Behavioral/Cognitive

Using IQ Discrepancy Scores To Examine the Neural Correlates of Specific Cognitive Abilities

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The underlying neural determinants of general intelligence have been studied intensively, and seem to derive from the anatomical and functional characteristics of a frontoparietal network. Little is known, however, about the underlying neural correlates of domain-specific cognitive abilities, the other factors hypothesized to explain individual performance on intelligence tests. Previous preliminary studies have suggested that spatially distinct neural structures do not support domain-specific cognitive abilities. To test whether differences between abilities that affect performance on verbal and performance tasks derive instead from the morphological features of a single anatomical network, we assessed in two independent samples of healthy human participants (N = 83 and N = 58; age range, 5–57 years) the correlation of cortical thickness with the magnitude of the verbal intelligence quotient (VIQ)-performance intelligence quotient (PIQ) discrepancy. We operationalized the VIQ-PIQ discrepancy by regressing VIQ onto PIQ (VIQ-regressed-on-PIQ score), and by regressing PIQ onto VIQ (PIQ-regressed-on-VIQ score). In both samples, a progressively thinner cortical mantle in anterior and posterior regions bilaterally was associated with progressively greater (more positive) VIQ-regressed-on-PIQ scores. A progressively thicker cortical mantle in anterior and posterior regions bilaterally was associated with progressively greater (more positive) PIQ-regressed-on-VIQ scores. Variation in cortical thickness in these regions accounted for a large portion of the overall variance in magnitude of the VIQ-PIQ discrepancy. The degree of hemispheric asymmetry in cortical thickness accounted for a much smaller but statistically significant portion of variance in VIQ-PIQ discrepancy.

Introduction

For over a century, psychologists have been interested in measuring individual intelligence. Modern intelligence quotient (IQ) tests are based either on a general model of intelligence (Wechsler, 2008) or on a multidimensional model of intelligence (Naglieri and Das, 1997). Neuroimaging studies have begun to investigate the neural determinants of intelligence. Most have focused on identifying the neural correlates of general intelligence (g) using the Wechsler scales of intelligence. The construct of general intelligence is validated by the positive intercorrelation of subtest scores in IQ tests, sometimes referred to as the “positive manifold” (Carroll, 1993; Colom and Thompson, 2011), and is thought to derive from the overall connectivity of multimodal association regions in the brain (Shaw et al., 2006; Jung and Haier, 2007; Narr et al., 2007; Karama et al., 2009). Lesion studies, however, demonstrate dissociable cognitive skills that are better explained by multidimensional models of intelligence. The neural determinants of these multiple dimensions, however, remain unknown.

Models of intelligence derive in large part from factor analyses, a mathematical technique that reduces correlational data to a smaller number of unobservable latent factors. The intercorrelation of performance on IQ subtests, shared variance, accounts for 50% of the total variance in performance. Roughly 70% of this shared variance represents g. The remaining 30% of variance is captured by factors representing domain-specific abilities (DSAs; Watkins, 2006). These DSAs vary with the differing IQ tests included in a factor analysis (Flanagan et al., 2000).

The few studies examining the neural correlates of DSAs have produced equivocal results (Andreasen et al., 1993; Gray et al., 2003; Witelson et al., 2006; Luders et al., 2007; Lange et al., 2010). In one study, g-adjusted verbal IQ (VIQ) and performance IQ (PIQ) scores did not correlate with cortical thickness (CT; Karama et al., 2011), suggesting that DSAs do not derive from spatially discrete, independent neural systems. The current study was designed to examine the neural systems that support intra-individual discrepancies in verbal- and performance-specific abilities. Following the methods used in recent studies (Karama et al., 2011), we regressed VIQ onto PIQ (VIQ-regressed-on-PIQ score) and PIQ onto VIQ (PIQ-regressed-on-VIQ score) and correlated the residual scores with CT. The shared variance between VIQ and PIQ represents variance associated with g, and the residual represents variance associated with domain-specific cognitive abilities and task-specific skills. The residual scores that we...
calculated index intraindividual variability in performance across VIQ- and PIQ-associated tasks. No other study, to our knowledge, has identified the neural correlates of the VIQ-PIQ discrepancy.

The VIQ-PIQ discrepancy may correlate with a single network of brain regions or with multiple networks. As prior studies did not identify discrete neural correlates of g-adjusted VIQ and PIQ, we did not expect the VIQ-PIQ discrepancy to depend on variation in multiple networks. We hypothesized instead that a single neural system underlies the imbalance between verbal and perceptual scores, and therefore that the VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores would correlate with CT in a single network of brain regions. A discrete network could explain discrepancies among DSAs by mediating competition between them, similar to explanations for the competition observed between memory systems in animals and humans (Packard, 2009). Moreover, detecting significant correlations in regions other than those associated with g would suggest that a neural system distinct from g governs the VIQ-PIQ discrepancy.

Materials and Methods

The Institutional Review Board at New York State Psychiatric Institute approved the studies from which these data are drawn. All parents gave consent for children to participate. Children gave their assent to participate. All participants were paid for their participation.

Participants

Two independent samples of healthy control individuals were included in this study. For both, we recruited healthy adults and children selected randomly from a telemarketing database. Recruitment methods for these studies have been reported previously (Goth et al., 2011). All participants were recruited as controls for imaging studies of adult and childhood neuropsychiatric disorders. Introductory letters were sent and then followed by screening telephone calls. A detailed clinical interview was performed that included the Kiddie-Schedule for Affective Disorders and Developmental Disorders (K-SADS) and the Structured Clinical Interview for the Diagnostic and Statistical Manual of Mental Disorders, fourth edition, Axis I Disorders (Spitzer et al., 1992) for adults. Two independent clinicians reviewed each file and rated each participant. Individuals with a current or past history of a psychiatric or neurological disorder were excluded.

Sample 1. This sample included 83 participants with a Full Scale IQ (FSIQ) of ≥80, who ranged in age from 5 to 54 years (mean age, 19.38 years; SD, 11.98 years). The sample was similarly distributed across gender (females, 40%) and was predominantly right handed (94%). Children under 18 years of age comprised 60% of the sample (Table 1).

Sample 2. This sample included 38 participants with an FSIQ of ≥80, who ranged in age from 7 to 57 years (mean age, 21.70 years; SD, 13.21 years). The sample was similarly distributed across gender (females, 50%) and was right handed (100%). Children under 18 years of age comprised 45% of the sample (Table 1).

Neuropsychological assessment

Because healthy participants were recruited as controls for several different studies across two university settings, intelligence testing was performed using different instruments depending on the research study protocol. Intelligence testing was performed in close temporal proximity to magnetic resonance imaging (MRI) scanning.

Sample 1. The majority of participants completed the Wechsler Abbreviated Scale of Intelligence (WASI; Wechsler, 1999) using the four-subtest format (n = 67). Sixteen children completed the Wechsler Intelligence Scale for Children, third edition (WISC-III; Wechsler, 1991). FSIQ correlations between the WASI and longer forms have been established at 0.87–0.92.

Sample 2. Fifteen children completed the WISC-III; abbreviated IQ indexes were estimated using established methods that correlate scores of ≥0.90 with scores from the full administration (Kaufman et al., 1996). The WISC-III four-subtest administration included Vocabulary and Information subtests to estimate VIQ, Block Design and Object Assembly to estimate PIQ, and the sum of these four tests to estimate Full Scale IQ. Thirty adults completed the Wechsler Adult Intelligence Scale (WAIS-R; Wechsler, 1981). Three children and two adults completed the WASI. Eight children completed the Kaufmann Brief Test of Intelligence (K-BIT; Kaufman and Kaufman, 1985). The K-BIT included Vocabulary for VIQ and Matrices for PIQ. Correlations of the K-BIT with WISC-III have been established at 0.89 (Canivez et al., 2005).

Although different instruments were used to calculate VIQ, PIQ, and FSIQ (in part, because of the wide age range of participants and, in part, because of differing study protocols), all instruments have been shown to correlate highly with one another and to test similar domains of cognitive function. Performance measures of the WASI (Block Design and Matrix Reasoning) correlate 0.60 and 0.54, respectively, with Perceptual Organization in the WISC-III, and 0.73 and 0.68, respectively, with the WAIS, third edition (WAIS-III; Kaufman et al., 2005). Verbal measures of the WASI (Vocabulary and Similarities) are included in the WISC-III and WAIS-III, and correlate 0.87 and 0.66, respectively, with the WISC-III and 0.90 and 0.75 with the WAIS-IIIVerbal Comprehension factor (Kaufman et al., 2005). We note that if all participants had been administered the same IQ test, measurement variance would have been reduced and our findings would likely have been stronger. That the findings were so strong and reproducible across datasets, despite the inclusion of some participants who were administered different IQ tests, is a testament to the robustness of our findings.

Residual scores

To create the VIQ-regressed-on-PIQ score, we regressed VIQ onto PIQ with an intercept of zero and saved the residual. To create the PIQ-regressed-on-VIQ score, we regressed PIQ onto VIQ with an intercept of zero and saved the residual.

Standardized difference scores

We assessed the correlation of CT with the VIQ-PIQ discrepancy score because discrepancy scores, as opposed to residual scores, are used frequently in clinical practice. Subtracting two standardized scores creates a difference score; the difference score will have an SE of difference (SEDiff) that is different from, and smaller than, the SD of the instrument. The SEDiff score is the SD of the difference score. SEDiff depends on the reliability of the scores being compared, in this case the IQ indices VIQ and PIQ. VIQ and PIQ have different reliabilities for each IQ instrument included in our study. Only a standardized discrepancy score makes discrepancy scores comparable across instruments. This standardized discrepancy score (z-score) is created using the unique
Cortical thickness.

IQ score distributions

The mean VIQ scores (112.7 in sample 1; 119.8 in sample 2) and PIQ scores (112.2 in sample 1; 118.6 in sample 2) in both samples were higher than in the standardization sample, wherein the mean was 100. Nevertheless, the mean VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores were essentially zero, and the distribution of the range of scores was close to the normal distribution. In samples 1 and 2, respectively, the correlations of VIQ with PIQ were 0.461 and 0.309; the correlations of (VIQ-PIQ)z with VIQ were 0.683 and 0.655; the correlations of (VIQ-PIQ)z with PIQ were −0.334 and −0.539; the correlations of VIQ-regressed-on-PIQ with VIQ were 0.684 and 0.639; the correlations of VIQ-regressed-on-PIQ with PIQ were −0.333 and −0.535; the correlations of PIQ-regressed-on-VIQ with VIQ were −0.672 and −0.626; and the correlations of PIQ-regressed-on-VIQ with PIQ were 0.347 and 0.549.

Image acquisition

Sample 1. Anatomical images were acquired on a 3 tesla whole-body scanner (Signa, General Electric) with a body transmitter coil and an eight-channel head receiver coil. High-resolution, T1-weighted brain images were acquired using a fast spoiled gradient recall sequence [inversion time, 500 ms; repetition time (TR), 4.7 ms; echo time (TE), 1.3 ms; field of view (FOV), 24 cm; image matrix, 256 × 256; acceleration factor, 2; 160 slices; slice thickness, 1 mm] encoded for sagittal slice reconstruction, providing voxel dimensions of 0.9375 × 0.9375 × 1.0 mm.

Sample 2. Anatomical images were acquired using a 1.5 tesla scanner (Signa; General Electric) for each participant. Head positioning was standardized using cantho-meatal landmarks. Images were acquired using a three-dimensional spoiled gradient recall sequence with the following parameters: TR, 24 ms; TE, 5 ms; 45° flip angle; matrix size 256 × 192; two excitations; FOV, 30 cm; 124 contiguous sagittal slices that were 1.2 mm thick without skip and reformatted with voxel dimensions 1.17 × 1.17 × 1.2 mm.

Image processing

Images were processed on Sun Ultra 10 workstations using ANALYZE 8.0 Biomedical Imaging Resource (Mayo Foundation, Rochester, MN) and software developed in-house; technicians were blind to subject characteristics and hemisphere (images were randomly flipped in the transverse plane before preprocessing). We resliced the MRI dataset to correct for any residual head rotation, tilt, or flexion/extension before performing morphometric analyses.

We removed inhomogeneities in image intensity before isolating cerebral tissue (Sled et al., 1998). Inter-rater reliabilities, calculated as intraclass correlation coefficients using a two-way random-effects model, were all >0.98 (Arntz et al., 1991).

Preprocessing. We corrected large-scale variations in image intensity using an algorithm developed at the Montreal Neurological Institute (Sled et al., 1998). Extracerebral tissues were removed using an automated tool for brain extraction (Shattuck and Leahy, 2002) that smooths image intensity with an anisotropic filter, detects three-dimensional (3-D) edges using a Marr-Hildreth edge detector (Marr and Hildreth, 1980), and selects as the brain the largest connected component with a closed boundary. We manually removed connecting dura on each slice in sagittal view and checked it in orthogonal views. The brainstem was transected at the pontomedullary junction.

Cortical gray matter segmentation. We sampled bilaterally gray-scale values of “pure” representations of cortical gray matter and white matter at four standard locations (frontal, temporal, occipital, and parietal lobes) using an 8 × 8 × 64 pixel array, which is sufficiently large to provide statistical stability but small enough to avoid including other tissue types, which can create partial volume effects. We averaged these four values for each tissue type, and computed a threshold value halfway between the mean gray and white matter values for a slice in the imaging volume. We used these threshold values on a slice-by-slice basis to provide an initial rough classification of cortical gray and white matter throughout the cerebrum. To provide the most accurate segmentation possible of the cortical mantle, we hand edited the classification in the coronal and transverse views. The intraclass correlation coefficient measuring the reliability of cortical gray matter volumes using a two-way random-effects model (Shrout and Fleiss, 1979) was 0.98.

Choice of template brain. To select a template brain that was most representative of our cohort, we identified a brain that was as representative as possible of the demographics of the sample (i.e., demographic data were closest to the mean as possible in terms of age, weight, and height). We then coregistered the brains of the remaining participants to this preliminary template. We determined point correspondences on the surfaces of their cortices and computed the distances of the corresponding points on the cerebral surface of the other participants from the surface of the template brain. The brain for which all points across the surface were closest (in terms of least squares) to the average of the computed distances was selected as the final template brain. We then performed a second coregistration of all brains to this most representative template. A single representative brain serves as a better template than does an averaged brain because a single brain has well defined tissue interfaces. Averaging images for a template blurs these boundaries and increases registration errors, which are important when distinguishing subtle effects across populations.

Sulcal overlay. We overlaid onto the statistical maps of our template brain the sulcal boundaries previously identified on the International Consortium for Brain Mapping (ICBM) high-resolution, single-participant template (http://www.loni.ucla.edu/ICBM/Downloads/Downloads_ICBMtemplate.shtml; Mazziotta et al., 2001) to aid visual identification of the locations of findings on the surface of the brain. We mapped the sulcal boundaries and 3-D labels of cortical gyri from the ICBM template onto the cortical surface of our template brain using our high-dimensional, nonrigid warping algorithm (see above).

Cortical thickness

For detailed descriptions of our procedures for surface morphologies, and related validation studies, see previous studies (Bansal et al., 2005). In short, random flips were reversed to provide their original correct orientation. Isolated brains were coregistered to the selected template brain (Plessen et al., 2006) using a similarity transformation, followed by a high-dimensional, nonrigid warping algorithm based on the dynamics of fluid flow. We removed the cortical mantle from each coregistered brain. We then used a three-dimensional morphological operator to distance transform this brain without the cortex from the coregistered brain of the same participant that contained the cortex (Rosenfeld and Pfaltz, 1968; Harilick and Shapiro, 1991). Thus, we computed CT as the smallest distance at each point on the external cortical surface from the outermost surface of the white matter in the coregistered brain. Because the brain and its local features, such as CT, were scaled during the similarity transformation of that brain to the template brain, these thickness values, measured in template space, inherently accounted for generalized scaling effects within the cerebrum.

Cortical asymmetry

To compare CTS across the right hemisphere with those across the left hemisphere, the right hemisphere of each brain was normalized to the template left hemisphere using a two-step process. First, the right hemisphere of each brain was flipped to create a mirror image so that it overlay the left hemisphere of the same brain. The right hemisphere was then warped to the left hemisphere by a high-dimensional, nonrigid warping algorithm based on the dynamics of fluid flow. Second, we used the deformation field that mapped the left hemisphere on the template left hemisphere to warp the right hemisphere to the template left hemisphere. This two-step process established correspondences across all points on the right hemisphere and those across the left hemisphere of the template brain, thereby allowing us to compute variations in CT at particular voxels in each hemisphere. We then assessed the correlation between the VIQ-regressed-on-PIQ score and the left-right CT differ-
ence at each voxel, as well as the PIQ-regressed-on-VIQ score and the left–right CT difference at each voxel.

Statistical analysis
We used multivariate linear regression at each point on the reference surface to examine associations of the VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores with CT. In addition to the main effects, we considered for inclusion in the model the two-way interactions of age (in years) and sex (male, female) with VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores by assessing their significance in the following linear regressions:

\[ CT_i = \beta_0 + \beta_1 \times \text{age} + \beta_2 \times \text{sex} + \beta_3 \times \text{VIQ} - \text{regressed on PIQ score} \times \text{sex} \] (1)

\[ CT_i = \beta_0 + \beta_1 \times \text{age} + \beta_2 \times \text{sex} + \beta_3 \times \text{PIQ} - \text{regressed on VIQ score} \times \text{sex} \] (2)

\[ CT_i = \beta_0 + \beta_1 \times \text{age} + \beta_2 \times \text{sex} + \beta_3 \times \text{VIQ} - \text{regressed on PIQ score} \times \text{age} \] (3)

\[ CT_i = \beta_0 + \beta_1 \times \text{age} + \beta_2 \times \text{sex} + \beta_3 \times \text{PIQ} - \text{regressed on VIQ score} \times \text{age} \] (4)

where \( CT_i \) is the set of CT measurements for all \( i \) participants.

The models were hierarchically well formulated (all lower-order component terms were included in the model, regardless of statistical significance). We used the false detection rate (FDR; Benjamini and Hochberg, 1995) to account for the multiple correlations computed across the cortical surface. We computed the correlation of CT with VIQ-regressed-on-PIQ scores and CT, with PIQ-regressed-on-VIQ score. We evaluated the \( p \) values of these correlations using Student’s \( t \) tests. Statistical maps were generated to display color-coded \( p \) values across the entire surface of the cortex. Statistical maps were also generated for the interactions of sex

Figure 1. Significance maps showing correlations of cortical thickness (CT) with VIQ-regressed-on-PIQ scores. Maps of correlation of VIQ-regressed-on-PIQ scores with CT, corrected for multiple comparisons using FDR. At each point on the cerebral surface, the statistical significance (probability values) of the correlation of the VIQ-regressed-on-PIQ score with CT is color coded. Warm colors (yellow, orange, red) represent a positive correlation (thicker cortex as the residual score increases), and cooler colors (blue and purple) represent a negative correlation (thinner cortex as the residual score increases). The color bar indicates the color coding of \( p \) values for testing of statistical significance at each point on the surface of the brain. Top, Maps of sample 1, \( N = 83 \). Bottom, Maps of sample 2, \( N = 58 \). CG, Cingulate gyrus; CUN, cuneus; FG, fusiform gyrus; GR, gyrus rectus; IFG, inferior frontal gyrus; IOG, inferior occipital gyrus; IP, inferior parietal lobe; ITG, inferior temporal gyrus; LG, lingual gyrus; MFG, middle frontal gyrus; MOG, middle occipital gyrus; MTG, middle temporal gyrus; OFG, orbital frontal gyrus; PreCu, precuneus; SFG, superior frontal gyrus; SMG, supramarginal gyrus; SOG, superior occipital gyrus; SPG, superior parietal gyrus; STG, superior temporal gyrus.
with VIQ-regressed-on-PIQ score, age with VIQ-regressed-on-PIQ score, sex with PIQ-regressed-on-VIQ score, and age with PIQ-regressed-on-VIQ score. As none of these interactions were statistically significant, they were removed from the final statistical model.

To assess hemispheric asymmetry, the differences between left and right CTs were regressed first on age (in years), sex (male, female), and VIQ-regressed-on-PIQ residual score, and then, in a separate analysis, the differences between left and right CTs were regressed on age, sex, and PIQ-regressed-on-VIQ residual score. The statistically significant results were color encoded and displayed on the surface of the template left hemisphere, with warm colors (yellow, orange, red) representing a positive correlation (thicker cortex as the residual score increases), and cooler colors (blue and purple) representing a negative correlation (thinner cortex as the residual score increases). The color bar indicates the color coding of p values for the testing of statistical significance at each point on the surface of the brain. Top, Maps of sample 1, N = 83. Bottom, Maps of sample 2, N = 58.

**Results**

We imaged 141 individuals, 83 (50 children, 33 adults) in sample 1 and 58 (26 children, 32 adults) in the sample 2. The mean FSIQ scores in the two samples were somewhat higher than the mean in the standardization sample (113 in sample 1; 121.7 in sample 2; 100 in standardization sample), but the ranges of the VIQ-regressed-on-PIQ and the PIQ-regressed-on-VIQ residual scores were normally distributed around a mean of zero in each sample. The VIQ-regressed-on-PIQ score and the PIQ-regressed-on-VIQ score correlated only weakly with FSIQ ($r^2$ = 0.305 and 0.291, respectively; p values <0.005). The proportion of participants who had at least 1 SD of difference (15 points) between VIQ and PIQ (34% in sample 1; 40% in sample 2) was similar to rates in normative samples (e.g., 26% in the WAIS-IV standardization sample; Wechsler, 2008). Similar to normative samples, approximately half of the participants who had at least 1 SD of difference between VIQ and PIQ had a VIQ greater than PIQ (57% in sample 1; 48% in sample 2), and half had a PIQ greater than VIQ (43% in sample 1; 52% in sample 2).

In both samples, CT correlated inversely with the magnitude of the VIQ-regressed-on-PIQ residual score and positively with the PIQ-regressed-on-VIQ residual score in both the right and left hemispheres within the anterior-most surface of the frontal lobe and within the occipital lobes (Figs. 1, 2). Specifically, ante-
rior regions included the inferior frontal (sample 2), middle frontal (both samples), superior frontal (both samples), orbitofrontal (both samples), and superior temporal (sample 2) gyri. Posterior regions included the inferior occipital gyrus (both samples), middle occipital gyrus (both samples), superior occipital gyrus (both samples), inferior parietal lobule (both samples), parietal gyrus (sample 2), superior parietal gyrus (sample 2), middle temporal gyrus (both samples), cuneus (both samples), precuneus (both samples), and supramarginal gyrus (sample 2). Additional correlations detected on the inferior surface of the brain included the inferior temporal (both samples), fusiform (both samples), lingual (both samples), and inferior orbitofrontal gyri (both samples). Additional regions detected on the mesial surface of the brain included the gyrus rectus (sample 1) and cingulate gyrus (sample 2).

The anterior regions where CT correlated with the VIQ-PIQ discrepancy encompassed Brodmann’s areas (BAs) 10, 11, and 47 in samples 1 and 2, and, additionally, include BAs 44, 45, and 46 in sample 2. BA 10 coordinates activity across multiple cognitive systems (Ramnani and Owen, 2004); BAs 11, 46, and 47 generally support behavioral inhibition (Schilling et al., 2012); and BAs 44 and 45 (Broca’s region) support expressive language functions (Foundas et al., 1996). The posterior regions where CT correlated with the VIQ-PIQ discrepancy encompassed BAs 18/19, 37, and 39/22 in sample 1; and BAs 17, 18, 19, 7, 20, 38, and 22 in sample 2. BAs 17, 18, 19, 20, 37, and 38 support visual processing of images and object recognition (Muller-Forell, 2005), and word recognition (Cohen et al., 2004); BA 7 supports the integration of multisensory data (Muller-Forell, 2005); and BAs 39 and 22 (Wernicke’s region) support receptive language (Binder et al., 1997). On the mesial surface, regions where CT correlated with the VIQ-PIQ discrepancy encompassed BAs 23, 24, and 32 in sample 2. These areas process feedback, detect changes in the environment, and allow for alterations of behavior in response to these signals (Pearson et al., 2011).

Scatterplots showing the association of CT with the VIQ-PIQ score within these regions of statistical significance show consistently in each sample that with progressive thinning of the cortical mantle, the VIQ-regressed-on-PIQ score becomes more positive (i.e., VIQ > PIQ). In addition, as the cortical mantle thickens, the PIQ-regressed-on-VIQ score becomes increasingly more negative (i.e., PIQ > VIQ; Fig. 3). Similarly, with progressive thickening of the cortical mantle, the PIQ-regressed-on-VIQ score becomes more positive (i.e., PIQ > VIQ), and as the
The cortical mantle thins, the PIQ-regressed-on-VIQ score becomes increasingly more negative (i.e., VIQ/H11022 PIQ; Fig. 4). Neither the VIQ-regressed-on-PIQ nor the PIQ-regressed-on-VIQ score interacted with age or sex in regions identified in the main effect in the regression model in either sample.

We compared the regions identified as correlating with VIQ-PIQ discrepancy in both the 1.5 T and 3 T samples with the regions previously identified as correlating with general intelligence. First, we compared our findings to the regions identified in a metareview of 37 studies of the neural correlates of general intelligence (Jung and Haier, 2007). The two networks have the following areas in common: BAs 10, 47, 18, and 19. Regions we identified as correlating with the VIQ-PIQ discrepancy but not with general intelligence are BAs 11 and 22. Regions that correlate with general intelligence but not with VIQ-PIQ discrepancy are BAs 6, 9, 32, 45, 46, 40, and 21. Second, we compared our findings to the regions identified as neural correlates of FSIQ within the same 1.5 T sample as in our present study (Goh et al., 2011) and identified only BAs 10 and 11.

The VIQ-PIQ discrepancy has long been considered a putative behavioral marker of psychiatric disorders, including speech-language impairment (Rice et al., 2005), autism spectrum disorders (Mayes, 2003; Charman, 2011), and nonverbal learning disability (Grodzinsky et al., 2010). Clinicians rely on the VIQ-PIQ difference score, not residual scores, to estimate the discrepancy between abilities. We therefore compared the regions identified as correlating with VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ residual scores with the regions identified as correlating with the VIQ-PIQ difference score (Fig. 5). This comparison demonstrated similar results for residual and difference scores within the anterior-most surface of the frontal lobe and within the occipital lobes, although the residual scores demonstrated stronger effects. The similarity across the type of measures indicates the validity of both residual scores and difference scores.

We also assessed whether the VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores correlate with measures of anatomical asymmetry in CT. To measure asymmetry in CT at each point along the cerebral surface of each participant, we established correspondences across all points on the surface of the right hemisphere with those on the surface of the left hemisphere and mapped these to the surface of the template brain, thereby allowing us to calculate asymmetry at each point as the CT in the left hemisphere minus the CT in the right hemisphere. We detected a significant inverse relationship between the magnitude of
the VIQ-regressed-on-PIQ score and the degree of CT asymmetry in inferior occipital (sample 1), middle occipital (sample 1), middle temporal (sample 1), superior temporal (both samples), precentral (sample 1), inferior frontal (both samples), and middle frontal (both samples) gyri (Fig. 6). We detected a significant positive relationship between the magnitude of the PIQ-regressed-on-VIQ score and the degree of CT asymmetry in the same regions listed above. Statistical significance in sample 2 is smaller than in sample 1, but the sample contained fewer subjects and the images were obtained on scanners of differing strength; overall, the direction of effect is similar in both samples, albeit sample 1 shows greater effects than sample 2. Scatterplots in all significant regions in both samples show that as VIQ-regressed-on-PIQ score increases (VIQ > PIQ), left–right hemisphere CT decreases. As PIQ-regressed-on-VIQ score increases, left–right hemisphere CT increases. Thus, thinner left hemisphere CT relative to right hemisphere CT was associated with higher verbal score relative to performance score (VIQ > PIQ), and thicker left hemisphere CT relative to right hemisphere CT was associated with higher performance relative to verbal score (PIQ > VIQ; Fig. 6).

Discussion
We have identified regional components of a putative neural system that underlies one well known but little-studied source of variance within and between individuals in domain-specific abilities, the VIQ-PIQ discrepancy. We detected significant thinning of anterior and posterior cortices associated with an increasing difference between verbal and performance IQ scores. We replicated these findings in two independent samples of healthy individuals, from youth to adulthood, on different age-appropriate versions of IQ tests and on two scanners of differing field strengths.

The brain regions we identified as underlying the VIQ-PIQ discrepancy support specific aspects of information processing relevant to performance on VIQ and PIQ tasks. In both samples, anterior regions encompassed BA 10, which coordinates activity across multiple cognitive systems (Ramnani and Owen, 2004), and BAs 11/47, supporting behavioral inhibition (Schilling et al., 2012). Posterior regions encompassed BAs 18/19, supporting object (Muller-Forell, 2005) and word recognition (Cohen and Dehaene, 2004), and BA 22, supporting receptive language (Binder et al., 1997). The regions where CT correlated with VIQ-PIQ discrepancy are generally distinct from other regions that correlate with general intelligence (Jung and Haier, 2007; Goh et al., 2011). Thus, the results support our hypothesis that a single neural system mediates the VIQ-PIQ discrepancy.

The entire range of values (for VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores) contributed to the correlations with CT. For VIQ-regressed-on-PIQ scores, positively valued differences (VIQ > PIQ) were associated with relative cortical thinning, and negatively valued differences (VIQ < PIQ) were associated with relative cortical thickening in the same regions. We detected effects in the opposite direction for PIQ-regressed-on-VIQ scores. We note
the opposing effects at each voxel in which thinning of the cortical mantle is associated with better performance on VIQ relative to PIQ tasks, and thickening of the cortical mantle at the same location is associated with better performance on PIQ relative to VIQ tasks.

The neural system we identified seems to operate like a CT rheostat that either improves or degrades performance of one domain-specific ability relative to the other as CT deviates either upwards or downwards from the population mean: when the cortical mantle is thin in an individual relative to the average CT of the population sample, VIQ is higher than PIQ; when the cortical mantle is thick, PIQ is higher than VIQ. In this model, the DSAs (measured with VIQ and PIQ) derive from two independent and distributed neural systems that compete, at least in part, for access to the limited capacity system, so that what access one system gains, the other loses, in a zero-sum game. The DSAs presumably rely on other circuits and brain regions in addition to the system that we have identified, but our system is presumably where the circuits that support these independent abilities physically interface with one another.

Alternatively, the opposing effects on performance of thinning and thickening at each voxel could arise if two relatively independent morphological features are present within the brain tissue imaged within each voxel, and if those two features relate in opposing ways to abilities that affect VIQ and PIQ scores. A smaller size of one feature (e.g., the thickness of one cortical layer or size of one cell type) would support better performance on VIQ tasks, and a larger size of another feature (a different cortical layer or a second cell type) would support better performance on PIQ tasks. Because CT at each voxel is a summation of the sizes of all the morphological elements within that voxel, these two morphological features and their differential associations with VIQ and PIQ tasks also would sum within each voxel, giving rise to the following four possible combinations of features that relate to the VIQ-PIQ discrepancy: (1) both morphological features are small (and thus one is contributing to higher VIQ scores and the other to lower PIQ scores): the cortical mantle is thin and VIQ is high relative to PIQ; (2) both features are large: the cortical mantle is thick, and PIQ is high relative to VIQ; (3) VIQ-related features are large and PIQ-related features are small: VIQ equals PIQ and both are low, and CT has an intermediate value nearer the population average; and (4) VIQ-related features are small and PIQ-related features are large: VIQ equals PIQ and both are high, and CT has an intermediate value near the population average. These combinations of morphological features and their relation to the VIQ-PIQ discrepancy can account for the linear relationship between the residual scores and CT that we detected. Therefore, the VIQ-PIQ

Figure 6. Correlations and scatterplots of VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores with asymmetry in cortical thickness (CT). Maps of correlation of VIQ-regressed-on-PIQ and PIQ-regressed-on-VIQ scores with asymmetry in CT (left hemisphere CT minus right hemisphere CT), corrected for multiple comparisons using FDR. Asymmetry correlated inversely with the magnitude of the VIQ-regressed-on-PIQ score in inferior occipital (sample 1), middle occipital (sample 1), middle temporal (sample 1), superior temporal (both samples), precentral (sample 1), inferior frontal (both samples) and middle frontal (both samples) gyri (Fig. 6). We detected a significant positive relationship between the magnitude of the PIQ-regressed-on-VIQ score and the degree of CT asymmetry in the same regions listed above. Top, Maps of sample 1, N = 83. Bottom, Maps of sample 2, N = 58.
discrepancy could derive from differences in morphology in two previously unidentified, independent systems supporting DSAs.

Most neuroimaging studies measure intelligence with the Wechsler scales and interpret findings in terms of verbal and performance scales. To the extent that verbal and performance scores share common requirements, they both measure general ability. In addition to this shared variance, however, each score represents unique variance associated with a domain-specific ability and variance in task-specific skills (Colom and Thompson, 2011). The constructs that these sources of variance represent have long been debated. Some IQ theorists argue that VIQ measures achievement, not ability, because performance on many of the IQ test items, such as vocabulary, depends on acquired knowledge (Naglieri and Bornstein, 2003). These theorists identify PIQ as simultaneous processing, the ability to understand the inter-relationships of variables as assessed, for example, with matrix tests (Kaufman and Kaufman, 1985; Naglieri and Das, 1997). Therefore, the VIQ-PIQ discrepancy can be interpreted as a difference between achievement and simultaneous processing. Achievement presumably represents experience and learning, whereas simultaneous processing involves a more innate rational ability. Our findings are consistent with the known differences in the neurobiological bases for these two cognitive processes. The association of cortical thinning with higher achievement relative to simultaneous processing scores suggests that achievement may benefit from consolidative processes like synaptic or dendritic pruning. The association of cortical thickening with higher simultaneous processing relative to achievement scores suggests that simultaneous processing may benefit from greater plasticity within the cortical mantle, which presumably requires greater dendritic and axonal arborization, and greater connectivity within and across cortical layers.

Lesion studies have demonstrated that unilateral damage to a hemisphere, depending on the location of the lesion, produces loss of a skill and a discrepancy between verbal and spatial ability (Warrington et al., 1986; Glaßcher et al., 2009), which at first consideration could imply a discrepancy with our findings of bilateral representation in the neural correlates of VIQ-PIQ discrepancy. Although lesion studies identify regions that are necessary for performing a particular cognitive skill, being necessary for a skill is not synonymous with being sufficient for it, as other regions may nevertheless contribute to performance measures. Human imaging studies are more likely than lesion studies to identify regions that support skills, even if not absolutely necessary for their performance. Moreover, persons with unilateral lesions often recover function through neureplastic responses in the opposite hemisphere, suggesting the importance of bilateral contributions to verbal and spatial skills (Vigneau et al., 2011). Additionally, various stimulation studies suggest that verbal and spatial skills are represented bilaterally in the brain (Pascual-Leone et al., 1994; Binder et al., 1996; Knecht et al., 2002). Finally, differences in conclusions from morphometric and lesion studies likely relate to fundamental differences in the measures and methods used to localize the effects of interest. CT, for instance, is a feature only of gray matter, which supports primarily local processing, whereas naturalistic lesions often affect more prominently white matter, which supports communication across brain regions.

In addition to bilateral correlations, we detected evidence in both samples for a less robust correlation of hemispheric asymmetry in CT with VIQ-PIQ discrepancy. A thinner left cortex relative to right was associated with higher VIQ relative to PIQ, and a thicker left cortex relative to right was associated with higher PIQ relative to VIQ. These findings are generally consistent with the hemisphere-specific findings from naturalistic human lesion studies (Glaßcher et al., 2009), and they are consistent with our primary finding that higher VIQ relative to PIQ is associated with cortical thinning and higher PIQ relative to VIQ is associated with cortical thickening.

Studies of patient populations suggest that the VIQ-PIQ discrepancy may be under genetic control. The VIQ-PIQ discrepancy in autism defines a particular phenotype that is linked to two genes, one of which is also implicated in speech–language impairment (Chapman et al., 2011). The presence of the VIQ-PIQ discrepancy in Prader-Willi syndrome depends on the disease genotype (either a deletion or uniparental disomy) and is associated with higher risk for associated autism spectrum behaviors (Copet et al., 2010).

Our study has limitations. To evaluate whether our results vary as a function of overall IQ would require a sufficiently large sample to assess these correlates separately in groups of people with low, medium, or high FSIQ. To assess the histological correlates of our findings would require a large postmortem study. Our participant samples spanned a wide range of ages, which potentially could add variability to the correlations that we assessed. Age did not interact with VIQ-PIQ discrepancy, however, suggesting that the association of VIQ-PIQ discrepancy with CT is consistent across ages. The inclusion of participants across age and developmental stages can be viewed as a relative strength of our study, as it increases the generalizability of our findings. The contributions of even the youngest children in our samples to the correlations of CT with VIQ-PIQ discrepancy (Figs. 3, 4) suggest that the morphological and cellular determinants of the discrepancies in these cognitive domains may be present early in life.

References


