Six months methylphenidate treatment improves emotion dysregulation in adolescents with attention deficit/hyperactivity disorder: a prospective study

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\textbf{Purpose:} Individuals with attention deficit/hyperactivity disorder (ADHD) may suffer from emotional dysregulation (ED), although this symptom is not listed among the diagnostic criteria. Methylphenidate (MPH) is useful in reducing emotional symptoms in ADHD. The aim of the present study was to determine both psychosocial risk factors and presence of ED in adolescents with ADHD before and after MPH treatment.

\textbf{Participants and methods:} Eighty-two patients aged 12–18 years with ADHD were included as participants. The Kiddie Schedule for Affective Disorders and Schizophrenia for School-Age Children – Present and Lifetime, the Difficulties in Emotion Regulation Scale (DERS), sociodemographic form, and the Inventory of Statements About Self-Injury were administered. Results were compared before and after 6 months MPH treatment.

\textbf{Results:} A significant improvement was detected on DERS for impulsivity (15.9±6.8 initial vs 14.2±6.5 final test, \textit{p}<0.01) and total score (88.4±23.3 initial vs 82.4±22.7 final test, \textit{p}<0.05) across all patients taking MPH regardless of subtype and sex. Despite treatment, a significant difference remained for impulsivity, strategies, and total score in patients with comorbid oppositional defiant disorder (ODD) compared with those without ODD, but no difference was detected for conduct disorder comorbidity. In patients who self-harm, scores for goals, impulsivity, strategies, clarity, and total score were higher before treatment: furthermore, impulsivity and total score remained high after treatment. In maltreated patients, goals, impulsivity, strategies, and total scores were significantly higher before treatment; however, their symptoms were ameliorated after treatment with MPH.

\textbf{Conclusion:} Individuals with severe ED may “self-medicate” by smoking and/or self-harming. MPH led to significant improvements in ED possibly owing, in part, to a decrease in impulsivity, so that individuals felt more able to supervise their emotions and engage in goal-directed behaviors. ED should be considered particularly in patients with additional psychosocial factors and ODD comorbidity, and included in the treatment plan.

\textbf{Keywords:} attention deficit, emotion, dysregulation, methylphenidate

\section*{Introduction}

Attention deficit/hyperactivity disorder (ADHD) is characterized by inattention, hyperactivity, and/or impulsivity inappropriate for the individual’s age and the developmental stage. It is among the most common neuropsychiatric disorders worldwide.\textsuperscript{1,2}

Untreated ADHD increases the risk of other psychiatric conditions and adverse life experiences such as smoking, substance misuse, suicidality, and/or non-suicidal self-harm. ADHD is one of the most comorbid conditions to other psychiatric disorders, with major depressive and anxiety disorders being the most common.\textsuperscript{3,4} A recent meta-analysis revealed a strong association between ADHD and the lifetime prevalence of suicide attempt, with a relative risk of 2.4.\textsuperscript{5}

In the current study, we aimed to assess the presence of ED in adolescents with ADHD before and after MPH treatment. We also aimed to determine both psychosocial risk factors and presence of ED in adolescents with ADHD.
ADHD has a significant adverse effect on an individual’s psychosocial functioning and quality of life, leading to more emotional and behavioral problems, lower self-esteem, and poorer outcomes in many domains. Executive function (EF) deficits play a major part in ADHD, affecting cognitive processes such as planning, problem-solving, and working memory. Problems with emotion regulation (ER) and/or modulating emotions also occur in individuals with EF deficits. ER is defined as the internal and external processes of observation, evaluation, and regulation of emotional reactions during goal-directed behaviors.

Impairment in ER is referred to as emotional dysregulation (ED), and is gradually gaining recognition as a core deficit in patients with ADHD. Emotional inhibition, understanding of emotions, and empathy have been found to be lessened in individuals with ADHD. These individuals may have difficulties in understanding social cues and regulating affect appropriately and may have frequent emotional swings, with an adverse effect on their social functioning. A recent meta-analysis showed that individuals with ADHD mostly had emotional reactivity, negativity, and lability.

Individuals with ADHD may also have difficulties in expressing appropriate emotional behaviors and may display negative emotionality. Regulating irritable and aggressive behaviors and/or emotions, in particular, are much more problematic in patients with ADHD. Moreover, individuals with persistent ADHD symptoms have higher rates of emotional lability, which is associated with poorer outcomes in many domains, adding support to the concept of emotional lability as another distinct dimension in ADHD.

As ED is not specific to ADHD, it may be related to or may serve as a suitable marker for comorbid conditions that frequently present with ADHD. One study found that a deficit in ER was related to depressive symptoms later in life in patients with childhood ADHD. In addition, high rates of comorbid oppositional defiant disorder (ODD) may help to explain emotional symptoms in children with ADHD. It has been suggested that ODD should be assessed as a disorder of ER rather than a disruptive behavior disorder. Patients with non-episodic irritability and persistent negative mood may be diagnosed with disruptive mood dysregulation disorder (DMDD), which is a new diagnosis introduced in DSM-5. ED and recurrent temper outbursts are frequent in these individuals. Nearly 75% of patients with DMDD have comorbid ADHD. Borderline personality disorder is another psychopathology with intense ED, and symptoms may overlap with ADHD. Therefore, it should be kept in mind that ED may be linked to or raise the risk of comorbid conditions in ADHD.

Adults with ADHD exhibit similar deficits to children with the disorder. Items related to ED are well-defined in the Wender Utah Rating Scale for ADHD. Moreover, treatment with methylphenidate (MPH) has shown to be effective for core symptoms and additional emotional symptoms in adult patients with ADHD.

In a minority of children with ADHD, stimulants may be associated with behavioral changes as side effects. In some patients, symptoms such as irritability, agitation, anxiety, and emotional lability may emerge with MPH usage. Moreover, clinicians should be aware of “rebound irritability”, which usually occurs alongside declining blood levels of MPH. However, based on the evidence, the association between MPH and changes in behavior is unclear. Further studies are needed to elucidate this association.

Atomoxetine (ATX), another agent approved for treating ADHD, has been found to be useful, with moderate effect size, for reducing emotional lability in adults with ADHD. Moreover, ATX has a similar effect on ED as on other ADHD symptoms in adults. However, data on ED and the effects of ATX in children and adolescents with ADHD are insufficient, and further studies are required.

Other therapeutic approaches, such as psychosocial interventions, can also be used in the treatment of ED. In a study, cognitive behavioral therapy-based group therapy and parent training interventions were shown to be useful in reducing mood lability in children with ADHD and DMDD. Dialectical behavioral therapy is another therapeutic intervention that has been shown to be effective for ER in college students and adults with ADHD. In addition, mindfulness-based training was found to significantly reduce self-reported ED in adults with ADHD. Although data regarding children and adolescents are limited, these interventions appear to be useful in reducing emotional symptoms in patients with ADHD. Therefore, they may be evaluated as an adjunct treatment model for ADHD.

The aim of this study was to determine the sociodemographic and clinical features and psychosocial risk factors of patients with untreated ADHD, thereby reflecting the early and problematic course of the disorder. An additional aim was to investigate difficulties with ER – especially in patients with additional adverse psychosocial risk factors – and measure the difference after MPH treatment for 6 months.
Emotional lability and/or dysregulation in ADHD have been well-studied, generally using parental reports. To the best of our knowledge, there have been no follow-up studies regarding the effect of MPH on ED in adolescents with ADHD using a self-report scale, although this has been well-studied in adult patients.  

**Participants and methods**

**Participants**

Ethical approval for the study was provided by Cukurova University School of Medicine Non-Invasive Clinical Research Ethics Committee (8 May, 2015; reference number: 42/11). Written and informed consent was obtained from the parents and the participants who participated voluntarily in the study.

Eighty-two patients who presented to Cukurova University School of Medicine Child and Adolescent Psychiatry Department were included in the study. Patients were included if they: 1) were aged between 12 and 18 years, 2) were diagnosed as having ADHD based on the Turkish Version of the Kiddie Schedule for Affective Disorders and Schizophrenia for School-Age Children – Present and Lifetime (K-SADS-PL), 3) were treatment naïve, and 4) had no organic pathology. Mental Retardation and Autism Spectrum Disorders were used as exclusion criteria.

Routine tests including measurement of biochemical parameters and electrocardiogram were performed before starting the treatment. Participants were examined at baseline, and 1, 3, and 6 months of treatment. After the participants were examined, MPH treatment was started according to guidelines. The scales and tests described in the following section were administered both before starting the treatment and at the 6-month examination.

**Measures**

Initially, sociodemographic information was obtained including the participant’s age, birth history, developmental stages, medical history, academic history, peer relationships, familial characteristics, and any adverse events. The form was completed by the physician, using responses from the parents. Participants were interviewed by a child and adolescent psychiatrist using the K-SADS-PL, and psychiatric diagnoses were made in accordance with the text of the DSM-IV-TR.

**K-SADS-PL**

K-SADS-PL is a semi-structured interview technique performed by physicians, which detects former and existing Axis I psychopathologies according to the criteria of the DSM-III and DSM-IV-TR. First, questions are asked about fundamental symptoms, and the answers are scored between 0 and 3 based on severity and/or frequency. Psychopathologies associated with higher scoring symptoms are reassessed, evaluated, and diagnosed. K-SADS-PL has been shown to be a reliable and valid method for diagnosing psychopathologies in children and adolescents. The validity and reliability of the Turkish version were established by Gökl er et al.

**Difficulties in Emotion Regulation Scale (DERS)**

DERS is a brief, self-report questionnaire designed to assess ED, developed by Grat z and Roemer in 2004. It consists of 36 items, scored on a scale ranging from 1 (almost never) to 5 (almost always); higher scores reflect greater problems with ER. DERS has six subscales and a total score. Although DERS was composed for adults, research has shown that it is also an appropriate tool for measuring ED in adolescents. The validity and reliability of the Turkish version were established by Sarıtaş et al.

**Inventory of Statements About Self-Injury (ISAS)**

ISAS is a self-report scale developed by Klonsky and Glenn in 2009. It consists of two parts: the frequency and accompanying features of NSSI behaviors are noted in the first part, and the function of self-harm is considered in the second. Only the first part was completed by the participants in the current study. The validity and reliability of the Turkish version were tested by Bildik et al.

**Statistical analysis**

The Statistical Package for the Social Sciences (SPSS), version 16.00, was used for statistical analyses (SPSS Inc.; Chicago, IL, USA). Categorical variables were analyzed using the chi-square test. As DERS is composed of ordinal data, nonparametric tests were used: the Mann–Whitney U-test for independent variables, and the Wilcoxon signed-rank test for dependent variables. In all analyses, 0.05 was accepted as the level of statistical significance.

**Results**

The study comprised 82 patients, including 39 (47.6%) girls and 43 (52.4%) boys. The mean age of the participants was 14.9±1.9 years. No significant differences were observed between girls and boys in terms of sociodemographic features such as parental education, occupational status, family structure, history of domestic violence, academic performance, and peer relationships.
All patients were treated with MPH. Psychometric measures were used both before and after treatment. A comparison of DERS scores for all patients before and after 6 months of treatment is given in Table 1.

Sixty-seven patients (81.7%) were diagnosed as having ADHD-combined subtype (ADHD-C), while 15 (18.3%) had the ADHD-inattention subtype (ADHD-I). The psychometric comparison of DERS among subtypes is shown in Table 2. There were no significant differences in initial and final test scores between patients with ADHD-C and those with ADHD-I subtypes.

Sixteen patients (19.5%) were diagnosed as having comorbid ODD, and 12 (14.6%) had conduct disorder (CD). For patients with comorbid CD, initial total test scores were significantly higher than those of patients without comorbid CD ($p<0.05$). For patients with comorbid CD, there was no significant difference between the initial and final test scores. However, there was a significant decrease in scores in patients without comorbid CD ($p<0.01$). Psychometric measures (DERS scores) of patients with ODD are presented in Table 3.

Twenty-three participants (28%) reported that they smoked regularly. Alcohol (n=9, 39.1%) and/or substance use (n=4, 100%) were significantly higher in smokers than in non-smokers ($p<0.05$). All smokers were diagnosed as having ADHD-C, and 16 of them (69.6%) had symptom onset before the age of 7 years. A comparison of psychometric properties in smokers and non-smokers is presented in Table 4.

Forty-eight (58.5%) patients reported at least one NSSI on ISAS, 30 (62.5%) of whom were females. The average initial NSSI age was 11.4±3.4 years with wound picking, scraping letters/shapes in the skin, and tearing hair out being the most common behaviors. Twenty-one (53.8%) were alone when self-harming, and four (8.3%) were under the effect of drugs. Twenty-six (60.5%) committed NSSI after a stressful event, and 25 (83.3%) harmed themselves within an hour of the impulse. Eight (16.3%) reported that there was somebody with NSSI within their immediate family or among their close friends. The two ADHD subtypes did not differ with respect to presence of NSSI, except that pinpricking was more frequent in ADHD-C ($p<0.05$). DERS measures of patients with and without NSSI are given in Table 5.

Twenty-three (28%) participants had a history of maltreatment. Among children with maltreatment exposure, ADHD-C subtype diagnoses were significantly more common than ADHD-I subtype diagnoses ($p<0.05$). Comorbidities were not found to be associated with maltreatment ($p>0.05$). A comparison of psychometric properties in participants reporting maltreatment is shown in Table 6.

### Discussion

ADHD may result in adverse consequences in many domains if undiagnosed or untreated. In this study, the participants were selected from patients who were treatment naive and aged older than 12 years, in order to describe the sociodemographic and clinical features of untreated ADHD. We also aimed to investigate the difficulties in ER that these patients experienced and reveal changes with MPH treatment using a self-report scale for adolescents.
### Table 5 Comparison of psychometric measures before and after treatment in patients with NSSI (mean ± SD)

<table>
<thead>
<tr>
<th>DERS</th>
<th>History of NSSI</th>
<th>Without history of NSSI</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Before treatment</td>
<td>After treatment</td>
</tr>
<tr>
<td>Non-acceptance</td>
<td>11.6±5.2</td>
<td>10.4±3.9</td>
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<tr>
<td>Goals</td>
<td>17.4±5.2</td>
<td>15.3±6.1</td>
</tr>
<tr>
<td>Impulsivity</td>
<td>18.2±5.8</td>
<td>16.2±6.4</td>
</tr>
<tr>
<td>Awareness</td>
<td>15.7±5.4</td>
<td>15.0±5.4</td>
</tr>
<tr>
<td>Strategies</td>
<td>20.6±8.2</td>
<td>18.1±8.3</td>
</tr>
<tr>
<td>Clarity</td>
<td>12.6±3.4</td>
<td>12.0±4.1</td>
</tr>
<tr>
<td>Total</td>
<td>96.3±20.7</td>
<td>87.7±24.5</td>
</tr>
</tbody>
</table>

**Notes:** Significantly different in comparison of initial and final test scores of the same group (*p<0.05, **p<0.01, ***p<0.001).** Significantly different in comparison of initial test scores of patients with or without a history of NSSI (§§§p<0.001). Significantly different in comparison of final test scores of patients with or without a history of NSSI (¶¶¶p<0.001).

**Abbreviations:** NSSI, non-suicidal self-injury; DERS, Difficulties in Emotion Regulation.

### Table 6 Comparison of the psychometric measures of patients with and without a history of maltreatment (mean ± SD)

<table>
<thead>
<tr>
<th>DERS</th>
<th>History of maltreatment</th>
<th>Without history of maltreatment</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Before treatment</td>
<td>After treatment</td>
</tr>
<tr>
<td>Non-acceptance</td>
<td>12.4±6.2</td>
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<td>Goals</td>
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<td>16.8±6.8*</td>
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<tr>
<td>Strategies</td>
<td>23.2±17.6</td>
<td>17.5±7.2*</td>
</tr>
<tr>
<td>Clarity</td>
<td>13.2±4.4</td>
<td>11.8±4.0</td>
</tr>
<tr>
<td>Total</td>
<td>104.2±20.1</td>
<td>86.5±23.5*</td>
</tr>
</tbody>
</table>

**Notes:** Significantly different in comparison of initial and final test scores of the same group (*p<0.05, **p<0.01, ***p<0.001).** Significantly different in comparison of initial test scores of patients with or without a history of maltreatment (¶&¶p<0.05, ¶¶&¶¶p<0.001). Significantly different in comparison of final test scores of patients with or without a history of maltreatment. (§p<0.05)

**Abbreviation:** DERS, Difficulties in Emotion Regulation.
Consistent with the literature, we found that MPH had a significant effect on ED, particularly in impulsivity and total scores. Symptoms such as poor inhibitory control, inattention, mood regulation, and working memory are thought to be associated with impaired prefrontal cortical “top-down” mechanisms. The prefrontal cortex and striatal regions can be targeted by stimulants that increase central dopamine and norepinephrine levels. Research has shown that medications that optimize catecholamine levels in these regions have positive effects on attention, behavior, and ER. In clinical presentation, this improvement may be partly connected to a decrease in impulsivity, so that individuals feel more successful in supervising their emotions and goal-directed behaviors, and are able to act in a more controlled way in general. Additionally, individuals with ADHD may experience improvements in terms of understanding, acceptance, monitoring negative emotions, planning adequate strategies, exhibiting appropriate behaviors, and controlling impulsivity as a result of the treatment process.

In our study, no significant associations were found between DERS scores and ADHD subtype. ED has been previously reported to be associated with ADHD, regardless of subtype, and hyperactive and/or impulsive behaviors decrease in individuals with ADHD during adolescence. Therefore, it has been postulated that clinical presentation of ED may be less similar to other domains in adolescents with both subtypes.

A significant association was determined only in impulsivity score before treatment, although there were associations of clarity, strategies, impulsivity, and total DERS scores with comorbid ODD during treatment. In the literature, it has been shown that ED and ODD are closely related, with ED having an impact on both social impairment and comorbid ODD. The difference that we detected in individuals with and without ODD comorbidity both before and after treatment, may support an association between ED and ODD and point to treatment resistance or permanence. Additional interventions may be helpful for these patients. On the other hand, no significant association was found between CD and ED.

Among our participants, the rate of smoking was 28%, consistent with the fact that individuals with ADHD are at heightened risk for smoking. In addition, it is widely accepted that both inattention and hyperactivity/impulsivity raise the risk of an individual trying tobacco and switching to regular smoking. DERS scores were significantly higher in smokers compared with non-smokers, consistent with the finding that adolescents with more severe ED may use smoking for mood regulation or stabilization. Smoking may serve as a way of coping with difficulties and/or be a means of self-medication. It has been reported in the literature that approximately one third of patients with ADHD use cigarettes as self-medication. During treatment, ED tended to decrease in smokers, although a statistical difference remained.

Notably, more than half of the participants (58.5%) in our study reported NSSI at least once. In the literature, ADHD is accepted as a risk factor for NSSI, suicidal thoughts, and suicide attempts. Hyperactivity symptoms in childhood are associated with self-damaging behaviors such as self-cutting and self-hitting in adolescence, and a higher number of ADHD symptoms is closely associated with earlier onset of self-harm and suicide attempts. The average age of the first NSSI was reported as 11 years in our study. Given that we accepted patients aged over 12 years, this may support the idea that NSSI may be one of the adverse consequences of untreated ADHD. In our study, NSSI was detected more frequently in girls than in boys, which is in accordance with the literature.

We did not find any significant association between NSSI and ADHD subtype. It has been previously reported that NSSI tends to occur in the ADHD-C subtype rather than ADHD-I, as NSSI is an impulsive behavior involving an urgent drive accompanied by poor inhibitory control. The majority of our patients who self-harmed (83.3%) did so immediately after experiencing the impulse to do so which supports the impulsive dimension. However, it has also been shown that having ADHD increases the risk of NSSI regardless of subtype, with ADHD-C having a greater risk compared with ADHD-I. Individuals with ADHD may perform NSSI owing to making frequent mistakes or, feeling unsuccessful and/or inefficient, hence NSSI can be a kind of help-seeking behavior. These findings support the idea that NSSI should be considered as both an impulsive behavior and a cognitive process.

In patients with NSSI, goals, impulsivity, strategies, clarity, and total scores in DERS were significantly higher compared with patients without NSSI. Adolescents who lack emotional clarity and ER strategies in general are prone to NSSI. It has been suggested that individuals with NSSI have difficulties in understanding, regulating, and expressing their emotions. This may cause them to respond in an exaggerated or sentimental fashion and have problems in tolerating negative emotions, potentially resulting in NSSI as a way of coping or as self-punishment. These findings are compatible with our study; participants with NSSI had difficulties in coping with daily events or negative experiences.
Problematic ER can be considered to be a predictive factor in addition to ADHD for future NSSI. On the other hand, the DERS impulsivity scores of individuals with and without NSSI did not differ during treatment. This supports the idea that treatment with MPH has a positive effect on coping with negative emotions and impulse control.

In our study, patients with a history of maltreatment were more severely affected by ER, particularly before treatment. Neglect and/or abuse may be associated with difficulties in recognizing, understanding, and expressing one’s emotional state; such difficulties could be included in the ED concept. Therefore, children who are exposed to inappropriate emotional interactions may suffer from problematic interpersonal relationships and social functioning. Individuals who are exposed to trauma have been shown to be at risk of further psychiatric disorders such as affective and personality disorders. Consequently, maltreatment appears to be an independent risk factor for ED and should be asked about. On the other hand, the difference before treatment largely diminished after MPH use in maltreated participants. This may be compatible with the fact that maltreatment exposure is an independent risk factor for ADHD, with a negative impact on the severity of symptoms. However, as demonstrated by our study, this difference may disappear during the treatment process.

Conclusion
In conclusion, as has been reported in follow-up studies, ADHD in childhood is associated with poor psychosocial functioning and additional psychiatric comorbidities. ED is among the main features of ADHD regardless of subtype and has a negative impact on individuals’ daily functioning, although it is not specific to ADHD. ED frequently occurs in patients with additional risk factors such as comorbid psychiatric conditions (ODD in particular), smoking, and a history of NSSI. Therefore, every individual with ADHD should be asked about ED and related problems such as irritability, temper outbursts, and mood swings. MPH is useful in reducing emotional symptoms in ADHD. The association between ED and ADHD determined here can be used to inform future studies with larger sample sizes.

Disclosure
The authors report no conflicts of interest in this work.

References


