



Inattentive and hyperactive traits differentially associate with interindividual functional synchrony during video viewing in young children without ADHD

Ryann Tansey ^{1,2,3,*}, Kirk Graff^{1,2,3}, Christiane S. Rohr^{1,2,3}, Dennis Dimond^{1,2,3}, Amanda Ip^{1,2,3}, Deborah Dewey^{1,2,3,4,5},
Signe Bray ^{1,2,3,4,6}

¹Child and Adolescent Imaging Research Program, University of Calgary, 28 Oki Drive NW, Calgary, AB, T3B 6A8, Canada,

²Alberta Children's Hospital Research Institute, University of Calgary, 28 Oki Drive NW, Calgary, AB, T3B 6A8, Canada,

³Hotchkiss Brain Institute, University of Calgary, 3330 Hospital Drive NW, Calgary, AB, T2N 4N1, Canada,

⁴Department of Pediatrics, Cumming School of Medicine, University of Calgary, 28 Oki Drive NW, Calgary, AB, T3B 6A8, Canada,

⁵Department of Community Health Science, Cumming School of Medicine, University of Calgary, 3280 Hospital Drive NW, Calgary, AB, T2N 4Z6, Canada,

⁶Department of Radiology, Cumming School of Medicine, University of Calgary, 3330 Hospital Drive NW, Calgary, AB, T2N 4N1, Canada

*Corresponding author: Ryann Tansey. Email: ryann.tansey@ucalgary.ca

Inattention and hyperactivity present on a spectrum and may influence the way children perceive and interact with the world. We investigated whether normative variation in inattentive and hyperactive traits was associated with differences in brain function, while children watched clips from an age-appropriate television program. Functional magnetic resonance imaging (fMRI) data and parent reports of inattention and hyperactivity traits were collected from 81 children 4–7 years of age with no parent-reported diagnoses. Data were analyzed using intersubject correlations (ISCs) in mixed effects models to determine if inattentive and hyperactive traits were associated with idiosyncrasy of fMRI response to the video. We hypothesized that pairs of children with higher average inattention and hyperactivity scores would show less interindividual brain synchrony to one another than pairs with lower average scores on these traits. Video watching engaged widespread visual, auditory, default mode and dorsal prefrontal regions. Inattention and hyperactivity were separably associated with ISC in many of these regions. Our findings suggest that the spectrum of inattention and hyperactivity traits in children without ADHD are differentially associated with neural processing of naturalistic video stimuli, which may have implications for understanding how children with different levels of these traits process audiovisual information in unconstrained conditions.

Key words: developmental neuroimaging; fMRI; hyperactivity; inattention; naturalistic paradigm.

Introduction

Inattention and hyperactivity are among the most common neurodevelopmental challenges affecting children. In children, nonclinical levels of inattention and hyperactivity have been associated with poorer outcomes such as higher rates of grade retention and graduation failure in adolescence (Bussing et al. 2010), less positive relationships with friends and parents (Rielly et al. 2006), and worse executive functioning (Brown and Casey 2016). Population-based studies have further shown that increased inattention symptoms in childhood are associated with lower academic performance in adolescence (Sayal et al. 2015; Salla et al. 2016) and reduced financial earnings in adulthood (Vergunst et al. 2019).

In adults without attention deficit/hyperactivity disorder (ADHD), inattentive and hyperactive/impulsive traits have been associated with interindividual synchrony of blood oxygen level dependent (BOLD) signal during

viewing of a naturalistic movie stimuli (Salmi et al. 2020). This suggests that inattention and hyperactivity may impact the way an individual perceives and interacts with their environment. However, the relationships between inattentive and hyperactive traits and brain function in typically developing children remain understudied. Here, we investigate whether inattentive and hyperactive traits in young children without ADHD are associated with neural processing of a complex audiovisual stimulus: clips from an age-appropriate television program.

To capture individual differences in the way the brain processes audiovisual (AV) media, we used intersubject correlation (ISC; Hasson et al. 2004). Functional magnetic resonance imaging (fMRI) studies show that naturalistic AV stimuli evoke synchronized brain activity across individuals (Hasson et al. 2004, 2008; Nastase et al. 2019) in visual, auditory, emotional, navigation, and language processing regions, as well as areas related to

Received: January 11, 2022. Revised: February 22, 2022. Accepted: February 24, 2022

© The Author(s) 2022. Published by Oxford University Press.

This is an Open Access article distributed under the terms of the Creative Commons Attribution License (<https://creativecommons.org/licenses/by/4.0/>), which permits unrestricted reuse, distribution, and reproduction in any medium, provided the original work is properly cited.

attentional control (Bottenhorn et al. 2018). By calculating the ISC (also referred to as “interindividual synchrony”; measured as the Pearson correlation between the BOLD time courses from corresponding voxels of individuals watching the same movie), we can quantify whether individuals with similar behavioral traits show synchronized processing of an AV stimulus. Using ISC, researchers have found that individuals with autism and elevated depressive symptoms can show idiosyncratic neural responses to movies (Byrge et al. 2015; Gruskin et al. 2020). Pairwise ISC measures allow researchers to investigate idiosyncrasy and granular individual differences of both brain function and behavior, as they can capture the ways in which 2 individuals may differ in their processing of a stimulus based on their specific continuous traits and phenotypes (Finn et al. 2020).

Previous work has examined the association between a clinical diagnosis of ADHD and ISC during video viewing. In 1 adult study (Salmi et al. 2020), controls displayed more synchronized brain activity than ADHD individuals in the lateral and medial occipital cortex, precuneus, temporoparietal junction, superior temporal cortex, and—when speech or music distractors were added to the movie—the posterior parietal cortex. However, they also found that in the control group, similarity of impulsivity scores was associated with greater ISC in the cuneus, dorsomedial prefrontal cortex, and temporoparietal junction, while similarity of inattention was associated with ISC in a small bilateral region of the precuneus. Contrary to these findings, in a case–control study that compared children with ADHD to non-ADHD controls (Tang et al. 2019), the brain activity of the ADHD group was more synchronized than the control group in widespread areas of the occipital and temporal lobes. To our knowledge, the specific dimensional relationships between inattention and hyperactivity in young children without ADHD and ISCs have not yet been investigated.

Video-watching offers attractive benefits in developmental neuroimaging research, as it probes brain function in a dynamic, multimodal, and arguably more “ecologically valid” context than traditional task-based or resting-state paradigms (Sonkusare et al. 2019), opening up an entirely new realm of unique questions and techniques. It also has the added benefit of reducing head motion of young children during MRI scans (Vanderwal et al. 2015; Greene et al. 2018). Furthermore, the centrality of screen media in many children’s lives warrants investigation into the ways it is processed by the brain and could offer important nuance regarding the potential benefits and disadvantages of this activity.

In this study, we investigate whether inattentive and hyperactive traits are associated with interindividual synchrony in young children without ADHD during the presentation of video clips from an educational television show. We hypothesized that greater inattention and hyperactivity scores would be associated with lower pairwise synchrony. Both inattention (Jonkman et al. 2017; Arabaci and Parris 2018) and hyperactivity traits

(Arabaci and Parris 2018) have been positively linked to mind-wandering in children and adults without ADHD (Frick et al. 2020), which could potentially direct attention away from the shared video stimulus, resulting in decreased synchrony (Nastase et al. 2019). Specifically, we test the hypothesis that brain function is more idiosyncratic in children with higher inattention and hyperactivity trait levels.

Materials and methods

Participants, study procedure, and stimuli

Participants were recruited from Calgary and the surrounding area in Southern Alberta as part of a longitudinal neuroimaging study in early childhood. Study procedures were approved by the University of Calgary Conjoint Health Research Ethics Board. Parents provided informed consent for their child’s participation and children provided assent. All data collection occurred at the Alberta Children’s Hospital. Children with a history of psychiatric or neurodevelopmental disorders were excluded, as were any children with a medical condition or other contraindications that prevented participation in an MRI scan. Prior to the scan, all participants underwent a mock scanner training session to prepare them for data collection. A total of 135 children (79 females, 56 males) aged 4–7 years participated. Structural MRI, 18-min of video-watching fMRI, and parent-report behavioral data were collected. Following data collection, children were excluded from analysis if they had more than 2 min of motion corrupted volumes (motion corruption defined as >0.2-mm framewise displacement, Jenkinson criteria; Jenkinson et al. 2002). This exclusion criterion was used to minimize differences in head motion among the participants, as head motion is very high in children in the scanner (Dosenbach et al. 2017). Furthermore, in ISC analysis, which considers participants as pairs, motion is additive within each pair. In total, 54 of the original sample of 135 children were excluded ($F=28$, $M=26$), leading to a final sample size of $n=81$ ($F=51$, $M=30$). The demographic characteristics of this final sample are presented in Table 1. The demographics of included vs. excluded children can be found in Supplementary Table S1.

During video-watching fMRI, participants passively viewed a selection of clips from the children’s television show “Elmo’s World.” This stimulus was selected as it contains content that is educational and gender neutral, human faces, depicts social interaction, and is similar to the naturalistic stimuli used in other developmental neuroimaging research (e.g. Cantlon and Li 2013). The video used in the scanner was also played for each participant during their mock scanner session to minimize between-participant effects of stimulus novelty.

MRI data acquisition

All neuroimaging data were acquired at the Alberta Children’s Hospital using a 3.0 T GE MR750w (Waukesha,

Table 1. Demographics.

	Age (years)	Censored volumes	Average relative FD (mm)	FSIQ	SNAP-I	SNAP-H	Sex
Range	4.14–7.89	1–57	0.035–0.17	80–139	0–2.89	0–2.33	F = 51
Mean	5.88	21.32	0.076	111.1	0.73	0.76	M = 30
SD	0.94	16.69	0.030	12.77	0.49	0.59	
Median	5.88	16.00	0.068	112	0.67	0.67	

Demographic summary for the entire sample ($n=81$).

WI) scanner with a 32-channel head coil. Structural images were acquired with a T_1 -weighted 3D BRAVO sequence with the following parameters: TR = 6.764 ms, TE = 2.908 ms, FA = 10°, voxel size $0.8 \times 0.8 \times 0.8$ mm, and matrix size of 300×300 . Functional images were acquired with a T_2^* -weighted gradient-echo EPI sequence with the following parameters: 34 axial slices, 433 volumes, TR = 2500 ms, TE = 30 ms, FA = 70°, voxel size of $3.5 \times 3.5 \times 3.5$ mm, and matrix size of 64×64 .

Inattention and hyperactivity measures

To measure inattention and hyperactivity traits, parents completed the SNAP-IV, a Likert-type rating scale measuring ADHD behaviors based on DSM-IV criteria (Bussing et al. 2008). We used a shortened version of the SNAP-IV (the MTA SNAP-IV), which has 2 separate subscales that measure the participant's level of inattention (9 items) or hyperactivity (9 items) (Bussing et al. 2008) and gives a final score for ADHD-hyperactive/impulsive (SNAP-H), ADHD-inattentive (SNAP-I), and ADHD-combined (SNAP-C). Scores can range from 0 to 3. A higher SNAP score indicates a higher level of ADHD-related behaviors. A parent-report SNAP score of greater than 1.2 is associated with an increased probability of concern, and scores above 1.8 are associated with a higher probability of ADHD diagnosis (Bussing et al. 2008). This study used the SNAP-I and SNAP-H scores.

fMRI preprocessing

fMRI data were preprocessed with an in-house, customized Nipype pipeline (Gorgolewski et al. 2011). Steps in the anatomical image preprocessing include bias correction via ANTs (Tustison et al. 2014) *n4BiasFieldCorrection*, removal of skull and nonbrain tissue via ANTs *antsBrainExtraction*, normalization to the NIHPD 4.5- to 8.5-year-old asymmetrical atlas in MNI space (Fonov et al. 2009, 2011) using ANTs *antsRegistration*, tissue segmentation using ANTs *Atropos*, and AFNI (Cox 1996)'s *3dmask_tool* to erode the tissue segmentations.

The functional image preprocessing pipeline generally followed recommendations in Ciric et al. (2018) and Graff et al. (2022). Head motion parameters were first estimated with FSL MCFLIRT (Jenkinson et al. 2002) (following recommendations from Power et al. (2017) to estimate head motion parameters for regression prior to slice time correction). Functional data then underwent slice time correction with FSL's *slicetimer*, and rigid body alignment was performed with FSL's MCFLIRT (Jenkinson et al. 2002).

Next, nonbrain tissue and skull stripping were performed with FSL's BET. We generated a study-specific EPI template in MNI 2-mm voxel space, following recommendations by Huang et al. (2010), and used ANTs *antsRegistration* (Tustison et al. 2014) to warp the EPI image to this template. Tissue segmentations from the T_1 image were warped to this functional image using FSL's FLIRT (Jenkinson et al. 2002; Greve and Fischl 2009). Linear and quadratic trends were removed, and a high-pass filter at 0.01 Hz was applied. Nuisance regressors included 6 head motion parameters, white matter, cerebrospinal fluid, and global signal. We also included the derivatives, quadratic terms, and quadratic term derivatives for each of the nuisance regressors into the regression model. Volumes with a framewise displacement exceeding 0.2 mm based on the Jenkinson criteria (Jenkinson et al. 2002) were censored (Power et al. 2012, 2014; Rohr et al. 2019). Finally, functional data were smoothed using a 8.0-mm FWHM Gaussian kernel, as previous studies have suggested that the optimal smoothing kernel for ISC data is slightly larger than twice the size of the voxels (Pajula and Tohka 2014; Nastase et al. 2019).

ISC analyses

Second-level group analyses were conducted using AFNI. Voxelwise, whole-brain ISCs were computed by calculating the Pearson correlation of the BOLD signal time course at corresponding voxels between each pair of subjects (Hasson et al. 2004). Pairwise r -maps were then entered into a linear mixed effects model with crossed random effects to address the statistical nonindependence in ISC data (Chen et al. 2016, 2017).

The pairwise inattention and hyperactivity terms included in models were calculated as the average SNAP-I and SNAP-H scores for each pair. When considering pairwise data for ISC analysis, it is possible to characterize the behavioral distance between individuals in a number of ways; typically, either by averaging the scores of a pair or calculating the absolute difference in scores between individuals in each pair. We selected the former metric as we are testing the hypothesis that idiosyncrasy of brain response increases with symptom scores (Finn et al. 2020). In other words, we expected that 2 children with low scores would show greater ISC to one another than 2 children with high scores, even if both pairs had the same absolute difference in scores between them. This hypothesis is grounded in the idea that children with high symptom scores will be more distractable and therefore

less likely to be attentive to the shared stimulus, and that it is unlikely that individuals will be distracted in a systematic way. Similarly, we included the average age of each pair as a control covariate, as distractibility decreases with age in childhood (Hoyer et al. 2021). To account for head motion, we included the total number of censored volumes per pair of subjects as a nuisance covariate. Sex was controlled for by including dummy covariates in the model representing sex makeup of each pair (female–female, female–male, and male–male). All continuous variables and covariates were mean-centered.

We created 3 models in total: (i) one that included inattention, (ii) one that included hyperactivity, and (iii) one that included both inattention and hyperactivity as variables in the model to account for both simultaneously. This allowed us to calculate a contrast for comparing associations between these measures. Statistical significance was determined through cluster-based thresholding using a voxelwise threshold of $P < 0.001$ and a cluster forming threshold corresponding to $\alpha = 0.05$ (Cox 1996). Figures visualizing surface projections of results were created using BrainNet Viewer (Xia et al. 2013).

To account for potential differences in attention to the stimulus, we calculated versions of all the models that included frontal eye field (FEF) synchrony as a control covariate. FEF signal has been used in previous studies as an approximate for visual attention and gaze behavior (Redcay et al. 2010; Moraczewski et al. 2018). We averaged the ISC values in all voxels within a bilateral FEF region of interest defined from the MIST parcellation (Urchs et al. 2019) for each pair and included this value as a covariate in the models.

Results

Sample characteristics

Demographic data for the final sample can be found in Table 1 and SNAP scores in Fig. 1.

ISCs across the whole sample

Significant ISC is seen in the entire sample across almost all of the brain. Due to high statistical power from the large number of pairs included in the analysis, nearly all gray matter voxels survive cluster thresholding. Figure 2 shows the unthresholded map of group average ISC (Pearson's r) of the whole sample.

Inattention model results

Higher ISC was associated with lower pairwise inattention scores in a number of areas in the occipital, temporal, and frontal cortices, including bilateral lateral occipital cortex, occipital pole, fusiform cortex, lingual gyri, supramarginal gyri, angular gyri, precuneus, precentral gyrus, parietal operculum cortex, and superior cerebellum; the right middle frontal gyrus and posterior cingulate; and the left temporal cortex, including the superior temporal gyrus and the medial temporal cortex

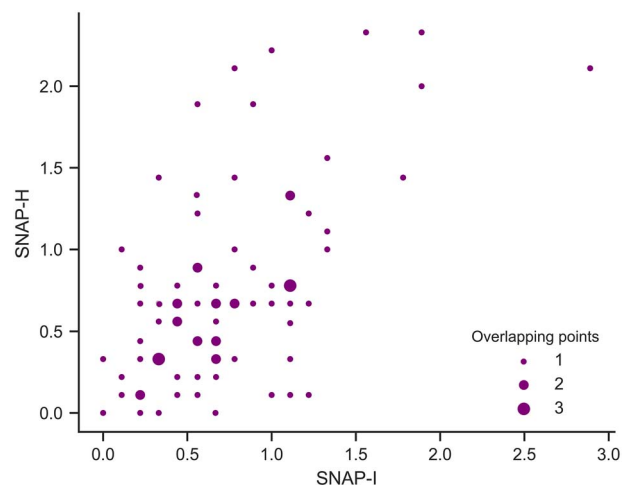


Fig. 1. Scatterplot of SNAP-I and SNAP-H scores. The size of points on the graph denotes the number of individuals located at the coordinates to capture overlapping data points. SNAP-I and SNAP-H are correlated at Spearman's $\rho = 0.612$, $P < 0.001$.

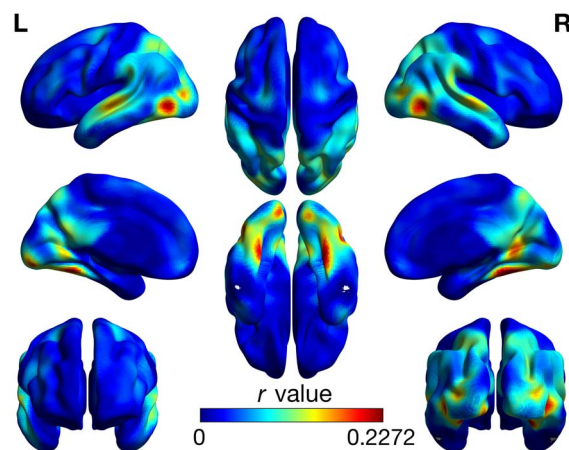


Fig. 2. Average groupwise ISC for the whole sample. Figure shows the unthresholded Pearson's r values for the whole-sample, average ISC.

(Fig. 3a; detailed cluster information can be found in Supplementary Table S2). Most associations were found in areas of the brain where the average group synchrony was relatively high (i.e. greater than $r = 0.05$; see Fig. 2).

Higher ISC was associated with greater pairwise inattention scores in the bilateral middle temporal gyri; the anterior right superior temporal gyrus and frontal pole; and the left occipital pole, lateral occipital cortex, supramarginal gyrus, temporal pole, orbital frontal cortex, and precentral gyrus. Models that controlled for FEF synchrony show similar results (Supplementary Fig. S1).

For all models, main findings remain generally unchanged after inclusion of the FEF covariate. Findings from these models are reported in Supplementary Figs. S1–S3.

Hyperactivity model results

Greater ISC was associated with decreased pairwise hyperactivity scores in bilateral visual occipital cortex, precentral gyri, middle frontal gyri, middle and superior

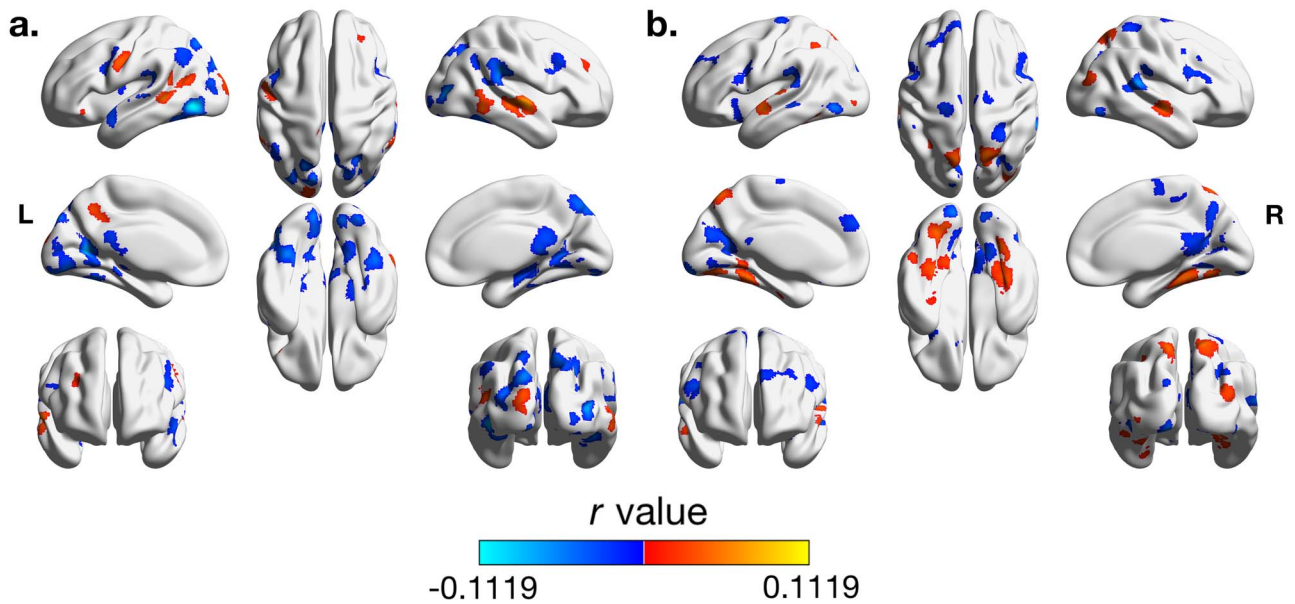


Fig. 3. Associations between ISC and average pairwise inattentive and hyperactive trait scores. a) The associations between inattention and ISC. b) The associations between hyperactivity and ISC. Color gradient indicates the β values, in units of Pearson's r . Cool colors denote negative associations (where ISC decreases as average trait score per pair increases) and warm colors denote positive associations (where ISC increases as average trait score per pair increases). For both inattention and hyperactivity, associations were seen in distributed areas of the occipital, temporal, parietal, and frontal lobes. Images are thresholded at a voxelwise threshold of $P < 0.001$ and a cluster-forming threshold of $\alpha = 0.05$.

temporal gyri, and superior cerebellum; right precuneus, posterior cingulate, intracalcarine cortex, postcentral gyrus, and superior parietal lobule; and left occipital pole, inferior and superior frontal gyri, supramarginal gyrus, frontal orbital cortex, and parietal operculum cortex (Fig. 3b; detailed cluster information can be found in Supplementary Table S3).

Greater ISC was associated with elevated pairwise hyperactivity scores in bilateral visual occipital cortex, superior temporal gyrus, fusiform gyri, lingual gyri, and superior cerebellum; right middle temporal gyrus; and left inferior temporal gyrus. Models controlling for FEF synchrony showed similar results to the original model (Supplementary Fig. S2).

Differences between inattention and hyperactivity

To determine whether there were dissociable relationships between ISC and inattention relative to hyperactivity, we created a third model that included both pairwise average inattention scores and pairwise average hyperactivity scores as variables. The contrast was calculated as inattention–hyperactivity, so that negative clusters are where the β for inattention scores was more negative than the β for hyperactivity scores, and vice versa for positive clusters (Fig. 4; detailed cluster information can be found in Supplementary Table S4). Pairwise inattention score had a more negative/less positive association with ISC in bilateral fusiform cortex and lateral occipital cortex; left inferior and superior gyri and supplementary motor area; and right temporoparietal junction. Hyperactivity score had a more negative/less positive association with ISC in bilateral precuneus, precentral gyrus, middle

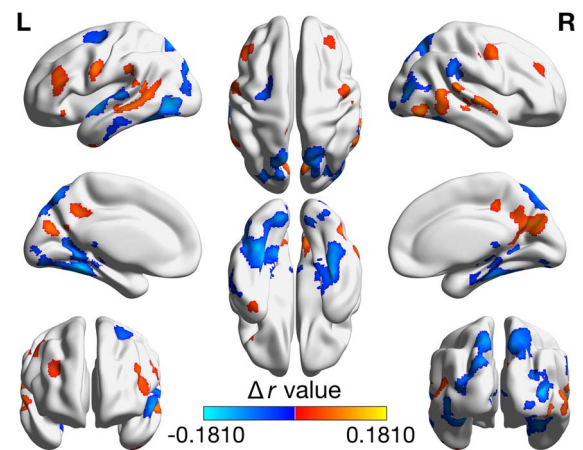


Fig. 4. Contrast between inattention and hyperactivity. Figure shows the difference in β values, in units of Pearson's r , corresponding to the inattention–hyperactivity contrast. Negative (cool) clusters indicate that pairwise inattention scores had a more negative/less positive association than pairwise hyperactivity scores, and positive (warm) clusters indicate that pairwise hyperactivity scores had a more negative/less positive association than pairwise inattention scores. Results are thresholded at a voxelwise threshold of $P < 0.001$ and a cluster-forming threshold of $\alpha = 0.05$.

temporal gyrus, and middle frontal gyrus; right superior gyrus; and left inferior frontal gyrus.

Discussion

This study investigated whether normative variation in inattentive and hyperactive traits in young children relates to interindividual brain synchrony during video viewing. We found that both average pairwise inattention and hyperactivity scores, as measured by the

SNAP-IV, show associations with interindividual synchrony in areas engaged by the video viewing task. For inattention, negative associations—where ISC was lower in pairs with higher average trait scores—were most notably found in both dorsal and ventral visual streams, temporal auditory processing areas, bilateral temporoparietal regions, and precuneus. Hyperactivity scores had a negative association with ISC in several areas including bilateral parietal association cortex, prefrontal cortex, and precuneus. Contrary to our hypothesis of primarily negative associations between ISC and trait scores, we found regions with significant positive associations for both inattention and hyperactivity. Our findings suggest that children's neural processing of complex audiovisual stimuli is associated with inattentive and hyperactive traits and highlight the importance of considering these traits separately and dimensionally in developmental neuroimaging research.

Relationships between inattention/hyperactivity and the brain in typically developing populations

Population-based studies have shown that across the spectrum of ADHD traits, subclinical symptoms of ADHD in childhood are linked to poorer academic performance at age 12 (Salla et al. 2016) and 16 (Sayal et al. 2015), as well as increased grade retention and failure to graduate (Galéra et al. 2009; Bussing et al. 2010). Higher inattention scores at 6 years old were also associated with a lower annual income 3 decades later (Vergunst et al. 2019). Despite the clear implication that subthreshold inattention and hyperactivity traits might affect future outcomes, the literature characterizing the dimensional relationships between trait inattention/hyperactivity and brain measures in undiagnosed individuals (especially in early childhood) has been limited. However, there has been some evidence from both population-based studies, as well as those specifically focusing on control samples with no reported diagnoses, that inattentive and hyperactive symptomology are related to brain structure and function in nonclinical cohorts. Measures such as cortical thinning (Shaw et al. 2011; Ducharme et al. 2012; Mous et al. 2014), putamen (Mous et al. 2015) and prefrontal gray matter volume (Albaugh et al. 2017, 2019), and functional connectivity (Hilger and Fiebach 2019; Rohr et al. 2019) have shown associations with inattention and/or hyperactivity in both children and adults in the general population and nonclinical samples. While none of these studies focused specifically on ISC, they are consistent with our finding of a relationship between the brain and normative attentive traits. We have expanded upon this previous work and shown that ISC also has an association with inattention and hyperactivity in the nonclinical population.

Separability of the inattention and hyperactivity trait dimensions in typically developing populations

In the clinical context, it has been suggested that ADHD is made up of a general underlying ADHD factor and

2 separable dimensions of inattention and hyperactivity/impulsivity (Toplak et al. 2009; Smith et al. 2013); however, it is unclear whether the separation of the inattention and hyperactivity dimensions extends into the normative spectrum of traits. Studies suggest that in the general population, inattention and hyperactivity have differential associations with cognitive measures (Kuntsi et al. 2014), symptom trajectories (Larsson et al. 2006, 2011), and educational attainment (Pingault et al. 2011). In terms of brain measures, inattention and hyperactivity have shown associations with gray matter volume in different areas of the brain in a population-based study (Albaugh et al. 2017), and Salmi et al. (2020) found different dimensional associations between ISC and inattention and hyperactivity in their adult control group. In our study, we also found different regional associations between ISC and inattention and hyperactivity in functionally diverse areas. Pairwise inattention scores had more negative/less positive relationships with ISC in areas related to sensory processing, motor planning (Abe and Hanakawa 2009), and attention/social cognition (Krall et al. 2015; Martin et al. 2019; Wilterson et al. 2021), relative to hyperactivity. This contrasts with the areas where we saw more negative/less positive associations between ISC and hyperactivity, which were found mostly in areas of language processing (Sliwinska et al. 2012; Liégeois et al. 2014; Yen et al. 2019), executive function (Friedman and Robbins 2022), and default mode (Utevsy et al. 2014; Li et al. 2019). This suggests that inattention and hyperactivity may preferentially affect different cognitive and processing systems, adding support that they are separable concepts. We note that because our sample did not include any children with an ADHD diagnosis, our findings may not generalize to children with clinically diagnosed ADHD.

Relationships between inattention/hyperactivity traits and interindividual synchrony

Previous work looking at ADHD and ISC in adults may also offer insight into how inattention and impulsivity relate to interindividual BOLD synchrony in the nonclinical population. In their study, Salmi et al. (2020) found that in their control group, increasing pairwise similarity in impulsivity scores was associated with greater ISC in the dorsomedial prefrontal cortex, while greater similarity in inattention scores was associated with higher ISC in a very small cluster in the precuneus. Our results—that there were widespread patterns of association between synchrony and both inattention and hyperactivity in young children without an ADHD diagnosis—reinforce the argument that the heterogeneity in inattention and hyperactivity in what researchers typically refer to as control groups in case-control studies could be reflective of a linked heterogeneity among brain structure, function, and behavioral traits. This has clear implications for conducting categorical studies of ADHD and its neural correlates.

Contrary to our hypotheses, we found several areas in both the inattention and hyperactivity analyses where

pairs with higher average symptoms levels are more synchronized to one another than pairs with low average scores. For inattention scores, this included temporal auditory processing areas, early visual occipital cortex, default mode network (precuneus), and left temporoparietal junction and precentral gyrus. Pairwise hyperactivity scores showed a positive association with ISC in downstream ventral visual areas, parietal association cortex, and bilateral temporal cortex. While these results were unexpected, we speculate that certain properties of the stimulus may be more salient to specific children depending on their attentive trait scores and capture their attention in a “bottom-up” way, resulting in a more stimulus-driven, synchronized response between individuals who score high on a trait. However, this interpretation will need to be confirmed in further research ideally integrating measures of attention such as eye-tracking.

Implications for screen media and education

Our study has implications for the use of AV media in educational settings. Recent work has suggested that interindividual synchrony (as measured with electroencephalography) can be used as a marker of “neural engagement” with an educational video stimulus, as an individual’s synchrony to the rest of the group was associated with better performance on a test of the video contents. Supporting the idea that engagement is reflected in interindividual synchrony, Song et al. (2021) found that ISC in the default mode network was higher during scenes in narrative movies that were deemed more “engaging” through participant self-report. Our findings, that both inattention and hyperactivity demonstrated associations with ISC, suggest that children’s engagement with educational video stimuli may vary with their inattentive and hyperactive traits. However, as we found both negative and positive associations with ISC for both traits, we cannot yet definitively characterize the nature of these relationships—for example, both positive and negative associations with inattention scores were found in the precuneus, an area often regarded as a core node of the DMN. Due to these mixed results, it is hard to surmise whether screen media, such as the television clips used in this study, are more or less engaging for children depending on their attentive traits. Further research should explore the relationships between inattentive and hyperactive traits, interindividual synchrony, and engagement with educational videos.

Limitations

This study had several strengths, including an early childhood age range, a dimensional perspective on inattentive and hyperactive traits, and a relatively long scan time (~18 min). The study also had several limitations to note. We found that children excluded for head motion had higher trait levels of inattention and hyperactivity than the children included in our sample

at the trend level, potentially limiting generalizability. Though we controlled for sex in our models, our sample had more females than males, which is not necessarily representative of the general population, where sex has a significant effect on type, prevalence, and severity of ADHD symptoms (Smidts and Oosterlaan 2007; Arnett et al. 2015). Finally, we were not able to collect accurate eye-tracking data, so we were unable to directly assess the impact of important confounds such as gaze behavior and visual attention to the stimulus. To partially mitigate this, we calculated models that used FEF synchrony as a covariate to approximate difference in visual attention and gaze behavior between individuals, as has been done in previous work (Redcay et al. 2010; Moraczewski et al. 2018). While the general findings remain unchanged after this control analysis, we cannot definitively conclude that patterns of visual attention to the stimulus were not a driving factor in the associations we found between inattention/hyperactivity and ISC. Further work needs to be done that investigates the extent to which visual attention and gaze behavior contribute to ISC.

Conclusions

This study found that inattentive and hyperactive traits are differentially associated with interindividual BOLD signal synchrony during the neural processing of videos by young children. An important future direction will be to ascertain whether these differences are linked with differences in understanding and retention, which could have implications for early childhood education. This work adds to a growing body of literature suggesting meaningful trait-linked variation in brain function in samples with no reported diagnoses.

Supplementary material

Supplementary material is available at *Cerebral Cortex Communications Journal* online.

Funding

This work was supported by Natural Sciences and Engineering Research Council of Canada (NSERC) Discovery Grant 435787 and Canadian Institutes of Health Research (CIHR) Project Grant 388385 awarded to SB, and a Cumming School of Medicine Graduate Scholarship and Hotchkiss Brain Institute Graduate Recruitment Scholarship awarded to RT.

Conflict of interest statement: The authors declare no competing interests.

References

- Abe M, Hanakawa T. Functional coupling underlying motor and cognitive functions of the dorsal premotor cortex. *Behav Brain Res.* 2009;198:13–23.
- Albaugh MD, Orr C, Chaarani B, Althoff RR, Allgaier N, D’Alborto N, Hudson K, Mackey S, Spechler PA, Banaschewski T, et al.

- Inattention and reaction time variability are linked to ventromedial prefrontal volume in adolescents. *Biol Psychiatry*. 2017;**82**:660–668.
- Albaugh MD, Ivanova M, Chaarani B, Orr C, Allgaier N, Althoff RR, D'Albeto N, Hudson K, Mackey S, Spechler PA, et al. Ventromedial prefrontal volume in adolescence predicts hyperactive/inattentive symptoms in adulthood. *Cereb Cortex*. 2019;**29**:1866–1874.
- Arabaci G, Parris BA. Probe-caught spontaneous and deliberate mind wandering in relation to self-reported inattentive, hyperactive and impulsive traits in adults. *Sci Rep*. 2018;**8**:1–10.
- Arnett AB, Pennington BF, Willcutt EG, Defries JC, Olson RK. Sex differences in ADHD symptom severity. *J Child Psychol Psychiatry Allied Discip*. 2015;**56**:632–639.
- Bottenhorn KL, Flannery JS, Boeving ER, Riedel MC, Eickhoff SB, Sutherland MT, Laird AR. Cooperating yet distinct brain networks engaged during naturalistic paradigms: a meta-analysis of functional MRI results. *Netw Neurosci*. 2018;**3**:27–48.
- Brown AJ, Casey BM. Subclinical ADHD-symptoms are associated with executive-functioning and externalizing problems in college students without ADHD-diagnoses. *J Educ Dev Psychol*. 2016;**6**:204.
- Bussing R, Fernandez M, Hardwood M, Hou W, Garvan C, Swanson J, Eyberg S. Parent and teacher SNAP-IV ratings of attention deficit/hyperactivity disorder symptoms: psychometric properties and normative ratings from a School District sample. *Assessment*. 2008;**15**:317–328.
- Bussing R, Mason DM, Bell L, Porter P, Garvan C. Adolescent outcomes of childhood attention-deficit/hyperactivity disorder in a diverse community sample. *J Am Acad Child Adolesc Psychiatry*. 2010;**49**:595–605.
- Byrge L, Dubois J, Tyszka JM, Adolphs R, Kennedy DP. Idiosyncratic brain activation patterns are associated with poor social comprehension in autism. *J Neurosci*. 2015;**35**:5837–5850.
- Cantlon JF, Li R. Neural activity during natural viewing of sesame street statistically predicts test scores in early childhood. *PLoS Biol*. 2013;**11**:1–13.
- Chen G, Shin YW, Taylor PA, Glen DR, Reynolds RC, Israel RB, Cox RW. Untangling the relatedness among correlations, part I: non-parametric approaches to inter-subject correlation analysis at the group level. *NeuroImage*. 2016;**142**:248–259.
- Chen G, Taylor PA, Shin YW, Reynolds RC, Cox RW. Untangling the relatedness among correlations, part II: inter-subject correlation group analysis through linear mixed-effects modeling. *NeuroImage*. 2017;**147**:825–840.
- Ciric R, Rosen AFG, Erus G, Cieslak M, Adebimpe A, Cook PA, Bassett DS, Davatzikos C, Wolf DH, Satterthwaite TD. Mitigating head motion artifact in functional connectivity MRI. *Nat Protoc*. 2018;**13**:2801–2826.
- Cox RW. AFNI: software for analysis and visualization of functional magnetic resonance neuroimages. *Comput Biomed Res*. 1996;**29**:162–173.
- Dosenbach NUF, Koller JM, Earl EA, Miranda-Dominguez O, Klein RL, Van AN, Snyder AZ, Nagel BJ, Nigg JT, Nguyen AL, et al. Real-time motion analytics during brain MRI improve data quality and reduce costs. *NeuroImage*. 2017;**161**:80–93.
- Ducharme S, Hudziak JJ, Botteron KN, Albaugh MD, Nguyen TV, Karama S, Evans AC. Decreased regional cortical thickness and thinning rate are associated with inattention symptoms in healthy children. *J Am Acad Child Adolesc Psychiatry*. 2012;**51**:18–27.e2.
- Finn ES, Glerean E, Khojandi AY, Nielson D, Molfese PJ, Handwerker DA, Bandettini PA. Idiosyncrasy: from shared responses to individual differences during naturalistic neuroimaging. *NeuroImage*. 2020;**215**:116828.
- Fonov V, Evans A, McKinstry R, Almlí C, Collins D. Unbiased non-linear average age-appropriate brain templates from birth to adulthood. *NeuroImage*. 2009;**47**:S102.
- Fonov V, Evans AC, Botteron K, Almlí CR, McKinstry RC, Collins DL. Unbiased average age-appropriate atlases for pediatric studies. *NeuroImage*. 2011;**54**:313–327.
- Frick MA, Asherson P, Brocki KC. Mind-wandering in children with and without ADHD. *Br J Clin Psychol*. 2020;**59**:208–223.
- Friedman NP, Robbins TW. The role of prefrontal cortex in cognitive control and executive function. *Neuropsychopharmacology*. 2022;**47**:72–89.
- Galéra C, Melchior M, Chastang JF, Bouvard MP, Fombonne E. Childhood and adolescent hyperactivity-inattention symptoms and academic achievement 8 years later: the GAZEL youth study. *Psychol Med*. 2009;**39**:1895–1906.
- Gorgolewski K, Burns CD, Madison C, Clark D, Halchenko YO, Waskom ML, Ghosh SS. Nipype: a flexible, lightweight and extensible neuroimaging data processing framework in Python. *Front Neuroinform*. 2011;**5**:1–15.
- Graff K, Tansey R, Ip A, Rohr C. Benchmarking common preprocessing strategies in early childhood functional connectivity and intersubject correlation fMRI. *Dev Cogn Neurosci*. 2022;**54**:101087.
- Greene DJ, Koller JM, Hampton JM, Wesevich V, Van AN, Nguyen AL, Hoyt CR, McIntyre L, Earl EA, Klein RL, et al. Behavioral interventions for reducing head motion during MRI scans in children. *NeuroImage*. 2018;**171**:234–245.
- Greve DN, Fischl B. Accurate and robust brain image alignment using boundary-based registration. *NeuroImage*. 2009;**48**:63–72.
- Gruskin DC, Rosenberg MD, Holmes AJ. Relationships between depressive symptoms and brain responses during emotional movie viewing emerge in adolescence. *NeuroImage*. 2020;**216**:116217.
- Hasson U, Nir Y, Levy I, Fuhrmann G, Malach R. Intersubject synchronization of cortical activity during natural vision. *Science (80-)*. 2004;**303**:1634–1640.
- Hasson U, Landesman O, Knappmeyer B, Vallines I, Rubin N, Heeger DJ. Neurocinematics: the neuroscience of film. *PRO*. 2008;**2**:1–26.
- Hilger K, Fiebach CJ. ADHD symptoms are associated with the modular structure of intrinsic brain networks in a representative sample of healthy adults. *Netw Neurosci*. 2019;**3**:567–588.
- Hoyer RS, Elshafei H, Hemmerlin J, Bouet R, Bidet-Caulet A. Why are children so distractible? Development of attention and motor control from childhood to adulthood. *Child Dev*. 2021;**92**:e716–e737.
- Huang CM, Lee SH, Hsiao IT, Kuan WC, Wai YY, Ko HJ, Wan YL, Hsu YY, Liu HL. Study-specific EPI template improves group analysis in functional MRI of young and older adults. *J Neurosci Methods*. 2010;**189**:257–266.
- Jenkinson M, Bannister P, Brady M, Smith S. Improved optimization for the robust and accurate linear registration and motion correction of brain images. *NeuroImage*. 2002;**17**:825–841.
- Jonkman LM, Markus CR, Franklin MS, Van Dalen JH. Mind wandering during attention performance: effects of ADHD-inattention symptomatology, negative mood, ruminative response style and working memory capacity. *PLoS One*. 2017;**12**:1–19.
- Krall SC, Rottschy C, Oberwelland E, Bzdok D, Fox PT, Eickhoff SB, Fink GR, Konrad K. The role of the right temporoparietal junction in attention and social interaction as revealed by ALE meta-analysis. *Brain Struct Funct*. 2015;**220**:587–604.
- Kuntsi J, Pinto R, Price TS, Van Der Meere JJ, Frazier-Wood AC, Asherson P. The separation of ADHD inattention and hyperactivity-impulsivity symptoms: pathways from genetic effects to cognitive impairments and symptoms. *J Abnorm Child Psychol*. 2014;**42**:127–136.

- Larsson H, Lichtenstein P, Larsson JO. Genetic contributions to the development of ADHD subtypes from childhood to adolescence. *J Am Acad Child Adolesc Psychiatry*. 2006;**45**:973–981.
- Larsson H, Dilshad R, Lichtenstein P, Barker ED. Developmental trajectories of DSM-IV symptoms of attention-deficit/hyperactivity disorder: genetic effects, family risk and associated psychopathology. *J Child Psychol Psychiatry Allied Discip*. 2011;**52**:954–963.
- Li R, Utevsky AV, Huettel SA, Braams BR, Peters S, Crone EA, van Duijvenvoorde ACK. Developmental maturation of the Precuneus as a functional Core of the default mode network. *J Cogn Neurosci*. 2019;**31**:1506–1519.
- Liégeois F, Mayes A, Morgan A. Neural correlates of developmental speech and language disorders: evidence from neuroimaging. *Curr Dev Disord Rep*. 2014;**1**:215–227.
- Martin AK, Kessler K, Cooke S, Huang J, Meinzer M. The right temporoparietal junction is causally associated with embodied perspective taking. *J Neurosci*. 2019;**40**:3089–3095.
- Moraczewski D, Chen G, Redcay E. Inter-subject synchrony as an index of functional specialization in early childhood. *Sci Rep*. 2018;**8**:1–12.
- Mous SE, Muetzel RL, El Marroun H, Polderman TJC, Van Der Lugt A, Jaddoe VW, Hofman A, Verhulst FC, Tiemeier H, Posthuma D, et al. Cortical thickness and inattention/hyperactivity symptoms in young children: a population-based study. *Psychol Med*. 2014;**44**:3203–3213.
- Mous SE, Hammerschlag AR, Polderman TJC, Verhulst FC, Tiemeier H, Van Der Lugt A, Jaddoe VW, Hofman A, White T, Posthuma D. A population-based imaging genetics study of inattention/hyperactivity: basal ganglia and genetic pathways. *J Am Acad Child Adolesc Psychiatry*. 2015;**54**:745–752.
- Nastase SA, Gazzola V, Hasson U, Keysers C. Measuring shared responses across subjects using intersubject correlation. *Soc Cogn Affect Neurosci*. 2019:669–687.
- Pajula J, Tohka J. Effects of spatial smoothing on inter-subject correlation based analysis of fMRI. *Magn Reson Imaging*. 2014;**32**:1114–1124.
- Pingault JB, Tremblay RE, Vitaro F, Carbonneau R, Genolini C, Falissard B, Cote SM. Childhood trajectories of inattention and hyperactivity and prediction of educational attainment in early adulthood: a 16-year longitudinal population-based study. *Am J Psychiatry*. 2011;**168**:1164–1170.
- Power JD, Barnes KA, Snyder AZ, Schlaggar BL, Petersen SE. Spurious but systematic correlations in functional connectivity MRI networks arise from subject motion. *NeuroImage*. 2012;**59**:2142–2154.
- Power JD, Mitra A, Laumann TO, Snyder AZ, Schlaggar BL, Petersen SE. Methods to detect, characterize, and remove motion artifact in resting state fMRI. *NeuroImage*. 2014;**84**:320–341.
- Power JD, Plitt M, Kundu P, Bandettini PA, Martin A. Temporal interpolation alters motion in fMRI scans: magnitudes and consequences for artifact detection. *PLoS One*. 2017;**12**:1–20.
- Redcay E, Dodell-Feder D, Parrow MJ, Mavros PL, Kleiner M, Gabrieli JDE, Saxe R. Live face-to-face interaction during fMRI: a new tool for social cognitive neuroscience. *NeuroImage*. 2010;**50**:1639–1647.
- Rielly NE, Craig WM, Parker KCH. Peer and parenting characteristics of boys and girls with subclinical attention problems. *J Atten Disord*. 2006;**9**:598–606.
- Rohr CS, Dimond D, Schuetze M, Cho IYK, Lichtenstein-Vidne L, Okon-Singer H, Dewey D, Bray S. Girls' attentive traits associate with cerebellar to dorsal attention and default mode network connectivity. *Neuropsychologia*. 2019;**127**:84–92.
- Salla J, Michel G, Pingault JB, Lacourse E, Paquin S, Galéra C, Falissard B, Boivin M, Tremblay RE, Côté SM. Childhood trajectories of inattention-hyperactivity and academic achievement at 12 years. *Eur Child Adolesc Psychiatry*. 2016;**25**:1195–1206.
- Salmi J, Metwaly M, Tohka J, Alho K, Leppämäki S, Tani P, Koski A, Vanderwal T, Laine M. ADHD desynchronizes brain activity during watching a distracted multi-talker conversation. *NeuroImage*. 2020;**216**:1–11.
- Sayal K, Washbrook E, Propper C. Childhood behavior problems and academic outcomes in adolescence: longitudinal population-based study. *J Am Acad Child Adolesc Psychiatry*. 2015;**54**:360–368.e2.
- Shaw P, Gilliam M, Liverpool M, Weddle C, Malek M, Sharp W, Greenstein D, Evans A, Rapoport J, Giedd J. Cortical development in typically developing children with symptoms of hyperactivity and impulsivity: support for a dimensional view of attention deficit hyperactivity disorder. *Am J Psychiatry*. 2011;**168**:143–151.
- Sliwiska MW, Khadilkar M, Campbell-Ratcliffe J, Quevenco F, Devlin JT. Early and sustained supramarginal gyrus contributions to phonological processing. *Front Psychol*. 2012;**3**:1–10.
- Smidts DP, Oosterlaan J. How common are symptoms of ADHD in typically developing preschoolers? A study on prevalence rates and prenatal/demographic risk factors. *Cortex*. 2007;**43**:710–717.
- Smith LC, Tamm L, Hughes CW, Bernstein IH. Separate and overlapping relationships of inattention and hyperactivity/impulsivity in children and adolescents with attention-deficit/hyperactivity disorder. *ADHD Atten Deficit Hyperact Disord*. 2013;**5**:9–20.
- Song H, Finn ES, Rosenberg MD. Neural signatures of attentional engagement during narratives and its consequences for event memory. *Proc Natl Acad Sci U S A*. 2021;**118**:1–12.
- Sonkusare S, Breakspear M, Guo C. Naturalistic stimuli in neuroscience: critically acclaimed. *Trends Cogn Sci*. 2019;**23**:699–714.
- Tang C, Huang Z, Zhou S, Wang Q, Yi F, Nie J. 2019. Movie-watching fMRI reveals inter-subject synchrony alteration in functional brain activity in ADHD. In: *Lecture notes in computer science (including subseries lecture notes in artificial intelligence and lecture notes in bioinformatics)*. Cham, Switzerland: Springer International Publishing, p. 104–111.
- Toplak ME, Pitch A, Flora DB, Iwenofu L, Ghelani K, Jain U, Tannock R. The unity and diversity of inattention and hyperactivity/impulsivity in ADHD: evidence for a general factor with separable dimensions. *J Abnorm Child Psychol*. 2009;**37**:1137–1150.
- Tustison NJ, Cook PA, Klein A, Song G, Das SR, Duda JT, Kandel BM, van Strien N, Stone JR, Gee JC, et al. Large-scale evaluation of ANTs and FreeSurfer cortical thickness measurements. *NeuroImage*. 2014;**99**:166–179.
- Urchs S, Armoza J, Moreau C, Benhajali Y, St-Aubin J, Orban P, Bellec P. MIST: a multi-resolution parcellation of functional brain networks. *MNI Open Res*. 2019;**1**:3.
- Utevsky AV, Smith DV, Huettel SA. Precuneus is a functional core of the default-mode network. *J Neurosci*. 2014;**34**:932–940.
- Vanderwal T, Kelly C, Eilbott J, Mayes LC, Castellanos FX. Inscapes: a movie paradigm to improve compliance in functional magnetic resonance imaging. *NeuroImage*. 2015;**122**:222–232.
- Vergunst F, Tremblay RE, Nagin D, Algan Y, Beasley E, Park J, Galera C, Vitaro F, Côté SM. Association between childhood Behaviors and adult employment earnings in Canada. *JAMA Psychiat*. 2019;**76**:1044–1051.
- Wilterson AI, Nastase SA, Bio BJ, Guterstam A, Graziano MSA. Attention, awareness, and the right temporoparietal junction. *Proc Natl Acad Sci U S A*. 2021;**118**:1–10.
- Xia M, Wang J, He Y. BrainNet viewer: a network visualization tool for human brain Connectomics. *PLoS One*. 2013;**8**:1–15.
- Yen M, DeMarco AT, Wilson SM. Adaptive paradigms for mapping phonological regions in individual participants. *NeuroImage*. 2019;**189**:368–379.