

# The Impact of Adult ADHD in the Quality of Life Profile

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3

## 4 Abstract

5 In this Multi-center, cross-sectional study we aimed to investigate the quality of life (QoL),  
6 and the neuropsychological and clinical characteristics of adults with attention-  
7 deficit/hyperactivity disorder (ADHD) with different developmental pathways. Our study  
8 sample included 25 control (healthy) subjects, 31 subjects with newly diagnosed ADHD  
9 without comorbidities (ADHD-C-D), 31 with newly diagnosed ADHD with comorbidities  
10 (ADHD+C-D), and 29 with previously diagnosed ADHD with comorbidities (ADHD+C+D).  
11 Compared with controls, ADHDs had little differences in the attentional performance but  
12 showed higher impulsivity, more severe symptoms of depression and anxiety, and lower  
13 QoL. The ADHD+C-D group showed more severe symptoms of depression and anxiety than  
14 the ADHD+C+D group ( $p = .037$  and  $p = .008$ , respectively), and poorer QoL in the  
15 psychological health sphere ( $p = .003$ ). In conclusion, differences between ADHD and  
16 control subjects were particularly remarkable in mood symptoms and QoL. Previous  
17 diagnosis might have a positive impact on mood symptoms and QoL in ADHD adults.

18

19 **Keywords:** ADHD, quality of life, adult ADD, comorbidity, developmental pathways

20

## 21 **Introduction**

22 Attention-Deficit/Hyperactivity Disorder (ADHD) has been thought to affect mainly during  
23 childhood and adolescence. However, there is increasing consensus in the fact that ADHD  
24 evolves throughout the patient's lifespan rather than ceasing in adulthood (L. Adler & Cohen,  
25 2004; Biederman, 2005; Jadidian, Hurley, & Taber, 2015; Young & Gudjonsson, 2008).

26 Overall, some ADHD core symptoms tend to decline over time, and they may manifest in  
27 different forms as patients adjust their social and personal environment to the  
28 symptomatology (L. Adler & Cohen, 2004; Biederman, Mick, & Faraone, 2000). In any case,  
29 ADHD will affect – to a greater or lesser extent – many aspects of the adult's life regardless  
30 of the degree of symptoms remission. Authors studying the impact of ADHD throughout the  
31 patient's lifespan observed a long-term persistence of the poor interpersonal skills, which  
32 resulted not only in fewer close friendships in the adulthood but also in a greater number of  
33 remarriages than control subjects (Bagwell, Molina, Pelham, & Hoza, 2001; Ingram,  
34 Hechtman, & Morgenstern, 1999; Murphy & Barkley, 1996; Wilson & Marcotte, 1996).  
35 Likewise, the reduced academic performance of ADHD patients, also characterized by  
36 increased disciplinary actions at school, results in a lower educational attainment (Ingram et  
37 al., 1999; Murphy & Barkley, 1996), limiting their access to qualified job positions. In the  
38 occupational area, ADHD patients have also shown greater chances to change jobs, either  
39 because they leave or they are dismissed (R A Barkley, 1998; Murphy & Barkley, 1996;  
40 Weiss, Hechtman, Milroy, & Perlman, 1985).

41 In addition to the multiple developmental pathways of ADHD patients, ADHD diagnosis and  
42 treatment during childhood influences the course of patients' life, leading to a broad diversity  
43 of clinical profiles in adult patients (L. Adler & Cohen, 2004; Biederman, 1998, 2005). In

44 some cases, patients succeed in coping with ADHD core symptoms, mostly by developing  
45 alternative behaviors, which results in a compensated psychological and cognitive function.  
46 However, the most common scenario is ADHD core symptoms persisting – more or less  
47 pervasively – during adulthood. Some of these patients followed an adaptive pathway, having  
48 a high rate of syndromic and symptomatic remission and a partially restored functioning. By  
49 contrast, others have to deal with a remarkable number of severe ADHD symptoms, which in  
50 most cases result in the emergence of comorbidities related to mood, anxiety, bipolar  
51 disorders, personality disorders, antisocial behavior and substance abuse disorders,  
52 particularly common in adult ADHD patients, with a prevalence that may reach up to 60%  
53 (L. Adler & Cohen, 2004; Fayyad et al., 2007). In these patients, comorbidities are more  
54 likely to be the actual therapeutic target, and the symptoms associated with the patient's  
55 comorbidity burden may even mask the inattention and hyperactivity symptoms, thus  
56 overlooking ADHD diagnosis and treatment (Ginsberg, Quintero, Anand, Casillas, &  
57 Upadhyaya, 2014).

58 Regardless of the developmental pathway of ADHD patients, both the evolution of the core  
59 symptoms during adulthood and the clinical complexity of adult ADHD patients limit the  
60 functional assessment and often hamper the identification of adult subjects with difficulties  
61 caused by an underlying ADHD. In this regard, some authors have highlighted the need to  
62 clarify ADHD symptoms beyond inattention and hyperactivity and to consider the multiple  
63 developmental pathways and the neuropathological heterogeneity of ADHD in adults  
64 (Russell A Barkley & Murphy, 2010; Nigg, 2005; Seidman, 2006; Sonuga-Barke, 2005).  
65 Thus, the quality of life (QoL) assessment may be considered a measure of the ADHD long-  
66 term outcomes, which encompasses the impact of both executive and emotional dysfunctions  
67 associated with the disorder. Some authors reported a negative correlation between different  
68 QoL measures and the severity of ADHD symptoms (L. A. Adler et al., 2009; Mattos, Louzã,

69 Palmi, de Oliveira, & Rocha, 2013). However, to our knowledge, QoL assessment in adult  
70 subjects with ADHD have been mainly focused on the outcomes of the various treatments (L.  
71 A. Adler et al., 2009; Mattos et al., 2013; Mick, Faraone, Spencer, Zhang, & Biederman,  
72 2008). Also, since ADHD has been traditionally considered a cognitive disorder, most trials  
73 including adult patients with ADHD have focused on the study of the executive and  
74 neuropsychological dysfunctions of these patients (Russell A. Barkley & Biederman, 1997;  
75 Seidman, 2006), while their emotional characteristics have been barely described.

76 Considering this background, the aim of this cross-sectional study was to assess the QoL and  
77 the neuropsychological, clinical and emotional characteristics of adult patients with ADHD,  
78 including those without previous ADHD diagnosis. To better understand the different profiles  
79 of adult ADHD patients, we grouped them according to various developmental pathways and  
80 investigated the behavior of each variable in all groups. In our analysis, we included a control  
81 group without a history of ADHD, and with confirmed absence of ADHD diagnosis.

## 82 **Material and methods**

### 83 *Study design*

84 In this multi-center, cross-sectional study, we assessed the neuropsychological performance,  
85 the clinical profile, and the QoL of adult ADHD patients and subjects without ADHD  
86 diagnosis at the study start. We recruited control and ADHD adults from two different mental  
87 health units in Madrid (Spain), and two support associations for patients affected by ADHD;  
88 control subjects included relatives of our ADHD patients who voluntarily agreed to  
89 participate in the study. Data were collected between October 2013 and December 2014. In  
90 addition to the scales related to the study outcomes, participants' clinical history was  
91 reviewed for previous mental disorders. The diagnosis interview for adult ADHD (DIVA)  
92 (Kooij, 2006) was used to confirm, rule out or diagnose ADHD for the first time in the study

93 subjects. This interview focuses on the 18 DSM-IV symptoms of ADHD and uses concrete  
94 and realistic examples to thoroughly investigate whether a symptom is currently present or  
95 was present in childhood. The assessment of ADHD symptoms and impairment in childhood  
96 included additional information retrieved from patients' relatives, when possible.

97 Accordingly, the Wender Utah Rating Scale (WURS) (Ward, Wender, & Reimherr, 1993)  
98 allowed for a retrospective assessment of the ADHD symptoms during childhood.

99 Assessments were performed by trained professionals: two experienced psychologists, R.V.  
100 and I.M. Before starting data collection, the investigators explained the methodology and the  
101 objectives of the study to eligible subjects, who were offered to freely participate in the study.  
102 All participants were native Spanish speakers and signed the informed consent approved by  
103 the research ethics committee of Gregorio Marañón Hospital (Madrid, Spain) (ref. 256/13)  
104 before entering the study. All data were managed in accordance with the local regulation on  
105 personal data protection (LOPD 13/1999).

### 106 *Subjects*

107 Male and female subjects aged between 18 and 65 years who were able to understand the  
108 instructions needed to carry out the proposed tests were included in the study. In order to  
109 minimize the bias associated with cognitive impairment, subjects with full-scale intelligence  
110 quotient below 70 on the Wechsler Adult Intelligence Scale (WAIS) were excluded from the  
111 study.

112 Participants were classified according to three characteristics: (a) current ADHD diagnosis  
113 (according to the DIVA assessment), (b) previous history of ADHD diagnosis in childhood or  
114 adolescence, and (c) presentation with comorbid psychiatric pathologies (according to the  
115 Structured Clinical Interview for DSM-IV Axis I Disorders [SCID-IV]) (First, Spitzer,  
116 Gibbon, & Williams, 1999). Consequently, participants were divided into four groups as

117 follows: 1. Non-ADHD healthy adults (control group); 2. ADHD adults without comorbidity  
118 and undiagnosed in childhood or adolescence (ADHD-C-D); 3. ADHD adults with  
119 comorbidity and undiagnosed in childhood or adolescence (ADHD+C-D); and 4. ADHD  
120 adults with comorbidity and previous ADHD diagnosis in their childhood or adolescence  
121 (ADHD+C+D). The putative group including patients without comorbidities and with  
122 previous ADHD diagnosis in their childhood or adolescence (ADHD-C+D) was discarded  
123 because all patients with previous ADHD diagnosis had at least one additional psychiatric  
124 comorbid included in the SCID-IV.

### 125 *Rating scales*

126 The severity of the ADHD symptoms in adulthood was assessed using the Conners' Adult  
127 Attention Rating Scale (CAARS) (C Keith Conners, 1999; La Malfa, Lassi, Bertelli, Pallanti,  
128 & Albertini, 2008). The presence of childhood symptoms was retrospectively assessed with  
129 the 61-item Spanish version of the WURS (Rodríguez-Jiménez et al., 2001). Participants'  
130 neuropsychological performance was assessed using three scales: the Conners' Continuous  
131 Performance Test (CPT), the Stroop Color-Word Interference Test (SCWT), and the  
132 Wechsler Adult Intelligence Scale (WAIS). The CPT (C K Conners, 1993) included the  
133 computerized measure of the number of omissions (missed targets), commissions (incorrect  
134 responses to non-targets), and hit reaction times. The SCWT measures the subject's ability to  
135 avoid the semantic interference when naming printed colors not matching the name of the  
136 color. The version used in this study contained 100 items corresponding to three different  
137 colors, and subjects were asked to read as many items as possible for 45 seconds (Golden &  
138 Freshwater, 1978). The WAIS test was used in its 4th edition of fifteen subtests grouped into  
139 four indexes: the verbal comprehension index, the perceptual reasoning index, the working  
140 memory index, and the processing speed index (Wechsler, 2014). For the purposes of this  
141 study, only the working memory index, the processing speed index, and the full-scale

142 intelligence quotient index (based on the combined performance of the four indexes) were  
143 considered for the analysis.

144 In addition to the SCID-IV, the clinical and emotional profile of patients and controls was  
145 defined in terms of impulsivity, anxiety, and severity of depressive symptoms. Impulsivity  
146 was assessed using the Barratt Impulsiveness Scale (BIS-11), including the attentional, non-  
147 planning, and motor subscales (Patton, Stanford, & Barratt, 1995). Anxiety was estimated as  
148 both anxiety trait and anxiety state using the State-Trait Anxiety Inventory (STAI)  
149 (Spielberg, Gorsuch, & Lushene, 2008). Finally, the severity of depressive symptoms was  
150 assessed using the Beck's Depression Inventory (BDI) (Beck, Ward, Mendelson, Mock, &  
151 Erbaugh, 1961).

152 The patients' QoL was measured using the Adult ADHD Quality-of-life (AAQoL) scale, a  
153 disease-specific tool for the assessment of QoL in adult ADHD patients (Brod et al., 2015).  
154 The AAQoL scale is based on a self-administered questionnaire of 29 items rated on a 5-  
155 point Likert scale. Items are grouped into four domains: (1) the life productivity subscore,  
156 regarding the ability to complete tasks and balance multiple projects; (2) the psychological  
157 health subscore, including feelings of being overwhelmed, anxious or fatigued; (3) the  
158 relationships subscore, concerned to the feelings of tension, annoyance, and frustration in  
159 relationships; and (4) the life outlook subscale, regarding the overall perception of  
160 satisfaction and success in life management. Finally, a QoL total score is obtained from the  
161 four subscale scores. In this study, participants completed the AAQoL questionnaire at home  
162 and returned it on the following visit.

### 163 *Statistical analysis*

164 Sociodemographic variables, as well as the scores of the various scales, were described for  
165 the study sample as a whole and for each individual group. Quantitative variables were

166 described as means and standard deviations or medians, while categorical variables were  
167 described as frequencies and percentages for each group. The outcome variables were  
168 compared using ANOVA or ANCOVA tests when the individual values were normally  
169 distributed, whereas the Kruskal-Wallis test was utilized for variables that failed to show a  
170 normal distribution (assessed using the Shapiro-Wilk test). The *post hoc* pairwise analyses of  
171 variables showing significant differences in the ANOVA or the ANCOVA tests were carried  
172 out using either the Bonferroni correction or the Dunnett's test. Accordingly, variables  
173 showing significant differences in the Kruskal-Wallis analyses were compared with the  
174 Dunn's test. The chi-square or Fisher's exact tests were used to compare the proportions of  
175 categorical variables among groups. The significance level for general hypothesis testing was  
176 set at  $\alpha = .05$ , and the analyses were conducted using the SPSS software (Version 22.0 for  
177 Windows. Armonk, NY: IBM Corp.) and MedCalc.

## 178 **Results**

### 179 *Subjects' characteristics*

180 A total of 119 subjects were considered for eligibility but 3 were excluded from control group  
181 due to a positive diagnosis on at least one DSM-IV axis I disorder, which provided a final  
182 study sample size of 116 subjects. After the initial assessment, 25 subjects were included in  
183 the control group, 31 in the ADHD-C-D group, 31 in the ADHD+C-D group, and 29 in the  
184 ADHD+D+C group.

185 Table 1 shows the subjects' main social and demographic characteristics in each group.

186 Overall, our sample was balanced in sex, but showed significant differences in most of the  
187 sociodemographic characteristics. The *post hoc* analysis of the mean age in the study groups  
188 revealed that subjects in the ADHD+C+D group were significantly younger than those

189 included in other ADHD groups ( $p < .05$  for both pairwise comparisons). Therefore, age was  
190 included as a covariate in the ANCOVA analyses when required.

191 Significant differences were also found regarding the level of independence and the  
192 relationship status. The proportion of subjects engaged in a relationship was significantly  
193 higher in the control group and lower among patients in the ADHD+C-D group. To further  
194 investigate the relationship between each group and the level of independence, patients were  
195 grouped into two main categories: independent and non-independent. The chi-square test  
196 revealed significant differences between study groups ( $p < .001$ ), with a greater proportion of  
197 independent subjects in the control group, and a greater proportion of non-independent  
198 subjects in the ADHD+C-D group.

199 Regarding the educational attainment, the control group had a greater proportion of subjects  
200 with higher education than the ADHD groups. Control subjects were also less likely to  
201 interrupt education before graduation and reported a lower incidence of signal events related  
202 to school history, such as absenteeism and disruptive behavior. The control group showed  
203 higher scores at secondary school and a lesser number of grade retentions than ADHD groups  
204 ( $p < .001$  and  $p < .05$  for all pairwise comparisons of scores and number of grade retentions,  
205 respectively) but no significant differences were observed between ADHD groups. Only  
206 subjects with comorbidities reported episodes of disruptive behavior during the school period,  
207 with a higher incidence in subjects with a previously diagnosed ADHD (ADHD+C+D).

208 The proportion of patients in each level of occupational attainment significantly differed  
209 among the study groups. To further investigate the relationship between study groups and  
210 working status, we regrouped the four initial categories into two main categories: employed  
211 and unemployed. The chi-square test for this new set of groups revealed a significantly  
212 greater proportion of employed subjects in the control group ( $p < .01$ ) than in any ADHD

213 group. Overall, study subjects reported a low number of job changes and layoffs, which were  
214 remarkably greater in the ADHD+C-D group.

### 215 *Neuropsychological and clinical characteristics*

216 Table 2 summarizes the neuropsychological mean scores and the subject clinical  
217 characteristics in each group. When assessing the neuropsychological performance, only the  
218 number of commissions in the CPT test and the full-scale intelligence quotient in the WAIS  
219 test were significantly different among study groups. For CPT commission score significant  
220 differences between the control group and ADHD+C-D and ADHD+C+D groups emerged in  
221 the Dunnett test *post hoc* analysis ( $p = .011$  and  $p = .039$ , respectively). For the full-scale  
222 intelligence quotient, the *post hoc* analysis revealed significant differences only between  
223 control subjects and those in the ADHD+C-D group ( $p = .003$ ). Finally, the overall  
224 comparison of the word-color index mean scores in the SCWT scale were close to the  
225 significance level ( $p = .086$ ), suggesting a trend to lower scores in control subjects as  
226 compared with all ADHD patients.

227 Study groups showed significant differences in the scores of all scales related to the clinical  
228 and emotional profile (Table 2). In the *post hoc* analysis of the BIS-11 scores, control  
229 subjects showed significantly lower scores than ADHD subjects in all subscales ( $p < .001$  for  
230 all pairwise comparisons); however, no significant differences were found among groups  
231 including ADHD patients. The corresponding analysis of STAI also revealed significantly  
232 lower trait and state anxiety scores in control subjects than in ADHD subjects ( $p < .01$  for all  
233 pairwise comparisons). For ADHD subjects, STAI mean scores were significantly higher in  
234 the ADHD+C-D group than in the ADHD+C+D group ( $p = .008$ ). Similarly, the severity of  
235 the depressive symptoms, assessed by the BDI scale, was significantly lower in control  
236 subjects than in ADHD subjects with comorbidities ( $p < .01$  for pairwise comparisons with

237 both ADHD+C-D and ADHD+C+D groups). When comparing ADHD groups, the BDI mean  
238 scores were significantly higher in the ADHD+C-D group than in the ADHD-C-D ( $p < .001$ )  
239 and ADHD+C+D ( $p = .037$ ) groups. The SCID-IV allowed identifying the presence of 8  
240 different comorbidities in the ADHD+C-D group, and 10 in the ADHD+C+D group (Figure  
241 1). However, the differences in the SCID outcomes between these two groups including  
242 patients with comorbidities were not significant.

243 Significant differences were seen in the severity of ADHD symptoms in the overall  
244 comparison (Table 2). The *post hoc* analysis of symptoms severity during adulthood,  
245 measured by the CARRS, showed significantly lower scores in the control group than in each  
246 of the ADHD groups ( $p < .001$  for all pairwise comparisons), but no significant differences  
247 between ADHD groups (pairwise comparisons). Similarly, the severity of symptoms during  
248 childhood, measured by the WURS, was significantly milder in control subjects than in  
249 ADHD subjects ( $p < .001$  for all pairwise comparisons). In this scale, subjects in the  
250 ADHD+C+D group showed a higher mean score than those in the ADHD+C-D group ( $p =$   
251  $.029$ ).

### 252 *Quality of life*

253 The self-perceived QoL, assessed by the AAQoL scale, was significantly different among the  
254 study groups (Figure 2). The *post hoc* analyses revealed significant differences between  
255 control subjects and those with ADHD diagnosis in all subscales, including the total AAQoL  
256 score ( $p < .05$  for all pairwise comparisons). Regarding the differences in the pairwise  
257 comparisons between ADHD groups, the ADHD+C-D group had significantly lower scores  
258 than the ADHD-C-D group in the productivity subscale ( $p = .01$ ), the psychological health  
259 subscale ( $p = .001$ ) and the total AAQoL scale ( $p = .006$ ). Subjects in the ADHD+C-D group

260 also exhibited a lower mean score in the psychological health subscale than those in the  
261 ADHD+C+D group ( $p = .003$ ).

## 262 **Discussion**

263 We investigated the QoL and the neuropsychological and clinical characteristics in different  
264 adult ADHD profiles and found that the developmental pathway of these patients exerts  
265 greater influence on the emotional manifestations than on the ADHD core symptoms. Among  
266 the different developmental pathways of ADHD subjects, we found that the diagnosis at an  
267 early age (childhood or adolescence) might have a positive impact on some QoL dimensions  
268 later on life.

269 In the last decades, only the most dysfunctional ADHD cases were properly identified,  
270 whereas subjects with less dysfunctional symptoms or with partially adaptive developmental  
271 pathways remained underdiagnosed (Russell A Barkley, Fischer, Smallish, & Fletcher, 2002;  
272 Russell A Barkley, Murphy, & Fischer, 2008; Setyawan et al., 2015). Moreover, the presence  
273 of comorbidities and their severity might increase the likelihood of receiving a specialized  
274 attention, thus favoring an increase in the diagnosis rate (Sayal, Mills, White, Merrell, &  
275 Tymms, 2015). In line with this retrospective scenario, our study subjects previously  
276 diagnosed with ADHD were significantly younger, showed more severe childhood symptoms  
277 in the retrospective interview, and reported more episodes of disruptive behavior during the  
278 school period than other ADHD subjects. In other indicators of school performance, such as  
279 school dropout rate, final grades, and the number of grade retentions, the comparisons  
280 between ADHD groups did not reveal significant differences. However, compared with  
281 control subjects, ADHD patients had lower grades and higher chances of early leaving school  
282 than control subjects. This observation is consistent with the temperament profile and the

283 greater likelihood of school dropout observed in previous studies with young ADHD patients  
284 (Flood et al., 2016; Ronna Fried et al., 2016; Willoughby, Gottfredson, & Stifter, 2016).

285 In addition to the time lapse in the diagnosis of ADHD, our results reflect an important risk of  
286 underdiagnosis in adults (Ginsberg et al., 2014). The modulation of the core symptoms  
287 (inattention, hyperactivity, and impulsivity) during adulthood has been pointed out as a major  
288 obstacle for ADHD diagnosis in adults. Indeed, whereas inattentiveness was central to  
289 ADHD diagnosis in the DSM-III, various authors have reported a low sensitivity of scales  
290 assessing attention in adults, particularly the CPT scale (Dulcan, 1997; Mcgee, Clark, &  
291 Symons, 2000). Likewise, the discrepancies observed when comparing the SCWT and WAIS  
292 scores between healthy and ADHD adult subjects (although being significant in most cases),  
293 led some authors to highlight the need for clarifying the executive dysfunctions associated  
294 with ADHD in adulthood (Boonstra, Oosterlaan, Sergeant, & Buitelaar, 2005; Ginsberg et al.,  
295 2014; Hervey, Epstein, & Curry, 2004; Seidman, 2006; Sergeant et al., 2002). In line with the  
296 observed modulation of attentional symptoms in adulthood, we found little differences  
297 between controls and ADHD subjects in the neuropsychological scales, with no significant  
298 results among study groups in the CPT omissions and hit reaction time subscales. We also  
299 failed to find significant differences in the WAIS working memory and processing speed  
300 indexes. These results suggest that even though neuropsychological tests are helpful tools for  
301 understanding the cognitive processes underlying ADHD, they do not fully describe the  
302 individual variability and might, therefore, be insufficient for ADHD diagnosis.

303 Symptoms related to impulsivity, rather than inattention, seem to have a greater influence on  
304 the difficulties that adult ADHD patients suffer in their working and relational spheres (Ogg,  
305 Bateman, Dedrick, & Suldo, 2016). However, there is no agreement on the extent of  
306 impulsivity persistence during adulthood, with some authors suggesting a faster decline in

307 impulsivity than in inattentive symptoms (Faraone et al., 2000), and others suggesting an  
308 intermediate persistence (up to 60%) (Young & Gudjonsson, 2008). In our study, healthy  
309 subjects displayed lower impulsivity scores in all BIS-11 subdomains. Accordingly, the CPT  
310 commission subscale, which is also considered a measure of impulsivity, was the only  
311 domain of the CPT scale that yielded significant differences among the study groups.

312 Notwithstanding the persistence of some cognitive symptoms and executive dysfunctions in  
313 adulthood, there is growing concern on the limitations of the executive and  
314 neuropsychological assessment in the diagnosis and management of adult ADHD patients  
315 (Russell A Barkley & Murphy, 2010; Ginsberg et al., 2014; Nigg, 2005; Seidman, 2006).

316 Some authors suggested that, along with the traditional ADHD symptoms, anxiety and  
317 depressive symptoms mediate the decline of QoL during adulthood (Yang, Tai, Yang, & Gau,  
318 2013). Comorbidities increase with age (Russell A Barkley et al., 2002; Murphy & Barkley,  
319 1996) and often arise when the subject leaves the controlled, familial environment (Russell A  
320 Barkley et al., 2002). It seems reasonable that the core symptoms of ADHD progress through  
321 an internalizing pattern as patients adapt themselves to the social environment, increasing the  
322 risk of developing anxiety. In our study, the differences between healthy and ADHD subjects  
323 were more remarkable in the scales assessing anxiety and depression than in those assessing  
324 the neuropsychological performance. Interestingly, among subjects with comorbidities  
325 (ADHD+C-D and ADHD+C+D groups), those without a previous ADHD diagnosis showed  
326 the highest anxiety scores. Likewise, patients in the ADHD+C-D group showed a  
327 significantly greater severity of depressive symptoms than other ADHD groups. Furthermore,  
328 depression has been related to the number and severity of conflicts and adverse events during  
329 life (Yang et al., 2013), but also to low school performance in adolescents with non-  
330 diagnosed cognitive dysfunctions (Schulte-Koerne, 2016). Considering these observations, it  
331 is not surprising that the ADHD+C-D group exhibited more frequent job changes and layoffs,

332 and a higher severity of depressive symptoms than other ADHD groups. Hence, in our  
333 sample, the absence of a previous ADHD diagnosis seemed to exert more influence on mood  
334 symptoms and adverse life events than the intensity of ADHD symptoms itself, since similar  
335 CAARS scores were observed in all ADHD groups. This observation is consistent with that  
336 of other authors who showed how the failure to diagnose ADHD prevents children and their  
337 parents from seeking the assistance they need to achieve their full potential in academic and  
338 psychosocial settings (Fredriksen et al., 2014; R. Fried et al., 2016).

339 Due to the psychological dysfunction and disability, ADHD has a strong impact on multiple  
340 dimensions of patients' life, including school performance, job success, friendship, and  
341 partner relationships. Therefore, a negative correlation between ADHD and the patients' QoL  
342 may be expectable (L. A. Adler et al., 2009; Mattos et al., 2013; Yang et al., 2013). For the  
343 assessment of the QoL we used an ADHD specifically designed scale, which provides  
344 separate information about four dimensions of the patients' life. As expected, control subjects  
345 exhibited better QoL than those with ADHD. Among ADHD subjects, the presence of  
346 comorbidities influenced QoL in most of the AAQoL domains. Interestingly, subjects in the  
347 ADHD+C-D group showed the lowest scores in all AAQoL subscales. These differences  
348 were statistically significant when compared with subjects without comorbidities in the total  
349 AAQoL score, and in the productivity and psychological health subscores. In the  
350 psychological health subscale, subjects with comorbidities and newly diagnosed ADHD  
351 (ADHD+C-D) also showed a significantly lower score than those with previous ADHD  
352 diagnosis. The psychological health subscale contains six items including feeling anxious,  
353 overwhelmed, and fatigued. Hence, the differences observed in the AAQoL scores between  
354 these two ADHD groups with comorbidities are consistent with the trend observed in the  
355 state and trait anxiety scores, which were also higher among newly diagnosed subjects than  
356 among those with a previous ADHD diagnosis.

357 The representativeness of our results might be limited by a selection bias potentially  
358 associated with the recruitment process. Considering the difficulties of recruiting non-  
359 diagnosed ADHD subjects, we screened subjects from two different settings: mental health  
360 units and support associations for ADHD patients. Thus, ADHD subjects who have followed  
361 an adaptive pathway and have compensated their difficulties could be more likely to be  
362 included in our study, whereas those with greater dysfunctionality may be underrepresented.  
363 Nonetheless, our study is innovative in the comprehensive description of different adult  
364 ADHD profiles, particularly those without a previous diagnosis. Finally, it would have been  
365 of great interest to include retrospective data about treatment in previously diagnosed ADHD  
366 subjects. However, we considered that retrospectively retrieving such information might have  
367 led to inaccurate records and a subsequent increase in the risk of bias.

368 In conclusion, our results show that despite the heterogeneity of adult ADHD subjects, the  
369 different developmental pathways – and most particularly the presence/absence of  
370 comorbidities and a previous diagnosis during childhood – display some consistent  
371 characteristics. The various profiles of adult ADHD subjects, rather than the  
372 neuropsychological scales, seem to differ in the emotional symptoms and the QoL. We also  
373 found that regardless of the limitations of ADHD management in the previous decades, the  
374 presence of a previous ADHD diagnosis might result in a better outcome, particularly in the  
375 emotional domain, which in turn has a relevant impact on the QoL of adult ADHDs. In this  
376 regard, future investigations should explore the individual influence of the various symptoms,  
377 dysfunctions, and comorbidities on the QoL of adult ADHD patients.

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**Table 1.** Social and demographic characteristics of subjects included in each study group. Percentages were calculated for each group.

	Overall (n = 116)	Subgroups			<i>p</i> value <sup>a</sup>	
		Control (n = 25)	ADHD-C- D (n = 31)	ADHD+C- D (n = 31)		ADHD+C+ D (n = 29)
Age (years), <i>mean (SD)</i>	38.3 (11.5)	43.6 (11.7)	37.8 (11.5)	30.5 (11.0)	<b>&lt; .001</b>	
Sex, <i>n (%) of females</i>	65 (56%)	14 (56%)	17 (55%)	19 (61%)	.900	
Relationship status, <i>n (%)</i>						
Never lived with a partner	19 (16%)	2 (8%)	4 (13%)	5 (16%)	8 (28%)	<b>.009</b>
Currently with a partner	86 (74%)	21 (84%)	27 (87%)	23 (74%)	15 (52%)	
Undetermined	11 (9%)	2 (8%)	0	3 (10%)	6 (21%)	
Independence level, <i>n (%)</i>						
Dependent of parents	23 (20%)	1 (4%)	2 (7%)	8 (26%)	12 (41%)	<b>&lt; .001</b>
Partially independent	9 (8%)	0	1 (3%)	1 (3%)	7 (24%)	
Independent, living alone	16 (14%)	1 (4%)	5 (16%)	7 (23%)	3 (10%)	
Living with a partner and/or children	66 (57%)	22 (88%)	23 (74%)	15 (48%)	6 (21%)	
Other	2 (2%)	1 (4%)	0	0	1 (3%)	
Educational level, <i>n (%)</i>						
Postgraduate studies	13 (11%)	4 (16%)	3 (10%)	2 (6%)	4 (14%)	.372
Bachelor's degree	25 (22%)	9 (36%)	4 (13%)	6 (19%)	6 (21%)	
Graduated/university studies of 1-3 years	30 (26%)	5 (20%)	10 (33%)	6 (19%)	9 (31%)	
≤ 12 academic years	37 (32%)	5 (20%)	12 (40%)	14 (45%)	6 (21%)	
No primary education	10 (9%)	2 (8%)	1 (3%)	3 (10%)	4 (14%)	
School performance, <i>median</i>						
Score in secondary school	6	7	6	6	6	<b>&lt; .001</b>
Grade retentions	1	0	1	1	1	<b>.008</b>
School events, <i>n (%)</i>						
School failure	38 (33%)	2 (8%)	13 (42%)	11 (36%)	12 (43%)	<b>.023</b>
School absenteeism/escape	15 (13%)	0	1 (3%)	8 (26%)	6 (25%)	<b>.006</b>
Disruptive behavior						
Non-violent	20 (17%)	0	5 (16%)	5 (16%)	11 (35%)	<b>.010</b>
Violent	7 (6%)	0	0	1 (3%)	6 (21%)	<b>.001</b>
Working status, <i>n (%)</i>						
Stable employment	39 (34%)	12 (48%)	14 (45%)	10 (32%)	3 (10%)	<b>&lt; .01</b>
Unstable employment	32 (28%)	9 (36%)	7 (23%)	10 (32%)	6 (21%)	
Unemployed	21 (18%)	2 (8%)	4 (13%)	8 (26%)	7 (24%)	
Retired	24 (21%)	2 (8%)	6 (20%)	3 (10%)	13 (45%)	
Working events, <i>median</i>						
Job changes	3	3	3	4	3	.338 <sup>b</sup>
Layoffs	0	0	0	1	0	<b>.047<sup>b</sup></b>

**ADHD-C-D**, patients without comorbidities and without previous ADHD diagnosis; **ADHD+C-D**, patients with comorbidities and without previous ADAH diagnosis; **ADHD-C+D**, patients without comorbidities and with previous ADHD diagnosis; **SD**, standard deviation.

<sup>a</sup>Significant *p* values (< .05) are in bold.

<sup>b</sup>Calculated considering only subjects with previous working experience.

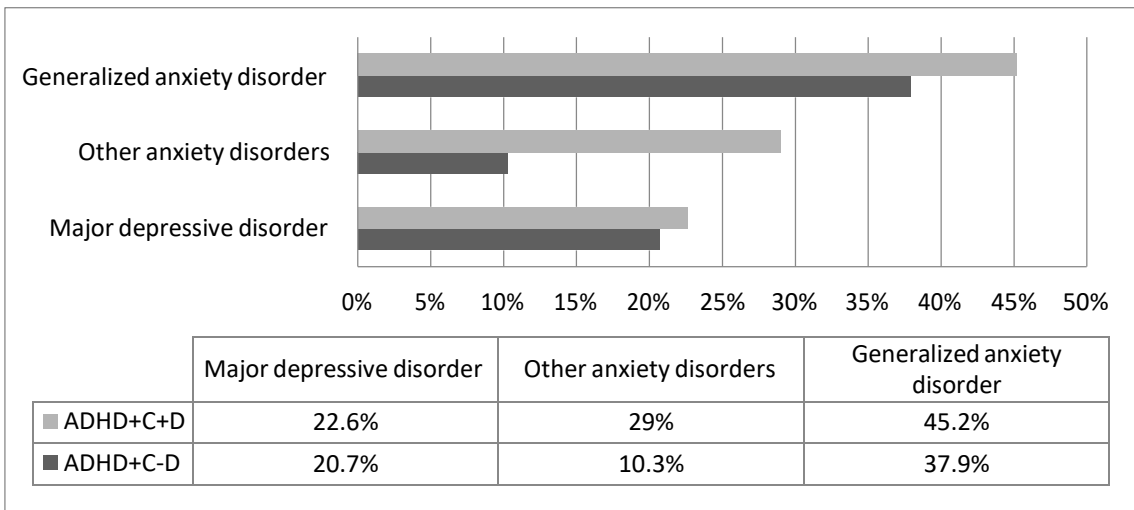


**Table 2.** Neuropsychological and clinical characteristics of the study subjects. Results are presented as the mean score (standard deviation) or median of each scale.

	Overall (n = 116)	Study groups				<i>p</i> value <sup>a</sup>
		Control (n = 25)	ADHD-C-D (n = 31)	ADHD+C-D (n = 31)	ADHD+C+D (n = 29)	
<b>Neuropsychological variables</b>						
Conners' continuous performance test						
Omissions	48.9 (27.1)	49.3 (28.6)	42.1 (24.1)	51.0 (27.7)	51.6 (27.8)	.444
Commissions	54.8 (31.6)	39.6 (29.1)	52.6 (32.8)	63.4 (27.8)	59.6 (33.0)	<b>&lt;.049</b>
Hit reaction time	62.2 (28.1)	65.0 (27.2)	61.8 (30.2)	64.2 (31.2)	59.8 (24.0)	.945
Stroop interference						
Word	1	1	1	1	1	.154
Color	1	1	1	2	1	.137
Word-Color	1	0	1	1	1	<b>.086</b>
Wechsler Adult Intelligence Scale						
Working memory index	98.6 (16.1)	102.3 (11.1)	100.0 (16.2)	93.4 (17.4)	97.7 (17.7)	.089
Processing speed index	102.9 (13.4)	108.2 (14.7)	104.5 (11.8)	99.9 (13.0)	98.6 (12.6)	.110
Full scale intelligence quotient	101.8 (16.5)	107.2 (13.0)	106.4 (13.2)	93.3 (20.4)	99.6 (14.5)	<b>.002</b>
<b>Clinical and emotional variables</b>						
Barratt impulsiveness scales						
Attentional	20.2 (8.3)	12.7 (3.8)	21.5 (4.7)	23.4 (8.7)	21.9 (9.6)	<b>&lt;.001</b>
Nonplanning	19.9 (8.6)	12.4 (5.0)	19.0 (7.7)	24.3 (8.6)	22.7 (7.7)	<b>&lt;.001</b>
Motor	20.3 (8.1)	12.1 (5.6)	21.4 (6.8)	23.6 (7.4)	22.5 (7.4)	<b>&lt;.001</b>
Total	56.5 (19.4)	35.8 (13.1)	59.8 (15.1)	66.2 (16.5)	60.3 (18.7)	<b>&lt;.001</b>
State-Trait Anxiety Inventory						
Trait	59.7 (28.7)	34.6 (25.3)	52.9 (25.4)	79.8 (16.1)	65.4 (27.9)	<b>&lt;.001</b>
State	59.3 (27.5)	38.4 (24.4)	55.4 (25.3)	75.2 (18.7)	64.6 (27.5)	<b>&lt;.001</b>
Beck's Depression Inventory	10.6 (8.5)	3.9 (2.9)	8.7 (6.3)	17.0 (8.5)	11.5 (9.0)	<b>&lt;.001</b>
<b>Intensity of ADAH symptoms</b>						
WURS	40.2 (19.5)	16.1 (11.3)	44.0 (12.5)	43.1 (15.5)	53.7 (17.1)	<b>&lt;.001</b>
Conners Adult Attention Rating Scale	13.8 (7.3)	5.4 (4.5)	15.4 (6.7)	15.8 (6.3)	17.2 (5.3)	<b>&lt;.001</b>

**ADHD-C-D**, patients without comorbidities and without previous ADHD diagnosis; **ADHD+C-D**, patients with comorbidities and without previous ADAH diagnosis; **ADHD-C+D**, patients without comorbidities and with previous ADHD diagnosis; **SD**, standard deviation.

<sup>a</sup>*p* value corresponding to given test statistic. Significant *p* values (< .05) are in bold.





Figure

