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ASD symptoms in adults with ADHD: a comparative study using ADOS-2

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Abstract

In this study, we examined autism spectrum disorder (ASD) symptoms in adults with attention-deficit hyperactivity disorder (ADHD) in comparison with normal controls using the Autism Diagnostic Observation Schedule, Second Edition (ADOS-2). Sixty-three adults with ADHD (mean age, 35.3 years; 38 men) and 31 controls (mean age, 38.7 years; 17 men) completed Module 4 of the ADOS-2, Autism Spectrum Quotient, Conners' Adult ADHD Rating Scale, and Wechsler Adult Intelligence Scale, Third Edition. Adults with ADHD were not clinically diagnosed with ASD, and their ADHD diagnosis was based on the Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition criteria. Between-group comparisons on the scoring patterns revealed significant ASD symptoms present in adults with ADHD, which was congruent with our previous study. Further, item level and correlation analyses showed the possibility that ASD symptoms in adult ADHD comprised of two distinct mechanisms, one qualitatively similar to ASD and the other arising from ADHD characteristics, highlighting the complex nature of ADHD-ASD symptom overlaps.

Keywords ADOS-2 · Adult · Attention-deficit/hyperactivity disorder · Autism spectrum disorder · Comparative study

Introduction

In the heat of ever-growing interests in adult attention-deficit/hyperactivity disorder (ADHD) among researchers, one of the unsolved questions is how ADHD is related to autism spectrum disorder (ASD). While the former is characterized by inattention, hyperactivity, and impulsivity, the latter presents deficits in social reciprocal communication and restricted and repetitive behaviors or interests [1]. Despite the long history of being considered as etiologically different and mutually exclusive disorders, literature now suggests that ADHD and ASD frequently co-occur [2-5], and their overlaps may not be limited to symptoms but extend to shared genetic underpinnings [3, 6-8]. In studies with children, up to 50% of individuals with ADHD were found to exhibit some form of ASD symptoms [3-5, 9, 10], and importantly, the co-existing ASD symptoms affect various clinical and functional outcomes of children with ADHD

☑ Wakaho Hayashi wakahoh@med.showa-u.ac.jp; wakaho15@gmail.com [11, 12]. For example, Cooper et al. [11] found that higher levels of autistic traits were associated with more severe ADHD symptoms, more psychiatric comorbidities, lower intelligence quotient (IQ), and greater motor and language problems in children with ADHD. Given the complex social pressure in adult life and the effect of developmental sequel, it is possible that adults with ADHD may be more affected by ASD symptoms compared to children, and their consequences can be profound. It is, therefore, crucial to evaluate the detailed profile and extent of ASD symptoms exhibited by adults with ADHD. However, only a handful of studies have examined such status in adults, and none used objective clinical measures. This led to our previous study, which examined ASD symptoms in adults with ADHD using the Autism Diagnostic Observation Schedule, Second Edition (ADOS-2) [13].

The ADOS-2 is a standardized, well-validated measure for assessing ASD symptoms and diagnosis [14]. Its uniqueness lies in the objectivity, thoroughness, and reliability of the assessment, in which a well-trained administrator performs a direct observation of an individual in a series of semi-structured plays and interviews. Adults with fluent speech receive Module 4, consisting of 15 activities that are rated using 32 items based on specific evaluation criteria.

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The scores of the specific items are converted into domain scores and then to a final algorithm score, which determines ASD or non-ASD classification. Both the original and recently revised Module 4 algorithms have good sensitivity (original, 89.6; revised, 90.5) and specificity (original, 72.2; revised: 82.2) [15]. While the original algorithm produces two domain scores: Communication and Reciprocal Social Interaction (RSI), the revised algorithm yields three domain scores: Language and Communication (LC), RSI, and Restricted and Repetitive Behaviors (RRB). The sum of LC and RSI is called Social Affect (SA), and together with RRB, it matches the two domain ASD criteria of the Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition (DSM-5) [1].

In our preliminary study, we examined ASD symptoms in a clinical sample of adults with ADHD using the ADOS-2 for the first time [13]. Though the participants showed no apparent ASD symptoms in childhood and therefore were not clinically diagnosed with ASD, 23.3% met the ASD cut-off on the ADOS-2. While the domain level analysis showed significant scoring on SA than RRB items, indicating the evident challenges in communication and reciprocal social interaction in adults with ADHD, item-level analysis revealed that (1) the quantitative rather than qualitative aspect of social communication is compromised in individuals with ADHD and (2) emotion processing and affective interaction may also be impaired in ADHD. These results were in line with previous studies, indicating that individuals with ADHD tend to be one-sided in their communication and lack relational reciprocity [9, 16–19]. Difficulty in affect recognition [20] and lack of emotional empathy [21] are also identified in individuals with ADHD. While our study successfully revealed the profile of more objective ASD symptoms present in adults with ADHD, only those with ADHD were involved, and as many as 11 items were endorsed by more than 30% of the sample. A comparative study was necessary to obtain a better picture of ASD symptoms in adults with ADHD. Therefore, we conducted this consecutive study in which the scoring patterns on the ADOS-2 were compared between adults with ADHD and healthy controls.

Methods

Participants

Participants were recruited from outpatients and inpatients at Showa University Karasuyama Hospital, central Tokyo, Japan, between October 2018 and December 2020. Only those fulfilling the DSM-5 criteria [1] for ADHD and with a full-scale IQ of over 85 on the Japanese version of the Wechsler Adult Intelligence Scale-Third Edition (WAIS-III) [22] were included. The exclusion criteria included age < 17 years and the presence of psychiatric disorders other than ADHD according to the DSM-5 criteria. The control group of neurotypical individuals was recruited from volunteer relatives or acquaintances of healthcare workers. In addition to the abovementioned exclusion criteria, those diagnosed with psychiatric disorders according to the DSM-5 criteria and those having first-order relatives with neurodevelopmental disorders were not included as controls. The final sample comprised 63 adults (56 from our previous study [13]) with ADHD and 31 neurotypical individuals.

Procedure

The detailed diagnostic process is explained elsewhere [13]. All participants were individually administered a comprehensive clinical assessment battery, which included (1) two sets of thorough interviews with participants and their caretakers on their developmental and medical history by psychiatrists, (2) the Japanese version of Conners' Adult ADHD Diagnostic Interview for DSM-IV (CAADID) [23, 24], and (3) reviews of the maternity record books and the elementary school (6-12 years old) reports. The final diagnosis of ADHD and comorbid mental disorders, if any, was determined by a consensus between psychiatrists and psychologists who were involved in this process based on the DSM-5 criteria. ASD diagnosis was a good clinical estimate (GCE), as neither the ADOS-2 nor the Autism Diagnostic Interview-Revised (ADI-R) [25] was employed in the assessment process. However, all psychiatrists and psychologists involved in this study are experts in neurodevelopmental disorders, and the clinical assessment was thorough with multiple resources. Individuals with a GCE of ASD diagnosis were excluded.

Control group participants underwent a 30-min interview with a psychiatrist to confirm the absence of mental disorders in themselves and any neurodevelopmental disorder in their first-order relatives.

The above diagnostic process was followed by the administration of the following assessments to all participants: (1) the ADOS-2 Module 4; (2) the Japanese version of the Conners' Adult ADHD Rating Scale-Self-Report: Long Version (CAARS-S:L-J), in which subjective ADHD symptoms were evaluated [26, 27]; (3) the autism-spectrum quotient (AQ) for measuring subjective ASD symptoms [28, 29], in which 33 is considered as the suitable cut-off [29]; and (4) the Japanese version of WAIS-III [22] to evaluate the IQ profile.

Autism diagnostic observation schedule, second edition

The ADOS-2 Module 4 (standard and all optional activities) was administered by trained and certified psychiatrists, and both the original (cut-off of 7) and revised algorithms (clinical cut-off of 8 and research cut-off of 10 [15]) were scored. Details of the administration and scoring processes are explained in the previous study, and our administration has a good inter-rater reliability [13].

Statistical analysis

All statistical analyses were performed using SPSS version 25.0 (IBM Corp., Armonk, NY, USA). The ADHD and control groups were compared on demographic (age, sex, and years of education) and clinical measurements (AQ, CAARS scores, and WAIS-III), as well as on the domain scores and proportions of individuals meeting the ASD cut-offs of the original and revised ADOS-2 algorithms. Independent samples t tests and chi-square tests were used for continuous and categorical variables, respectively. For detailed item-level analysis, the proportion of participants with each score (0, 1, 1)2, 3, 7, and 8) on every ADOS-2 item was calculated for the two groups. Additionally, the proportion of non-zero scorers (those who scored 1, 2, 3, or 8 for B3) on each ADOS-2 item was compared between the ADHD and control groups using chi-square tests. To assess the relationship between each measurement, Pearson product-moment correlation coefficients were calculated between AQ, CAARS, WAIS-III, and ADOS-2 (revised algorithm) scores for the groups. Further, we created a "typical ADHD" group by excluding participants who met the ASD cut-off (clinical cut-off of the revised algorithm) from the ADHD group and compared the characteristics between this group and the control group by performing the abovementioned analyses. The significance level was set at 0.05, except for the correlations, for which 0.01 was set to account for the possibility of a type I error.

Ethics

This study was evaluated and approved by the Medical Ethics Committee of Showa University School of Medicine, and the protocols were carried out in accordance with the Declaration of Helsinki. Written informed consent was obtained from all included participants after information was provided about the study.

Results

Demographic and clinical characteristics

A total of 94 participants, 63 adults with ADHD (mean [M] age, 35.3 years, standard deviation [SD] = 10.9 years, 38 men) and 31 controls (M age, 38.7 years, SD = 6.7 years, 17 men) were analyzed and compared. Their demographics and clinical characteristics are shown in Table 1. The background demographics showed no group differences, in which both groups had relatively high average years of education and slight male preponderance. The average full, verbal, and performance IQs were all above 100 and showed no group

 Table 1
 Demographic and clinical characteristics of the ADHD and control groups

	ADHD $(n=63)$		Control $(n=31)$		t test	p
	Mean (SD)	Range	Mean (SD)	Range	t	
Age	35.3 (10.9)	17–64	38.7 (6.7)	29–52	1.86	ns
Sex (male) <i>N</i> (%)	38	60.3	17	54.8	0.26*	ns**
Years of education	15.3 (1.8)	12-18	15.8 (1.3)	12–18	1.54	ns
AQ:Total	33.4 (8.0)	11-48	13.4 (8.1)	4–35	- 11.27	< 0.001
CAARS:Inattention/Memory Problems	77.4 (12.0)	37–90	51.6 (10.1)	35-78	- 10.90	< 0.001
CAARS:Hyperactivity/Restlessness	64.3 (16.7)	36–90	53.8 (13.0)	34–90	- 3.34	0.001
CAARS:Impulsivity/Emotional Lability	70.0 (15.1)	40–90	49.3 (10.9)	34-70	- 7.61	< 0.001
CAARS: Problems with Self-Concept	69.8 (11.1)	39-87	49.0 (10.7)	34–73	- 8.71	< 0.001
CAARS:DSM-IV Inattentive	78.6 (12.6)	44–90	51.6 (9.5)	37–77	- 11.55	< 0.001
CAARS:DSM-IV Hyperactive-Impulsive	68.4 (18.1)	38–90	51.4 (12.1)	36 - 86	- 5.37	< 0.001
CAARS:DSM-IV Total ADHD Symptoms	75.9 (14.0)	46–90	51.7 (10.6)	36–79	- 9.34	< 0.001
CAARS: ADHD Index	75.1 (11.3)	43–90	48.5 (8.5)	34–72	- 12.69	< 0.001
WAIS-III:FIQ	106.2 (11.4)	85-136	107.4 (9.5)	91-129	0.50	ns
WAIS-III:VIQ	107.3 (12.2)	81-137	106.1 (10.1)	85-131	- 0.53	ns
WAIS-III:PIQ	104.0 (14.1)	72–134	107.6 (11.1)	87–128	1.33	ns

p < 0.05 by independent samples t test (**chi-square test)

ns non-significant, AQ Autism spectrum quotient, CAARS Conners' Adult ADHD Rating Scale, FIQ full-scale intelligence quotient, PIQ performance intelligence quotient, VIQ verbal intelligence quotient, WAIS-III Wechsler Adult Intelligence Scale-Third Edition

*Chi-square value

differences. The ADHD group scored significantly higher than the control group not only on all CAARS scales but also on AQ total score. The typical ADHD group comprised of 48 individuals (M age, 35.3 years, SD=11.1 years, 29 men), and its comparison against controls took the same pattern as that of the ADHD group, with no background differences but significantly higher AQ and CAARS scores than the control group (Table 2).

ADOS-2 domain and total scores

Table 3 shows the final scores of the original and revised ADOS-2 algorithms and the percentage of individuals who met the ASD cut-off. While no one in the control group met the ASD cut-off, a total of 11 (17.5%), 15 (23.8%), and 11 (17.5%) individuals met the ASD cut-off by the original, revised clinical, and revised research algorithms, respectively, in the ADHD group. All average domain and total scores of the original and revised algorithms were significantly higher in the ADHD group than those in controls.

Comparison between adults with ADHD and controls on ADOS-2 scoring

Table 4 compares the proportion of non-zero scorers on each ADOS-2 item between the ADHD and control groups. Eleven items differentiated the two groups, in which adults with ADHD scored significantly higher than controls on A2 (Speech Abnormalities Associated With Autism), A8 (Conversation), B1 (Unusual Eye Contact), B2 (Facial Expressions Directed to Examiner), B3 (Language Production and Linked Nonverbal Communication), B4 (Shared Enjoyment in Interaction), B7 (Insight Into Typical Social Situations and Relationships), B8 (Responsibility), B9 (Quality of Social Overtures), B11 (Quality of Social Response), and B12 (Amount of Reciprocal Social Communication). No item in C (Imagination), D (Stereotyped Behaviors and Restricted Interests), and E (Other Abnormal Behaviors) domains showed a significant group difference.

Table 5 compares the proportion of non-zero scorers on each ADOS-2 item between the typical ADHD and control groups. Only six items differentiated the two groups: A2, B2, B3, B4, B8, and B12. A8, B1, B7, B9, and B11 no longer showed significant differences between the typical ADHD and control groups.

The detailed scoring patterns on each ADOS-2 item by the groups are depicted in Fig. 1a (A: Language and Communication), Fig. 1b (B: Reciprocal Social Interaction), and Fig. 1c (C: Imagination, D: Stereotyped Behaviors and Restricted Interests, E: Other Abnormal Behaviors). Fiftytwo (82.5%) individuals in the ADHD group scored on at least two of A, B, and D domains, which are relatively important for the algorithm.

 Table 2
 Demographic and clinical characteristics of the typical ADHD and control groups

	Typical ADHD	(n = 48)	Control $(n=31)$	1	t test	р
	Mean (SD)	Range	Mean (SD)	Range	t	
Age	35.3 (11.1)	17–64	38.7 (6.7)	29 -52	1.68	ns
Sex (male) <i>N</i> (%)	29	60.4	17	54.8	0.24*	ns**
Years of education	15.2 (1.8)	12-18	15.8 (1.3)	12-18	1.74	ns
AQ:Total	32.6 (7.6)	17–47	13.4 (8.1)	4–35	- 10.53	< 0.001
CAARS:Inattention/Memory Problems	77.3 (12.4)	37–90	51.6 (10.1)	35-78	- 10.07	< 0.001
CAARS:Hyperactivity/Restlessness	64.3 (17.2)	36–90	53.8 (13.0)	34–90	- 3.08	0.003
CAARS:Impulsivity/Emotional Lability	70.8 (14.7)	42–90	49.3 (10.9)	34–70	- 7.43	< 0.001
CAARS:Problems with Self-Concept	69.0 (11.6)	39–87	49.0 (10.7)	34–73	- 7.81	< 0.001
CAARS:DSM-IV Inattentive	78.2 (13.4)	44–90	51.6 (9.5)	37–77	- 10.26	< 0.001
CAARS:DSM-IV Hyperactive-Impulsive	69.6 (17.9)	39–90	51.4 (12.1)	36 - 86	- 5.38	< 0.001
CAARS:DSM-IV Total ADHD Symptoms	76.0 (14.5)	46–90	51.7 (10.6)	36–79	- 8.59	< 0.001
CAARS:ADHD Index	74.7 (12.0)	43–90	48.5 (8.5)	34–72	- 11.35	< 0.001
WAIS-III:FIQ	105.3 (11.5)	85-136	107.4 (9.5)	91–129	0.85	ns
WAIS-III:VIQ	107.2 (12.3)	81-134	106.1 (10.1)	85-131	- 0.43	ns
WAIS-III:PIQ	102.7 (15.2)	72–134	107.6 (11.1)	87-128	1.65	ns

p < 0.05 by independent samples t test (**chi-square test)

ns non-significant, AQ Autism spectrum quotient, CAARS Conners' Adult ADHD Rating Scale, FIQ full-scale intelligence quotient, PIQ performance intelligence quotient, VIQ verbal intelligence quotient, WAIS-III Wechsler Adult Intelligence Scale-Third Edition

*Chi-square value

	ADHD $(n=63)$		Control $(n=31)$		t test	р
	Mean (SD)	Range	Mean (SD)	Range	t	
ADOS-2: Orig COM	1.35 (1.43)	0–6	0.74 (0.82)	0–3	- 2.62	0.010
ADOS-2: Orig RSI	2.40 (2.30)	0–9	0.45 (0.57)	0–2	- 6.34	< 0.001
ADOS-2: Orig Total	3.75 (3.43)	0-15	1.19 (0.95)	0–3	- 5.50	< 0.001
ADOS-2: Rev LC	0.92 (0.99)	0–4	0.52 (0.63)	0–2	- 2.41	0.018
ADOS-2: Rev RSI	3.52 (2.87)	0-11	1.19 (1.08)	0–4	- 5.69	< 0.001
ADOS-2: Rev SA	4.44 (3.68)	0-14	1.71 (1.24)	0–6	- 5.31	< 0.001
ADOS-2: Rev RRB	0.65 (1.00)	0–5	0.10 (0.30)	0-1	- 4.03	< 0.001
ADOS-2: Rev Total	5.10 (4.18)	0-15	1.81 (1.33)	0–6	- 5.69	< 0.001
	N	%	N	%	X^2	
Orig Autism Spectrum/Autism	11	17.5	0	0.0	6.13	0.013
Rev Clinical cut-off ASD	15	23.8	0	0.0	8.78	0.003
Rev Research cut-off ASD	11	17.5	0	0.0	6.13	0.013

Table 3 Mean domain scores and the percentages of individuals meeting the ASD cut-offs of the original and revised ADOS-2 algorithms for the ADHD and the control groups

p < 0.05 by independent samples t test or chi-square test

ADOS-2 Autism Diagnostic Observation Schedule, Second Edition, ASD Autism spectrum disorder, Com communication, LC language and Communication, Orig Original Algorithm, Rev revised Algorithm, RRB restricted and repetitive behaviors, RSI reciprocal social interaction, SA social affect

Relationship between the ADOS-2 and other clinical measures

Tables 6, 7, and 8 show the correlations between AQ, CAARS, WAIS-III, and ADOS-2 scores in the ADHD, control, and typical ADHD groups, respectively. While ADOS-2 scores did not correlate with AQ, CAARS, or WAIS-III in the ADHD and control groups, ADOS-2 RSI, SA, and total scores positively correlated with CAARS Impulsivity/Emotional Lability score in the typical ADHD group. ADOS-2 intra-scale correlations also differed between the groups, in which RRB scores were correlated with LC, RSI, and SA scores only in the ADHD group.

Discussion

This is a consecutive study to our first attempt in evaluating ASD symptoms in clinical adults with ADHD using the ADOS-2. Herein, we compared the scoring patterns on the ADOS-2 between adults with ADHD who were not clinically diagnosed with ASD and typically developing controls. The overall analysis clearly indicated that adults with ADHD have significant ASD symptoms, particularly impairment in reciprocal social interaction. Item-level and correlational analyses revealed impairment in both qualitative and quantitative aspects of social communication; however, ASD symptoms in ADHD may not be a uniform construct.

Consistent with our preliminary study, approximately 20% of adults with ADHD without apparent childhood

ASD symptoms met the ASD cut-off on the ADOS-2 as opposed to none among the controls. Moreover, all domain and final algorithm scores were significantly higher in individuals with ADHD than the controls. These results clearly indicate that adults with ADHD have symptoms, similar to ASD, of both impairment in social communication and restricted and repetitive behaviors and interests. It is possible that, as some researchers postulate, ADHD and ASD are overarching disorders [30, 31], and since the manifestation of symptoms change along with development [32], one might exhibit ASD symptoms later in adulthood despite the absence of overt ASD symptoms in childhood. It also adds support to the shared biological underpinnings between the two disorders as reported by previous researchers [3, 6-8]. Additionally, another possibility is that ADHD symptoms and their related issues, such as being rejected by others [33–38] or being involved in bullying [39], can give rise to ASD-like symptoms later in adulthood as they had been deprived of the appropriate opportunity and experience of developing healthy social interaction skills.

The item-level analysis between the ADHD and control groups suggested that adults with ADHD have difficulties not only in quantitative but also qualitative aspects of social communication. Most RSI items and some LC items were significantly more frequently endorsed by the ADHD group than the control group. These included A2 (Speech Abnormalities Associated With Autism), A8 (Conversation), B1 (Unusual Eye Contact), B2 (Facial Expressions Directed to Examiner), B3 (Language Production and Linked Nonverbal Communication), B7 (Insight Into Typical Social Situations

Table 4 Non-zero scorers on each ADOS-2 item for the ADHD and control groups

	ADHD $(n=63)$	Control $(n=31)$	X^2	р
A Language and Communication	N (%)			
A1 Overall Level of Non-Echoed Spoken Language	0 (0.0)	0 (0.0)	_	_
A2 Speech Abnormalities Associated With Autism ^c	21 (33.3)	1 (3.2)	10.51	0.001
A3 Immediate Echolalia	2 (3.2)	0 (0.0)	1.01	0.316
A4 Stereotyped/Idiosyncratic Use of Words or Phrases ^b	7 (11.1)	1 (3.2)	1.66	0.198
A5 Offers Information	9 (14.3)	2 (6.5)	1.23	0.267
A6 Asks for Information	52 (82.5)	22 (71.0)	1.66	0.197
A7 Reporting of Events	16 (25.4)	8 (25.8)	0.00	0.966
A8 Conversation ^b	17 (27.0)	2 (6.5)	5.43	0.020
A9 Descriptive, Conventional, Instrumental, or Informational Gestures ^a	17 (27.0)	6 (19.4)	0.65	0.419
A10 Emphatic or Emotional Gestures ^b	32 (50.8)	13 (41.9)	0.65	0.419
B Reciprocal Social Interaction				
B1 Unusual Eye Contact ^b	9 (14.3)	0 (0.0)	4.90	0.027
B2 Facial Expressions Directed to Examiner ^b	29 (46.0)	2 (6.5)	14.73	< 0.001
B3 Language Production and Linked Nonverbal Communication	20 (31.7)	0 (0.0)	12.50	< 0.001
B4 Shared Enjoyment in Interaction	17 (27.0)	0 (0.0)	10.21	0.001
B5 Communication of Own Affect ^c	26 (41.3)	8 (25.8)	2.15	0.142
B6 Comments on Others' Emotions/Empathy ^a	23 (36.5)	8 (25.8)	1.08	0.299
B7 Insight Into Typical Social Situations and Relationships ^c	48 (76.2)	15 (48.4)	7.27	0.007
B8 Responsibility ^a	21 (33.3)	3 (9.7)	6.12	0.013
B9 Quality of Social Overtures ^b	18 (28.6)	0 (0.0)	10.96	0.001
B10 Amount of Social Overtures/Maintenance of Attention ^c	27 (42.9)	9 (29.0)	1.68	0.195
B11 Quality of Social Response ^b	17 (27.0)	0 (0.0)	10.21	0.001
B12 Amount of Reciprocal Social Communication ^b	23 (36.5)	1 (3.2)	12.10	0.001
B13 Overall Quality of Rapport	7 (11.1)	2 (6.5)	0.52	0.470
C Imagination				
C1 Imagination/Creativity	30 (47.6)	10 (32.3)	2.01	0.157
D Stereotyped Behaviors and Restricted Interests				
D1 Unusual Sensory Interest in Play Material/Person ^c	1 (1.6)	0 (0.0)	0.50	0.481
D2 Hand and Finger and Other Complex Mannerisms ^c	3 (4.8)	0 (0.0)	1.53	0.217
D3 Self-Injurious Behavior	0 (0.0)	0 (0.0)	-	-
D4 Excessive Interest in or References to Unusual or Highly Specific Top- ics or Objects or Repetitive Behaviors ^c	4 (6.3)	1 (3.2)	0.40	0.526
D5 Compulsions or Rituals	4 (6.3)	1 (3.2)	0.40	0.526
E Other Abnormal Behaviors				
E1 Overactivity/Agitation	9 (14.3)	1 (3.2)	2.67	0.102
E2 Tantrums, Aggression, Negative or Disruptive Behavior	1 (1.6)	0 (0.0)	0.50	0.481
E3 Anxiety	9 (14.3)	1 (3.2)	2.67	0.102

p < 0.05 by chi-square test, in bold: statistically significant

^aIn original algorithm only

^bIn both original and revised algorithms

^cIn revised algorithm only

and Relationships), B9 (Quality of Social Overtures), B11 (Quality of Social Response), and B12 (Amount of Reciprocal Social Communication). While the same reasons described in our previous study were behind the high scoring on A2, A8, and B12 (they tended to be loud and communicate in a one-sided manner such as expressing their own

interest without paying much attention to the examiner), the current study also revealed that in comparison to controls, they were also impaired in the appropriate use of eye contact and its coordination in social communication (B1, B2, B3, B9, and B11). Further, these impairments are also reflected in the high B7 scoring. The vicious circle of peer rejection,

	Typical ADHD $(n=48)$	Control $(n=31)$	X^2	р
A Language and Communication	N (%)			
A1 Overall Level of Non-Echoed Spoken Language	0 (0.0)	0 (0.0)	-	-
A2 Speech Abnormalities Associated With Autism ^c	9 (18.8)	1 (3.2)	4.11	0.043
A3 Immediate Echolalia	1 (2.1)	0 (0.0)	0.65	0.419
A4 Stereotyped/Idiosyncratic Use of Words or Phrases ^b	5 (10.4)	1 (3.2)	1.39	0.239
A5 Offers Information	2 (4.2)	2 (6.5)	0.21	0.651
A6 Asks for Information	37 (77.1)	22 (71.0)	0.37	0.542
A7 Reporting of Events	6 (12.5)	8 (25.8)	2.29	0.130
A8 Conversation ^b	5 (10.4)	2 (6.5)	0.37	0.545
A9 Descriptive, Conventional, Instrumental, or Informational Gestures ^a	9 (18.8)	6 (19.4)	0.00	0.947
A10 Emphatic or Emotional Gestures ^b	19 (39.6)	13 (41.9)	0.04	0.835
B Reciprocal Social Interaction				
B1 Unusual Eye Contact ^b	1 (2.1)	0 (0.0)	0.65	0.419
B2 Facial Expressions Directed to Examiner ^b	17 (35.4)	2 (6.5)	8.65	0.003
B3 Language Production and Linked Nonverbal Communication	7 (14.6)	0 (0.0)	4.96	0.026
B4 Shared Enjoyment in Interaction	7 (14.6)	0 (0.0)	4.96	0.026
B5 Communication of Own Affect ^c	14 (29.2)	8 (25.8)	0.11	0.745
B6 Comments on Others' Emotions/Empathy ^a	13 (27.1)	8 (25.8)	0.02	0.900
B7 Insight Into Typical Social Situations and Relationships ^c	33 (68.8)	15 (48.4)	3.28	0.070
B8 Responsibility ^a	14 (29.2)	3 (9.7)	4.24	0.040
B9 Quality of Social Overtures ^b	4 (8.3)	0 (0.0)	2.72	0.099
B10 Amount of Social Overtures/Maintenance of Attention ^c	14 (29.2)	9 (29.0)	0.00	0.990
B11 Quality of Social Response ^b	5 (10.4)	0 (0.0)	3.45	0.063
B12 Amount of Reciprocal Social Communication ^b	10 (20.8)	1 (3.2)	4.87	0.027
B13 Overall Quality of Rapport	1 (2.1)	2 (6.5)	0.98	0.321
C Imagination				
C1 Imagination/Creativity	19 (39.6)	10 (32.3)	0.44	0.510
D Stereotyped Behaviors and Restricted Interests				
D1 Unusual Sensory Interest in Play Material/Person ^c	1 (2.1)	0 (0.0)	0.65	0.419
D2 Hand and Finger and Other Complex Mannerisms ^c	1 (2.1)	0 (0.0)	0.65	0.419
D3 Self-Injurious Behavior	0 (0.0)	0 (0.0)	-	-
D4 Excessive Interest in or References to Unusual or Highly Specific Topics or Objects or Repetitive Behaviors ^c	2 (4.2)	1 (3.2)	0.05	0.831
D5 Compulsions or Rituals	1 (2.1)	1 (3.2)	0.10	0.752
E Other Abnormal Behaviors				
E1 Overactivity/Agitation	5 (10.4)	1 (3.2)	1.39	0.239
E2 Tantrums, Aggression, Negative or Disruptive Behavior	1 (2.1)	0 (0.0)	0.65	0.419
E3 Anxiety	5 (10.4)	1 (3.2)	1.39	0.239

p < 0.05 by chi-square test, in bold: statistically significant

^aIn original algorithm only

^bIn both original and revised algorithms

^cIn revised algorithm only

reduced interaction, and poor social skills [40–42] likely damages the quality and understanding of social relationships. Theory of mind deficits [43, 44] may also contribute to the difficulties in social communication and grasping the nature of typical relationships. Interestingly however, when we excluded those with overt ASD symptoms (i.e., those who met ASD cut-off on ADOS-2) and compared "typical ADHD" against controls, most of the qualitative items were no longer endorsed more frequently in the ADHD than control group. Such items include

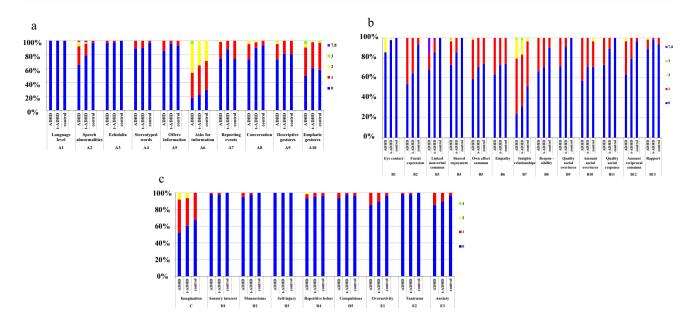


Fig. 1 Percentage of scores on each ADOS-2 item for the ADHD, typical ADHD, and control groups of **a** Language and Communication (A); **b** Reciprocal Social Interaction (B); and **c** Imagination (C),

B1 (Unusual Eye Contact), B9 (Quality of Social Overtures), and B11 (Quality of Social Response). The quantitative item; B12 (Amount of Reciprocal Social Communication), affect-related items; B2 (Facial Expressions Directed to Examiner) and B4 (Shared Enjoyment in Interaction), and the sense of responsibility (B8) remained significantly different than controls in endorsement frequency. This possibly suggests that ASD symptoms present in ADHD are not on a quantitative continuum to ASD but rather a mixture of distinct mechanisms. One is qualitatively the same as those with ASD, in which the integration of different forms of social communication is significantly impaired. The other is the resulting symptom resemblance of ADHD to ASD, in which inattentive or impulsive communication wrecks reciprocal interaction, rather more quantitatively (one-sided communication style). The correlation patterns support this point as only the "typical ADHD" group showed CAARS-ADOS-2 correlations, especially between impulsivity and reciprocal social interaction, whereas the overall ADHD group did not. Further, non-identical SA-RRB relationships in these two groups may describe the heterogeneity of ASD symptoms in adults with ADHD. It is possible that the entanglement of these mechanisms may become more prominent in adulthood than in childhood as they mutually potentiate along development and experiences. Our study highlights the importance of careful and detailed assessment of ASD symptoms and co-occurrence in ADHD as they may not be homogenous in nature. Recent neuroimaging studies have identified several overlapping and distinctive cortical and subcortical structural changes between ADHD and Stereotyped Behaviors and Restricted Interests (D), and Other Abnormal Behaviors (E) domains. *t-ADHD* typical ADHD, commun: communication; behav: behaviors. *See Tables 4, 5 for full item names

ASD [32, 45, 46]. In particular, O'Dwyer et al. [47] studied the neural structural correlates of ASD-like symptoms in ADHD, in which the volumes of caudate nucleus and globus pallidus were found to play a critical role. Although the results remain inconsistent and are subject to longitudinal evaluation, it is crucial to examine these biological markers in combination with clinical symptoms in future studies.

In contrast to our previous work, we did not find as much evidence as we did before, for the difficulties with affective interaction in adults with ADHD. The only emotion-related items that were scored significantly more likely by the ADHD than control groups were B2 (Facial Expressions Directed to Examiner) and B4 (Shared Enjoyment in Interaction). Other items such as A10 (Emphatic or Emotional Gestures), B5 (Communication of Own Affect), and B6 (Comments on Others'' Emotions/Empathy), which were frequently endorsed by adults with ADHD in the preliminary work, did not show any group differences in the scoring patterns. Given the frequent scoring by controls on B5, B6, and B10, cultural factors may have skewed these results. In Japan, it is often considered impolite to ask personal questions to examiners or others at the first meeting, and the Japanese have fewer and more covert expressions of emotions and body language than the Western culture [48, 49]. It is possible that these factors masked the group differences, including A6 (Asks for Information), and further cross-cultural studies are necessary to evaluate affectrelated difficulties in ADHD. Further, given the relatively lower percentage of adults with ADHD scoring positive on all D domain (Stereotyped Behaviors and Restricted

	AQ:Total CAARS:Inatt CAARS:Hype	CAARS:Hype	CAARS:Imp	CAARS:ADHD Index	WAIS-III: VIQ	WAIS-III: PIQ	WAIS-III: FIQ	ADOS- 2: Rev LC	ADOS- 2: Rev RSI	ADOS-2: Rev SA	ADOS-2: Rev RRB	ADOS-2: Rev Total
AQ:Total	0.22	0.30	0.52*	0.55*	0.12	0.08	0.16	0.10	0.23	0.21	0.10	0.21
CAARS:		0.39*	0.53*	0.64*	-0.01	- 0.11	- 0.06	0.03	0.10	0.09	0.14	0.11
Inatt												
CAARS: Hvne			0.66*	0.78*	- 0.09	- 0.26	- 0.17	0.13	0.14	0.14	0.00	0.13
CAARS:				0.86^{*}	- 0.16	0.01	- 0.04	0.10	0.15	0.14	0.01	0.13
Imp												
CAARS:					- 0.13	- 0.16	-0.13	0.13	0.25	0.23	0.09	0.22
ADHD												
Index												
-SIA-						0.37*	0.80^{*}	0.05	0.05	0.05	- 0.13	0.02
III:VIQ												
-SIA-							0.76^{*}	0.17	0.17	0.18	0.10	0.18
DI4:III												
WAIS-								0.19	0.18	0.19	0.01	0.17
III:FIQ												
ADOS-2:									0.77*	0.87*	0.36^{*}	0.85*
Rev LC												
ADOS-2:										0.99*	0.39*	0.96*
Rev RSI												
ADOS-2:											0.40*	0.98*
Rev SA												
ADOS-2:												0.59*
Rev RRB												
ADOS2:												
Rev Total												

activity/Restlessness. Imp Impulsivity/Emotional Lability, Inatt Inattention/Memory Problems, LC language and communication, PIQ Performance intelligence quotient, Rev revised algorithm, RRB restricted and repetitive behaviors, RSI reciprocal social interaction, SA social affect, VIQ verbal intelligence quotient, WAIS-III Weehsler adult intelligence scale, third edition *Correlation is significant at the 0.01 level (2-tailed)

Table 7 Correl	lations be	tween AQ, CA.	ARS, WAIS-III, ¿	and ADOS-2 in a	Correlations between AQ, CAARS, WAIS-III, and ADOS-2 in the control group $(n=31)$	n = 31)							
V	AQ:Total	CAARS:Inatt	CAARS:Inatt CAARS:Hype	CAARS:Imp	CAARS:ADHD Index	WAIS-III: VIQ	WAIS-III: PIQ	WAIS-III: FIQ	ADOS- 2:Rev LC	ADOS-2: Rev RSI	ADOS-2: Rev SA	ADOS- 2:Rev RRB	ADOS-2: Rev Total
AQ:Total		0.15	- 0.16	0.14	0.23	- 0.12	- 0.24	- 0.21	- 0.05	0.32	0.25	0.38	0.32
CAARS: Inatt			0.55*	0.63*	0.68*	0.28	0.13	0.26	0.12	0.35	0.36	0.20	0.39
CAARS:				0.51^{*}	0.67*	0.07	- 0.20	- 0.06	- 0.24	- 0.16	- 0.26	- 0.08	- 0.26
Hype													
CAARS:					0.76^{*}	0.49*	- 0.09	0.28	- 0.06	0.09	0.05	0.10	0.07
Imp													
CAARS: ADHD						0.25	- 0.24	0.04	- 0.03	0.08	0.05	0.01	0.05
Index													
WAIS-							0.36	0.86^{*}	0.23	0.16	0.25	0.10	0.26
III: VIQ													
-SIAN								0.79*	0.13	0.27	0.30	0.13	0.31
DI4:III													
WAIS-									0.22	0.26	0.34	0.14	0.35
III:FIQ													
ADOS-2: Rev LC										- 0.01	0.50*	- 0.28	0.41
ADOS-2:											0.86^{*}	0.35	0.89*
Rev RSI													
ADOS-2:												0.17	0.98*
Rev SA													
ADOS-2:													0.38
Rev RRB													
ADOS2:													
Rev Total													
ADOS-2 Autis Hvneractivity/I	tm Diagn Restlessne	ostic Observati ss. Inn Imnili	ion Schedule, Se sivitv/Emotional	cond Edition, A Lability. Inatt I	ADOS-2 Autism Diagnostic Observation Schedule, Second Edition, AQ Autism spectrum quotient, CAARS Conners' Adult ADHD rating scale, FIQ Full scale intelligence quotient, Hype Hyneractivity/Restlessness Impulsivity/Emotional Lability. Inattention/Memory Problems. I.C.I. anouage and Communication. PIO performance intelligence quotient. Rev Revised	m quotient, (v Prohlems, /	CAARS Conn	ers' Adult A.	DHD rating	scale, <i>FIQ</i> F 0 nerformanc	ull scale inte se intelligenc	elligence qu	otient, Hype Rev Revised

atyperactivity resuccessions, *imp* impute wrythingtonia realing, *indu* matering interaction, *SA* social affect, *VIQ* verbal intelligence quotient, *WAIS-III* Wechsler adult intelligence scale, third edition *Correlation is significant at the 0.01 level (2-tailed)

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				•									
	AQ:Tota]	l CAARS:Inatt	AQ:Total CAARS:Inatt CAARS:Hype	CAARS:Imp	CAARS:ADHD Index	WAIS-III: VIQ	WAIS-III: PIQ	WAIS-III: FIQ	ADOS- 2:Rev LC	ADOS- 2: Rev RSI	ADOS-2: Rev SA	ADOS- 2:Rev RRB	ADOS-2: Rev Total
AQ:Total		0.26	0.32	0.56*	0.54*	0.14	0.02	0.14	0.07	0.08	0.08	0.01	0.08
CAARS:Inatt			0.49*	0.61^{*}	0.70*	- 0.06	- 0.12	- 0.10	0.13	0.25	0.24	- 0.03	0.20
CAARS:Hype				0.64*	0.80^{*}	- 0.14	- 0.34	- 0.24	0.02	0.15	0.12	0.09	0.14
CAARS:Imp					0.88*	- 0.22	- 0.02	- 0.08	0.22	0.39^{*}	0.38*	0.15	0.38*
CAARS:ADHD Index						- 0.15	- 0.23	- 0.18	0.16	0.35	0.33	0.08	0.32
WAIS-III: VIQ							0.36	0.77*	0.09	0.04	0.06	- 0.17	- 0.01
WAIS-III: PIQ								0.78*	0.09	0.05	0.07	0.04	0.07
WAIS-III: FIQ									0.20	0.12	0.16	- 0.05	0.12
ADOS-2:Rev LC										0.57*	0.75*	0.22	0.74*
ADOS-2:Rev RSI											0.97*	0.19	0.92*
ADOS-2:Rev SA												0.22	0.95*
ADOS-2:Rev RRB													0.51^{*}
ADOS2:Rev Total													
ADOS-2 Autism Hyperactivity/Re	Diagnostic estlessness,	Cobservation Solution International Construction Solution	chedule, Second	Edition, AQ Au ity, Inatt Inatter	ADOS-2 Autism Diagnostic Observation Schedule, Second Edition, AQ Autism spectrum quotient, CAARS Conners' Adult ADHD rating scale, FIQ Full scale intelligence quotient, Hype Hyperactivity/Restlessness, Imp Impulsivity/Emotional Lability, Inant Inatention/Memory Problems, LC Language and Communication, PIQ performance intelligence quotient, Rev revised	otient, CAAR blems, LC L	S Conners' A anguage and	Adult ADHD Communica	rating sca tion, PIQ	ale, FIQ Fu performanc	ill scale inte te intelligene	lligence que te quotient,	otient, <i>Hype</i> Rev revised

algorithm, RRB restricted and repetitive behaviors, RSI reciprocal social interaction, SA social affect, VIQ verbal intelligence quotient, WAIS-III Wechsler adult intelligence scale, third edition ^{*}Correlation is significant at the 0.01 level (2-tailed)

Table 8 Correlations between AQ, CAARS, WAIS-III, and ADOS-2 in the typical ADHD group (n=48)

particularities in behavior and language may play a critical role in differentiating between ADHD and ASD. However, the absence of group differences on these items may be due to the small number of positive cases. Since RRB may not be fully observed during the limited time in the ADOS-2 [15], the combined use of other RRB evaluations is ideal in exploring these possible distinctive symptoms in ADHD and ASD.

The current study has some limitations. First, since the exclusion of ASD was based on the clinical diagnosis without the ADOS-2 and ADI-R, adults with ASD may have been included amiss. However, this effect should be small as we conducted a thorough examination of developmental history from multiple resources. Second, we did not use medication-naïve individuals, and psychotropic medications could have altered the ADOS-2 performance. A further study on a larger non-medicated ADHD sample is currently planned. Finally, there are reports on a rather limited specificity of the ADOS-2 Module 4 [50]. Although we believe that the ADOS-2 was appropriate in our study to identify the details of ASD symptoms, further evaluations of validity, especially on items that may have cultural effects, are warranted.

In conclusion, the present comparative study using the ADOS-2 demonstrated that adults with ADHD have significant ASD symptoms compared with neurotypical controls. While both qualitative and quantitative aspects of social communication were found impaired, similar to in those with ASD, the results highlighted the heterogeneous nature of ASD symptoms present in adult ADHD.

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Availability of data and material Research data will not be shared as the participants did not consent for open data sharing.

Code availability Not applicable.

Declarations

Conflict of interest The authors declare that they have no conflicts of interest.

Ethics approval This study was evaluated and approved by the Medical Ethics Committee of Showa University School of Medicine, and the protocols were carried out in accordance with the Declaration of Helsinki.

Consent to participate Written informed consent was obtained from all participants included in the study after information was provided about the study.

Consent for publication Not applicable.

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