

The broad autism phenotype in real-life: clinical and functional correlates of autism spectrum symptoms and rumination among parents of patients with autism spectrum disorder

Barbara Carpita ¹, Claudia Carmassi ¹, Sara Calderoni ², Dario Muti ¹, Alessia Muscarella ¹, Gabriele Massimetti ¹, Ivan M. Cremone ¹, Camilla Gesi ¹, Eugenia Conti ², Filippo Muratori ², Liliana Dell'Osso ¹

¹ Department of Clinical and Experimental Medicine, University of Pisa, Pisa, Italy

² Department of Developmental Neuroscience, IRCCS Fondazione Stella Maris, University of Pisa, Pisa, Italy

Abstract

Objective

Increasing literature reported higher rates of psychiatric disorders in parents of children with autism spectrum disorder (ASD), as well as of autistic-like features in social and cognitive functioning. However, little attention has been paid to the association between autistic traits (AT) and global functioning in this population. The aim of the present work was to investigate clinical and functional correlates of AT among parents of ASD children, with a specific focus on ruminative thinking.

Methods

One hundred and twenty parents of ASD children were assessed by the Structured Clinical Interview for Diagnostic and Statistical Manual of Mental Disorders (DSM-5), the Adult Autism Subthreshold Spectrum (AdAS Spectrum), the Ruminative Response Scale (RRS), the Social and Occupational Functioning Assessment Scale (SOFAS).

Results

Subjects with at least 1 psychiatric disorder (39.2%) showed significantly higher AdAS Spectrum and RRS scores. Subjects with a history of school difficulties and with language development alterations scored significantly higher on specific AdAS Spectrum domains. A significant negative correlation was found between SOFAS and AdAS Spectrum scores, as well as between SOFAS and RRS scores. AdAS Spectrum nonverbal communication domain score was identified as a statistically predictive variable for the presence of psychiatric disorders and lower SOFAS scores. Finally, we found a significant indirect effect of AdAS total score on SOFAS score, which was fully mediated by RRS total score.

Conclusions

AT in parents of ASD children seem to be associated with a higher vulnerability toward psychopathology and with a lower global functioning. Ruminative thinking may play a role in the relationship between AT and functional outcome.

Introduction

Autism spectrum disorder (ASD) is a condition characterized by impaired social communication and interactions, associated with a pattern of repetitive behaviors and narrow interests, with or without intellectual disability.¹ A wide number of studies focusing on autism etiopathogenesis highlighted the role of genetic heritability, which seems to interact with environmental, mostly intrauterine, factors.²⁻⁵ Although a family aggregation is reported in ASD since 1977,⁶ an increasing number of studies stressed the presence of a wide spectrum of milder symptoms, such as cognitive, social, and language impairments, among parents and siblings of ASD subjects. This evidence led to the concept of broad autism phenotype (BAP), a label employed to describe the wide spectrum of subthreshold autistic traits (AT) that can be frequently detected among first-degree relatives of ASD probands.⁶⁻¹⁵ Parents of ASD patients often show little interest in social interactions, and they usually have few, shallow friendships.^{14,16} In this framework, Sucksmith et al reported, among fathers of ASD probands, the presence of empathy deficits.¹⁷ Moreover, parents and siblings of ASD children frequently show difficulties related to verbal and nonverbal communications¹⁸⁻²² as well as impaired executive functions²³ and a ASD-like pattern of narrow interests.^{19-21, 24, 25} Taken together, all these elements—impaired social skills, language difficulties, restricted interests, and behaviors, similar, although less severe, to those of ASD patients—represent the typical features of BAP, which has been interpreted as a mild phenotypic variant of the disorder.^{14, 26-28} Some authors reported that even the Theory of Mind, the ability to assess information related to other people's mental states, seems to be compromised in parents of ASD children,²⁹⁻³² although this result has not been confirmed in all the studies.^{21, 33} In this framework, other studies pointed out that some specific personality traits can be frequently found among family members of ASD subjects, such as shyness, suspiciousness, hypersensitivity to criticism, aloofness, insensitivity, and rigidity.^{14, 34} A wide number of studies investigated neurobiological features of first-degree relatives of ASD children, further supporting the conceptualization of BAP.^{2, 4, 15} In particular, neuroimaging has been variously employed in this population, reporting neurostructural and neurofunctional alterations in parents of ASD probands, also in those not

behaviorally impaired (although more significant abnormalities were reported in subjects with behavioral correlates of BAP).¹⁵ Although genetic epidemiological studies stressed a genetic basis and heritability not only of ASD features, but also of BAP-like personality traits,¹⁵ abnormalities in different kinds of biochemical parameters have been found among parents of ASD children, similar to those reported in their probands.^{2,4} Higher blood levels of serotonin (5-HT) is one of the traits more consistently associated with autism spectrum,⁴ but most recent studies highlighted also altered levels of inflammatory markers, oxidative stress, and microbiota profiles in both ASD and BAP subjects.^{2,4} These findings are of particular interest because they may contribute to explaining the higher prevalence of autoimmune diseases and gastrointestinal symptoms among ASD children and their relatives, further characterizing BAP also from a neuroimmunological point of view.^{2,4}

More recently, increasing literature is stressing that subthreshold autism spectrum symptoms seem to be continuously distributed in general population, and in particular in some high-risk groups, such as university students, as well as among clinical samples of patients with other psychiatric disorders, where AT are generally associated with higher severity and with increased suicide ideation and behaviors.^{35–47} Although in literature the impact of AT in global functioning has been investigated in general population,^{48–51} studies among parents of ASD children generally focused mainly on work and social adjustment related to the burden to raise a child with ASD, rather than on the functional impairment related to their own AT.⁵² On the other hand, the prevalence of psychiatric conditions has been investigated also among relatives of ASD probands, highlighting an increased risk for mood disorders: one of the firsts studies in the field was led by DeLong and Dwyer,⁵³ who found a higher rate of mood disorders in relatives of ASD children with respect to the general population. Other studies^{54,55} reported an increased rate of anxiety and mood disorders among parents of ASD children when compared with parents of children with Down's syndrome. Although the burden of raising an autistic child may partially explain the higher rate of psychiatric conditions in this group, it has been observed that this issue cannot account for the overall increased rate of mood disorders.^{56,57} Moreover, according to another study, the majority of depressed mothers of ASD children reported their first depressive episode before the birth of the child.^{58,59} In this framework, Ingersoll and Hambrick, highlighting an increased rate of depression in mothers of ASD children, hypothesized a role of AT in the development of mood symptoms.⁵⁹ Several studies stressed also a high rate of social anxiety and obsessive–compulsive disorder among relatives of autistic probands,^{56,60–62} leading to hypothesize that social anxiety and obsessive–compulsive traits may be considered as core features of BAP.^{58,60} It is interesting to note that social anxiety is a disorder which implies an impairment in social functioning, with strong neurobiological underpinnings,^{63,64} and which is supposed to mask autistic-like symptoms especially among females, where ASD frequently remain undiagnosed.⁶⁵ In this framework, it is noteworthy that a large body of evidence is stressing that mental rumination, as a maladaptive pattern of repetitive thinking which often impairs problem solving and negative emotion processing, may play a key role in the onset of different kinds of psychiatric disorders,^{66–68} increasing also the risk of suicide ideation and behaviors.^{36,39,46,69,70} Ruminative thinking is a dimension frequently associated with ASD and AT,^{46,71,72} which has been reported to possibly mediate the relationship between autistic-like traits and psychiatric symptoms.⁴² However, this dimension has been poorly investigated among parents of children with ASD.

Although, as reported above, previous studies reported a higher frequency of different kinds of psychiatric disorders among parents of ASD probands, as well as autistic-like deficits in social communication and executive functioning, no study to the best of our knowledge has investigated the association between AT, ruminative thinking and global functioning (including clinical correlates) in parents of subjects with ASD.

In this framework, the aim of the present work was to investigate clinical and functional correlates of AT in a sample of parents of children with ASD, with a special focus on ruminative thinking. We hypothesized to find a high correlation between AT and rumination and an association of AT and ruminative thinking with a lower global functioning, as well as with a higher presence of psychiatric disorders. We also hypothesized that ruminative thinking may mediate the association between AT and the impairment in global functioning.

Methods

Although, as reported above, previous studies reported a higher frequency of different kinds of psychiatric disorders among parents of ASD probands, as well as autistic-like deficits in social communication and executive functioning, no study to the best of our knowledge has investigated the association between AT, ruminative thinking and global functioning (including clinical correlates) in parents of subjects with ASD.

In this framework, the aim of the present work was to investigate clinical and functional correlates of AT in a sample of parents of children with ASD, with a special focus on ruminative thinking. We hypothesized to find a high correlation between AT and rumination and an association of AT and ruminative thinking with a lower global functioning, as well as with a higher presence of psychiatric disorders. We also hypothesized that ruminative thinking may mediate the association between AT and the impairment in global functioning.

Instruments

AdAS Spectrum

The AdAS Spectrum is an instrument developed and validated by Dell'Osso et al,⁷² with the aim to assess autism spectrum symptoms in adults with average intelligence and without language impairment. It has been tailored to evaluate not only overthreshold manifestations, but also a wide range of subthreshold, isolated or "atypical" features (including female-specific manifestations) such as those included in the BAP. The questionnaire is composed by 7 domains, for a total of 160 dichotomous items. The AdAS Spectrum has been proved to be highly correlated with other instruments in the field, such as the Autism Spectrum Quotient (Pearson's correlation = 0.77). The instrument demonstrated also an excellent reliability (Kuder Richardson's coefficient = 0.964).

RRS

The RRS is an instrument composed by 22 items, widely utilized in literature to investigate the presence of ruminative thinking. The items are grouped in 3 dimensions: reflection, brooding, and

depression, and the answers are organized in a 4-point Likert scale. The RRS demonstrated an excellent internal consistency (Cronbach's alpha = 0.89).^{73, 74}

SOFAS

The SOFAS is an observational instrument widely employed in literature to assess the level of social and occupational functioning in a continuum, with scores ranging from 1 to 100 (higher scores are associated with better functioning). It is specified that the impairment must be directly related to the psychiatric (or somatic) symptoms in order to be rated.⁷⁵

Statistical analyses

We utilized Student's t tests to compare the mean scores on AdAS Spectrum and RRS between males and females, between subjects with/without a psychiatric disorder, as well as between subjects with/without a history of school difficulties or a history of language development alterations. We performed a Pearson's correlation coefficient in order to evaluate the association between AdAS Spectrum and RRS scores, between AdAS Spectrum and SOFAS scores, as well as between RRS and SOFAS scores. A multiple logistic regression analysis, with a single block entry (sex, AdAS Spectrum, and RRS domain scores were the independent variables), was employed to identify the best predictors of the presence of a psychiatric disorder. Moreover, multiple linear regression analysis was performed to identify the factors most predictive of SOFAS score, with sex, AdAS Spectrum, and RRS domain scores as independent variables (also in this case, all variables were entered as a single block). Finally, we performed a mediation analysis with AdAS Spectrum total score as predictor, SOFAS score as dependent variable, and RRS total score as mediator. The Hayes's PROCESS tool was utilized; bootstrap confidence intervals and Sobel test for indirect effect were computed. All analyses were performed using SPSS version 25 (IBM Corp., 2017).

Results

Our sample was composed by couples of parents of 60 ASD children, for a total of 120 subjects (60 males, 60 females) with a mean age of 39.82 ± 6.02 . A total of 39.2% of the subjects reported a full-blown psychiatric disorder, whereas 15.8% of the subjects reported several comorbid psychiatric conditions. In particular, 11.7% of the subjects showed a bipolar disorder (N = 14), 17.5% (N = 21) a panic disorder, 25% (N = 30) other anxiety disorders, whereas 5.8% (N = 7) reported feeding and eating disorders. According to our data, 22.5% (N = 27) of the sample was employing psychopharmacological therapies. In particular, 11.6% of the total sample (N = 14) was taking only 1 drug: among them, 64.29% (N = 9) employed benzodiazepines, 7.14% (N = 1) antiepileptics and 28.57% (N = 4) antidepressants. Moreover, 10 subjects (8.33% of the whole sample) were taking both antidepressants and benzodiazepines. Finally, 1 subject was taking antidepressant and antipsychotics, and 2 subjects were treated with antidepressant, antipsychotics, and benzodiazepines. We found an overall rate of 5.8% (N = 7) of previous school difficulties, while 8 subjects (6.7%) reported a history of alterations in language development. The mean score on AdAS Spectrum was 37.04 ± 20.35 , while the mean score on RRS was 36.76 ± 9.65 . No gender differences were found on AdAS Spectrum total and domain scores, while females

scored significantly higher than males on RRS reflection subscale (8.10 ± 2.36 vs 7.18 ± 2.50 ; $t = -1.998$; $p = .048$). Subjects with a psychiatric disorder showed significantly higher AdAS Spectrum total and domain scores as well as significantly higher RRS total and domain scores (Table 1). Subjects with a history of school difficulties when compared to those without, reported significantly higher AdAS Spectrum total scores (61.14 ± 31.05 vs 35.45 ± 18.54 , $t = -3.38$, $p = .001$), as well as significantly higher AdAS Spectrum childhood/adolescence (10 ± 4.69 vs 5.11 ± 3.49 , $t = -3.51$, $p = .001$), nonverbal communication (11.86 ± 6.09 vs $6.601 \pm 4.10.49$, $t = -3.18$, $p = .002$), and inflexibility/adherence to routine (15.00 ± 9.02 vs 9.60 ± 5.95 , $t = -2.24$, $p = .027$) domain scores, while they did not report higher RRS scores. Subjects with a language development alteration scored significantly higher only on AdAS Spectrum childhood/adolescence domain score (5.18 ± 3.50 vs 8.50 ± 5.45 ; $t = -2.474$; $p = .015$). We found significant correlations between all AdAS Spectrum and RRS total and domain scores (Table 2). Moreover, we found a significant negative correlation between SOFAS score and AdAS Spectrum total and domain scores, with the exception of AdAS Spectrum empathy domain score. The highest correlation was found between SOFAS and AdAS Spectrum nonverbal communication domain score (Table 2). Significant negative correlations were found also between SOFAS score and RRS total and domain scores (Table 2). We performed a multiple logistic regression analysis in order to identify the best predictors for the presence of psychiatric disorders, which is a categorical variable. Although multiple independent variables (in particular: sex, AdAS Spectrum, and RRS domain scores) were entered as possible predictors, our results identified only the female gender and AdAS Spectrum nonverbal communication domain score as predictive variables for the presence of a psychiatric disorder (Table 3). Moreover, we performed a multiple linear regression analysis with SOFAS score as the dependent continuous variable in order to evaluate possible statistically significant predictors of social and occupational functioning, including sex, AdAS Spectrum, and RRS domain scores as independent variables. The regression equation was significant ($F[11, 93] = 3.676$, $p < .001$), with $R^2 = 0.303$, and the model identified the AdAS Spectrum nonverbal communication domain score as the only significant predictive factor for a lower SOFAS score (Table 4). Finally, the significant negative correlations of both AdAS Spectrum and RRS total scores with SOFAS score ($r = -0.38$, $p < .001$, and $r = -0.40$, $p < .001$, respectively) as well as the significant positive correlation between AdAS Spectrum and RRS, suggested us to verify the eventual presence of a mediating effect of RRS total score on the relationship between AdAS Spectrum and SOFAS score. The mediation analysis showed a significant indirect effect of AdAS Spectrum total score on SOFAS score through RRS total score, $b = -0.12$, 95% bootstrapped 95% CI $[-0.23, -0.04]$; Sobel test showed $b = -0.12$, $Z = -2.64$, $p = .008$ for RRS total score. It is noteworthy that the mediation appears to be full because there was a significant total effect ($b = -0.220$, $p = .001$), but not a significant direct effect ($b = -0.10$, $p = .152$) of AdAS Spectrum total score on SOFAS score (Figure 1).

Discussion

The aim of the present work was to investigate clinical and functional correlates of AT and ruminative thinking in a sample of parents of children with ASD. The rates of psychiatric disorders in our sample are in line with those highlighted in previous literature, which stressed a higher presence of mood and anxiety disorders in parents of ASD children than in general population.⁵⁷ In particular, our sample showed a 11.7% rate of bipolar disorders, while the reported prevalence in

the general Italian population is about 1%.^{76,77} However, we did not find any case of major depressive disorder (MDD), which is the most common mood disorder in the general Italian population, with a prevalence about 10%.⁷⁷ Although previous studies reported higher rates of different kinds of mood disorders, including MDD, in parents of ASD children,^{57,59} our results are somewhat in line with recent findings that suggested a specific association of autism spectrum with bipolar disorders, also from a genetic point of view.⁴⁶ Moreover, while in the general Italian population the reported prevalence of anxiety disorders is about 11%, mainly represented by specific phobia (5.7%),⁷⁷ we found in our sample a rate of 17.5% only for panic disorder (vs a prevalence of 2% in the general Italian population),⁷⁷ and a 25% rate of other anxiety disorders. Finally, according to other studies that stressed higher rates of feeding and eating disorders among BAP subjects,³⁹ in our sample parents of ASD children showed a 5.8% rate of feeding and eating disorders, while in the general Italian population, the reported prevalence is about 0.6%.⁷⁸ Globally our results, in line with other studies,^{39,40,42,46} seem to suggest an association between BAP and specific kinds of psychiatric disorders, although further researches in wider samples are needed to clarify this point.

The relatively high rates of school difficulties and language development alterations that we found in our sample, associated with significantly higher AT during childhood and adolescence, are in line with other studies that reported increased rates of delays in language development and pragmatic language alterations among relatives of ASD children.^{13,24,79} Moreover, we found that higher AT were associated with the presence of psychiatric disorders, as well as to a lower global functioning. This result adds to previous data, which reported high levels of AT in different kinds of psychiatric disorders,⁴⁰⁻⁴⁷ as well as a higher frequency of psychiatric conditions in parents of subjects with ASD.^{56,57} Some studies in this field ascribed the higher rates of psychopathology in parents of ASD children to the extremely stressful experience of raising a child with ASD, which may be more severe than the stress of raising a child with different kinds of disorders, such as mental retardation.^{57,80-82} In this framework, our data are in line to those of Ingersoll and Hambrick⁵⁹ who reported that the presence of depressive mood in mothers of ASD children was directly related to their own BAP score (as measured by the Autism Spectrum Quotient), rather than to parental distressing. Previous studies variously highlighted also an impairment in social and occupational functioning among parents of ASD children, focusing in particular on the stress related to the diagnosis of their child.⁵² When the child receives a diagnosis of ASD, the family unit must adjust to their new circumstances in a way that changes their life, including marital relationships, work arrangement, coping styles, and future perspectives.⁵² However, according to our results, it seems that lower SOFAS scores were associated with higher AT: in this framework, we could also hypothesize a role of the BAP in the impairment of global functioning observed in this group, as reported in different populations.⁴⁸⁻⁵¹ Both poor functional outcome and the presence of psychiatric conditions among parents of ASD children might be, at least partially, considered as a consequence of their own BAP traits.^{59,83-85} BAP may act as a risk factor for the development of psychiatric symptoms, either directly, because mood and anxiety disorders may be considered as an associated feature of BAP, with which they share also a genetic liability, or indirectly: the presence of AT may be considered as a vulnerability factor for developing post-traumatic spectrum symptoms after stressful events, including also those related to the role of caregivers.^{39,40} Moreover, BAP features may imply more difficulties in reaching social support due to the impairment in social skills.^{39,42,85} Other features associated with BAP, in particular

intolerance of uncertainty and sensory sensitivity, have been considered as a potential risk factor for the development and maintenance of affective problems in the general ⁸⁶⁻⁸⁸ as well as in ASD.^{89,90} Our results identify in the nonverbal communication problems (as measured by AdAS Spectrum), the statistically predictive dimension for the presence of psychiatric conditions, and poorer social and occupational functioning. These data are in line with a previous study, which stressed the link between autistic-like nonverbal communication impairment and suicide risk among university students.⁴⁷ It is noteworthy that social support seems to be a strong predictor of adjustment for parents of children with ASD,^{91,92} while communication impairment may result in higher social isolation, with more difficulties in externalizing and processing emotions, leading to reach lower social support, thus increasing the vulnerability to psychopathology.^{42, 47, 93, 94} In our sample, we did not find gender differences with respect to AT. Several investigations reported an increased expression of some (but not all) BAP features among male relatives: in particular higher aloofness, rigidity, irritability, and sensitive traits¹¹ as well as pragmatic language alterations.⁹⁵ Despite that, not all the studies have reported such differences,^{10, 14, 96} and our results seem to be in line with these latter. However, according to our data, female gender was a statistically predictive factor for the presence of psychiatric disorders, together with autistic-like nonverbal communication features. These data reflect results from other studies, which highlighted higher rates of psychopathology among females, including in samples composed by parents of ASD children.⁵⁷ In our sample, gender was not a predictive factor for the presence of a lower social and occupational functioning. Previous studies reported, among parents of ASD children, a lower functioning in mothers than in fathers, which was generally associated with higher levels of stress.^{97, 98} Although the role of gender differences on the impact of parenting stress is still controversial,^{83, 99} females often show, in different kinds of samples, a higher vulnerability to trauma- and stress-related symptoms.^{100, 101}

Our data reported also a significant association between ruminative thinking and the presence of psychiatric disorders, as well as between ruminative thinking and lower global functioning. Mental rumination involves perseverative thoughts that revolve around a negative emotion or situation. As an unintended and involuntary process, rumination is often long lasting and may consume cognitive resources, leading to poorer outcome.^{38, 66} These results confirm the wide amount of studies which stressed the role of rumination in promoting different kinds of psychopathology.⁶⁶⁻⁶⁸ On the other hand, ruminative thinking is a feature frequently associated with AT.^{42, 71, 72} ASD patients frequently show also features like perseverance and repetitiveness, including pervasive rumination on specific subjects.³⁸ In a recent meta-analysis, it has been stressed that ASD patients often show difficulties in inhibitory control,¹⁰² a feature that seems to contribute, in this population, to increasing the tendency to rumination, in particular about negative emotional experiences, eventually implying a higher vulnerability toward depressive symptomatology.¹⁰³ In this framework, our results, which stressed a significant correlation between AT and rumination as respectively measured by AdAS Spectrum and RRS, add to previous literature,⁴ confirming the strong link between those 2 dimensions. Moreover, it is noteworthy that, according to the mediation analysis, we found a significant effect of AT on the social and occupational functioning score; however, this effect was totally mediated by the presence of ruminative thinking. This result is in line with another study, which reported a mediating role of rumination in the relationship between AT and mood symptoms.⁴² Globally, these data seem to further suggest a

transnosographic role of rumination in the development of psychopathology and functional impairment in subjects with higher vulnerability.

This study suffers from several limitations. First, the cross-sectional design prevented us from clarifying the temporal, and eventually causal, relationship among AT, rumination, and global functioning or psychiatric symptoms development. Moreover, the AdAS Spectrum and the RRS are self-reported instruments, and patients may have over- or underestimated their own symptoms, eventually leading to the presence of biases in our results. This is an exploratory study conducted in a small sample and without a control group, mainly focused on psychopathological features chosen due to their known relationship with BAP. As a consequence, many demographic variables, other than sex, potentially involved in the relationship between AT and global functioning were not included in the study. In particular, according to previous literature, features such as marriage quality, access to social support, as well as the child symptom severity, may play a significant role in the level of adjustment among parents of ASD children.^{81, 91, 92} We did not collect any kind of neuroimaging, genetic, or biochemical data, although literature stressed the role of different kinds of biological variables in determining BAP features. In particular, according to the most recent studies, specific attention should be provided in clarifying the role of neurostructural and neurofunctional characteristics, as well as the role of oxidative stress, immune system activation and gut microbiota profiles, in shaping specific BAP psychopathological features and trajectories.^{2, 4, 15, 104, 105} Finally, pharmacological factors were not included in the analysis due to the small sample size and the high heterogeneity of treatments reported among the subjects. Further studies in wider samples and with a longitudinal design are warranted to clarify the relationship between AT, rumination, and both clinical and functional outcomes among parents of ASD patients.

References

1. American Psychiatric Association (APA). Diagnostic and Statistical Manual of Mental Disorders. 5th ed. Arlington, VA: American Psychiatric Association; 2013.
2. Carpita, B, Muti, D, Dell’Osso, L. Oxidative stress, maternal diabetes, and autism spectrum disorders. *Oxid Med Cell Longev*. 2018;2018:3717215.
3. Bai, D, Yip, BHK, Windham, GC, et al. Association of genetic and environmental factors with autism in A5-country cohort. *JAMA Psychiatry*. 2019 (Epub ahead of print). doi:10.1001/jamapsychiatry.2019.1411.
4. Carpita, B, Marazziti, D, Palego, L, et al. Microbiota, immune system and autism spectrum disorders. an integrative model towards novel treatment. *Curr Med Chem*. 2019 (Epub ahead of print). doi:10.2174/0929867326666190328151539
5. Kim, JY, Son, MJ, Son, CY, et al. Environmental risk factors and biomarkers for autism spectrum disorder: an umbrella review of the evidence. *Lancet Psychiatry*. 2019;6(7):590–600.
6. Folstein, S, Rutter, M. Infantile autism: a genetic study of 21 twin pairs. *J Child Psychol Psychiatry*. 1977;18(4):297–321.
7. Piven, J, Gayle, J, Chase, GA, et al. A family history study of neuropsychiatric disorders in the adult siblings of autistic. *J Am Acad Child Adolesc Psychiatry*. 1990;29(2):177–183.

8. Bolton, PF, Macdonald, H, Pickles, A, et al. A case-control family history study of autism. *J Child Psychol Psychiatry*. 1994;35(5):877-900.
9. Le Couteur, A, Bailey, A, Goode, S, et al. A broader phenotype of autism: the clinical spectrum in twins. *J Child Psychol Psychiatry*. 1996;37(7):785-801.
10. Piven, J, Palmer, P, Landa, R, et al. Personality and language characteristics in parents from multiple incidence autism families. *Am J Med Genet*. 1997;74(4):398-411.
11. Murphy, M, Bolton, P, Pickles, A, et al. Personality traits of the relatives of autistic probands. *Psychol Med*. 2000;30(6):1411-1424.
12. Pickles, A, Starr, E, Kazak, S, et al. Variable expression of the autism broader phenotype: findings from extended pedigrees. *J Child Psychol Psychiatry*. 2000;41(4):491-502.
13. Szatmari, P, MacLean, JE, Jones, MB, et al. The familial aggregation of the lesser variant in biological and nonbiological relatives of PDD probands: a family history study. *J Child Psychol Psychiatry*. 2000;41(5):579-586.
14. Losh, M, Childress, D. Defining key features of the broad autism phenotype: a comparison across parents of multiple- and single-incidence autism families. *Am J Medical Genet B Neuropsychiatr Genet*. 2008;147B(4):424-433.
15. Billeci, L, Calderoni, S, Conti, E, et al. The broad autism (endo)phenotype: neurostructural and neurofunctional correlates in parents of individuals with autism spectrum disorders. *Front Neurosci*. 2016;10:346.
16. Piven, J, Palmer, P, Jacobi, D, et al. Broader autism phenotype: evidence from a family history study of multiple-incidence autism families. *Am J Psychiatry*. 2008;154(2):185-190.
17. Sucksmith, E, Allison, C, Baron-Cohen, S, et al. Empathy and emotion recognition in people with autism, first-degree relatives, and controls. *Neuropsychologia*. 2013;51(1):98-105.
18. Bishop, DV, Maybery, M, Maley, A, et al. Using self-report to identify the broad phenotype in parents of children with autistic spectrum disorders: a study using the autism-spectrum quotient. *J Child Psychol Psychiatry*. 2004;45(8):1431-1436.
19. Whitehouse, AJ, Barry, JG, Bishop, DV. The broader language phenotype of autism: a comparison with specific language impairment. *J Child Psychol Psychiatry*. 2007;48(8):822-830.
20. Wheelwright, S, Auyeung, B, Allison, C, et al. Defining the broader, medium and narrow autism phenotype among parents using the Autism Spectrum Quotient (AQ). *Mol Autism*. 2010;1(1):10.
21. Sucksmith, E, Roth, I, Hoekstra, RA. Autistic traits below the clinical threshold: re-examining the broader autism phenotype in the 21st century. *Neuropsychol Rev*. 2011;21(4):360-389.
22. Ruta, L, Mazzone, D, Mazzone, L, et al. The autism-spectrum quotient—Italian version: a cross-cultural confirmation of the broader autism phenotype. *J Autism Dev Disord*. 2012;42(4):625-633.
23. Delorme, R, Gousse, V, Roy, I, et al. Shared executive dysfunctions in unaffected relatives of patients with autism and obsessive compulsive disorder. *Eur Psychiatry*. 2007;22(1):32-38.
24. Briskman, J, Happe, F, Frith, U. Exploring the cognitive phenotype of autism: weak "central coherence" in parents and siblings of children with autism: II. Real-life skills and preferences. *J Child Psychol Psychiatry*. 2001;42(3):309-316.

25. Robel, L, Rousselot-Pailley, B, Fortin, C, et al. Subthreshold traits of the broad autistic spectrum are distributed across different subgroups in parents, but not siblings, of probands with autism. *Eur Child Adolesc Psychiatry*. 2014;23(4):225–233.
26. Bailey, A, Palferman, S, Heavey, L, et al. Autism: the phenotype in relatives. *J Autism Dev Disord*. 1998;28(5):369–392.
27. Yirmiya, N, Shaked, M. Psychiatric disorders in parents of children with autism: a meta-analysis. *J Child Psychol Psychiatry*. 2005;46(1):69–83.
28. Ruzich, E, Allison, C, Smith, P, et al. Subgrouping siblings of people with autism: identifying the broader autism phenotype. *Autism Res*. 2016;9(6):658–665.
29. Happé, F. An advanced test of Theory of Mind: understanding of story characters' thoughts and feelings by able autistic, mentally handicapped, and normal children and adults. *J Autism Dev Disord*. 1994;24(2):129–154.
30. Baron-Cohen, S, Jolliffe, T, Mortimore, C, et al. Another advanced test of Theory of Mind: evidence from very high functioning adults with autism or Asperger syndrome. *J Child Psychol Psychiatry*. 1997;38(7):813–822.
31. Baron-Cohen, S, Wheelwright, S. The “reading the mind in the eyes” test revised version: a study with normal adults, and adults with Asperger Syndrome or high-functioning autism. *J Child Psychol Psychiatry*. 2001;42(2):241–251.
32. White, S, Hill, E, Happé, F, et al. Revisiting the strange stories: revealing mentalizing impairments in autism. *Child Dev*. 2009;80(4):1097–1117.
33. Ozonoff, S, Rogers, SJ, Farnham, JM, et al. Can standard measures identify subclinical markers of autism? *J Autism Develop Disord*. 1993;23(3):429–441.
34. Piven, J. The biological basis of autism. *Curr Opin Neurobiol*. 2008;7(5):708–712.
35. Dell’Osso, L, Dalle Luche, R, Cerliani, C, et al. Unexpected subthreshold autism spectrum in a 25-year-old male stalker hospitalized for delusional disorder: a case report. *Compr Psychiatry*. 2015;61:10–14.
36. Dell’Osso, L, Gesi, C, Carmassi, C. Suicide and autism spectrum disorder: the role of trauma. *J Psychopathol*. 2016;22(2):107–109.
37. Dell’Osso, L, Luche, RD, Gesi, C, et al. From Asperger’s Autistischen psychopathen to DSM-5 autism spectrum disorder and beyond: a subthreshold autism spectrum model. *Clin Pract Epidemiol Ment Health*. 2016;12:120–131.
38. Dell’Osso, L, Abelli, M, Carpita, B, et al. Historical evolution of the concept of anorexia nervosa and relationships with orthorexia nervosa, autism, and obsessive–compulsive spectrum. *Neuropsych Dis Treat*. 2016;12:1651–1660.
39. Carbone, M, Miniati, M, Simoncini, M, et al. Undetected autism subthreshold spectrum as risk factor for suicidal gestures in adulthood: a case report. *J Psychopathol*. 2018;24:37–40.
40. Dell’Osso, L, Muti, D, Carpita, B, et al. The Adult Autism Subthreshold Spectrum (AdAS) model: a neurodevelopmental approach to mental disorders. *J Psychopathol*. 2018;24:118–124.
41. Dell’Osso, L, Cremone, IM, Carpita, B, et al. Correlates of autistic traits among patients with borderline personality disorder. *Compr Psychiatry*. 2018;83:7–11.
42. Dell’Osso, L, Carpita, B, Cremone, IM, et al. The mediating effect of trauma and stressor related symptoms and ruminations on the relationship between autistic traits and mood spectrum. *Psychiatry Res*. 2018;279:123–129.

43. Dell'Osso, L, Conversano, C, Corsi, M, et al. Polysubstance and behavioral addictions in a patient with bipolar disorder: role of lifetime subthreshold autism spectrum. *Case Rep Psychiatry*. 2018;2018:1547975.
44. Dell'Osso, L, Carpita, B, Muti, D, et al. Prevalence and characteristics of orthorexia nervosa in a sample of university students in Italy. *Eat Weight Disord*. 2018;23(1):55–65.
45. Dell'Osso, L, Corsi, M, Gesi, C, et al. Adult Autism Subthreshold Spectrum (AdAS Spectrum) in parents of pediatric patients with epilepsy: correlations with post-traumatic stress symptoms. *Compr Psychiatry*. 2018;83:25–30.
46. Dell'Osso, L, Carpita, B, Muti, D, et al. Mood symptoms and suicidality across the autism spectrum. *Compr Psychiatry*. 2019;91:34–38.
47. Dell'Osso, L, Bertelloni, CA, Di Paolo, M, et al. Problematic Internet use in university students attending three superior graduate schools in Italy: is autism spectrum related to suicide risk? *Int J Environ Res Public Health*. 2019;16(7):1098.
48. Jobe, LE, White, SW. Loneliness, social relationships, and a broader autism phenotype in college students. *Pers Individ Diff*. 2007;42(8):1479–1489.
49. Trevisan, D, Birmingham, E. Examining the relationship between autistic traits and college adjustment. *Autism*. 2016;20(6):719–729.
50. Wallace, GL, Budgett, J, Charlton, RA. Aging and autism spectrum disorder: evidence from the broad autism phenotype. *Autism Res*. 2016;9(12):1294–1303.
51. Suzuki, T, Miyaki, K, Eguchi, E, et al. Distribution of autistic traits and their association with sociodemographic characteristics in Japanese workers. *Autism*. 2018;22(8):907–914.
52. Ooi, KL, Ong, YS, Jacob, SA, et al. A metasynthesis on parenting a child with autism. *Neuropsychiatr Dis Treat*. 2016;12:745–762.
53. DeLong, RT, Dwyer, JT. Correlation of family history with specific autistic subgroups: Asperger's syndrome and bipolar affective disease. *J Autism Develop Disord*. 1988;12(4):593–60.
54. Piven, J, Chase, GA, Landa, R, et al. Psychiatric disorders in the parents of autistic individuals. *J Am Acad Child Adolesc Psychiatry*. 1991;30(3):471–478.
55. Dumas, JE, Wolf, LC, Fisman, SN, et al. Parenting stress, child behaviour problems, and dysphoria in parents of children with autism, Down syndrome, behaviour disorders, and normal development. *Exceptionality*. 1991;2(2):97–110.
56. Bolton, PF, Pickles, A, Murphy, M, et al. Autism, affective and other psychiatric disorders: patterns of familial aggregation. *Psychol Med*. 1998;28(2):385–395.
57. Goussé, V, Galéra, C, Bouvard, M, et al. Aggregation of social deficits and psychiatric disorders in parents of children with autism: toward a temperamental link? *Encephale*. 2011;37(2):119–126.
58. Micali, N, Chakrabarti, S, Fombonne, E. The broad autism phenotype. *Autism*. 2004;8(1):21–37.
59. Ingersoll, B, Hambrick, DZ. The relationship between the broader autism phenotype, child severity, and stress and depression in parents of children with autism spectrum disorders. *Res Autism Spectr Disord*. 2011;5(1):337–344.
60. Piven, J, Palmer, P. Psychiatric disorder and the broad autism phenotype: evidence from a family study of multiple-incidence autism families. *Am J Psychiatry*. 1999;156(4):557–563.
61. Wilcox, JA, Tsuang, MT, Schnurr, T, et al. Case-control family study of lesser variant traits in autism. *Neuropsychobiology*. 2003;47(4):171–177.

62. O'Neill, LP, Murray, LE. Anxiety and depression symptomatology in adult siblings of individuals with different developmental disability diagnoses. *Res Dev Disabil.* 2016;51–52:116–125.
63. Marazziti, D, Baroni, S, Giannaccini, G, et al. A link between oxytocin and serotonin in humans: supporting evidence from peripheral markers. *Eur Neuropsychopharmacol.* 2012;22(8):578–583.
64. Marazziti, D, Abelli, M, Baroni, S, et al. Recent findings on the pathophysiology of social anxiety disorder. *Clin Neuropsychiatry.* 2014;11(2):91–100.
65. Attwood, T. *Asperger's and Girls.* Arlington, TX: Future Horizons; 2006.
66. Nolen-Hoeksema, S, Wisco, BE, Lyubomirsky, S. Rethinking rumination. *Perspect Psychol Sci.* 2008;3(5):400–424.
67. Watkins, ER. Depressive rumination and co-morbidity: evidence for brooding as a transdiagnostic process. *J Ration Emot Cogn Behav Ther.* 2009;27(3):160–175.
68. Dell'Osso, L, Cremone, IM, Carpita, B, et al. Rumination, posttraumatic stress disorder, and mood symptoms in borderline personality disorder. *Neuropsychiatr Dis Treat.* 2019;15:1231–1238.
69. Teismann, T, Forkmann, T. Positive mental health moderates the association between depression and suicide ideation: a longitudinal study. *Int J Clin Health Psychol.* 2018;18(1):1–7.
70. Law, KC, Tucker, RP. Repetitive negative thinking and suicide: a burgeoning literature with need for further exploration. *Curr Opin Psychol.* 2018;22:68–72.
71. Mayes, SD, Gorman, AA, Hillwig-Garcia, J, et al. Suicide ideation and attempts in children with autism. *Res Autism Spectr Disord.* 2013;7(1):109–119.
72. Dell'Osso, L, Gesi, C, Massimetti, E, et al. Adult Autism Subthreshold Spectrum (AdAS Spectrum): validation of a questionnaire investigating subthreshold autism spectrum. *Compr Psychiatry.* 2017;73:61–83.
73. Nolen-Hoeksema, S, Morrow, J. A prospective study of depression and posttraumatic stress symptoms after a natural disaster: the 1989 Loma Prieta Earthquake. *J Pers Soc Psychol.* 1991;61(1):115–121.
74. Parola, N, Zendjidjian, XY, Alessandrini, M, et al. Psychometric properties of the Ruminative Response Scale-short form in a clinical sample of patients with major depressive disorder. *Patient Prefer Adherence.* 2017;11:929–937.
75. Goldman, HH, Skodol, AE, Lave, TR. Revising axis V for DSM-IV: a review of measures of social functioning. *Am J Psychiatry.* 1992;149(9):1148–1156.
76. Faravelli, C, Guerrini Degl'Innocenti, B, Aiazzi, L, et al. Epidemiology of mood disorders: a community survey in Florence. *J Affect Disord.* 1990;20(2):135–141.
77. De Girolamo, G, Polidori, G, Morosini, P, et al. Prevalence of common mental disorders in Italy: results from the European Study of the Epidemiology of Mental Disorders (ESEMeD). *Soc Psychiatry Psychiatr Epidemiol.* 2006;41(11):853–861.
78. Faravelli, C, Abrardi, L, Bartolozzi, D, et al. The Sesto Fiorentino Study: background, methods and preliminary results. *Psychother Psychosom.* 2004;73(4):216–225.
79. August, G, Stewart, M, Tsai, L. The incidence of cognitive disabilities in the siblings of autistic children. *Br J Psychiatry.* 1981;138(5):416–422.
80. Hastings, RP. Parental stress and behaviour problems of children with developmental disability. *J Intellect Dev Disabil.* 2002;27(3):149–160.

81. Benson, PR. The impact of child symptom severity on depressed mood among parents of children with ASD: the mediating role of stress proliferation. *J Autism Dev Disord*. 2006;36(5):685–695.
82. Hayes, SA, Watson, SL. The impact of parenting stress: a meta-analysis of studies comparing the experience of parenting stress in parents of children with and without autism spectrum disorder. *J Autism Dev Disord*. 2013;43(3):629–642.
83. Ingersoll, B, Meyer, K, Becker, MW. Increased rates of depressive mood in mothers of children with ASD associated with the presence of the broader autism phenotype. *Autism Res*. 2010;4(2):143–148.
84. Lau, WY, Gau, SS, Chiu, YN, et al. Autistic traits in couple dyads as a predictor of anxiety spectrum symptoms. *J Autism Dev Disord*. 2014;44(11):2949–2963.
85. Pruitt, MM, Rhoden, M, Ekas, NV. Relationship between the broad autism phenotype, social relationships and mental health for mothers of children with autism spectrum disorder. *Autism*. 2018;22(2):171–180.
86. Aron, E, Aron, A, Jagiellowicz, J. Sensory processing sensitivity: a review in the light of the evolution of biological responsiveness. *Pers Soc Psychol Rev*. 2012;16(3):262–282
87. Carleton, RN. Into the unknown: a review and synthesis of contemporary models involving uncertainty. *J Anxiety Disord*. 2016;39:30–43.
88. McEvoy, P, Mahoney, AEJ. Achieving certainty about the structure of intolerance of uncertainty in a treatment-seeking sample with anxiety and depression. *J Anxiety Disord*. 2011;25(1):112–122.
89. Wigham, S, Rodgers, J, South, M, et al. The interplay between sensory processing abnormalities, intolerance of uncertainty, anxiety and restricted and repetitive behaviours in autism spectrum disorder. *J Autism Develop Disord*. 2015;45(4):943–952.
90. Uljarević, M, Carrington, S, Leekam, S. Brief report effects of sensory sensitivity and intolerance of uncertainty on anxiety in mothers of children with autism spectrum disorder. *J Autism Develop Disord*. 2016;46(1):315–319.
91. Boyd, B. Examining the relationship between stress and lack of social support in mothers of children with autism. *Focus Autism Other Dev Disabil*. 2002;17(4):208–215.
92. Benson, PR, Kersh, J. Marital quality and psychological adjustment among mothers of children with ASD: cross-sectional and longitudinal relationships. *J Autism Dev Disord*. 2011;41(12):1675–1685.
93. Pelton, MK, Cassidy, SA. Are autistic traits associated with suicidality? A test of the interpersonal-psychological theory of suicide in a non-clinical young adult sample. *Autism Res*. 2018;10(11):1891–1904.
94. Carmassi, C, Bertelloni, CA, Salarpi, G, et al. Is there a major role for undetected autism spectrum disorder with childhood trauma in a patient with a diagnosis of bipolar disorder, self-injuring, and multiple comorbidities? *Case Rep Psychiatry*. 2019;2019:4703795.
95. Ruser, TF, Arin, D, Dowd, M, et al. Communicative competence in parents of children with autism and parents of children with specific language impairment. *J Autism Develop Disord*. 2007;37(7):1323–1336.
96. Landa, R, Piven, J, Wzorek, MM, et al. Social language use in parents of autistic individuals. *Psychol Med*. 1992;22(1):245–254.

97. Dabrowska, A, Pisula, E. Parenting stress and coping styles in mothers and fathers of pre-school children with autism and Down syndrome. *J Intellect Disabil Res.* 2010;54(3):266–280.
98. Al Kandari, S, Alsalem, A, Abohaimed, S, et al. Brief report: social support and coping strategies of mothers of children suffering from ASD in Kuwait. *J Autism Dev Disord.* 2017;47(10):3311–3319.
99. Carmassi, C, Corsi, M, Bertelloni, CA, et al. Mothers and fathers of children with epilepsy: gender differences in post-traumatic stress symptoms and correlations with mood spectrum symptoms. *Neuropsychiatr Dis Treat.* 2018;14:1371–1379.
100. Dell’Osso, L, Dalle Luche, R, Carmassi, C. A new perspective in post-traumatic stress disorder: which role for unrecognized autism spectrum? *Int J Emerg Mental Health Hum Resil.* 2015;17:436–438
101. Carmassi, C, Corsi, M, Bertelloni, CA, et al. Post-traumatic stress and major depressive disorders in parent caregivers of children with a chronic disorder. *Psychiatry Res.* 2019;279:195–200.101
102. Geurts, HM, Bergh, SF, Ruzzano, L. Prepotent response inhibition and interference control in autism spectrum disorders: two meta-analyses. *Autism Res.* 2014;7(4):407–420.
103. Siegle, GJ, Moore, PM, Thase, ME. Rumination: one construct, many features in healthy individuals, depressed individuals, and individuals with lupus. *Cognit Ther Res.* 2004;28(5):645–668.
104. Dell’Osso, L, Del Grande, C, Gesi, C, et al. A new look at an old drug: neuroprotective effects and therapeutic potentials of lithium salts. *Neuropsychiatr Dis Treat.* 2016;12:1687–1703.
105. Carmassi, C, Palagini, L, Caruso, D, et al. Systematic review of sleep disturbances and circadian sleep desynchronization in autism spectrum disorder: toward an integrative model of a self-reinforcing loop. *Front Psychiatry.* 2019;10:366.

Table 1. Comparison between subjects with or without a psychiatric disorder on AdAS spectrum and RRS score

	Subjects with psychiatric disorders (N= 47) Mean ± SD	Subjects without psychiatric disorders (N= 73) Mean ± SD	<i>t</i>	<i>P</i>
<i>AdAS spectrum</i>				
Childhood/Adolescence	6.60 ± 4.06	4.60 ± 3.29	-2.90	.005
Verbal communication	5.13 ± 2.95	3.39 ± 2.07	-3.46	.001
Nonverbal communication	9.04 ± 4.66	5.48 ± 3.58	-4.60	<.001
Empathy	3.80 ± 2.42	2.84 ± 2.27	-2.17	.032
Inflexibility / Adherence to routine	11.76 ± 6.76	8.69 ± 5.61	-2.63	.010
Restricted interests / Rumination	6.07 ± 3.82	3.73 ± 3.06	-3.45	.001
Hyper/Hyporeactivity to sensory input	3.34 ± 2.49	2.34 ± 2.05	-2.35	.021
Total	45.76 ± 21.22	31.06 ± 17.50	-4.02	<.001
<i>RRS</i>				
Reflection	8.46 ± 2.50	7.06 ± 2.29	-3.03	.003
Brooding	10.50 ± 2.93	8.41 ± 2.39	-4.09	<.001
Depression	21.82 ± 5.78	18.52 ± 4.83	-3.26	.001
Total	40.78 ± 10.12	33.83 ± 8.20	-3.96	<.001

Results statistically significant for $p < .05$ are in bold.

Table 2. Pearson's correlation between AdAS Spectrum, RRS, and SOFAS scores.

	SOFAS	RRS reflection	RRS brooding	RRS depression	RRS total
<i>AdAS spectrum</i>					
Childhood/Adolescence	-0.284*	0.355*	0.370*	0.338*	0.403*
Verbal communication	-0.399*	0.462*	0.540*	0.581*	0.615*
Nonverbal communication	-4.55*	0.439*	0.535*	0.546*	0.592*
Empathy	-0.159	0.265*	0.457*	0.406*	0.442*
Inflexibility / Adherence to routine	-0.245#	0.371*	0.525*	0.496*	0.539*
Restricted interests / Rumination	-0.287*	0.481*	0.550*	0.544*	0.598*
Hyper/Hyporeactivity to sensory input	-0.200#	0.366*	0.401*	0.423*	0.459*
Total	-0.376*	0.491*	0.612*	0.599*	0.658*
<i>RRS</i>					
Reflection	-0.345*		-		
Brooding	-0.330*		-		
Depression	-0.437*		-		
Total	-0.430*		-		

* $p < .005$.

$p < .05$.

Table 3. Multiple logistic regression analysis for predictors of psychiatric disorders. Results statistically significant for $p < .05$ are in bold.

Model	B(SE)	OR	95% CI	<i>p</i>
K	-4.46 (1.26)	0.01	-	<.001
Gender	1.23 (0.49)	3.43	1.32-8.90	.011
<i>AdAS spectrum</i>				
Childhood/Adolescence	0.00 (0.08)	1.00	0.86-1.16	.973
Verbal communication	0.06 (0.14)	1.06	0.80-1.40	.678
Nonverbal communication	0.18 (0.09)	1.20	1.01-1.42	.041
Empathy	0.02 (0.12)	1.02	0.81-1.29	.877
Inflexibility / Adherence to routine	0.08 (0.07)	1.93	0.82-1.05	.238
Restricted interests / Rumination	0.08 (0.11)	1.09	0.87-1.35	.463
Hyper/Hyporeactivity to sensory input	0.04 (0.15)	0.94	0.72-1.30	.802
<i>RRS</i>				
Reflection	0.02 (0.14)	0.98	0.74-1.29	.883
Brooding	0.21 (0.13)	1.24	0.96-1.59	.100
Depression	0.02 (0.07)	0.98	0.86-1.13	.805

Cox $R^2 = 0.246$; Nagelkerke $R^2 = 0.331$.

Results statistically significant for $p < .05$ are in bold.

Table 4. Multiple linear regression with SOFAS score as dependent variable.

Model	B (SE)	β	95% CI	<i>t</i>	<i>p</i>
K	104.49 (4.84)	-	94.87 to 114.11	21.57	<.001
Gender	-3.41 (2.05)	0.15	-7.47 to 0.66	0.167	.099
<i>AdAS spectrum</i>					
Childhood/Adolescence	0.13 (0.35)	0.04	0.83 to 0.58	0.36	.724
Verbal communication	0.56 (0.63)	0.13	-1.81 to 0.68	0.90	.371
Nonverbal communication	0.96 (0.37)	0.37	-1.70 to -0.22	-2.57	.012
Empathy	0.12 (0.50)	0.02	0.87 to 1.10	0.23	.818
Inflexibility / Adherence to routine	0.20 (0.28)	0.11	0.36 to 0.75	0.70	.486
Restricted interests / Rumination	0.24 (0.51)	0.07	0.77 to 1.24	0.47	.641
Hyper/Hyporeactivity to sensory input	0.44 (0.64)	0.08	0.83 to 1.67	0.69	.494
<i>RRS</i>					
Reflection	0.01 (0.62)	0.00	-1.23 to 1.21	0.01	.992
Brooding	0.13 (0.57)	0.03	0.099 to 1.25	0.23	.820
Depression	0.59 (0.33)	0.28	-1.25 to 1.21	-1.76	.082

Results statistically significant for $p < .05$ are in bold.

Figure 1. Mediation analysis results

