyses revealed significant thinning in the posterior superior temporal gyrus, canonical Wernicke’s area, t(8) = 2.42, p ≤ 0.05. See Fig. 2. While 8 of the 9 children with apraxia showed thinning in Wernicke’s following therapy, only 1 of the 3 Controls showed thinning over the same period. Decreasing thickness in Wernicke’s over the course of therapy was not significantly correlated to change scores on any of the standardized speech measures, p > 0.05.

Discussion
This is the first study to investigate cortical thickness correlates of idiopathic apraxia of speech and changes in cortical thickness associated with brief intensive speech therapy in children. We found that children with apraxia had thicker left supramarginal gyrus than Controls. In the absence of appreciable lesions, the quantitative neuroanatomical approach reveals a subtle morphological atypicality associated with motor speech deficits. The study also provides support for the benefits of PROMPT and speech motor intervention in children. Over the course of therapy, children with apraxia experienced thinning of the left posterior superior temporal gyrus (Wernicke’s area). This focal change evidences a potential for rapid and robust experience-dependent structural plasticity in childhood.

Table 2 Changes in speech performance following PROMPT therapy

| Measure                        | Magnitude of mean difference | Within-subjects t-test |
|--------------------------------|-----------------------------|------------------------|
| GFTA-2 sounds—words—raw score  | 9.1                         | (11) = 3.6, p = 0.004  |
| HCAPP Phonological deviations—total | 39.7                       | (11) = 7.3, p ≤ 0.001  |
| VMPAC focal motor control—% correct | 11.5                       | (11) = 3.7, p = 0.004  |
| VMPAC sequencing—% correct    | 7.7                         | (11) = 3.1, p = 0.010  |

GFTA Goldman Fristoe test of articulation, VMPAC Verbal motor production assessment for children, HCAPP Hodson computerized analysis of phonological patterns
The clinical significance of thicker left supramarginal gyri in children with idiopathic apraxia is not clear. The effect is apparent only at the group level, and thickness in the region does not correlate to degree of speech impairment. A thicker left supramarginal gyrus may reflect a subtle pathology; if sustained throughout childhood, a thick left supramarginal gyrus may indicate immaturity or altered development, as the region is expected to overgrow in the first years of life and prune back in childhood and early adolescence (Shaw et al. 2008; Sowell et al. 2004; see also, Porter et al. 2011). In adults, the left supramarginal gyrus is known to play an important role in speech production; injury to this region is associated with an apraxic presentation characterized by phonemic discrimination and speech planning deficits (for a recent review, see Gow 2012). However, the relationship between left supramarginal integrity and speech production in childhood has not been established. In their recent literature review, (Liegeois and Morgan 2012) found that childhood apraxia of speech and dysarthria were associated with bilateral, but not unilateral perisylvian insult. Our findings suggest that subtle unilateral atypicalities may underlie observed speech deficits in the absence of neurological injury.

Perhaps the most interesting finding in this study is that children with apraxia in our study experienced significant thinning of Wernicke’s, and all experienced speech gains. The small subset of Controls with serial imaging showed a tendency toward increasing thickness (2 of 3 participants) in the same area, over the same period. Given the brief interval between scans and the spatial specificity of changes, it is likely that
participation in the therapy program drove the observed
cortical thinning in the clinical group. With a longer block
of therapy, we may expect more robust changes and/or
involvement of additional regions, particularly those
known to support oral-motor control and expressive lan-
guage (i.e., Rolandic and inferior frontal cortex).

The location of between- and within-group effects
speaks to the deficits observed in idiopathic apraxia of
speech, and a possible mechanism for gains enjoyed by
participants receiving PROMPT. The left supramarginal
gyrus and neighboring posterior superior temporal gyrus
are each involved in sensorimotor integration, and are
necessary for accurate speech comprehension and produc-
tion (Gow 2012). Children with motor speech disorders
are known to have deficits in phonological processes (McNeill
et al. 2009; see also, Tkach et al. 2011), which are indi-
cally addressed in PROMPT therapy. During each session,
PROMPT therapists provide tactile and kinesthetic cues to
help clients produce target phonemes. Feedback is pro-
vided externally by the therapist (i.e., with touch and
speech, with the provision of auditory models), and inter-
naturally through somatosensory and auditory information.
Collectively, the multimodal and multisensory feedback
serves to reinforce the training. In PROMPT, development
of phonological processing skills provides an internal val-
dicator for subsequent articulation attempts. Wernicke’s
area undergoes thinning (possibly neuronal pruning) during
participation in PROMPT therapy, which may reflect the
rapid development of sensorimotor processes necessary for
accurate speech production.

One of the major challenges in conducting this sort of
research is obtaining homogenous clinical samples that are
sufficiently large for the study of subtle structural effects.
To minimize the number of comparisons conducted, we
confined our analyses to a small set of ROIs established on
theoretical grounds. We observed both between-groups and
within-subjects effects with modest samples, demonstrat-
ing a sensitivity of the quantitative neuroanatomical
approach; however, we may have lacked sufficient power
to fully characterize the structural correlates of idiopathic
verbal apraxia and PROMPT therapy. In the future, large-
scale studies of children with apraxia will permit whole-
brain analyses, potentially revealing relatively subtle or
focal correlates of the disorder or of therapy that are not
easily detected in relatively coarse ROI analyses. The
inclusion of protracted follow-up scans and behavioral
assessments will speak to the stability and functional rel-
ervance of the documented short-term changes. Future
studies will also benefit from the inclusion of a comparably
sized control group that is scanned serially. Longitudinal
study of control participants is required for distinction of
normal developmental changes and those occurring as a
result of intervention. This is particularly the case when
investigating structural change over long intervals (i.e.,
with sustained long-term speech therapy), as the pediatric
brain is known to undergo substantial morphological
change as part of normal development.

The quantitative neuroanatomical approach described in
this study can be easily implemented with other popula-
tions, including children with complex or neurogenic forms
of apraxia or dysarthria. Group analyses permit assessment
at a much finer scale than possible through reading of
individual scans. Obtaining high-quality MRIs of young
children without sedation can be challenging, but is worth
attempting, particularly for idiopathic disorders. There is
some evidence that the potential for experience-dependent
structural change decreases with advancing age (Wenger
et al. 2012); the study of pediatric populations over time, or
over the course of therapy, may be of relatively high yield.
In the current study, serial assessment of a modest sample
of children receiving PROMPT for idiopathic apraxia
informed of a possible mechanism and neural target for
therapeutic action—the intervention promoted develop-
ment of sensory-motor systems controlling speech pro-
duction, associated with thinning and possible maturation
of Wernicke’s area.

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References

American Speech-Language-Hearing Association (2007a) Childhood
apraxia of speech [Position Statement]. www.asha.org/policy
American Speech-Language-Hearing Association (2007b) Childhood
apraxia of speech [Position Statement]. www.asha.org/policy
Bogen JE, Bogen GM (1976) Wernicke’s region—where is it? Ann N
Y Acad Sci 280:834–843
Chumpelik DA (1984) The PROMPT system of therapy: theoretical
framework and applications for developmental apraxia of
speech. Semin Speech Lang 5(5):139–156
Collins DL, Neelin P, Peters TM, Evans AC (1994) Automatic 3D
intersubject registration of MR volumetric data in standardized
talairach space. J Comput Assist Tomogr 18(2):192–205
Dragninski B, Gaser C, Busch V, Schuierer G, Bogdahn U, May A
(2004) Neuroplasticity: changes in grey matter induced by
training. Nature 427(6972):311–312
Dunn LM, Dunn DM (2007) Peabody picture vocabulary test, 4th
dn. Pearson, San Antonio

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