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Experiential Avoidance Predicts Persistence of Major Depressive Disorder and Generalized Anxiety Disorder in Late Adolescence

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ABSTRACT

Objective: Experiential avoidance (EA) is a transdiagnostic construct that may underlie the high comorbidity between major depressive disorder (MDD) and generalized anxiety disorder (GAD). This analysis used data from a longitudinal study (conducted September 2010–April 2016) to examine whether adolescent EA varies by MDD and GAD symptomatology trajectory and predicts said trajectories. Longitudinal associations between EA, anxiety, and depression symptoms were also examined.

Methods: Adolescents aged 15 to 20 years (N = 183) were followed for 2 years using a comprehensive assessment battery. Symptom trajectory modeling, using weekly symptom ratings, identified 4 MDD and 4 GAD trajectories that were collapsed to form combined MDD/GAD trajectory groups: Persistent (n = 81), High-Decreasing (n = 44), Normal-Increasing (n = 37), and Minimal (n = 21). Group-based trajectory modeling, analyses of covariance, structural equation modeling, and linear regression analyses were performed. *DSM-IV-TR* criteria were used for MDD and GAD diagnoses.

Results: The Persistent adolescents had higher EA than other groups (*P* values $\leq .001$), with greater EA stability versus High-Decreasing adolescents (*P* = .008). EA predicted anxiety and depressive symptoms alike (*P* values $\leq .005$), which in turn did not predict EA (*P* values $\geq .188$). EA, at both time points, predicted combined MDD/GAD trajectories after adjustment for depressive and anxiety symptoms and other confounders (*P* values $< .001$).

Conclusions: EA appears to be an important predictor of MDD and GAD symptomatology in older adolescents, potentially serving as a treatment target. Findings suggest a possible trait-like nature for EA, perhaps increasing risk for the emergence and persistence of MDD and/or GAD.

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Major depressive disorder (MDD) and generalized anxiety disorder (GAD) are prevalent psychiatric disorders and leading causes of disability worldwide.¹ MDD and GAD are highly comorbid, particularly among adolescents, which results in more significant distress and functional impairment as compared to either disorder alone.^{2,3} Identifying factors that underlie their comorbidity may further our understanding of MDD and GAD, leading to improved treatments.

Experiential avoidance (EA) is a multifaceted transdiagnostic construct that offers promise in this pursuit. Specifically, EA refers to an “unwillingness to remain in contact with uncomfortable private events by escaping or avoiding these experiences.”^{4(p1154)} Avoidance of distressing experiences reduces immediate contact with the distressing experiences, delivering short-term relief. Long-term, however, it leads to greater dysfunction and increased distress.^{5,6} EA is posited to relate to various forms of psychopathology through both implicit (ie, classically conditioned) and explicit (ie, deliberate avoidance) pathways.^{7,8}

Cross-sectional adolescent studies^{8,9} have found positive associations between EA, depression, and anxiety at diagnostic (categorical) and symptom (dimensional) levels. These studies demonstrated high EA among participants with current MDD and GAD. High EA has also been identified in adolescents at high risk for MDD who have not yet developed a major depressive episode.¹⁰ It is unknown, however, whether EA changes with disorder onset (or recurrence) and remittance in adolescents.

A few longitudinal studies in adults have examined EA in depressive and anxiety disorder occurrence and remittance, as well as symptom increase. For example, using 6-year data from the Netherlands Study of Depression and Anxiety (NESDA),¹¹ Spinhoven and colleagues¹² found an association between EA and current depressive and anxiety disorders (collectively termed *distress disorders*). While EA was mostly stable within individuals, it did increase and decrease with distress disorder occurrence and remittance, respectively. EA also predicted changes in diagnostic status.¹² A separate study¹³ found reduction in EA was significantly associated with reduction in depressive symptoms over the course of 1 year’s treatment of borderline personality disorder. Finally, EA positively predicted daily levels of social anxiety and emotional distress over 3 weeks in a sample of college undergraduates.¹⁴ While these findings highlight the role EA might play in the onset and course of distress disorders, other adult studies^{15,16} failed to support it. To the

Clinical Points

- Experiential avoidance (EA) is a multifaceted transdiagnostic construct implicated in major depressive disorder (MDD) and generalized anxiety disorder (GAD). However, investigation of the role of EA in MDD and GAD symptom trajectories, as well as the longitudinal associations of EA with depressive and anxiety symptoms, is lacking in adolescence.
- EA was found to associate with persistence of MDD and/or GAD symptomatology. Moreover, longitudinal analyses showed EA to predict subsequent depressive and anxiety symptoms as well as severity of combined MDD/GAD symptoms.
- Findings suggest EA is an important predictor of MDD and GAD symptomatology in older adolescents, perhaps serving as a treatment target for extant evidence-based psychotherapeutic interventions.

best of our knowledge, there has yet to be a longitudinal study of adolescent EA in relation to distress disorders (ie, MDD and GAD). Moreover, no study has jointly examined longitudinal relations between EA, depression, and anxiety symptom severity.

The present investigation is a secondary data analysis from a 2-year longitudinal study investigating the skeletal effects of selective serotonin reuptake inhibitors (SSRIs) in adolescents.¹⁷ The primary aim of the present study was to examine whether adolescent EA varies by combined MDD/GAD symptomatology trajectory and predicts trajectories over and above depressive and anxiety symptoms.

We hypothesized that adolescents with persistent MDD and/or GAD would endorse significantly elevated EA^{9,10} and exhibit more stability of EA severity longitudinally compared to adolescents of other combined MDD/GAD trajectory groups.¹² We also expected that EA would predict combined MDD/GAD trajectories after controlling for anxiety and depression. In an exploratory analysis, we also sought to examine the directionality of longitudinal associations between EA, anxiety, and depression symptoms.

METHODS

Procedures and Participants

The University of Iowa Institutional Review Board approved this longitudinal, observational study,¹⁷ which was conducted from September 2010 to April 2016 (ClinicalTrials.gov identifier: NCT02147184). Adult participants and parents or guardians of minor participants provided written informed consent, and minors provided assent. Older adolescents aged 15 to 20 years were recruited through inpatient and outpatient clinical settings, advertisements, and word of mouth. They were enrolled either psychotropic-free or within a month of starting an SSRI, regardless of diagnostic status excluding eating disorders and substance dependence.¹⁷ Additional exclusion criteria included significant medical history; pregnancy;

concomitant treatment with other antidepressants, mood stabilizers, or antipsychotics; chronic use of medications affecting bone metabolism; or plans to soon move out of state.

At the baseline visit, demographic data were collected, including self-reported race/ethnicity. Participants completed the Beck Depression Inventory-II (BDI-II)¹⁸ and the Beck Anxiety Inventory (BAI),¹⁹ and trained research staff administered the Longitudinal Interval Follow-Up Evaluation for Adolescents (A-LIFE).²⁰ Participants returned for follow-up visits every 4 months to complete this battery, and post-visit meetings were held to reach clinical consensus on A-LIFE ratings. EA was assessed using the Acceptance and Action Questionnaire, second version (AAQ-II).²¹ This questionnaire was administered twice during the study, the first time being at the first follow-up visit. A mean (SD) of 342.1 (56.7) days passed between the initial (V1) and second (V2) visits at which AAQ-II data were collected.

Measures

Experiential avoidance. The AAQ-II is a self-completed questionnaire consisting of 7 items rated on a 7-point Likert scale with higher scores indicating greater EA (range, 7–49).²¹ The AAQ-II has demonstrated good psychometrics in late-adolescent and young-adult samples.²² Internal consistencies, as measured by Cronbach α , were 0.92 and 0.93 for V1 and V2, respectively.

Depressive symptoms. On the self-completed, widely used 21-item BDI-II,¹⁸ responses are scored on a 4-point Likert scale with higher scores indicating greater depression (range, 0–63). Internal consistencies were 0.93 and 0.94 for V1 and V2, respectively.

Anxiety symptoms. On the self-completed 21-item BAI,¹⁹ responses are scored on a 4-point Likert scale with higher scores indicating greater anxiety (range, 0–63). Internal consistencies were 0.91 at both V1 and V2.

MDD and GAD weekly symptoms. The clinician-administered A-LIFE²⁰ was used to track weekly MDD and GAD symptoms. Clinician ratings on the A-LIFE range from 0 for no symptoms, to 2 to 4 for varying levels of symptom severity and impairment, to 5 and 6 for meeting full *DSM-IV-TR* criteria.

Data Analytic Strategy

Consistent with Spinhoven and colleagues,¹² MDD and GAD were subsumed under the broader term *distress disorders*. This approach was motivated by significant overlap in symptom presentation and high rates of MDD-GAD co-occurrence.^{23–26} A-LIFE-based weekly MDD and GAD ratings were analyzed using group-based trajectory modeling (GBTM), a statistical method used to identify clusters of individuals following similar patterns of progression over time.²⁷ Models with 3- and 4-group solutions were generated using the censored normal distribution and the polynomial function of time (linear, quadratic, or cubic) that best fit the data. In each model, individuals were assigned to the group in which the GBTM-determined probability of membership was highest. The choice of best-fitting model was informed

Table 1. Participants' Combined MDD and GAD Trajectory Group Assignments^a

Group	n	Baseline Symptom Severity	Follow-Up Symptom Severity
Persistent	81	Elevated MDD and/or GAD	Elevated MDD and/or GAD
High-Decreasing	44	Elevated MDD and/or GAD	Decreasing MDD and GAD
Normal-Increasing	37	Low MDD and GAD	Increasing MDD and/or GAD
Minimal	21	Low to absent MDD and GAD	Low to absent MDD and GAD

^aClinical significance of symptomatology determined via weekly psychiatric status scores on the A-LIFE.

Abbreviations: A-LIFE = Longitudinal Interval Follow-Up Evaluation for Adolescents, GAD = generalized anxiety disorder, MDD = major depressive disorder.

by model fit indices²⁸ and ultimately made by maximizing the number of non-redundant MDD and GAD patterns. Four-group, rather than alternative, solutions for MDD and GAD trajectories were decided upon because they provided better fits to the data and were of greater clinical utility, allowing for more meaningful interpretation of findings. The GBTM analysis was conducted using PROC TRAJ in SAS 9.4.²⁹

Bivariate relations were examined with Pearson correlations. Chi-square (χ^2) tests of independence and Bonferroni-corrected analyses of covariance (ANCOVAs) were used for group comparisons. Longitudinal associations between EA, anxiety, and depressive symptoms were investigated using cross-lagged autoregressive structural equation modeling (SEM) analyses allowing for residual correlations. Acceptable model fit required the root mean square error of approximation (RMSEA) ≤ 0.08 , the comparative fit index (CFI) ≥ 0.90 , and the Tucker-Lewis index (TLI) ≥ 0.90 . Excellent model fit required RMSEA ≤ 0.06 , CFI ≥ 0.95 , and TLI ≥ 0.95 ; in both instances, χ^2 was to be large and nonsignificant ($P \geq .05$).^{30,31} Finally, hierarchical linear regression models were tested with AAQ-II scores predicting combined MDD/GAD symptom trajectories. Variance inflation factor and tolerance indices indicated degree of predictor multicollinearity.³² Covariates included age, sex, and cumulative SSRI exposure (defined as cumulative time [in days] taking an SSRI) given potential confounding effects.^{33,34}

Supplemental SEM and regression analyses were performed with an abbreviated, 5-item AAQ-II score to examine whether results would replicate when the items strongly resembling anxiety symptoms ("I worry about not being able to control my worries and feelings" and "Worries get in the way of my successes") were excluded. This approach was taken to reduce potential predictor-outcome overlap.³⁵ SPSS version 19³⁶ and MPlus version 7.2³⁷ were utilized.

RESULTS

Participant Disposition

A total of 279 participants were recruited; however, 40 were missing all EA data, 8 were excluded for bipolar disorder and/or psychosis, and 2 were excluded for SSRI use prior to starting the study. Forty-six participants provided EA data only at V1 and were retained for the relevant analyses. This left 183 participants with complete data, who did not significantly differ from those with partial EA data on age ($P = .131$), sex ($P = .507$), ethnicity ($P = .523$), EA ($P = .747$),

anxiety symptoms ($P = .484$), or depressive symptoms ($P = .871$), though they differed on race ($P \leq .001$).

Identification of MDD and GAD Trajectories Within the Sample

A 4-group trajectory model offered the best fit for both MDD and GAD regardless of whether linear, quadratic, or cubic modeling was used (see Supplementary Figures 1 and 2). For MDD, group 1 exhibited no symptoms, group 2 exhibited declining symptoms from a relatively low level, group 3 exhibited remitting symptoms from a clinically significant baseline, and group 4 exhibited persistent clinically significant symptoms. As for GAD, group 1 initially reported minimal symptoms that increased slightly over time, group 2 had moderate baseline symptoms that further increased during follow-up, group 3 had declining symptoms from a clinically significant level, while group 4 appeared to maintain a clinically significant symptom level over the study period. Subsequently, per these MDD and GAD trajectories, participants were assigned to 1 of 4 combined MDD/GAD trajectory categories (Table 1).

Descriptive Statistics and Group Comparison Results

Table 2 lists demographic and clinical characteristics of included participants. Several significant group differences were observed across variables. With respect to EA, Persistent adolescents reported significantly greater symptoms at both V1 and V2 than adolescents in other trajectory groups, who did not differ from one another (P values $\geq .05$). Adjusting for age, sex, and cumulative SSRI exposure did not substantially alter the findings.

Bivariate Correlations

EA, depression, and anxiety symptoms were significantly correlated at each of the 2 visits when EA was assessed, as well as across visits (Table 3).

Within-Subject Change in EA Between V1 and V2

Groups significantly differed in EA change between V1 and V2 ($F_{3, 175} = 4.23$, $P = .006$, $\eta^2 = 0.068$) after adjustment for age ($P > .90$), sex ($P > .50$), cumulative SSRI exposure ($P > .40$), and V1 EA ($\beta = -0.469$; $P < .001$; 95% CI, -0.602 to -0.335). Post hoc analyses showed that the effect was driven by a significant difference in EA change between the High-Decreasing and the Persistent groups (mean difference in Δ EA = 4.38; 95% CI, 0.815 to 7.950; $P = .008$).

Table 2. Participant Characteristics and Group Comparison Results on Data From Visits at Which EA Data Were Collected^a

Variable ^b	Complete Data Sample, N = 183	Persistent (P), n = 81	High-Decreasing (HD), n = 44	Normal-Increasing (NI), n = 37	Minimal (M), n = 21	Test Statistic (P Value)	Group Difference
Age, y	18.95 (1.61)	19.22 (1.39)	18.41 (1.87)	19.22 (1.40)	18.52 (1.86)	3.40 (.019)	P > HD
Female, %	61.7	77.8	50.0	54.1	38.1	17.28 (<.001)	P > HD, NI, M; M < HD, NI
White, %	77.6	70.4	81.8	78.4	95.2	6.73 (.077)	...
Non-Hispanic, %	81.4	74.1	88.6	78.4	100.0	3.51 (.136)	...
EA Visit 1 ^c							
SSRI (% yes)	38.3	53.2	48.1	9.1	0.0	39.13 (<.001)	P > NI, M
AAQ-II score	17.02 (8.33)	21.57 (8.37)	15.10 (6.31)	11.45 (4.75)	10.71 (4.56)	27.86 (<.001)	P > HD > NI, M
BAI score	4.68 (6.29)	7.42 (7.44)	3.25 (4.20)	1.98 (3.11)	0.29 (0.62)	16.80 (<.001)	P > HD, NI, M
BDI-II score	5.83 (7.55)	9.93 (8.56)	4.21 (6.24)	2.00 (2.47)	0.42 (0.78)	17.97 (<.001)	P > HD, NI, M
EA Visit 2							
SSRI (% yes)	21.7	28.6	26.2	5.9	0.0	15.45 (.001)	P > NI, M
AAQ-II score	16.56 (8.68)	21.51 (9.54)	13.80 (5.56)	12.57 (5.22)	10.24 (4.41)	22.40 (<.001)	P > HD, NI, M
BAI score	4.25 (6.38)	7.36 (7.82)	2.00 (3.11)	1.89 (3.89)	1.14 (2.57)	14.09 (<.001)	P > HD, NI, M
BDI-II score	5.13 (7.62)	9.30 (9.39)	2.61 (3.92)	1.59 (2.68)	0.57 (0.87)	19.31 (<.001)	P > HD, NI, M

^aData are mean (SD) unless noted otherwise.^bTwenty-six participants elected not to self-report ethnicity.^cFigures for EA visit 1 include participants with partial EA data (n = 229): Persistent (n = 109), High-Decreasing (n = 52), Normal-Increasing (n = 44), and Minimal (n = 24).

Abbreviations: AAQ-II = Acceptance and Action Questionnaire, second version; BAI = Beck Anxiety Inventory; BDI-II = Beck Depression Inventory-II; EA = experiential avoidance; SSRI = selective serotonin reuptake inhibitor.

Table 3. Pearson Correlations Among Primary Study Variables

Variable	1	2	3	4	5	6	7	8	9	10
1. Age ^a	...									
2. Sex ^b	0.027	...								
3. AAQ-II at V1 ^c	0.163*	-0.231***	...							
4. BAI at V1 ^c	0.063	-0.183**	0.564***	...						
5. BDI-II at V1 ^c	0.081	-0.154*	0.645***	0.759***	...					
6. SSRI at V1 ^{c,d}	-0.101	-0.076	0.262***	0.398***	0.309***	...				
7. AAQ-II at V2	0.085	-0.170*	0.649***	0.516***	0.527***	0.243***	...			
8. BAI at V2	0.011	-0.208**	0.399***	0.529***	0.421***	0.312***	0.522***	...		
9. BDI-II at V2	-0.033	-0.202**	0.528***	0.584***	0.631***	0.273***	0.655***	0.710***	...	
10. SSRI at V2	-0.095	-0.057	0.224**	0.369***	0.288***	0.931***	0.207**	0.318***	0.274***	...

^aAge (y) at baseline visit.^bSex: female = 0.^cParticipants with partial EA data (EA collected at V1 only) included (n = 229).^dCumulative SSRI exposure (days).

*P < .05. **P < .01. ***P < .001.

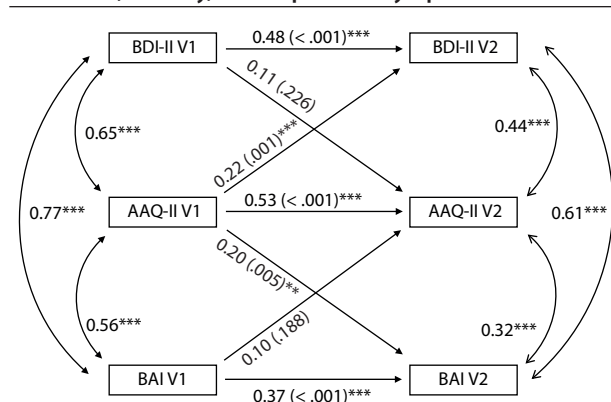
Abbreviations: AAQ-II = Acceptance and Action Questionnaire, second version; BAI = Beck Anxiety Inventory; BDI-II = Beck Depression Inventory-II; EA = experiential avoidance; SSRI = selective serotonin reuptake inhibitor; V1 and V2 = first and second visits at which EA data were collected.

Association of EA, Depression, and Anxiety Over Time

SEM analyses were performed to examine the association between V1 and V2 EA, depression, and anxiety symptoms. Interrelations were controlled for with residual correlations (Figure 1), while adjusting for age, sex, and cumulative SSRI exposure. The specified model fit the data excellently ($\chi^2_5 = 8.16$, $P > .10$, RMSEA = 0.059, CFI = 0.993, TLI = 0.951). EA at V1 predicted V2 anxiety and depressive symptoms, with V1 EA being the sole significant predictor of V2 EA. Therefore, EA held significant associations with both anxiety and depression across time points. Notably, the model yielded the same pattern of results when using abbreviated AAQ-II scores.

EA Predicts MDD/GAD Symptom Trajectories

Hierarchical linear regression models examined V1 and V2 EA as predictors of combined MDD/GAD trajectories (1 = Minimal, 2 = Normal-Increasing, 3 = High-Decreasing, and 4 = Persistent). Findings were similar whether the

Figure 1. Associations Between V1 and V2 Experiential Avoidance, Anxiety, and Depressive Symptoms^a^aData are standardized parameter estimates (P values).

P < .01. *P < .001.

Abbreviations: AAQ-II = Action and Acceptance Questionnaire-II, BAI = Beck Anxiety Inventory, BDI-II = Beck Depression Inventory-II, V1 and V2 = first and second study visits at which all 3 assessments were administered.

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Table 4. Hierarchical Regression Results Showing That EA at V1 and V2 Predicts Combined MDD/GAD Trajectories

Variable	B	SE	β	t	P	F	R ²	Adjusted ^a R ²	ΔR^2
EA V1^b									
Step 1	24.234	0.248	0.238	...
Constant	1.247	0.709	...	1.760	.080				
Age	0.091	0.037	0.145	2.464	.015*				
Sex	-0.558	0.125	-0.261	-4.454	<.001***				
SSRI ^c	0.004	0.001	0.389	6.609	<.001***				
Step 2	23.748	0.396	0.380	0.148
Constant	1.283	0.641	...	2.003	.046*				
Age	0.049	0.034	0.078	1.443	.151				
Sex	-0.381	0.116	-0.179	-3.285	.001***				
SSRI ^c	0.003	0.001	0.259	4.448	.001***				
AAQ-II	0.041	0.009	0.325	4.541	<.001***				
BAI	-0.001	0.014	-0.003	-0.036	.972				
BDI-II	0.018	0.012	0.132	1.485	.139				
EA V2									
Step 1	18.056	0.232	0.219	...
Constant	1.191	0.827	...	1.439	.152				
Age	0.093	0.043	0.141	2.147	.033*				
Sex	-0.557	0.142	-0.258	-3.930	<.001***				
SSRI ^c	0.002	0.000	0.383	5.819	<.001***				
Step 2	18.008	0.380	0.359	0.148
Constant	0.896	0.752	...	1.191	.235				
Age	0.072	0.040	0.110	1.826	.070				
Sex	-0.397	0.132	-0.184	-3.012	.003**				
SSRI ^c	0.002	0.000	0.284	4.557	<.001***				
AAQ-II	0.036	0.010	0.284	4.504	<.001***				
BAI	-0.001	0.014	-0.008	-0.088	.930				
BDI-II	0.021	0.013	0.154	1.595	.112				

^aAdjusted for the number of model predictors.

^bParticipants with partial EA data (EA collected at V1 only) included (n = 229).

^cCumulative SSRI exposure (days).

*P < .05. **P < .01. ***P < .001.

Abbreviations: AAQ-II = Acceptance and Action Questionnaire, second version; B = unstandardized β ; BAI = Beck Anxiety Inventory; BDI-II = Beck Depression Inventory-II; GAD = generalized anxiety disorder; MDD = major depressive disorder; ΔR^2 = change in explained variance; R² = variance; SE = standard error; SSRI = selective serotonin reuptake inhibitor; V1 and V2 = first and second visits at which EA data were collected.

outcome was treated as an ordinal or as a dimensional variable, and therefore data are presented with dimensional outcomes for ease of interpretation. Age, sex, and cumulative SSRI exposure were entered as covariates in step 1, and the model accounted for 25% of outcome variance (Table 4). V1 EA, anxiety, and depressive symptoms were entered at step 2, predicting an additional 15% of outcome variance with V1 EA as the only significant symptom-level predictor. V2 analyses yielded similar results, with EA the only significant symptom-level predictor of combined MDD/GAD trajectories. Variance inflation factor (median ≤ 1.154) and tolerance (median ≤ 0.867) indices revealed no threat of multicollinearity in either model.³²

After imputing for missing data, increasing the sample to n = 239, V2 analyses yielded the same pattern of results (data available on request). Moreover, V1 and V2 results were little changed when analyses were performed with abbreviated AAQ-II scores.

DISCUSSION

The present investigation is the first longitudinal study to examine the role of EA in late-adolescence distress disorders. Moreover, this study is the first to examine directionality of

longitudinal relations between EA, depression, and anxiety symptoms in any age group. Levels of EA were consistently elevated among adolescents with Persistent combined MDD/GAD symptom trajectories relative to other groups. Moreover, among symptom trajectory groups, EA was significantly less likely to change in the Persistent versus the High-Decreasing group, suggesting that consistently elevated EA levels may portend chronic MDD and/or GAD symptoms. Notably, EA was the only significant symptom-level predictor of combined MDD/GAD trajectories, even after accounting for potential confounders. In fact, SEM analyses showed EA to predict future anxiety and depressive symptoms but not vice versa.

Significantly greater EA among adolescents with persistent MDD and/or GAD symptoms echoes prior findings^{8–10,12} showing EA to be elevated among adults and adolescents with current MDD and GAD. These adolescents may more consistently utilize emotion and thought suppression strategies, which can perpetuate psychopathology.⁶ Patterns of inner experience and behavior (ie, thought suppression, emotional suppression, and avoidance coping)⁸ among these adolescents may be enduring, similarly present during times of persistent MDD and/or GAD symptoms and remission. Experimental evidence and laboratory studies on emotional

and thought avoidance strategies support the idea that experiential avoidance may be a core feature associated with mood and anxiety disorders, including MDD and GAD.^{38–40} In fact, efforts to suppress an unwanted thought can lead to a temporary relief, followed by a period of increased thought frequency and emotional distress.⁴¹ Thus, although these strategies may be reinforcing in the short term, they can result in a vicious cycle of cognitive and emotional avoidance followed by more intense emotional distress associated with reemergence of the unwanted thoughts.^{38,39} Interrupting the cycle of avoidance is a key component of several psychotherapeutic interventions.

Prospectively, EA significantly predicted subsequent anxiety and depressive symptoms, but the opposite was not the case. This finding is consistent with prior research⁴² demonstrating that EA in adults predicts depressive symptom reduction. The fact that earlier depression and anxiety failed to predict subsequent EA suggests unidirectional relations between EA and depressive and anxiety symptoms.

EA, rather than anxiety or depressive symptoms, predicted MDD/GAD symptomatology consistently across longitudinal SEM and hierarchical regression analyses. This result aligns with the finding of higher EA in adolescent girls at risk for MDD compared to healthy controls¹⁰ and provides support for EA's being conceptualized as a transdiagnostic process in late-adolescent distress disorders. An important consideration deals with the degree to which EA is a multifaceted construct, operationally overlapping with anxiety- and depression-related symptoms (ie, worry, neuroticism, and rumination).^{8,43} To mitigate this concern, we performed supplemental analyses with abbreviated AAQ-II scores, omitting GAD-related items to avoid predictor-criterion contamination; the results remained largely unchanged. Remaining AAQ-II items on this

abbreviated version includes items relating to dysfunctional distress (ie, "It seems like most people are handling their lives better than I am").⁴⁴ Therefore, dysfunctional distress, regardless of excessive worrying, appears to have significant cross-sectional and longitudinal associations with depressive and anxiety symptomatology.

Despite its strengths, this study is not without limitations. Assessing baseline EA would have been preferable; however, the AAQ-II was a post hoc addition to the study battery. Social phobia may have been relevant but was excluded due to oversensitivity of diagnostic measures to subthreshold symptoms. Additionally, a significant portion of participants were taking SSRIs, which treat MDD and GAD effectively.^{45,46} Although accounted for in the analyses, it is not fully clear how SSRI use may have moderated the results. Moreover, whether the findings apply to early adolescents is to be determined. Identifying which EA processes, if any, are more dominant than others given stage of childhood development may also be worth investigating. Finally, generalizability of the findings could have been hampered by the limited ethnic/racial diversity of the participants.

The findings of the current study suggest that an intervention targeting EA in adolescents may lead to valuable outcomes. Acceptance and commitment therapy (ACT) is a behavioral intervention that aims to help patients overcome EA with acceptance, mindfulness, and behavioral change strategies.³⁹ ACT has been identified as an empirically supported treatment for MDD and anxiety disorders as well as other chronic health problems.⁴⁷ Moreover, ACT treatment trials in adults have shown that decreases in experiential avoidance are associated with depressive and anxiety symptom reduction.^{48,49} Future studies should examine whether changes in EA among adolescents mediate treatment outcomes in depressive and anxiety symptoms.

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Supplementary material: Available at PSYCHIATRIST.COM

REFERENCES

- World Health Organization. *Depression and Other Common Mental Disorders: Global Health Estimates*. Geneva, Switzerland: World Health Organization; 2017.
- Garber J, Weersing VR. Comorbidity of anxiety and depression in youth: implications for treatment and prevention. *Clin Psychol (New York)*. 2010;17(4):293–306.
- González-Tejera G, Canino G, Ramírez R, et al. Examining minor and major depression in adolescents. *J Child Psychol Psychiatry*. 2005;46(8):888–899.
- Hayes SC, Wilson KG, Gifford EV, et al. Experiential avoidance and behavioral disorders: a functional dimensional approach to diagnosis and treatment. *J Consult Clin Psychol*. 1996;64(6):1152–1168.
- Lynch TR, Robins CJ, Morse JQ, et al. A mediational model relating affect intensity, emotion inhibition, and psychological distress. *Behav Ther*. 2001;32(3):519–536.
- Wenzlaff RM, Wegner DM. Thought suppression. *Annu Rev Psychol*. 2000;51(1):59–91.
- Blackledge JT, Hayes SC. Emotion regulation in acceptance and commitment therapy. *J Clin Psychol*. 2001;57(2):243–255.
- Chawla N, Ostafin B. Experiential avoidance as a functional dimensional approach to psychopathology: an empirical review. *J Clin Psychol*. 2007;63(9):871–890.
- Venta A, Sharp C, Hart J. The relation between anxiety disorder and experiential avoidance in inpatient adolescents. *Psychol Assess*. 2012;24(1):240–248.
- Mellick W, Vanwoerden S, Sharp C. Experiential avoidance in the vulnerability to depression among adolescent females. *J Affect Disord*. 2017;208:497–502.
- Licht CM, de Geus EJ, Zitman FG, et al. Association between major depressive disorder and heart rate variability in the Netherlands Study of Depression and Anxiety (NESDA). *Arch Gen Psychiatry*. 2008;65(12):1358–1367.
- Spinhoven P, Drost J, de Rooij M, et al. A longitudinal study of experiential avoidance in emotional disorders. *Behav Ther*. 2014;45(6):840–850.
- Berking M, Neacsiu A, Comtois KA, et al. The impact of experiential avoidance on the reduction of depression in treatment for borderline personality disorder. *Behav Res Ther*. 2009;47(8):663–670.
- Kashdan TB, Barrios V, Forsyth JP, et al. Experiential avoidance as a generalized psychological vulnerability: comparisons with coping and emotion regulation strategies. *Behav Res Ther*. 2006;44(9):1301–1320.
- Bjornsson A, Carey G, Hauser M, et al. The effects of experiential avoidance and rumination on depression among college students. *Int J Cogn Ther*. 2010;3(4):389–401.
- Shallcross AJ, Troy AS, Boland M, et al. Let it be:

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- accepting negative emotional experiences predicts decreased negative affect and depressive symptoms. *Behav Res Ther.* 2010;48(9):921–929.
17. Calarge CA, Butcher BD, Burns TL, et al. Major depressive disorder and bone mass in adolescents and young adults. *J Bone Miner Res.* 2014;29(10):2230–2237.
18. Beck AT, Steer RA, Brown GK. *Manual for the Beck Depression Inventory-II.* San Antonio, TX: Psychological Corp; 1996.
19. Steer RA, Beck AT. Beck Anxiety Inventory. In: Zalaquett CP, Woods RJ, eds. *Evaluating Stress: A Book of Resources.* Lanham, MD: Scarecrow Education; 1997:23–40.
20. Birmaher B, Axelson D, Strober M, et al. Clinical course of children and adolescents with bipolar spectrum disorders. *Arch Gen Psychiatry.* 2006;63(2):175–183.
21. Hayes SC, Strosahl K, Wilson KG, et al. Measuring experiential avoidance: a preliminary test of a working model. *Psychol Rec.* 2004;54(4):553–578.
22. Bond FW, Hayes SC, Baer RA, et al. Preliminary psychometric properties of the Acceptance and Action Questionnaire-II: a revised measure of psychological inflexibility and experiential avoidance. *Behav Ther.* 2011;42(4):676–688.
23. Judd LL, Akiskal HS, Maser JD, et al. Major depressive disorder: a prospective study of residual subthreshold depressive symptoms as predictor of rapid relapse. *J Affect Disord.* 1998;50(2–3):97–108.
24. Gorwood P. Generalized anxiety disorder and major depressive disorder comorbidity: an example of genetic pleiotropy? *Eur Psychiatry.* 2004;19(1):27–33.
25. Kendler KS. Reflections on the relationship between psychiatric genetics and psychiatric nosology. *Am J Psychiatry.* 2006;163(7):1138–1146.
26. Lahey BB, Rathouz PJ, Van Hulle C, et al. Testing structural models of DSM-IV symptoms of common forms of child and adolescent psychopathology. *J Abnorm Child Psychol.* 2008;36(2):187–206.
27. Nagin DS, Odgers CL. Group-based trajectory modeling in clinical research. *Annu Rev Clin Psychol.* 2010;6(1):109–138.
28. Akaike H. A new look at the statistical model identification. *IEEE Trans Automat Contr.* 1974;19(6):716–723.
29. Jones BL, Nagin DS, Roeder K. A SAS procedure based on mixture models for estimating developmental trajectories. *Social Methods Res.* 2001;29(3):374–393.
30. Hu L, Bentler PM. Cutoff criteria for fit indexes in covariance structure analysis: conventional criteria versus new alternatives. *Struct Equ Modeling.* 1999;6(1):1–55.
31. Barrett P. Structural equation modelling: adjudging model fit. *Pers Individ Dif.* 2007;42(5):815–824.
32. Cohen P, West SG, Aiken LS. *Applied Multiple Regression/Correlation Analysis for the Behavioral Sciences.* Mahwah, New Jersey: Psychology Press; 2014.
33. Rudolph KD. The interpersonal context of adolescent depression. In: Nolen-Hoeksema S, Hilt LM, eds. *Handbook of Depression in Adolescents.* New York, NY: Routledge/Taylor & Francis Corp; 2009:377–418.
34. Price J, Cole V, Goodwin GM. Emotional side-effects of selective serotonin reuptake inhibitors: qualitative study. *Br J Psychiatry.* 2009;195(3):211–217.
35. Wolgast M. What does the Acceptance and Action Questionnaire (AAQ-II) really measure? *Behav Ther.* 2014;45(6):831–839.
36. IBM SPSS Statistics for Windows, Version 19.0 [computer program]. Armonk, NY: IBM Corp; 2010.
37. Muthén LK, Muthén BO. *Mplus User's Guide. Sixth Edition.* Los Angeles, CA: Muthén & Muthén; 1998–2011.
38. Barlow DH, Allen LB, Choate ML. Toward a unified treatment for emotional disorders. *Behav Ther.* 2004;35(2):205–230.
39. Hayes SC, Luoma JB, Bond FW, et al. Acceptance and commitment therapy: model, processes and outcomes. *Behav Res Ther.* 2006;44(1):1–25.
40. Moses EB, Barlow DH. A new unified treatment approach for emotional disorders based on emotion science. *Curr Dir Psychol Sci.* 2006;15(3):146–150.
41. Wegner DM, Zanakos S. Chronic thought suppression. *J Pers.* 1994;62(4):616–640.
42. Boylan KR, Bieleing PJ, Marriott M, et al. Impact of comorbid anxiety disorders on outcome in a cohort of patients with bipolar disorder. *J Clin Psychiatry.* 2004;65(8):1106–1113.
43. Spinhoven P, Drost J, de Rooij M, et al. Is experiential avoidance a mediating, moderating, independent, overlapping, or proxy risk factor in the onset, relapse and maintenance of depressive disorders? *Cognit Ther Res.* 2016;40(2):150–163.
44. Gámez W, Chmielewski M, Kotov R, et al. The brief experiential avoidance questionnaire: development and initial validation. *Psychol Assess.* 2014;26(1):35–45.
45. Goodyer I, Dubicka B, Wilkinson P, et al. Selective serotonin reuptake inhibitors (SSRIs) and routine specialist care with and without cognitive behaviour therapy in adolescents with major depression: randomised controlled trial. *BMJ.* 2007;335(7611):142.
46. Walkup JT, Albano AM, Piacentini J, et al. Cognitive behavioral therapy, sertraline, or a combination in childhood anxiety. *N Engl J Med.* 2008;359(26):2753–2766.
47. Gauntlett-Gilbert J, Connell H, Clinch J, et al. Acceptance and values-based treatment of adolescents with chronic pain: outcomes and their relationship to acceptance. *J Pediatr Psychol.* 2013;38(1):72–81.
48. Hayes SC, Levin ME, Plumb-Villardaga J, et al. Acceptance and commitment therapy and contextual behavioral science: examining the progress of a distinctive model of behavioral and cognitive therapy. *Behav Ther.* 2013;44(2):180–198.
49. Forman EM, Herbert JD, Moitra E, et al. A randomized controlled effectiveness trial of acceptance and commitment therapy and cognitive therapy for anxiety and depression. *Behav Modif.* 2007;31(6):772–799.

Editor's Note: We encourage authors to submit papers for consideration as a part of our Focus on Childhood and Adolescent Mental Health section. Please contact Karen D. Wagner, MD, PhD, at kwagner@psychiatrist.com.

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Supplementary Material

Article Title: Experiential Avoidance Predicts Persistence of Major Depressive Disorder and Generalized Anxiety Disorder in Late Adolescence

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List of Supplementary Material for the article

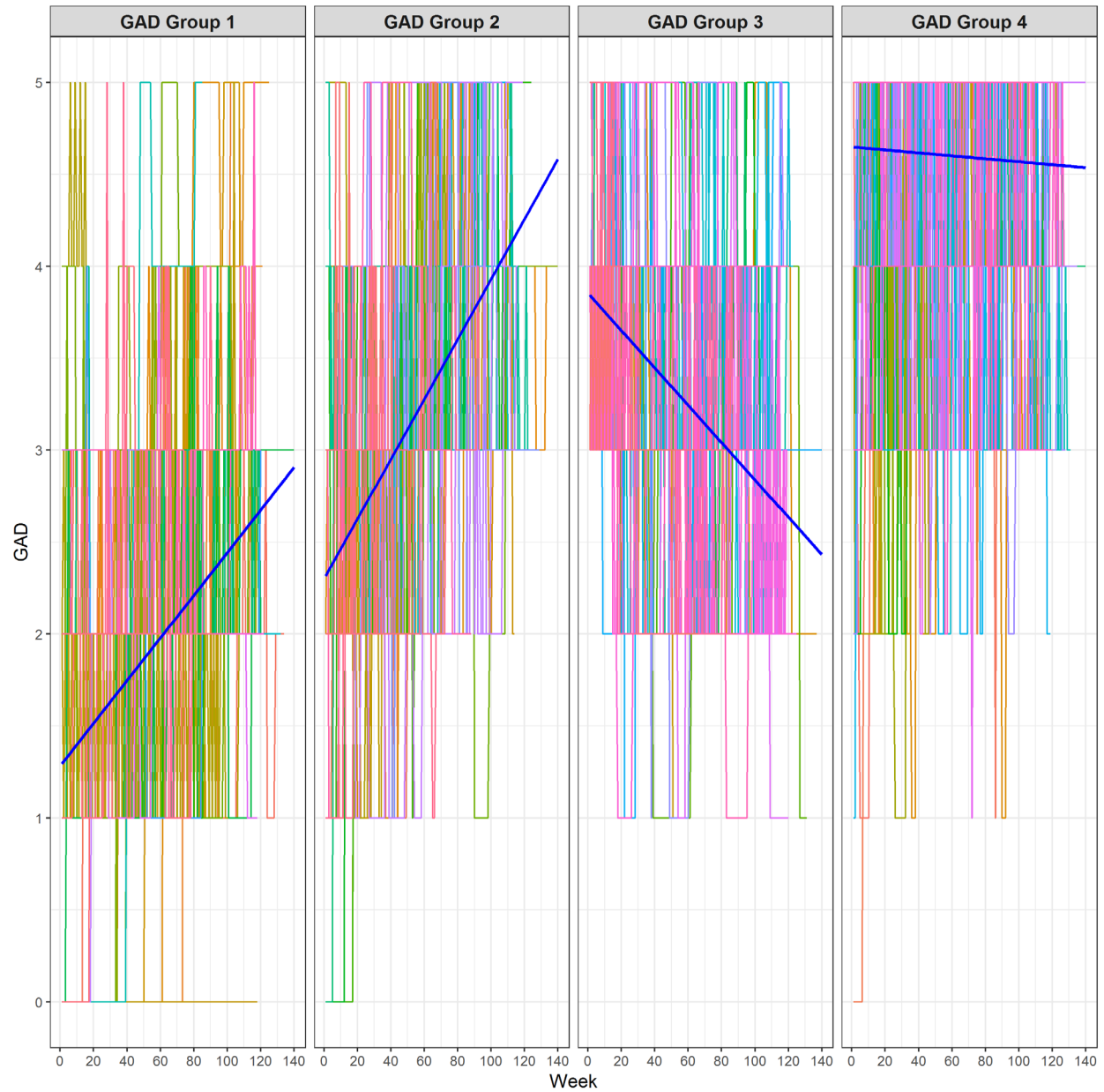
1. [Figure 1](#) 4-group solution for MDD symptom trajectory, with each column representing group depressive symptoms
2. [Figure 2](#) 4-group solution for GAD symptom trajectory, with each column representing group generalized anxiety symptoms

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Supplementary Figure 1. 4-group solution for MDD symptom trajectory, with each column representing group depressive symptoms. Depression = A-LIFE MDD symptom weekly ratings.



Supplementary Figure 2. 4-group solution for GAD symptom trajectory, with each column representing group generalized anxiety symptoms. GAD = A-LIFE GAD symptom weekly ratings.