# **SUPPLEMENT**

# Relationship between quality of life and psychopathological profile: data from an observational study in children with ADHD

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**Abstract** Although ADHD significantly affects the quality of life (QoL) of patients and their families, QoL in children with ADHD has rarely been investigated in association with psychopathological profile, and the relationship remains unclear. The open-label OBSEER study evaluated the effectiveness and tolerability of Equasym XL<sup>®</sup>, a modified-release methylphenidate, in routine care of children and adolescents (aged 6-17 years) with ADHD. At baseline, questionnaires assessing psychopathological profile (Strengths and Difficulties Questionnaire, SDQ; parental ratings) and QoL (KINDL; parent, child or adolescent versions) were completed; OoL was reassessed at final visit. We analysed the relationship between psychopathology and parent/patient-rated QoL in ADHD at baseline. Data from 721 consecutively referred children and adolescents were analysed. QoL was similarly low from parent and self-ratings and independent of severity on the SDQ subscale hyperactivity/inattention. Self-ratings indicated that additional conduct disorder was associated with further reduction in QoL. Similarly, children with high scores from parent and adolescent ratings on the SDQ subscale conduct problems had reduced QoL on some KINDL subscales. Adolescents with ADHD not receiving

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D. Breuer · M. Döpfner Department of Child and Adolescent Psychiatry, University of Cologne, Cologne, Germany medication at baseline reported lower QoL than those already on medication. Results show that children and adolescents with ADHD have low QoL, independent of core symptom severity. Additional conduct problems may further impact QoL negatively, while ADHD medication use may show a trend towards improved QoL. Not all psychopathological problems associated with ADHD affect QoL similarly. As parents appear to have a less critical view of QoL compared with children's self-ratings, both parent and child ratings should be included in clinical assessments.

 $\begin{tabular}{ll} \textbf{Keywords} & SDQ \cdot Quality \ of \ life \cdot \ KINDL \ question naire \cdot \ Methylphenidate \cdot \ ADHD \end{tabular}$ 

## Introduction

Over the past few years, clinical work and research on quality of life (QoL) has become increasingly important in improving both physical and mental health in children. In the field of attention deficit hyperactivity disorder (ADHD), several studies on QoL have been published and were included in a systematic review [7]. This level of interest is not surprising due to the complexity of the relationships; QoL is not only influenced by the disorder itself, but also by many proximal (i.e. family, friendship) and distal (socioeconomic and cultural) factors. In addition to its core symptoms of attention deficit, hyperactivity and impulsivity, ADHD is associated with numerous developmental, cognitive, emotional, social and academic impairments [12, 17, 25]. One possible reason for this might be that in ADHD, more than 80% of children and adults are likely to have at least one other psychiatric disorder and more than 50% are likely to have two [10, 23]. Thus, in



ADHD, psychiatric comorbidity is the rule rather than the exception [24]. These comorbidities, as well as the other proximal and distal factors, affect the QoL of both patients and their families [7].

There is significant potential for overlap between the instruments designed to measure QoL and those used to measure psychopathology or functional impairment. Clear distinctions between symptoms (e.g. low mood or poor concentration) and their potential effects (i.e. functional impairment and/or reduced QoL), and also between functional impairment on the one hand and QoL on the other, are desirable [7]. Otherwise, there is a clear risk that apparent QoL effects are so closely related to symptoms and functional impairment that their association with the disorder will become a tautology [7]. In this context, it becomes evident that examining not only the relationship between QoL and categorical psychopathology (i.e. an ADHD diagnosis), but also the relationship with dimensional psychopathology (i.e. functional impairment and symptom scores), would result in improved understanding of the complex interplay of one or more disorders as well as of proximal and distal factors.

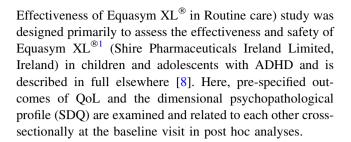
For an outcome as subjective as QoL, the influence of the rater's role should also be highlighted. A parent's rating of their child's QoL gives only one perspective of the overall impact of the illness on QoL. This may limit the validity of studies that do not ask children for their view on their own QoL. Such a difference is reflected, for example, by the only modest levels of agreement between the child's and other informants' ratings of QoL [13, 15], particularly with regard to non-observable aspects (such as emotional or social functioning). Parent ratings may, however, provide an important alternative perspective. Therefore, in studies on ADHD, measures of QoL as well as measures of psychopathology should include both a child- and a parent-rated version.

The aim of this analysis was to have a detailed look at the relationships between dimensional psychopathology and both parent- and child-rated QoL in ADHD. This analysis also evaluated the validity of the QoL concept, as described by Coghill et al. [5]. By studying correlative considerations, the relationship between QoL and psychopathological problems may be better understood. The effects of medication on QoL in ADHD are reported in a companion paper [8].

# Methods

Study design

The open-label, prospective, multicentre, observational, post-marketing OBSEER (OBservation of Safety and



## Patients and treatment

Patients with ADHD (diagnosed according to the Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition, Text Revision [DSM]-IV-TR [1] or the International Classification of Diseases [ICD]-10 criteria [26]) aged 6–17 years and attending school were included if treatment with Equasym XL® was planned by the treating physician. Patients either could have received prior medication or were medication naive. The study was conducted in accordance with local regulations and under the therapeutic responsibility of the attending physicians; ethics or institutional review board approval was not required for this study. Written informed consent was obtained from parents.

Assessments

QoL

QoL was assessed using the Kinder Lebensqualitätsfragebogen (KINDL) questionnaire for the assessment of health-related QoL in childhood and adolescence [18]. This is a validated tool comprising 24 items, with six subscores (physical well-being, emotional well-being, self-esteem, family, friends and school). Three versions were used according to age group: KID-KINDL was used for children aged 6–11 years; the self-reported KIDDO-KINDL for adolescents aged 12–17 years; and KINDL for parents of patients aged 6–17 years. Scores were transformed so that the range of possible values for the subscores and the total score was from 0 (most negative state) to 100 (most positive state).

SDQ

Parents were asked to complete a version of the Strengths and Difficulties Questionnaire (SDQ) that supplements the 25 core items on specific strengths and difficulties with an



<sup>&</sup>lt;sup>1</sup> Equasym XL is the UK trade name, and is registered and marketed by Shire in the following countries under the following trademarks: Denmark, Equasym Depot; Finland, Equasym Retard; France, Quasym LP; Germany, Equasym Retard; Ireland, Equasym XL; Netherlands, Equasym XL; Norway, Equasym Depot; Sweden, Equasym Depot; South Korea, Metadate CD; Mexico, Metadate CD. Information correct at August 2011.

overall rating of whether their child has emotional or behavioural problems [11, 22]. Each of the 25 items is rated as being not true (0), somewhat true (1) or certainly true (2); each of the five subscales consists of five items, thus yielding scores between 0 and 10 for each subscale. Although the wording chosen for 10 of the 25 SDQ questions addresses positive behavioural attributes, five of these 10 item scores were inverted before being summed up. Thus, four of the SDQ subscales represent problem scores (emotional symptoms, conduct problems, hyperactivity/ inattention and peer problems), which were added together to obtain a total difficulties score ranging from 0 to 40. The fifth subscale assesses the positive aspect of prosocial behaviour. Scores for each subscale were considered to be normal, borderline or abnormal. Total difficulties scores were considered to be normal (range 0-12), borderline (range 13-15) or abnormal (range 16-40).

## Statistical analyses

The following outcomes were assessed: parent-rated dimensional psychopathological severity according to the SDQ and parent- and self-rated QoL at baseline.

Results presented are for the intent-to-treat population. Analysis of variance (ANOVA) modelling was performed to determine the effects of parent-rated SDQ subscale scores and total difficulties score on QoL. In subgroup analyses, patients were assigned to groups according to their diagnosis (ADHD only versus ADHD with conduct disorder) and treatment prior to starting Equasym XL<sup>®</sup> (no treatment; or treatment with modified-release methylphenidate [MPH], immediate-release MPH [MPH-IR] administered once daily, MPH-IR administered several times per day or 'other', for which treatment was unspecified [atomoxetine, amphetamine or insufficiently specified]). Finally, correlation coefficients between the parent-reported SDQ subscales and total difficulties score and the parent- and self-rated QoL scales in the study (KINDL, KID-KINDL and KIDDO-KINDL) were calculated.

Scale means for parent ratings of both the SDQ and the KINDL scales were compared with those from the BELLA study, which assessed the prevalence of general and specific mental health problems in a representative sample of children and adolescents in Germany [19, 21].

# Results

#### Study population

In total, 852 patients were enrolled in the study. Thirty patients were excluded from the analysis due to invalid data; a further 101 patients were excluded because their

SDQ data were missing or incomplete. Therefore, 721 patients were included in the analysis population. Baseline demographics are described elsewhere [8].

Relationship between SDQ scores in the normal range and QoL

Self-ratings of QoL from children aged 6–11 years showed that those with normal (range 0–5) scores on the *hyperactivity/inattention* subscale of the SDQ tended to achieve higher, albeit not significantly, scores on the KID-KINDL *friends* subscale than if they had borderline (6) or abnormal (range 7–10) SDQ scores (mean [SD] 68.4 [19.8] vs. 61.3 [20.7] and 65.5 [22.7]; F(2, 474) = 2.445;  $0.05 < P \le 0.1$ ).

Children with normal (range 0–12) or borderline (range 13–15) SDQ total difficulties scores had higher scores, indicating better QoL, on the KINDL *friends* subscale than children with abnormal (range 16–40) SDQ scores (mean [SD] 63.2 [19.4] and 63.4 [20.8] vs. 58.7 [21.2]; F (2, 718) = 3.817;  $P \le 0.05$ ). Compared with children with borderline or abnormal SDQ total difficulties scores, those with normal scores tended to rate themselves as more content on the KID-KINDL *school* subscale (mean [SD] 61.8 [21.3] vs. 53.6 [23.0] and 58.2 [22.5]; F (2, 474) = 2.589;  $0.05 < P \le 0.1$ ).

Adolescents with normal SDQ total difficulties scores tended to show the highest QoL on the KIDDO-KINDL *family* subscale compared with those with borderline or abnormal scores (mean [SD] 73.5 [20.9] vs. 59.3 [23.6] and 66.6 [22.9]; F (2, 164) = 2.545; 0.05 < P < 0.1).

Correlation between severity of SDQ scores and QoL

Correlation analyses showed that children with high SDQ total difficulties scores had significantly lower QoL on the parent-rated KINDL total score (r = -0.091;  $P \le 0.05$ ) and on the subscales *emotional well-being* (r = -0.106;  $P \le 0.01$ ) and *friends* (r = -0.131;  $P \le 0.01$ ; Table 1). There was also a similar trend for the subscale *self-esteem* (r = -0.063;  $0.05 < P \le 0.1$ ).

Children with high scores on the SDQ *emotional problems* subscale had significantly lower values on the parent-rated KINDL total score (r = -0.097;  $P \le 0.01$ ) and on the subscales *emotional well-being* (r = -0.096;  $P \le 0.05$ ) and *friends* (r = -0.118;  $P \le 0.01$ ). A similar trend was found for the subscale *school* (r = -0.073;  $0.05 < P \le 0.1$ ).

For the SDQ subscale *conduct problems*, children with higher scores tended to have lower QoL based on the parent-rated KINDL scale *emotional well-being* (r = -0.069;  $0.05 < P \le 0.1$ ), although this was not significant.

There were no significant correlations between the SDQ subscale *hyperactivity/inattention* and any of the parent-rated KINDL subscales.



**SDO** KINDL Total Subscales difficulties Emotional Conduct Hyperactivity/ Peer Prosocial score problems problems inattention problems behaviour  $-0.067^{T}$ Total score -0.091\*-0.097\*\*-0.055 ns -0.019 ns 0.017 ns Subscales Physical well-being -0.038 ns 0.005 ns0.031 ns 0.00 ns -0.008 ns -0.003 ns -0.106\*\*-0.096\* $-0.069^{T}$ -0.084\* $0.063^{T}$ Emotional well-being -0.032 ns Self-esteem  $-0.063^{T}$ -0.057 ns -0.050 ns-0.003 ns -0.055 ns 0.013 ns -0.032 ns -0.012 ns 0.040 ns Family -0.025 ns -0.055 ns 0.009 ns Friends -0.131\*\*-0.118\*\*-0.039 ns -0.053 ns -0.134\*\*0.003 ns $-0.073^{\mathrm{T}}$ -0.043 ns -0.024 ns -0.026 ns 0.009 ns -0.038 ns School

Italics indicate significant or borderline significant results \*  $P \le 0.05$ ; \*\*  $P \le 0.01$ ; The  $0.05 < P \le 0.1$ ; is a not significant (two-tailed test, Spearman's rank correlation)

**Table 2** Correlation between the SDQ (parent ratings) and the KIDDO-KINDL total scores and subscales (patient ratings, age 12-17 years) (N=167)

KIDDO-KINDL	SDQ						
	Total difficulties score	Subscales					
		Emotional problems	Conduct problems	Hyperactivity/ inattention	Peer problems	Prosocial behaviour	
Total score	-0.159*	-0.089 ns	-0.042 ns	-0.174*	-0.119 ns	-0.025 ns	
Subscales							
Physical well-being	-0.056  ns	-0.091  ns	0.014 ns	-0.021 ns	-0.047  ns	-0.104 ns	
Emotional well-being	-0.076  ns	-0.030  ns	0.004 ns	-0.109 ns	-0.069  ns	-0.079 ns	
Self-esteem	-0.170*	$-0.143^{\mathrm{T}}$	-0.054  ns	-0.098 ns	-0.153*	-0.014 ns	
Family	$-0.128^{\mathrm{T}}$	-0.024  ns	-0.120  ns	-0.155*	-0.045  ns	0.100 ns	
Friends	-0.190*	$-0.137^{\mathrm{T}}$	0.021 ns	-0.202**	-0.143  ns	-0.120 ns	
School	-0.032  ns	0.023 ns	0.050 ns	$-0.132^{\mathrm{T}}$	-0.029  ns	0.033 ns	

Italics indicate significant or borderline significant results  $*P \le 0.05$ ;  $**P \le 0.01$ ;  $^T 0.05 < P \le 0.1$ ; ns = not significant (two-tailed test, Spearman's rank correlation)

Children scoring high on the SDQ subscale *peer problems* had significantly lower QoL on the KINDL subscales *emotional well-being* (r = -0.084;  $P \le 0.05$ ) and *friends* (r = -0.134;  $P \le 0.01$ ). A similar trend was observed for the KINDL total score (r = -0.067;  $0.05 < P \le 0.1$ ).

There were no significant correlations between psychopathology and QoL self-ratings from children aged 6–11 years (KID-KINDL; data not shown).

Adolescents who had high SDQ total difficulties scores had a significantly lower QoL on the KIDDO-KINDL subscales self-esteem  $(r=-0.170;\ P\leq 0.05)$  and friends  $(r=-0.190;\ P\leq 0.05)$ , with a trend for lower QoL on the family subscale  $(r=-0.128;\ 0.05 < P\leq 0.1;\ Table\ 2)$ . Children with high scores on the SDQ subscale emotional problems tended to have lower scores on the KIDDO-KINDL subscales self-esteem  $(r=-0.143;\ 0.05 < P\leq 0.1)$  and friends  $(r=-0.137;\ 0.05 < P\leq 0.1)$ .

Adolescents with high scores on the SDQ subscale hyperactivity showed a significantly lower QoL on the KIDDO-KINDL total score  $(r = -0.174; P \le 0.05)$ , family  $(r = -0.155; P \le 0.05)$  and friends  $(r = -0.202; P \le 0.01; Table 2)$ . Furthermore, a similar trend was found

for the KINDL subscale *school* (r = -0.132; 0.05 <  $P \le 0.1$ ). Adolescents with high scores on the SDQ subscale *peer problems* tended to have a lower QoL on the KIDDO-KINDL subscale *self-esteem* (r = -0.153;  $P \le 0.05$ ). There were no significant correlations between the SDQ subscales *conduct problems* or *prosocial behaviour* and any of the KIDDO-KINDL subscales.

Compared with the recent normative reference values for children and adolescents in Germany obtained in the epidemiological BELLA study [19, 21], children and adolescents with ADHD, as seen in the OBSEER study, were significantly more severely affected on every SDQ subscale and had a significantly lower QoL on every KINDL subscale (Table 3).

# Comorbid conduct disorder and QoL

One-way ANOVA was used to test whether there were differences between scores on each scale associated with having an additional conduct disorder (Table 4). Of note, SDQ scores, including those for the subscale *conduct problems*, did not differ between diagnostic groups.



Table 3 Comparison of scale means and effect sizes for parent-rated SDQ and KINDL scores at baseline in the OBSEER study with a reference population from the BELLA study

Parent SDQ ratings	Study	Cohen's d	t-test		
	BELLA [21]; <i>n</i> = 2,406 Mean (SD)	OBSEER; $n = 721$ Mean (SD)		t	Significance
Total difficulties	7.8 (5.2)	18.6 (4.3)	-2.3	45.59	***
Emotional symptoms	1.8 (1.8)	3.8 (2.5)	-0.9	23.20	***
Conduct problems	1.9 (1.5)	4.2 (1.7)	-1.4	31.30	***
Hyperactivity/ inattention	3.0 (2.2)	6.9 (1.6)	-2.0	23.41	***
Peer problems	1.4 (1.6)	3.7 (1.4)	-1.5	50.10	***
Prosocial behaviour	7.9 (1.7)	6.7 (2.1)	0.6	15.54	***
Parent KINDL ratings	Study		Cohen's d	t-test	
	BELLA [19]; <i>n</i> = 2,863 Mean (SD)	OBSEER; $n = 721$ Mean (SD)		t	Significance
Total score	76.3 (10.1)	62.9 (13.3)	1.1	30.51	***
Physical well-being	76.5 (17.3)	71.4 (18.3)	0.3	6.52	***
Emotional well-being	80.8 (12.8)	67.6 (17.6)	0.9	22.51	***
Self-esteem	68.8 (14.2)	55.1 (18.5)	0.8	21.84	***

62.2 (19.4)

60.2 (20.9)

60.9 (18.7)

The Cohen's d effect size was calculated as small  $(0.20 \ge d < 0.50)$ , medium  $(0.50 \ge d < 0.80)$  or large  $(0.80 \ge d)$  [6] SD standard deviation \*\*\* P < 0.001

Similarly, there were no differences in QoL between children with or without additional conduct disorder from KINDL parent ratings.

Family

Friends

School

In contrast, self-ratings from children aged 6–11 years (KID-KINDL) showed that those with an additional conduct disorder had lower QoL than those without (Table 4). This applied to the total score ( $F(1,427)=6.246; P \leq 0.05$ ) and the subscales self-esteem ( $F(1,432)=6.264; P \leq 0.05$ ), family ( $F(1,430)=11.683; P \leq 0.001$ ) and friends ( $F(1,431)=4.230; P \leq 0.05$ ). A similar trend was found for the subscale school ( $F(1,430)=3.015; 0.05 < P \leq 0.1$ ), although this was not significant.

Adolescents aged 12–17 years (KIDDO-KINDL) showed a similar pattern to children for the QoL evaluation. The most impaired were the adolescents with an additional conduct disorder, with an effect on the total score (F (1, 135) = 11.715;  $P \le 0.001$ ) and the scales physical well-being (F (1, 137) = 5.011;  $P \le 0.05$ ), emotional well-being (F (1, 137) = 6.366;  $P \le 0.05$ ), self-esteem (F (1, 137) = 4.639;  $P \le 0.05$ ), family (F (1, 136) = 10.291;  $P \le 0.01$ ) and friends (F (1, 137) = 4.660;  $P \le 0.05$ ).

## Medication subgroups and QoL

When subgroups of patients were compared according to whether they had received prior medication or not, no statistically significant differences were found either for the KINDL or for the KID-KINDL (Table 5). For the KIDDO-

KINDL, adolescents receiving any prior medication scored significantly higher on the KIDDO-KINDL subscale *school* (F (1, 166) = 2.448;  $P \le 0.05$ ) than those who had not received prior medication. A similar trend was found for the subscales self-esteem (F (1, 166) = 2.376; 0.05 <  $P \le 0.1$ ) and family (F (1, 166) = 1.997; 0.05 <  $P \le 0.1$ ). There were no significant differences in SDQ scores according to whether or not the patient had received prior medication.

0.9

1.0

0.9

24.95

24.31

20.10

\*\*\*

## Discussion

77.7 (14.3)

78.0 (13.4)

76.0 (16.0)

ADHD is associated with a variety of behavioural problems; therefore, knowledge of the complete psychopathological profile is important for treatment decisions and follow-up. Danckaerts et al. [7] showed that a robust negative effect on QoL was reported by the parents of children with ADHD across a broad range of psychopathology symptoms. Additional comorbid disorders, such as oppositional defiant disorder [16], and increased symptom levels also seem to predict reduced QoL [14]. For this reason, it is useful to know what influence psychopathological profile has on QoL. We investigated the relationship between these two parameters in this analysis of the observational OBSEER trial. An advantage of this study is that self-assessment of children was included and thus findings are not limited to just the views of parents or caregivers, unlike the majority of QoL studies [7].



**Table 4** Diagnosis and QoL/behavioural problems

Rating scale	N	Diagnosis	ANOVA		
		F 90.0; n = 370 Mean (SD)	F 90.1; n = 268 Mean (SD)	F	Significance
QoL					
Parent KINDL ratings					
Total score	638	63.1 (13.3)	62.6 (13.4)	0.269	ns
Physical well-being	638	72.1 (17.7)	70.5 (19.2)	1.157	ns
Emotional well-being	638	67.6 (17.9)	67.2 (17.0)	0.102	ns
Self-esteem	638	55.6 (18.4)	54.8 (18.6)	0.229	ns
Family	638	63.4 (18.7)	60.8 (20.5)	2.677	ns
Friends	638	59.5 (21.4)	61.1 (20.6)	0.915	ns
School	684	60.7 (18.8)	61.0 (18.7)	0.054	ns
Patient KID-KINDL ratings	(age 6–1	1 years)			
Total score	429	66.6 (12.6)	63.2 (15.0)	6.246	*
Physical well-being	429	72.2 (17.0)	73.4 (17.9)	0.754	ns
Emotional well-being	429	71.4 (16.5)	70.6 (17.8)	0.159	ns
Self-esteem	429	59.7 (19.2)	54.2 (22.6)	6.264	*
Family	429	70.1 (18.6)	63.5 (22.7)	11.683	***
Friends	429	66.6 (20.6)	62.1 (23.7)	4.230	*
School	429	59.6 (21.8)	55.7 (23.5)	3.015	T
Patient KIDDO-KINDL ratio	ngs (age	12-17 years)			
Total score	137	69.7 (10.7)	62.4 (14.1)	11.715	***
Physical well-being	137	76.3 (15.6)	70.5 (16.3)	5.011	*
Emotional well-being	137	76.8 (15.5)	69.3 (16.8)	6.366	*
Self-esteem	137	62.0 (17.9)	55.9 (22.6)	4.639	*
Family	137	71.7 (20.2)	58.8 (26.1)	10.291	**
Friends	137	72.5 (17.2)	65.6 (22.4)	4.660	*
School	137	58.7 (16.9)	54.2 (16.4)	2.046	ns
Behavioural problems					
Parent SDQ ratings					
Total difficulties	638	18.3 (6.3)	18.9 (6.3)	0.857	ns
Emotional symptoms	638	3.8 (2.3)	3.8 (2.6)	0.365	ns
Conduct problems	638	4.0 (2.2)	4.3 (2.3)	1.044	ns
Hyperactivity/inattention	638	7.0 (2.2)	7.0 (2.2)	0.183	ns
Peer problems	638	3.6 (2.5)	3.8 (2.4)	1.163	ns
Prosocial behaviour	638	6.8 (2.0)	6.6 (2.1)	2.501	ns

ANOVA analysis of variance, F90.0 disturbance of activity and attention, F90.1 hyperkinetic conduct disorder, QoL quality of life, SD standard deviation

Our results show that classifying scores on the SDQ subscale *hyperactivity/inattention* and the SDQ total problem score as normal, borderline or abnormal did not sufficiently differentiate between children's QoL as rated by their parents. This may be due to the fact that, in this fairly homogeneous clinical sample, most children with ADHD had low QoL. Indeed, this was reflected in both parent and self-ratings. Thus, it is probable that an apparent 'bottom effect' did not allow QoL to be adequately differentiated using severity-categorized SDQ values in this sample.

Although ADHD symptom severity appeared to have little influence on QoL, having a comorbid conduct disorder had a substantial impact. However, this was evident

only on self-ratings; ratings from parents did not differ between QoL of children with ADHD only and those with ADHD plus a conduct disorder. The findings indicate that adolescents, in particular, experience a significant decrease in QoL if they have a comorbid conduct disorder. Therapists should consider this aspect when planning multimodal treatment and adjust the sequence of interventions according to the subjective view of impairment given by the patient.

Parent and child ratings did not identify any significant differences in QoL due to prior medication. However, adolescents not taking medication at baseline had significantly lower QoL on the KIDDO-KINDL subscale *school* compared with adolescents taking any MPH medication. A



<sup>\*</sup>  $P \le 0.05$ ; \*\*  $P \le 0.01$ ; \*\*\*  $P \le 0.001$ ;

T  $0.05 < P \le 0.1$ ; ns = not significant

Table 5 Medication subgroups and QoL/behavioural problems

Rating scale	N	Prior treatment					ANOVA	
		None $n = 187$ Mean (SD)	MPH-IR once-daily $n = 82$ Mean (SD)	MPH-IR repeated $n = 239$ Mean (SD)	MPH-MR $n = 179$ Mean (SD)	Other/no specification $n = 34$ Mean (SD)	F	Significance
QoL								
Parent KINDL ratings								
Total score	721	63.4 (13.3)	61.0 (12.9)	62.8 (13.9)	63.4 (12.5)	63.3 (12.0)	0.577	ns
Physical well-being	721	71.7 (18.5)	70.1 (18.6)	72.3 (18.3)	70.4 (18.8)	75.0 (13.9)	0.723	ns
Emotional well-being	721	68.7 (15.9)	65.0 (17.8)	66.2 (18.9)	69.7 (17.5)	67.5 (15.2)	1.584	ns
Self-esteem	721	54.9 (19.1)	53.2 (20.7)	55.6 (18.5)	55.6 (16.8)	54.4 (18.4)	0.303	ns
Family	721	63.0 (19.0)	58.8 (19.8)	62.8 (20.1)	62.4 (18.8)	61.8 (19.7)	0.784	ns
Friends	721	61.3 (21.3)	59.8 (20.0)	59.3 (20.8)	61.4 (21.2)	55.7 (20.8)	0.770	ns
School	721	61.5 (19.5)	58.9 (17.7)	60.5 (19.4)	61.0 (17.7)	65.6 (17.2)	0.852	ns
Patient KID-KINDL ratings	(age 6-	11 years)						
Total score	477	63.8 (13.3)	65.8 (14.3)	67.2 (14.1)	65.4 (12.6)	62.9 (16.1)	1.349	ns
Physical well-being	477	74.4 (18.3)	70.4 (15.3)	71.9 (18.0)	72.8 (16.9)	74.0 (14.3)	0.682	ns
Emotional well-being	477	70.3 (16.7)	69.3 (17.3)	73.2 (17.8)	70.7 (16.5)	69.1 (17.7)	0.951	ns
Self-esteem	477	73.7 (21.4)	61.5 (19.4)	59.6 (21.5)	56.6 (18.7)	58.0 (23.3)	2.171	T
Family	477	66.0 (19.5)	69.8 (20.8)	70.1 (20.4)	67.4 (20.1)	59.0 (23.9)	1.806	ns
Friends	477	64.4 (20.2)	64.6 (21.6)	67.0 (23.2)	66.5 (20.6)	60.8 (25.9)	0.576	ns
School	477	53.9 (23.1)	59.2 (23.4)	61.3 (21.7)	58.1 (20.2)	56.6 (29.0)	2.109	T
Patient KIDDO-KINDL ration	ngs (age	12-17 years)						
Total score	167	63.3 (14.3)	66.0 (10.3)	69.0 (11.3)	67.3 (14.0)	66.2 (7.9)	1.250	ns
Physical well-being	167	71.7 (18.1)	76.2 (17.1)	75.1 (18.0)	73.0 (14.4)	73.3 (17.5)	0.340	ns
Emotional well-being	167	75.6 (18.1)	68.8 (11.4)	73.9 (15.4)	73.8 (17.5)	71.6 (10.2)	0.571	ns
Self-esteem	167	51.7 (20.5)	60.6 (15.4)	61.5 (16.3)	62.6 (19.4)	61.9 (21.0)	2.376	T
Family	167	61.6 (27.3)	63.5 (18.7)	72.9 (21.9)	68.1 (21.5)	58.5 (13.2)	1.997	Т
Friends	167	69.1 (20.7)	66.5 (15.6)	70.7 (18.5)	68.8 (20.8)	70.5 (16.3)	0.165	ns
School	167	50.1 (17.7)	60.2 (15.3)	59.7 (16.6)	57.6 (17.5)	61.4 (14.2)	2.448	*
Behavioural problems								
Parent SDQ ratings								
Total difficulties	721	19.0 (6.5)	17.4 (5.4)	18.6 (6.7)	18.7 (6.0)	18.2 (5.6)	0.993	ns
Emotional symptoms	721	3.9 (2.4)	3.6 (2.4)	3.9 (2.6)	3.7 (2.4)	3.5 (2.6)	0.498	ns
Conduct problems	721	4.4 (2.4)	3.7 (2.2)	4.1 (2.4)	4.2 (2.1)	4.4 (1.8)	1.494	ns
Hyperactivity/inattention	721	7.0 (2.2)	6.8 (1.9)	6.7 (2.1)	7.3 (2.3)	6.9 (2.4)	1.778	ns
Peer problems	721	3.8 (2.5)	3.3 (2.2)	3.9 (2.6)	3.5 (2.3)	3.4 (2.4)	1.270	ns
Prosocial behaviour	721	6.6 (2.0)	6.6 (2.0)	6.6 (2.1)	6.8 (2.1)	7.1 (1.9)	0.777	ns

*ANOVA* analysis of variance, *IR* immediate release, *MPH* methylphenidate, *MR* modified release, *QoL* quality of life, *SD* standard deviation  $*P \le 0.05$ ;  $^{\rm T}0.05 < P \le 0.1$ ; ns = not significant

similar trend was found for the subscales *self-esteem* and *family*, but the majority of scales showed no differences. Hence, the positive effect of medication on QoL seen in some previous trials [4, 9] was observed only as an overall trend in this study.

Scores for the SDQ subscales and total difficulties had varying associations with the QoL assessment scales used in the OBSEER study. For parental ratings, the highest correlation (negative) was seen between the SDQ subscale peer problems and the KINDL subscale friends. Thus, children with lower QoL according to the subscale friends had significantly more problems with their peers. It is likely that these scales measure similar, but opposing, aspects of behaviour. Furthermore, it was found that a high psychopathological value on the SDQ subscales peer problems, emotional problems and total problems is accompanied by lower QoL on the KINDL subscales emotional well-being, friends and total score. However,



these significant relationships are based on low correlation coefficients and thus need to be interpreted with caution. In addition, adolescents with high scores on the SDQ scale *hyperactivity* had a significantly lower QoL (KIDDO-KINDL). This association was most notable for the total score and the subscales *family* and *friends*.

The results of this study are consistent with the results of another observational trial, the ADORE study [2], which recruited a comparable sample (with regard to sample size and demographic characteristics) of children with ADHD across Europe. In that study, baseline SDQ values were similar to those in the current study, and the QoL of children with ADHD in the ADORE study was markedly lower at baseline [20] than the norms for children in the community [3].

However, according to SDQ scales, a relatively high proportion of children and adolescents in the OBSEER study were classified as having impaired psychopathology. A total of 67.1% of children were classified as 'abnormal' on the SDQ total score and 57.1% on the SDQ subscale *hyperactivity/inattention*, compared with 6–9% of children in the normative BELLA study [21].

A limitation of our analysis is that the clinical diagnoses were not based on structured interviews, which may limit the reliability of the findings. Furthermore, for the ANOVA, the multiple testing of correlations was not adjusted and therefore should be regarded as exploratory. Finally, the prior medication status of patients was not identified at randomization, which makes it difficult to interpret the presence or absence of meaningful differences.

In addition to providing further evidence that QoL is low in children with ADHD, our results show that different psychopathological problems influence QoL in different ways. According to child and adolescent self-ratings, having a comorbid conduct disorder was associated with significantly lower QoL. Based on our results, it is recommended that when assessing QoL, children's ratings should be used in conjunction with parental ratings for a complete picture.

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