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Lower Cardiac Vagal Activity Predicts Self-Reported Difficulties With Emotion Regulation in Adolescents With ADHD

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Objective: To investigate the relation between cardiac vagal activity (CVA), a measure of autonomic nervous system (ANS) flexibility, and self-reported emotion regulation (ER) difficulties in adolescents with attention-deficit/hyperactivity disorder (ADHD) and controls.

Methods: The sample comprised 11–17-year-old adolescents with ADHD (n=34) and controls (n = 33). Multiple linear regression analyses investigated the relation between CVA, as indexed by high frequency heart rate variability (HF-HRV), and ER difficulties as assessed by the Difficulties in Emotion Regulation Scale (DERS). Supplemental analyses were performed in ADHD and control groups separately. Analyses assessed effects of body mass index (BMI), physical activity levels, and HF peak as a surrogate of respiration on CVA.

Results: Lower CVA was associated with ER difficulties, and specifically with limited access to effective ER strategies. When investigating the relation between CVA and ER in the ADHD and control groups separately, there was a tendency of lower CVA predicting limited access to effective ER strategies in the ADHD group, and not in the control group.

Conclusion: The results suggest that lower CVA, i.e., reduced ANS flexibility, in adolescents with ADHD and controls is associated with self-reported ER difficulties, and specifically with limited access to effective ER strategies. There was a tendency for lower CVA to predict limited ER strategies only in the adolescents with ADHD and not controls.

Keywords: attention-deficit/hyperactivity disorder, heart rate variability, cardiac vagal activity, emotion regulation, autonomic nervous system, difficulties in emotion regulation scale
INTRODUCTION

Individuals with attention-deficit/hyperactivity disorder (ADHD) are characterized by difficulties with self-regulation (1), including emotion regulation [ER; (2)]. ER is the ability to influence which emotions one displays, at what time, and how they are experienced and expressed (3). In children and adolescents with ADHD, it is estimated that 25–45% have difficulties with ER (4). In healthy individuals, such ER difficulties are linked to inflexibility of the autonomic nervous system (ANS), often indexed by lower resting state cardiac vagal activity [CVA; (5–7)]. In ADHD, the link between ER difficulties and lower CVA is not as clear. This might be due to the reliance on parent-reports of ER difficulties in previous studies (8, 9). In the current study, we aimed to investigate the relation between self-reported ER difficulties and level of CVA in adolescents with ADHD.

Emotions—multifaceted processes involving changes in experiential, behavioral, and physiological responses (10)—are not always appropriate and in accordance with environmental demands, emphasizing the importance of ER for adaptive behavior and achievement of long-term goals (3). ER difficulties might be due to “bottom-up” psychological mechanisms such as orienting to emotional, salient stimuli and reward processing, or due to poorer “top-down” regulation (2). This might manifest as atypical allocation of attention to emotionally valenced stimuli, emotional lability (11, 12), and socially unsuitable behavior (2). The experience of emotions are associated with varying degrees of physiological arousal generated by the ANS (13). Flexibility of the ANS increases the capacity to rapidly modulate physiological and emotional states in accordance with situational demands. This flexibility is thought to depend on vagal modulation and to be reflected by CVA, derived from the variance in consecutive heart beat intervals (heart rate variability). CVA has been suggested as a transdiagnostic marker of ER difficulties (14). In healthy adults, self-reported ER difficulties have been associated with lower CVA, and specifically with difficulties with emotional impulse control, emotional clarity (15), and acceptance of negative emotions (16). Further, in a group of adolescents with and without psychiatric disorders (not ADHD), self-reports of difficulties with emotional awareness associated with lower CVA (17). Also of relevance, tendencies of worrying (18), anxiety disorders (19), and clinical depression in children and adolescents (20) are associated with lower CVA.

In ADHD, ER difficulties are shown to have a negative influence on well-being and self-esteem, peer relationships, school performance (21), and social skills (22). Further, such difficulties are linked to general anxiety disorder (23) and major depressive disorder (MDD; (24)), which are comorbid in about 20–30% of children and adolescents with ADHD [(22, 25)]. Only few studies have investigated the association between ER difficulties and CVA in children and adolescents with ADHD, and these have used parent-reports of ER difficulties. One such study found no association between CVA and ER difficulties (8), however another study found a dichotomous index of ER difficulties to be associated with lower CVA (9). Further, Griffiths et al. (26) found correlations of CVA with parent-reported symptoms of anxiety, social difficulties, and oppositional behavior in children and adolescents with ADHD. They concluded that CVA seem to reflect level of ER and adaptive behavior. However, no study has investigated whether CVA is associated with self-reported ER difficulties in children or adolescents with ADHD. As self-reports might capture internal, non-observable aspects of ER, they could provide information that parent reports do not detect. This is supported by a study in children and adolescents with ADHD, where self-reported symptoms of anxiety associated better with observed neurocognitive dysfunctions than parent- and teacher reports (27). Further, children with ADHD and controls have been found to report higher health-related quality of life (HRQoL) than HRQoL assessed by parent reports. This highlights the importance of also investigating self-reports in the study of psychophysiological aspects in children and adolescents with ADHD.

In the current study, we investigated the association between CVA and self-reported ER difficulties in adolescents with ADHD, and without ADHD (i.e. control group). We applied a naturalistic study design by including control participants with mental health disorders that often are comorbid to ADHD, such as anxiety disorders and Tourette Syndrome (TS). We hypothesized that lower CVA would be associated with ER difficulties, and expected this effect to be more prominent in adolescents with ADHD. We further explored the association between CVA and specific aspects of ER: acceptance of emotional responses, goal-directed behavior, and impulse control in relation to emotions, emotional awareness, access to effective ER strategies, and emotional clarity. We hypothesized that ER strategies would be the domain most closely associated with CVA, as CVA has been found to associate with neural activity during reappraisal and response modulation (28). This suggests that individuals displaying low CVA have difficulties with recruiting prefrontal brain areas necessary for modulation of amygdala activity during explicit ER. Further supporting our hypothesis, lower CVA has been associated with less use of constructive coping strategies (29) and interpersonal ER strategies such as support seeking (30).

METHODS

Design

The current study is a cross-section of a follow-up project on ER in children with ADHD [studies from first wave: (31–33)]. The retention rate from the first assessment was 66.2% (ADHD= 60.7%, controls=71.7%); **Supplemental Figure 1**. The study protocol was approved by the Regional Committee for Medical Research Ethics of Western Norway (Study number: 2014/1304).

Participants

The sample comprised 34 adolescents with ADHD and 33 controls between 11 and 17 years of age (**Table 1**). The majority of participants were male (n= 43; 64.2%). Children with suspected ADHD were originally referred from child and...
TABLE 1 | Descriptive statistics for ADHD and control groups.

| Variable                       | ADHD (n = 34) | Controls (n = 33) | t    | χ²  | df |
|-------------------------------|---------------|------------------|------|-----|----|
| **Background information**    |               |                  |      |     |    |
| Gender (n, %)                 |               |                  | 0.008 | 1   |    |
| Male                          | 22 (64.7)     | 21 (63.6)        |      |     |    |
| Female                        | 12 (35.3)     | 12 (36.4)        |      |     |    |
| **Age (years)**               | 14.34 (1.52)  | 14.62 (1.15)     | 0.86 | 66  |    |
| **IQ**                        | 93.66 (10.27) | 106.94 (10.63)   | 5.21*| 66  |    |
| **BMI (kg/m²)**               | 22.62 (5.73)  | 21.11 (2.91)     | -1.37| 49  |    |
| **Physical activity**         | 1.68 (1.18)   | 2.70 (1.18)      | 16.68*| 5  |
| **(times/week)**              |               |                  |      |     |    |
| **HF peak (Hz)**              | -1.41 (0.25)  | -1.36 (2.1)      | 0.85 | 65  |    |
| **Questionnaire scores**      |               |                  |      |     |    |
| DERS TOTAL                    | 84.56 (23.83) | 67.60 (15.36)    | -3.47*| 57  |    |
| NONACCEPTANCE                 | 11.05 (4.07)  | 11.21 (4.42)     | 0.29 | 65  |    |
| GOALS                         | 14.41 (5.57)  | 14.21 (3.29)     | -0.59*| 60  |    |
| IMPULSE                       | 13.82 (4.48)  | 9.33 (4.29)      | -3.35*| 57  |    |
| AWARENESS                     | 19.02 (6.83)  | 18.25 (6.10)     | 0.76 | 66  |    |
| STRATEGIES                    | 15.00 (6.75)  | 12.24 (4.09)     | -2.40*| 55  |    |
| CLARITY                       | 10.79 (3.71)  | 8.33 (2.79)      | -3.06*| 65  |    |
| ADHD-RS                       | 25.81 (10.86) | 25.21 (6.67)     | -9.38*| 55  |    |
| CVA (HF-HRV (n.u.))           | 1.78 (1.12)   | 1.82 (1.11)      | 1.55 | 65  |    |

**ADHD**, Attention Deficit Hyperactivity Disorder; **BMI**, Body Mass Index; **HF peak**, Peak of high frequency heart rate variability; **DERS TOTAL**, Difficulties in Emotion Regulation Scale total score; **NONACCEPTANCE**, Non-acceptance subscale; **GOALS**, Goals subscale; **IMPULSE**, Impulse control subscale; **AWARENESS**, Emotional awareness subscale; **STRATEGIES**, Emotion regulation strategies subscale; **CLARITY**, Emotional clarity subscale; **ADHD-RS**, ADHD Rating Scale; **CVA**, Cardiac Vagal Activity; IQ scores were collected in the first assessment of the study, when participants were 8-12 years old. *= p.<.05 **= p.<.01.

TABLE 2 | ADHD subtypes and comorbid diagnoses.

| ADHD subtype             | ADHD (n, %) | Controls (n, %) |
|--------------------------|-------------|----------------|
| IA                       | 13 (38.2)   | -             |
| HI                       | 1 (2.9)     | -             |
| C                        | 20 (58.8)   | -             |
| TS                       | 12 (35.3)   | 5 (15.2)      |
| Anxiety disorders        | 10 (29.4)   | 2 (6.1)       |
| ODD                      | 7 (20.6)    | -             |
| OCD                      | 2 (5.9)     | 1 (3.0)       |
| MDD                      | 1 (2.9)     | -             |
| Tourette Syndrome        | 1 (2.9)     | -             |
| Oppositional Defiant Disorder | 1 (2.9) | - |
| OCD, Obsessive-Compulsive Disorder | 1 (2.9) | - |
| Major Depressive Disorder | -          | 1 (3.0)       |

Barbara, CA). The signal was conducted through three adhesive Ag/AgCl electrodes (T815 Dia. 55).

The first 5 minutes of the ECG recordings were subject to preparation and analysis in Kubios HRV version 2.2 (38). Interbeat intervals (IBI) were calculated, and, if necessary, corrected. A total of 11 IBI corrections were made (.4–1.5% of IBIs in the corrected recordings), in six recordings (ADHD, n = 3; controls, n = 3). Additionally, one R-R interval containing an extra systole was removed from a recording.

Recordings were subject to a frequency analysis with the Fast Fourier Transformation, yielding a power spectrum displaying activity in the very low frequency (VLF, < .04 Hz), low frequency (LF,.04–.15 Hz) and high frequency (HF,.15–.4 Hz) domain (39). HF power was chosen as the applied measure of CVA (40). We assessed HF power in normalized units (n.u.), which corrects for the influence of VLF-HRV, and thus can be regarded as a measure of pure vagal influence (41). We used HF peak to indirectly control for respiratory frequency (42).

**Emotion Dysregulation**

The Difficulties in Emotion Regulation Scale [DERS; (43)] is a 36-item self-report questionnaire assessing ER difficulties. Items are rated on a five-point Likert-type scale from 1 (“almost never”) to 5 (“almost always”). In addition to a total score (DERS TOTAL), the questionnaire yields information on ER difficulties relating to different facets: (i) Non-acceptance of emotional responses (NONACCEPTANCE; e.g. “When I’m upset, I become embarrassed for feeling that way”; Cronbach’s α in the current sample =.88); (ii) Difficulties engaging in goal-directed behavior (GOALS; e.g. “When I’m upset, I have difficulty focusing on other things”; α=.89); (iii) Impulse control difficulties (IMPULSE; e.g. “When I’m upset, I feel out of control”; α=.92); (iv) Lack of emotional awareness (AWARENESS; e.g. “I am attentive to how I’m feeling”; α=.75); (v) Limited access to ER strategies perceived as effective (STRATEGIES; e.g. “When I’m upset, it takes me a long time to feel better”; α=.85); and (vi) Lack of emotional clarity...
(CLARITY; e.g. “I am confused about how I feel”; α=.73). High scores reflect a high level of emotional dysregulation.

A Norwegian translation of the DERS was used (44) Although originally developed for adults, the DERS has been validated in adolescent samples, where it has shown satisfactory to excellent internal consistency (Cronbach’s α=.72-.89) and adequate construct validity (45, 46). Further, it has been applied in a study of an adolescent ADHD sample (47).

In the current sample, six participants (ADHD: n=3, Controls: n=3) had one through five missing item scores, in total 14 missing scores.

**Physical Activity and Body Mass Index (BMI)**

Engagement in physical activity was assessed during the K-SADS, as the adolescents reported in “Other activities; Sport and Exercise”. Physical activity levels were scored on a six-point Likert-type scale from 0 (“zero times a week”) to 5 (“seven times or more a week”), as for Question 1 of the Physical Activity Questionnaire for Adolescents [PAQ-A; (48)]. The PAQ-A scoring norm was applied due to its convergent validity (49). Physical activity levels in between two categories were lowered to the nearest category. Further, BMI was calculated from measured height and weight as weight/height2 (kg/m²). BMI could not be calculated for four participants (ADHD, n=2; Controls, n=2), as they declined to get weighed or due to technical errors.

**Procedure**

Adolescents and their parents participated in assessments conducted over two days. Test administrators were blinded to group status. On the first day, participants received information about the project and the assessments they would undergo. Then, ECG recordings were conducted between 9 a.m. and 1 p.m., to control for the effect of circadian variation on CVA (50). To rule out short-term effects of centrally stimulating medication for ADHD on heart rate (51) and ER, 89.5% of the adolescents with short-term effects of centrally stimulating medication for ADHD on stimulants (52) had a washout period of at least 24 h. This equals minimum five half-lives of the stimulant medication (52). On the second day, adolescents and their parents were separately interviewed with the K-SADS and/or DERS and/or CVA. Missing ADHD-RS data from one participant with ADHD was imputed with the sample mean. Missing BMI values were calculated for four participants (ADHD, n=2; Controls, n=2), as they declined to get weighed or due to technical errors.

**Statistical Analysis**

Statistical analyses were performed in the Statistical Package for the Social Sciences version 24.0 (SPSS; IBM Corp., Armonk, NY, 2016). CVA data were log-transformed to approximate a normal distribution (53). We performed preliminary analyses with independent samples t-tests of group differences in distribution of age, gender, ADHD-RS scores, DERS scores, CVA, HF peak, Full-Scale IQ, and on HRV power spectrum values. Bivariate correlations were conducted between age, BMI, physical activity, HF peak, CVA, and DERS scores. No outliers, defined as values ± 3SD from the mean, were detected for DERS TOTAL, ADHD-RS, or CVA. Missing ADHD-RS data from one participant with ADHD was imputed with the sample mean. Missing BMI values were replaced with the sample mean and missing DERS values by the sample means of the respective items.

To test the hypothesis that lower CVA would be related to higher self-reported ER difficulties, we conducted multiple linear regression analyses. The DERS TOTAL score and the DERS subscale scores were used as dependent variables, respectively. CVA and diagnostic status were included as predictors. Analyses were adjusted for effects of age, gender, and HF peak. The regression model were adjusted for BMI [see (54)] and physical activity [see (55)] by including these as predictors, when a significant correlation between these variables and CVA and/or DERS subscale scores were found in the preliminary analyses. To adjust for multiple testing, we performed Bonferroni correction (56), yielding a corrected alpha level of p =.007 (.05/7). In follow-up regression analyses, an interaction term of ADHD×CVA (CVA scores were centralized with z scores for the interaction term) was added as a predictor to test for interaction effects between ADHD and CVA on ER difficulties (57, 58). As supplementary analyses, we conducted the same multiple linear regression analyses separately for the ADHD and control groups. We also performed follow-up analyses excluding participants in the control group with psychiatric disorders, to control for effects of a mixed participants control group.

**RESULTS**

**Preliminary Analyses**

There were no differences in age nor gender between the ADHD and control group (Table 1). Parents reported higher ADHD-RS scores in the ADHD group compared to the control group. There were no significant differences in ADHD-RS scores between samples from the first and second assessment of the study. Further, control participants reported significantly higher physical activity levels than adolescents with ADHD did. There were no significant differences in BMI or HF peak between the groups. Bivariate correlation analyses showed that lower CVA was associated with higher DERS TOTAL, STRATEGIES, and CLARITY scores (Supplemental Table 1). Higher CVA also correlated with higher physical activity. CVA did not correlate with age nor BMI. Further, independent samples t-test showed no group difference in CVA. The ADHD group reported higher total scores and subscale scores on the DERS, except for on AWARENESS. No gender effects were found on CVA (Supplemental Table 2). The only gender effect that appeared in relation to DERS scores was that girls reported higher CLARITY scores than boys (Mean values: girls = 11.04; boys = 8.77; t = 2.67, p=.009). There were no group differences in CVA power spectrum values (Supplemental Table 3).

**Predicting Emotion Dysregulation by CVA**

To test the effect of CVA on ER difficulties, we conducted multiple linear regression analyses (Table 3), which showed that lower CVA predicted higher STRATEGIES scores (Figure 1). There was a tendency for lower CVA to also predict higher DERS TOTAL scores. No effects of CVA were detected on NONACCEPTENCE, GOALS, IMPULSE, AWARENESS, or...
CLARITY. The diagnostic status of ADHD predicted higher scores on DERS TOTAL and all DERS subscales except for AWARENESS. Gender covaried with DERS TOTAL, STRATEGIES, and CLARITY, in that girls had higher scores. Higher HF peak covaried with higher CLARITY and GOALS. Neither age nor physical activity covaried with any of the DERS scores. Follow-up regression analyses adding the interaction term ADHD*CVA did not detect an interaction effect between ADHD and CVA on any of the DERS scores. Conducting the same multiple regression analyses separately in the diagnostic groups showed that lower CVA predicted higher STRATEGIES in the ADHD group (Supplemental Table 4: Supplemental Figure 2). CVA did not predict any other DERS scores in the ADHD group, and did not predict any DERS scores in the control group. Lastly, excluding controls with comorbid disorders, we found that CVA still predicted STRATEGIES (\( p=.004 \)) in multiple linear regression analyses.

**DISCUSSION**

The aim of the present study was to investigate whether CVA, an index of autonomic flexibility, was associated with self-reported ER difficulties in adolescents with ADHD and controls. Results showed that, as expected, lower CVA significantly predicted higher STRATEGIES scores. There was also a tendency for lower CVA to predict higher DERS TOTAL scores. This indicates that level of CVA reflects the ability to use effective strategies for ER. Results also showed that the ADHD diagnosis associated with higher DERS scores, and thus higher ER difficulties, except for in relation to lack of emotional awareness. However, no interaction effects between level of CVA and ADHD on the DERS scores were found. When analyzing the association between CVA and DERS separately in the diagnostic groups, we found that lower CVA was associated with higher STRATEGIES scores in the ADHD group.
group and not in the controls. Thus, despite the non-significant interaction effect between CVA and ADHD, there was a tendency for CVA to be associated with less access to effective ER strategies specifically in the ADHD group.

Our results suggest that level of CVA is important for the use of ER strategies perceived as effective by the individual experiencing and expressing the emotions. The STRATEGIES subscale measures the flexible use of situationally appropriate strategies (43). This finding is in line with a study of young adults, where higher CVA was associated with the use of constructive coping strategies during stressful events (29). Further, a study of adolescents found higher CVA to be associated with a tendency of seeking support from others in times of suffering, an important way to modulate and alleviate emotions (30). Lower CVA has been suggested to be associated with difficulties in recruiting prefrontal brain areas necessary for modulation of activity in amygdala during explicit ER, such as reappraisal and response modulation (28). Such prefrontal cortical regions have been found to display altered structural and functional maturation in ADHD (2, 4), and activity in these areas are hypothesized to be reflected in CVA (7). Through this network, inappropriate timing and magnitude of emotional experiences and behavioral reactions might be associated with reduced vagal modulation (5).

In our study, the small sample size, and as such low statistical power, may restrict the detection of interaction effects between ADHD and level of CVA on DERS scores. Despite this restriction, when looking into the relation between CVA and ER for the ADHD and control groups separately, we found a significant effect of CVA on ER (i.e., STRATEGIES) only in the adolescents with ADHD. This indicates that despite a lack of a significant interaction effect, there is a tendency of lower CVA to associate with higher ER difficulties specifically in the ADHD group. Since there are few studies of CVA and ER in ADHD, it is still not clear whether lower CVA predicts ER difficulties specifically in ADHD. As far as we know, only one previous study has found parent-reported emotion dysregulation to be associated with lower CVA in children and adolescents with ADHD (9). Further, another study found lower CVA in children and adolescents with ADHD to be associated with higher levels of anxiety, social difficulties, and oppositional behavior (26). It is possible that lower CVA could be related to ER difficulties in general, and not specifically for individuals with ADHD. In the current study, adolescents with ADHD reported higher levels of ER difficulties than controls, as also reported by parents in previous studies (2). It is therefore important to be aware of the association between ER difficulties and inflexible autonomic functioning among adolescents with ADHD, whether this association is specific for ADHD or not. This is because lower CVA might have implications for not only psychological, but also somatic health, morbidity, and longevity (59).

Previous studies of CVA and DERS in young, healthy adults have found CVA to be associated with less emotional impulse control and emotional clarity (15), and less acceptance of negative emotions (16). A longitudinal study over three years of children with and without psychiatric conditions found increasing CVA to be associated with fewer ER difficulties as measured by the DERS, and specifically with higher emotional awareness (17). The inconsistency in results between studies of CVA and DERS shows that it is not clear on a subscale level which specific ER domain are associated with CVA. In our study, we investigated a different age group than two of the previous studies. Our results are therefore not directly comparable to the previous findings.

Our results, indicating that lower CVA and thus inflexibility of the ANS is associated with limited access to ER strategies among adolescents with ADHD and controls, might have important theoretical and clinical implications. Our study contributes to the understanding of which specific domains of ER are related to CVA, which could support theoretical models of the relation between neural and psychological processes. Further, this might lead to increased focus on biofeedback treatment, which has been suggested to increase ER abilities [e.g. (60)]. Also, our finding of a significant association between level of CVA and self-reported ER difficulties shows that adolescents with ADHD seem to have insight into their own problems, which could be a good starting point for psychotherapeutic interventions. Further, future studies of adolescent ADHD samples might benefit from using self-reports instead of only observational reports from parents or teachers when investigating relations between ER difficulties and neurobiological variables such as CVA.

The current study has several strengths and limitations. One strength is that we have included possible confounders on CVA such as physical activity levels (55), BMI (54), and HF peak (42). Further, we controlled for circadian influences on CVA. Regarding limitations, we only used one questionnaire to assess ER difficulties, which may not capture the entire complexity of the concept. In addition, although we asked participants to conduct a washout period of ADHD medications to rule out short-term effects, we cannot rule out possible longer-term effects of such medication on CVA (51, 61, 62) and/or ER (63). Further, the mixed control group of adolescents with and without psychiatric diagnoses other than ADHD may have contributed to lower statistical power in detecting group differences. Finally, the study is cross-sectional in nature. Although there is evidence suggesting that lower CVA prospectively predicts emotion dysregulation (64), there is a possibility that difficulties in ER could influence CVA. Future research on the field would benefit from longitudinal designs, as well as studies designed to manipulate ER, to examine this notion.

DATA AVAILABILITY STATEMENT

Datasets are available on request. The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation, to any qualified researcher.

ETHICS STATEMENT

This study was carried out in accordance with the recommendations of Regional Committee for Medical Research Ethics of Western Norway, study number 2014/1304. All subjects
gave written informed consent in accordance with the Declaration of Helsinki. The protocol was approved by the Regional Committee for Medical Research Ethics of Western Norway.

**AUTHOR CONTRIBUTIONS**

LS and KP designed the study. LS, SA, EK, and HE were involved in data collection. EK and LS, and BO performed data acquisition and analyses. EK and LS wrote the first draft, and JK, OF, BO, HE, and KP contributed to manuscript editing. All authors have read and approved the final manuscript.

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**SUPPLEMENTARY MATERIAL**

The Supplementary Material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fpsyg.2020.00244/full#supplementary-material
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