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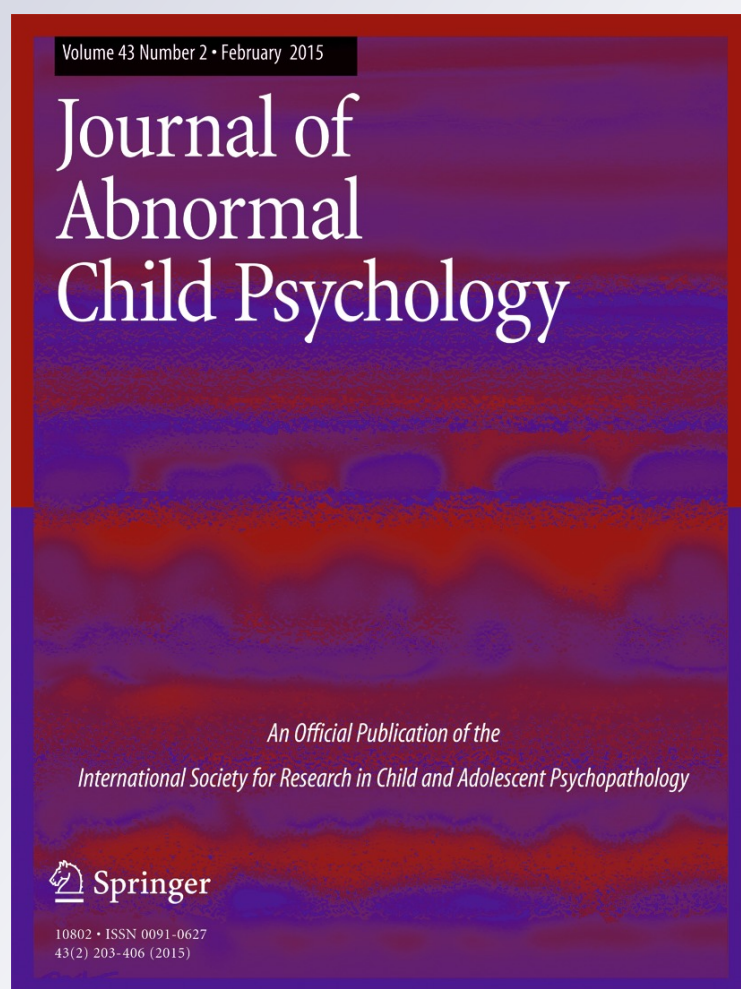
Katie A. McLaughlin & Kevin King

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Developmental Trajectories of Anxiety and Depression in Early Adolescence

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Abstract Adolescence is a period of heightened vulnerability for the onset of internalizing psychopathology. Characterizing developmental patterns of symptom stability, progression, and co-occurrence is important in order to identify adolescents most at risk for persistent problems. We use latent growth curve modeling to characterize developmental trajectories of depressive symptoms and four classes of anxiety symptoms (GAD, physical symptoms, separation anxiety, and social anxiety) across early adolescence, prospective associations of depression and anxiety trajectories with one another, and variation in trajectories by gender. A diverse sample of early adolescents ($N=1065$) was assessed at three time points across a one-year period. All classes of anxiety symptoms declined across the study period and depressive symptoms remained stable. In between-individual analysis, adolescents with high levels of depressive symptoms experienced less decline over time in symptoms of physical, social, and separation anxiety. Consistent associations were observed between depression and anxiety symptom trajectories within-individuals over time, such that adolescents who experienced a higher level of a specific symptom type than would be expected given their overall symptom trajectory were more likely to experience a later deflection from their average trajectory in other symptoms. Within-individual deflections in GAD, physical, and social symptoms predicted later deflections in depressive symptoms, and deflections in depressive symptoms predicted later deflections in GAD and separation anxiety symptoms. Females had higher levels of

symptoms than males, but no evidence was found for variation in symptom trajectories or their associations with one another by gender or by age.

Keywords Adolescence · Anxiety · Depression · Development · Trajectories · Growth curve model

Adolescence is a period of heightened vulnerability for the onset of internalizing psychopathology. The prevalence of depression is only 2.8 % in children under the age of 13 and doubles to 5.6 % among adolescents aged 13–18 (Costello, Erkanli, and Angold 2006). The incidence of major depression peaks between the ages of 15 and 18 (Hankin et al. 1998), and an increase in rates of depressive symptoms among females occurs between the ages of 13 and 15 (Nolen-Hoeksema and Girgus 1994; Twenge and Nolen-Hoeksema 2002). The gender difference in depression is first observable during the early adolescent period and becomes pronounced by late adolescence (Hankin et al. 1998). Onset of many anxiety disorders also occurs during this time period, including social anxiety disorder, panic disorder, and generalized anxiety disorder (GAD) (Beesdo et al. 2007; Cambell, Brown, and Grisham 2003; Last et al. 1992; Wittchen et al. 2000). Adolescent anxiety disorders and depression portend a wide range of negative consequences across the life-course, including risk of recurrent episodes in adulthood (Copeland et al. 2009; Fombonne et al. 2001a; Pine et al. 1998), suicidality (Brent et al. 1993; Fombonne et al. 2001b), and poor psychosocial functioning (Fombonne et al. 2001b; Gotlib, Lewinsohn, and Seeley 1998). Sub-threshold symptoms of anxiety and depression are also associated with heightened risk of later psychopathology (Fergusson et al. 2005; Georgiades, Lewinsohn, Monroe, and Seeley 2006; Pine et al. 1999).

Although adolescence is a period of vulnerability for internalizing psychopathology, most adolescents do not become

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K. A. McLaughlin (✉) · K. King
Department of Psychology, University of Washington, Box351525,
Seattle, WA 98195, Washington
e-mail: mclaughk@uw.edu

clinically anxious or depressed. Characterizing developmental patterns of symptom stability, progression, and co-occurrence is important in order to identify adolescents most at risk for persistent problems. In a five-year prospective study of adolescents followed from age 10 to 18, Hale et al. (2008) found evidence for decreases in symptoms of panic disorder, separation anxiety, GAD, and phobias, with the exception of social phobia symptoms which remained relatively constant (Hale, Raaijmakers, Muris, van Hoof, and Meeus 2008). Similar findings have been observed in several prospective community studies, indicating that most types of anxiety symptoms decrease across early to middle adolescence (Burstein, Ginsburg, Petras, and Ialongo 2010; Crocetti, Klimstra, Keijsers, Hale, and Meeus 2009; Van Oort, Greaves-Lord, Verhulst, Ormel, and Huizink 2009). In contrast, symptoms of depression increase developmentally across this period, particularly for girls (Ge, Conger, and Elder 2001; Twenge and Nolen-Hoeksema 2002; Van Oort et al. 2009), suggesting that anxiety and depression symptoms follow different developmental trajectories during adolescence.

These divergent patterns are perplexing, because symptoms of anxiety and depression are highly co-occurring, and both concurrent and sequential comorbidity between anxiety disorders and major depression are common in children and adolescents (Angold, Costello, and Erkanli 1999; Wittchen et al. 2000). This raises important questions about the relationship between symptoms of anxiety and depression during adolescence. Numerous studies have attempted to disentangle the temporal sequencing of anxiety and depression using a cross-lagged approach to examine whether the presence of anxiety predicts subsequent increases in depression, whether the presence of depression predicts subsequent increases in anxiety, or both. Virtually all of these studies find that children and adolescents with an anxiety disorder are at elevated risk for the onset of major depression and that prediction from anxiety to depression is stronger than the reverse (Costello et al. 2003; Merikangas et al. 2003; Pine et al. 1998; Wittchen et al. 2000). Similar findings have been observed in studies that examine cross-lagged associations between anxiety and depression symptoms. For example, Cole et al. (1998) found that symptoms of anxiety at one time point predicted increases in depressive symptoms at a subsequent time point, but depressive symptoms did not predict increases in anxiety in a sample of third- and sixth-grade children. However, beginning in adolescence major depression is associated in some studies with future onset of anxiety disorders (Fergusson and Woodward 2002; Kim-Cohen et al. 2003), and specifically predicts future onset of GAD (Moffitt et al. 2007; Pine et al. 1998). Bidirectional associations between anxiety and depression symptom trajectories have also been observed during adolescence, such that adolescents with increasing symptoms of anxiety tend to exhibit increasing symptoms

of depression across time, and vice versa (Hale et al. 2009; Leadbeater et al. 2012).

Prior studies have utilized diverse methods for examining the prospective association between anxiety and depression. For example, Cole et al. (1998) tested an auto-regressive cross-lagged panel model, which tested how anxiety and depression at one time point predicted anxiety and depression at the next time point. This model presumes that associations between anxiety and depression occur as a result of a cascading, point-to-point process, such that change in anxiety at a later time point is related to prior status in depression (over and above the prior level of anxiety), or vice versa. On the other hand, Hale et al. (2009) and Leadbeater et al. (2012) tested associations between latent trajectories of anxiety and depression with parallel process growth curve models, which take the same repeated observations of anxiety and depression that could be used in a cross-lagged panel model as indicators of an unobserved latent developmental trajectory. This model assumes that associations between anxiety and depression occur as a result of larger developmental processes, and that anxiety influences depression either because the level of one is associated with the rate of development in the other, or because the rates of development in both are correlated. However, both of these approaches have limitations. Auto-regressive models do not distinguish between and within-individual variance, which means that associations between variables represent a mixture of between-individual differences in symptoms (i.e. because some adolescents have higher anxiety than others over time in general), and within-individual differences (i.e. how deviations from an adolescent's typical anxiety symptoms influence later depression). Parallel process growth models focus *only* on between-individual differences in symptom associations, and ignore within-individual associations.

We argue that examining within-individual associations is equally important as examining between-individual associations. It is often of interest to not only understand *for whom* symptoms change, but also *when* are they expected to change and *why*. Frequently, decisions about when to intervene are drawn from between-individual studies. For example, a trajectory study might show that higher baseline social anxiety is associated with increases in depressive symptoms over time, and an intervention might be developed to reduce social anxiety as a means of preventing increases in depression. However, such studies are making conclusions about within-individual processes (i.e. that decreasing social anxiety will reduce depressive symptoms) from between-individual data. However, analyses that examine how associations vary across time within the same individual provide an opportunity to examine these types of questions more directly (Fleeson 2007). Auto-regressive latent trajectory models (also called ALT models) allow researchers to simultaneously test hypotheses about between- and within-individual differences in

change over time (Bollen and Curran 2004; Curran and Hussong 2003). In other words, they may be used to test whether between-individual differences in the level of anxiety or depression influences the rate of change in the other, as well as how within-individual differences in point-to-point levels of each are associated over time. Prior research has demonstrated the utility of these types of models to examine how environmental or behavioral factors predict trajectory deflections in psychopathology (e.g., King, Molina, and Chassin 2009). Although prior studies have examined latent trajectory models to identify groups of adolescents who follow similar patterns of anxiety disorder and major depression comorbidity over time (Olino et al. 2010) as well as latent growth models to determine how between-individual differences in anxiety and depression symptom trajectories are associated with one another (Hale et al. 2009; Leadbeater et al. 2012), we are unaware of studies that have examined *both* the between-individual and within-individual associations of symptom trajectories with one another. Thus our study was designed to compare these approaches (i.e., approaches examining only between-individual effects, including an autoregressive cross-lagged panel model and a parallel process growth curve model, and the ALT model which examines both between- and within-individual effects) to studying change over time in anxiety and depression in early adolescence. Characterizing anxiety and depression symptom trajectories and their associations with one another over time is a necessary first step before substantive predictors of between- and within-individual differences in these trajectories can be identified.

Importantly, trajectories of anxiety and depression symptoms in early adolescence, as well as their associations with one another, may differ by gender. Evidence from several prospective community studies suggests that cross-lagged associations between anxiety disorders and major depression are stronger for females than males. The continuity of depression, GAD, social phobia, and specific phobia was significantly greater for females than males from age 9 to 16 in the Great Smoky Mountain Study (Costello et al. 2003). Moreover, the presence of an anxiety disorder predicted subsequent onset of depression, and the presence of depression predicted later onset of anxiety disorder across this age range, but only for females (Costello et al. 2003). In the Dunedin Study, strong continuity was observed between internalizing disorders at age 11 and age 15 for females but not males (McGee et al. 1992). A similar pattern was found in a longitudinal community study of late adolescents and early adults, such that the association between anxiety and later onset of depression and between depression and later onset of anxiety was stronger for females than males (Merikangas et al. 2003). Existing evidence thus suggests stronger continuity and symptom progression of anxiety and depression for females than males. However, we are unaware of previous research examining gender differences in the relationship between symptom

trajectories, particularly during the early adolescent risk period.

We address this gap in the literature using data from a three-wave prospective study of early adolescents. First, we examine developmental trajectories of depressive symptoms and four classes of anxiety symptoms (GAD, physical symptoms, separation anxiety, and social anxiety) across early adolescence. Second, we examine how trajectories of anxiety symptoms are related to trajectories of depressive symptoms and whether these relationships vary across anxiety symptom types. We examine three modeling approaches for characterizing longitudinal associations between symptoms of depression and anxiety: auto-regressive, parallel process, and ALT models. Finally, we investigate gender differences in trajectories of anxiety and depressive symptoms and in the covariation of symptom trajectories.

Method

Participants

Adolescents were recruited from the total enrollment of two middle schools (Grades 6–8; ages 10–15) in a small, urban community in central Connecticut with the exception of students in self-contained special education classrooms and technical programs who did not attend school for the majority of the day. The parents of all eligible children ($N=1,567$) in the participating middle schools were asked to provide active consent for their children to participate in the study. Parents who did not return written consent forms to the school were contacted by telephone. We received IRB permission to obtain verbal consent from parents over the phone for their child to participate in the study, as this was the only feasible method for obtaining active parental consent. No differences were observed between children whose parents returned the consent form compared to those who provided verbal consent. Twenty-two percent of parents did not return consent forms and could not be reached to obtain consent, and 6 % of parents declined to provide consent. All youths provided assent for participation. The overall participation rate in the study at Time 1 was 72 % ($N=1,065$). Of participants who were present at Time 1, 217 (20.4 %) did not participate at the Time 2 assessment, and 330 (31.0 %) did not participate at Time 3, largely due to transient student enrollment in this district. Data from the school district indicate that over the 4-year period from 2000–2004, 22.7 % of students had left the district (Connecticut Department of Education, 2006). An additional 121 participants were present at the Time 2 assessment and 40 at the Time 3 assessment who did not participate at Time 1. The total sample at each wave is as follows: Time 1 ($N=1,065$), Time 2 ($N=1,034$), Time 3 ($N=811$). Of these,

1,437 had data on anxiety or depression from at least one observation and were included in analyses.

The Time 1 sample included 51.2 % ($N=545$) boys and 48.8 % ($N=520$) girls. Participants were evenly distributed across grade level (mean age=12.2, $SD=1.0$). The race/ethnicity composition of the sample was as follows: 13.2 % ($N=141$) non-Hispanic White, 11.8 % ($N=126$) non-Hispanic Black, 56.9 % ($N=610$) Hispanic/Latino, 2.2 % ($N=24$) Asian/Pacific Islander, 0.2 % ($N=2$) Native American, 0.8 % ($N=9$) Middle Eastern, 9.3 % ($N=100$) Biracial/Multiracial and 4.2 % ($N=45$) Other racial/ethnic groups. Twenty-seven percent ($N=293$) of participants reported living in single-parent households. The community in which the participating middle schools reside is relatively low SES, with a per capita income of \$18,404 (Connecticut State Department of Education, 2005 based on data from 2001). School records indicated that 62.3 % of students qualified for free or reduced lunch in the 2004–2005 school year. There were no differences across the two schools in demographic variables.

Measures

Anxiety symptoms Anxiety symptoms were assessed using the Multidimensional Anxiety Scale for Children (MASC; March, Parker, Sullivan, Stallings, and Conners 1997) and the Penn State Worry Questionnaire for Children (PSWQ; Chorpita et al. 1997). The MASC is a 39-item widely used measure of anxiety in children. The MASC assesses physical symptoms of anxiety, harm avoidance, social anxiety, and separation anxiety and is appropriate for children ages 8 to 19. Each item presents a symptom of anxiety, and participants indicate how true each item is for them on a four-point Likert scale ranging from never true (0) to very true (3). The MASC has high internal consistency and test-retest reliability across 3-month intervals, and established convergent and divergent validity (Muris et al. 2002). The MASC sub-scales demonstrated good reliability in this sample for physical symptoms ($\alpha=0.78$) and social anxiety ($\alpha=0.81$) and demonstrated adequate reliability for separation anxiety ($\alpha=0.68$).

The PSWQ-C is a 14-item measure that assesses the frequency, severity, and controllability of worry. Items are rated on a 4-point Likert scale ranging from never true (0) to always true (3), with higher scores reflecting greater engagement in worry. This measure was adapted from the adult version of the PSWQ (Meyer et al. 1990) and demonstrates sound psychometric properties, including high internal consistency, excellent test-retest reliability and high convergent and discriminant validity (Chorpita et al. 1997). The PSWQ-C successfully differentiates children with GAD from children without anxiety or mood disorders (Chorpita et al. 1997) and demonstrated good reliability in this sample ($\alpha=0.85$).

Depressive symptoms The Children's Depression Inventory (CDI; Kovacs 1992) is a widely used self-report measure of depressive symptoms in children and adolescents. The CDI includes 27 items consisting of three statements (e.g., *I am sad once in a while, I am sad many times, I am sad all the time*) representing different levels of severity of a specific symptom of depression. The CDI has sound psychometric properties, including internal consistency, test-retest reliability, and discriminant validity (Kovacs 1992; Reynolds 1994). The item pertaining to suicidal ideation was removed from the measure at the request of school officials and the Institutional Review Board (IRB). The 26 remaining items were summed to create a total score ranging from 0 to 52. The CDI demonstrated good reliability in this sample ($\alpha=.82$).

Procedure

Participants completed study questionnaires during their homeroom period. Seven months elapsed between the Time 1 (November 2005) and Time 2 (June 2006) assessments, and an additional 5 months elapsed between Time 2 and Time 3 (November 2006) assessments. Participants who contributed data at any time point were included in the data analysis, and missing data were handled during model estimation (see Analytic Strategy for details). Participants were assured of the confidentiality of their responses and the voluntary nature of their participation. The study was approved by the IRB at Yale University.

Analytic Strategy

All analyses were conducted using MPlus 6.1 (Muthén and Muthén 2008) with the MLR estimator, accounting for missing data by using full information maximum-likelihood estimation (FIML) (Little and Rubin 1987). FIML utilizes all available data, rather than the covariance matrix, to obtain parameter estimates. It is superior to complete case analysis because it provides superior power to detect effects due to the increased sample size, and is robust to missing data when data are missing at random (Schafer and Graham 2002). To test whether varying permutations of each model improved model fit, we used Satorra's chi-square difference tests for MLR chi-square (Satorra 2000). In terms of assessing model fit, researchers frequently cite Hu & Bentler (Hu and Bentler 1999) as providing rules for model fit, when their work more appropriately provides a starting point for determining adequate model fit (Marsh et al. 2004). There has been robust debate about how to utilize different relative fit indices to determine whether a structural equation model reproduces the covariance matrix (Barrett 2007; Millsap 2007). Moreover, recent research has suggested that even the more

stringent relative criteria suggested by Hu and Bentler (1999), such as $RMSEA < .05$, can suggest that a model is well-fitting even when it is not (Chen et al. 2008). Thus we assessed model fit using a combination of chi-square, relative fit indices (including the Tucker Lewis Index (TLI), comparative fit index (CFI), the root-mean square error of approximation (RMSEA), the Bayesian Information Criteria (BIC), model residuals, and model modification indices.

We analyzed the associations between anxiety and depression symptoms in four series of models, one for each class of anxiety symptoms (GAD, physical symptoms, separation anxiety, and social anxiety). Model fitting proceeded as follows. First, we first estimated unconditional growth curve models to understand how symptoms of anxiety and depression changed across the year of assessments. Next, we estimated a series of bivariate models to understand how anxiety and depression symptoms covaried across time. The first model was an autoregressive cross-lagged panel model, with the cross-lagged associations first fixed, and then freed across time. Next we tested a parallel process growth curve model, with the intercept fixed to Time 1. Finally, we tested an ALT model (Bollen and Curran 2004), which added cross-lagged associations among the residual variances. We did not add autoregressive associations among the residuals because not all authors agree that these are necessary, in part because these effects imply some change process over and above what is captured by the latent growth trajectory (Voelkle 2008). We compared these models to determine which produced the best fit to the data. Then we added gender and single-parent household status to the final model to control for their effects on the model, and added age to control for cohort effects and tested whether the estimated parameters in each final model were equal across gender using the Wald test for parameter constraints.

Results

Attrition

Analyses comparing participants who completed all assessments to those who did not revealed that participants who completed the Time 1 but not the Time 2 assessment were more likely to be female, $\chi(1)^2 = 6.85$, $p < .01$, and from a single-parent household, $\chi(1)^2 = 8.93$, $p < .01$, but did not differ in grade level, race/ethnicity, Time 1 anxiety, or Time 1 depression (p -values > 0.10) from those who completed both assessments. Participants who were lost to follow-up by the Time 3 assessment were more likely to be from a single-parent household, $\chi(1)^2 = 26.92$, $p < .001$, and had higher Time 1 symptoms of depression, $F(1, 1065) = 9.63$, $p = .002$, than adolescents who were present at all three assessments but

did not differ in gender, race/ethnicity, or Time 1 anxiety symptoms.

Descriptive Statistics

The means and standard deviations of each type of anxiety symptom and depressive symptoms are shown in Table 1, separately for males and females.

Unconditional Growth Curve Models

We first examined average patterns of change in anxiety and depression symptoms across one year of early adolescence (measured at three time points: month 0, month 7 and month 12), and included age and gender as predictors of slopes and intercepts. Table 2 summarizes these findings.

Depressive Symptoms On average, although the slope of depressive symptoms was positive, we did not observe statistically significant change in symptoms over the observed time period. However, there was significant variability across adolescents in the rate of change over time, with some adolescents exhibiting increases in depressive symptoms and others exhibiting decreases over time. An adolescent whose growth in depressive symptoms was 1 standard deviation above the mean rate of change would have reported a CDI score by the end of the year that was nearly 12 points higher than their Time 1 score. Finally, the slope and intercept of depression were uncorrelated, such that initial level of depressive symptoms was unrelated to the rate of change over time.

Anxiety Symptoms All four types of anxiety symptoms decreased, on average, across the year of assessment. Adolescents differed in their rate of decline, however, such that some adolescents declined more than others, and some adolescents experienced increases in symptoms over the course of the year. The intercepts and slopes of each type of anxiety symptoms were negatively correlated, meaning that adolescents with higher initial levels of anxiety symptoms at Time 1 exhibited greater declines over the course of the year. The one exception was social anxiety symptoms, where the intercept and slope were only marginally negatively associated.

Prospective Associations Between Anxiety and Depressive Symptoms

We next tested a sequence of models examining the prospective association between anxiety and depressive symptoms across the one year study period, estimating an unconditional autoregressive panel model (AR), a parallel process growth model, and an ALT model in turn, separately for the association between depression and each type of anxiety symptom.

Table 1 Means and standard deviations of anxiety and depressive symptoms by gender

	Total sample		Males		Females		Gender difference
	Mean	SD	Mean	SD	Mean	SD	t-value
Time 1							
Depressive Symptoms	9.67	(6.44)	9.11	(6.37)	10.25	(6.47)	2.88*
GAD Symptoms	14.62	(7.38)	13.12	(6.84)	16.10	(7.61)	6.50*
Social Anxiety Symptoms	8.48	(5.63)	7.22	(5.31)	9.83	(5.67)	7.75*
Separation Anxiety Symptoms	6.61	(4.51)	5.71	(4.20)	7.56	(4.64)	6.80*
Physical Anxiety Symptoms	9.22	(5.64)	8.25	(5.36)	10.25	(5.77)	5.88*
Time 2							
Depressive Symptoms	10.64	(8.15)	9.70	(8.18)	10.73	(7.74)	1.90 ⁺
GAD Symptoms	14.20	(7.21)	12.85	(6.94)	15.03	(7.33)	4.37*
Social Anxiety Symptoms	7.55	(5.96)	6.26	(5.69)	8.67	(5.95)	6.05*
Separation Anxiety Symptoms	5.76	(4.77)	5.08	(4.92)	6.28	(4.58)	3.71*
Physical Anxiety Symptoms	8.32	(6.51)	7.20	(6.45)	9.10	(6.38)	4.33*
Time 3							
Depressive Symptoms	9.41	(7.85)	9.07	(8.24)	9.70	(7.62)	0.95
GAD Symptoms	13.05	(6.41)	12.00	(6.10)	13.99	(6.74)	3.67*
Social Anxiety Symptoms	6.74	(5.91)	5.70	(5.46)	7.81	(6.28)	4.39*
Separation Anxiety Symptoms	5.48	(4.73)	5.07	(5.01)	5.84	(4.67)	1.93
Physical Anxiety Symptoms	7.69	(6.35)	6.67	(6.29)	8.32	(6.45)	3.12*

Significant at the .05 level, 2-sided independent samples *t*-test, Coefficient approached significance at the .10 level, 2-sided independent samples *t*-test

The autoregressive panel models and the ALT models both provided adequate to good fit to the data, while the fit of the parallel process models were poor. Moreover, the fit of the ALT models were all substantially improved with the addition of the set of covariates, while the fit of the AR models were

somewhat less improved. We also tested whether freeing the cross-lags in the ALT and AR models improved model fit, and in all cases model fit was not significantly improved. By multiple fit criteria, both the unconditional and conditional ALT models fit the data better than the AR models and the parallel process models (see Supplementary Online Table 1). Thus, we describe the ALT model throughout the remainder of the results. For parsimony, the final ALT models were specified with covariance of slope and intercept of depression and the effects of the covariates on the slope of depression (all non-significant) fixed to zero. Doing this did not result in worsened model fit by BIC, model residuals or any other indicators of fit, and provided increased degrees of freedom. Results from the final models are shown in Table 3.

Table 2 Unconditional Univariate Growth Model Mean and Variance Estimates

	Mean	SE	Variance	SE
Depressive Symptoms Intercept	9.85	0.19	32.59	4.21
Depressive Symptoms Slope	0.10 ⁺	0.14	9.01	2.31
Intercept-Slope Covariance	-4.16 ⁺	2.70	–	–
General Anxiety Disorder Intercept	14.78	0.22	28.67	4.37
General Anxiety Disorder Slope	-0.80	0.14	4.07	2.19
Intercept-Slope Covariance	-5.48	2.76	–	–
Social Anxiety Symptoms Intercept	8.46	0.16	20.01	2.12
Social Anxiety Symptoms Slope	-0.90	0.10	4.39	1.15
Intercept-Slope Covariance	-2.37 ⁺⁺	1.28	–	–
Separation Anxiety Symptoms Intercept	6.58	0.13	13.88	1.39
Separation Anxiety Symptoms Slope	-0.65	0.09	3.91	0.78
Intercept-Slope Covariance	-3.28	0.87	–	–
Physical Anxiety Symptoms Intercept	9.27	1.36	21.56	2.24
Physical Anxiety Symptoms Slope	-0.79	0.11	4.67	1.36
Intercept-Slope Covariance	-2.73	1.36	–	–

All coefficients were significant except, Coefficient was not significant, $p > .10$, 2-sided test, Coefficient approached significance, $p < .10$, 2-sided test

GAD Variance in the slope of GAD symptoms was marginally related to initial levels of depressive symptoms, suggesting that adolescents who reported greater depression symptoms at the first interview increased somewhat less on GAD symptoms over time.

Within time, the residual variances of GAD and depressive symptoms were moderately correlated ($r = .32 - .50$, $p < .001$), indicating that within-individual differences, or deviations from trajectories at specific time points, were associated. There were also bidirectional associations among these residuals, such that elevations in depressive symptoms over and above an adolescent's average trajectory at one time point were associated with time-specific increases in GAD

Table 3 Trajectory level and time specific associations between anxiety symptoms and depressive symptoms

Depressive Symptoms	GAD		Physical Symptoms		Separation Anxiety		Social Anxiety	
	Estimate	SE	Estimate	SE	Estimate	SE	Estimate	SE
Between-individual regression effects								
Depression Intercept predicting Anxiety slope	−0.08+	0.05	0.15*	0.05	0.13*	0.04	0.11*	0.04
Anxiety intercept Depression predicting slope	−0.04*	0.05	0.03	0.07	−0.07	0.06	−0.03	0.06
Within-individual regression effects								
Depression Anxiety	0.10*	0.04	0.04	0.04	0.02	0.03	0.04	0.03
Anxiety Depression	0.11*	0.03	0.17*	0.05	0.19*	0.06	0.10*	0.04
Between-individual residual covariances								
Anxiety slope/intercept	−4.55+	2.51	−2.54	1.17	−2.87*	0.90	−2.58*	1.21
Depression slope/intercept	0.00	0.00	0.00	0.00	0.00	0.00	0.00	0.00
Intercepts covariance	9.59*	2.54	7.18*	7.56	0.47	1.88	4.30*	2.00
Slope covariance	−1.32	1.03	−0.69	1.27	−0.52	0.81	−0.16	0.96
Within-individual covariance								
Anxiety with depression	9.21*	1.06	8.52*	2.20	5.21*	1.62	5.94*	1.59

Significant at the .05 level, 2-sided test, controlling for age, gender and single parent household, Coefficient approached significance, $p < .10$ level, 2-sided test, controlling for age, gender and single parent household

symptoms in the next assessment ($\beta = .10, p < .01$), and vice-versa ($\beta = .09, p < .01$). Fig. 1 (Panel C) illustrates this effect.

Age ($\beta = .13, p < .01$), gender ($\beta = .10, p < .01$) and household ($\beta = .12, p < .01$) were all associated with the intercept of depression, such that older adolescents, females, and those from single-parent households reported more depressive symptoms at the initial time point. The covariates were unrelated to the slope or intercept of GAD symptoms.

Social anxiety symptoms Among the latent growth factors, the intercepts of depression and social anxiety symptoms were correlated ($r = .20, p < .01$), but the correlation between their growth factors was not significant ($r = -.04, p = .87$). Higher initial levels of depressive symptoms predicted growth in social anxiety symptoms over time ($\beta = 0.28, p = .01$). Because social anxiety symptoms declined on average over time, we estimated the predicted slope of social anxiety symptoms at low, mean and high levels of initial depressive symptoms to better understand this effect. An intercept is estimated for each parameter in a growth curve model, providing an estimate of the mean level when all predictors of that parameter are at zero. Thus, for this model, the intercept of the slope of social anxiety is interpreted as the expected rate of change in social anxiety symptoms for an adolescent with no initial depressive symptoms. This intercept was -1.14 (which corresponds to a 2.28 point decline in social anxiety across the course of the year). We estimated the predicted slope of social anxiety at mean levels of depressive symptoms and at 1 standard deviation above the mean. The predicted slope of social anxiety at mean levels of depression was estimated at -0.47 and at 1 SD above the mean of initial depression was -0.03 . In other words, higher initial depressive

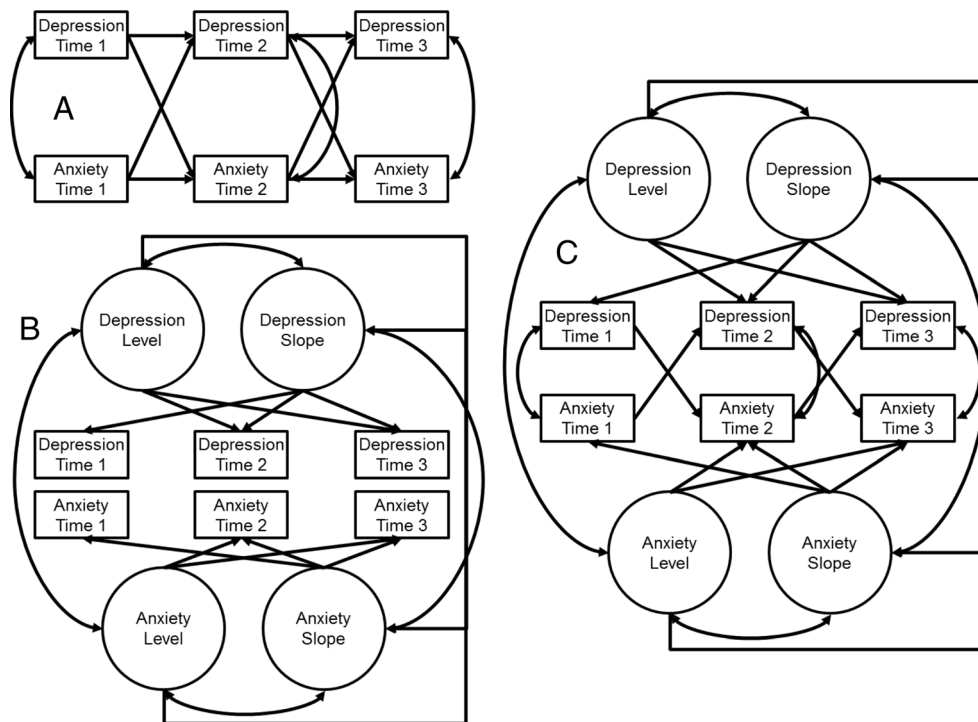
symptoms were related to a trend towards a slower rate of decline in social anxiety symptoms over time.

Within time, the residual variances of social anxiety symptoms and depressive symptoms were moderately correlated ($r = .23 - .47, p < .001$). There were also prospective associations among the residuals, such that elevations in social anxiety symptoms over and above the average trajectory in one year were associated with time-specific increases in depressive symptoms in the next year ($\beta = .07, p < .01$), but not vice-versa (See Fig. 2).

Females reported higher levels of social anxiety symptoms at the first time point, ($\beta = .28, p < .01$), and adolescents from single-parent households exhibited greater increases in social anxiety over time compared to those living with two caregivers ($\beta = .12, p < .05$).

Separation anxiety symptoms Among the latent growth factors of depression and separation anxiety symptoms, the intercepts were uncorrelated ($r = .02, p = .80$), as were the slopes ($r = -.14, p = .55$). Higher depressive symptoms at Time 1 were associated with changes in separation anxiety symptoms over time ($\beta = .36, p < .01$), but the initial level of anxiety was not associated with changes in depressive symptoms over time. As above, we estimated the predicted separation anxiety slope at low, average and high levels of initial depressive symptoms. At low levels of initial depressive symptoms, separation anxiety symptoms declined the most, and the slope became less steep as depressive symptoms increased. Finally, initial levels of separation anxiety symptoms were associated with the rate of change in separation anxiety, ($r = -.48, p < .001$), such that adolescents with the highest levels of initial symptoms showed the greatest declines over time.

Fig. 1 Illustration of the three unconditional models tested: **a)** autoregressive cross-lagged panel model; **b)** parallel process growth curve model; and **c)** autoregressive latent trajectory (ALT) model

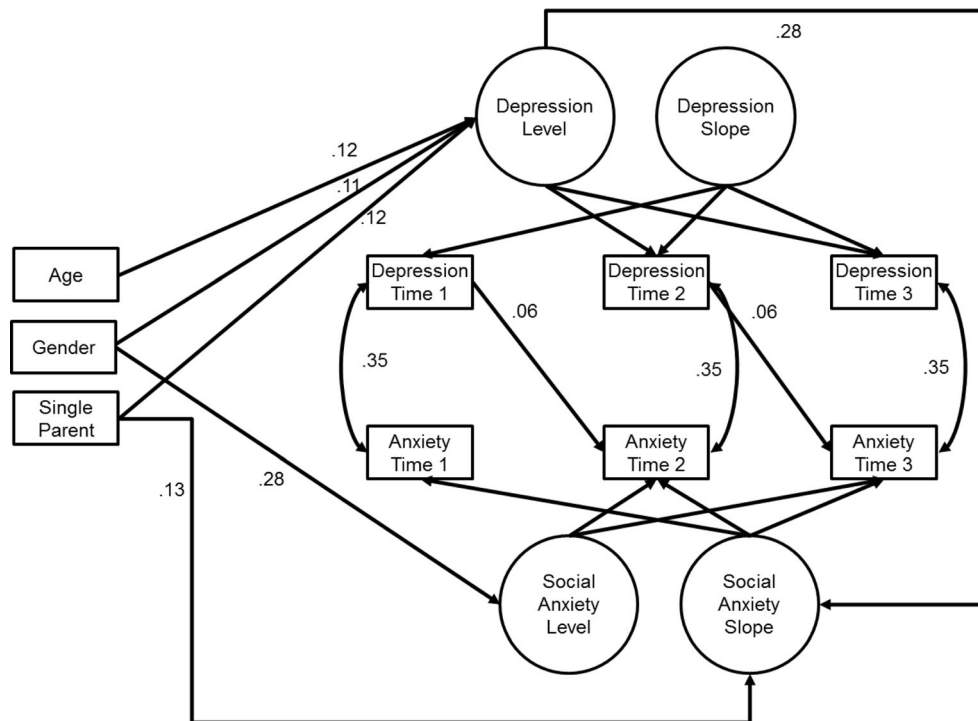


Within time, the residual variances of separation anxiety and depressive symptoms were moderately correlated ($r=.28 - .51, p<.001$), indicating that within-individual differences were associated. Moreover, elevations in separation anxiety symptoms over and above the average trajectory were associated with deflections

from depressive trajectories ($\beta=.11, p<.01$), but not vice versa.

Females reported higher levels of separation anxiety symptoms at baseline, ($\beta=.24, p<.01$), as did younger adolescents ($\beta=-.17, p<.001$). Moreover, males reported slower decreases (because the average slope was

Fig. 2 Illustration of the conditional ALT model of depression and social anxiety over 1 year. Non- significant paths and residual covariances among latent variables among latent variables are not shown for the sake of simplicity



negative) in separation anxiety over time ($\beta = -.18$, $p < .01$).

Physical anxiety symptoms Among the latent growth factors of depression and physical anxiety symptoms, the intercepts were correlated ($r = .37$, $p < .001$), but the slopes were not ($r = -.26$, $p = .64$). Higher initial levels of depressive symptoms predicted slower declines in physical anxiety symptoms over time ($\beta = 0.43$, $p < .05$). We estimated predicted growth in physical anxiety symptoms in a manner similar to our approach for social and separation anxiety symptoms. High initial depressive symptoms were related to a slower rate of decline in physical anxiety symptoms over time.

Within time, the residual variances of GAD and depressive symptoms were moderately correlated ($r = .34 - .45$, $p < .001$), indicating that deviations from average trajectories at specific time points were associated. There were also prospective associations among the residuals, such that elevations in physical anxiety symptoms over and above the average trajectory in one year were associated with time-specific increases in depressive symptoms in the next year ($\beta = .13$, $p < .001$), but not vice-versa.

Females reported higher levels of separation anxiety symptoms at the first time point, ($\beta = .21$, $p < .001$), but covariates were unassociated with change in physical anxiety over time.

Moderation by Gender

Finally, we tested whether any of the associations among the latent growth factors or the residuals differed by gender. For these analyses, we allowed the factor means and variances to differ across gender (to allow for gender differences in trajectories, reported above), and compared models where all correlational and regression parameters of interest were constrained to be equal across gender with a fully freed model. We followed an omnibus model testing strategy, comparing the fit of models where all parameters were allowed to vary across gender with models where all parameters were forced to be equal. This reduced the number of statistical tests and the chance for Type 1 error. When the omnibus test was significant, we examined each specific parameter in turn to determine which parameter differed by gender. Controlling for age and single parent status, we found no evidence of gender moderation that reached the .05 significance level at either the between or within individual level.

Discussion

Adolescence is a period of substantial vulnerability with regards to internalizing psychopathology, particularly major

depression. Although anxiety and depression are strongly associated with one another during this developmental period, few studies have characterized the dynamics of anxiety and depression symptom trajectories and their associations with one another across early adolescence. Previous research suggests that adolescents with an anxiety disorder are at elevated risk for future depression (Costello et al. 2003; Merikangas et al. 2003; Pine et al. 1998; Wittchen et al. 2000), and in some studies, major depression is associated with later risk for anxiety disorder onset, particularly GAD (Fergusson and Woodward 2002; Kim-Cohen et al. 2003; Moffit et al. 2007; Pine et al. 1998). Recent work has also examined latent growth trajectories of anxiety and depressive symptoms over time, using approaches that examine between-individual cross-lagged associations (Cole et al. 1998; Hale et al. 2009; Leadbeater et al. 2012). We extend these previous studies by using latent growth curve modeling to characterize trajectories of anxiety and depression symptom change across early adolescence, investigate the associations between anxiety and depressive symptom trajectories across time, and evaluate the degree to which within-individual deviations from average trajectories in anxiety and depression are related to one another. Our findings reveal a more complex set of interrelationships between anxiety and depressive symptoms in early adolescence than has previously been described.

Our first goal was to examine developmental trajectories of anxiety and depressive symptoms. We found that most adolescents experienced declines in anxiety symptoms across early adolescence. This pattern was observed for symptoms of GAD, social anxiety, separation anxiety, and physical anxiety. These findings are consistent with several recent prospective studies that have also observed declines in anxiety symptoms over the early adolescent period (Burstein et al. 2010; Crocetti et al. 2009; Hale et al. 2008; Van Oort et al. 2009). Declines in symptoms of separation anxiety over time were particularly pronounced for adolescents with high initial levels of separation anxiety symptoms. In contrast, we found no change in the mean level of depressive symptoms over the one-year study period. Although developmental increases in symptoms of depression during early to mid-adolescence have been frequently documented (Ge et al. 2001; Twenge and Nolen-Hoeksema 2002; Van Oort et al. 2009), a similar pattern of non-significant changes in adolescent-reported depressive symptoms was reported in a school-based sample of early adolescents followed for several years (Garber, Keiley, and Martin 2002), and other studies of early adolescents have also found stability in adolescent reports of depressive symptoms (Garrison et al. 1990). One explanation for the stability in depressive symptoms observed here is that our early adolescent sample has yet to enter the risk window when depressive symptoms begin to increase more drastically in mid-adolescence (Ge et al. 2001; Hankin et al. 1998).

Our second objective was to investigate associations between anxiety and depressive symptoms, both between individuals and within individuals, over time. In between-individual analysis, adolescents with higher initial levels of depressive symptoms exhibited slower declines in physical, separation and social anxiety symptoms over time, above and beyond the effects of age and gender. This means that experiencing depressive symptoms appears to arrest declines in multiple forms of anxiety symptoms over time. Slower declines in physical anxiety symptoms among adolescents with high levels of depressive symptoms may reflect overlap in symptoms of anxiety and depression (e.g., psychomotor agitation) or be the result of cognitive processes, such as anxiety sensitivity. High levels of anxiety sensitivity have been observed in adolescents with depressive symptoms (Weems et al. 1997), and consistent evidence suggests that anxiety sensitivity is associated with vulnerability to anxiety pathology in youths, particularly to panic attacks and other physical manifestations of anxiety (Lau, Calamari, and Waraczynski 1996; Pollock et al. 2002; Weems, Hammond-Laurence, Silverman, and Ginsburg 1998). Depressive symptoms may influence social anxiety trajectories in adolescence because of the strong influence of peer relationships on social anxiety during this developmental period. Affiliation with a peer group, positive relationships with a best friend, and engagement in dating relationships all protect against social anxiety in early adolescence (LaGreca and Harrison 2005), and adolescents with high levels of depressive symptoms engage in a variety of negative interpersonal behaviors—such as reassurance seeking—that generate stressors in their interpersonal relationships and result in diminishing friendship quality over time (Joiner et al. 1992; Prinstein et al. 2005). Similar social mechanisms may also explain the relationship between symptoms of depression and slower decline in separation anxiety symptoms during early adolescence, although this possibility remains to be examined empirically. The identification of cognitive (e.g., rumination; McLaughlin and Nolen-Hoeksema 2011), affective (e.g., chronic negative affect; Chorpita 2002), environmental (e.g., trauma exposure, stressful life events; Rudolph et al., 2005), and neurobiological factors (e.g., pubertal timing; Graber et al. 2004) that underlie transitions between anxiety and depression symptoms during early adolescence is an important direction for future research.

A more nuanced pattern emerged when we examined the associations between anxiety and depression symptom trajectories within individuals over time, an approach that has not previously been used to investigate the development of internalizing psychopathology during early adolescence. Our findings argue for the importance of such an approach. The most consistent associations between symptoms of anxiety and depression in our sample were observed in within-individual analysis—in other words, when adolescents experienced a

higher level of a specific type of symptom at a particular time than would be expected given their overall symptom trajectory, they were also likely to experience a later deflection from their average trajectory in other symptoms. Adolescents who experienced an elevation in symptoms of social anxiety and physical anxiety relative to their average trajectory experienced subsequent deflections from their average trajectory of depressive symptoms, but not vice versa. The association between elevations in social anxiety and future elevations in depressive symptoms is consistent with previous between-individual prospective studies showing that adolescent social anxiety disorder predicts the future onset of major depression and poor prognosis of depression recovery (Beesdo et al. 2007; Stein et al. 2001). Panic disorder is also associated with the later onset of major depression (Gorman and Coplan 1996; Roy-Byrne et al. 2000). However, we are unaware of previous studies linking within-individual elevations in social and physical anxiety symptoms to later elevations in depressive symptoms. In contrast, deflections from one's average trajectory of depressive symptoms predicted later deflections in separation anxiety symptoms, but not vice versa, suggesting greater avoidance of social situations outside the family among adolescents with higher levels of depressive symptoms at a particular point in time than is typical for them. Youths with high levels of depressive symptoms are more likely to be the targets of peer victimization and bullying than those with low levels of depression (Hodges et al. 1997; Vernberg 1990), which might be one mechanism explaining greater reluctance to engage in social activities outside the family among adolescents with elevated depressive symptoms. Future research is needed to identify mechanisms linking depressive symptoms to future elevations in separation anxiety. Finally, we observed bidirectional associations between within-individual elevations in symptoms of GAD and depression. Previous studies in both adolescents and adults indicate that depression is associated more strongly with future GAD than with other anxiety disorders (Moffitt et al. 2007; Pine et al. 1998), and our results are consistent with this pattern. Together, these findings highlight the importance of identifying social, environmental, and intrapersonal factors that result in changes in symptom levels for an adolescent relative to his/her typical symptom trajectory and developing strategies to determine when such deflections have occurred in order to deliver interventions at those points when adolescents are most vulnerable to accelerating increases in symptomatology. They further suggest that even small increases in depression and anxiety symptoms during early adolescence might be important clinically, as they can be early signs of a trajectory of increasing symptom over time.

Our third goal was to examine gender differences in both developmental trajectories of anxiety and depressive symptoms and associations of symptoms with one another across

time. In this sample, gender differences in overall levels of symptoms were more pronounced than differences in the associations of anxiety and depression across time. Generally, females reported higher levels of both depression and anxiety of all types, but there were no differences in the rates of change in depression or anxiety over time. Thus, across at least this one-year period in early adolescence, males and females had parallel trajectories of internalizing symptoms, with females reporting higher levels of symptoms. Several previous prospective epidemiological studies of youth psychopathology have reported greater heterotypic continuity between anxiety disorders and major depression (i.e., stronger prospective associations between anxiety disorders and subsequent major depression) in females as compared to males (Costello et al. 2003; McGee et al. 1992; Merikangas et al. 2003). At the level of symptom trajectories, we found no gender differences in the prospective associations between anxiety and depression, in contrast to what has been reported in previous studies. This might be due to the fact that our follow-up period was shorter than in previous longitudinal studies or because our sample was younger, on average, than in some previous studies that have observed stronger associations between depression and anxiety for females than males (e.g., Merikangas et al. 2003). Perhaps most notably, these previous studies focused on diagnostic outcomes, whereas our analyses focus on associations of symptom trajectories.

These findings should be interpreted in light of several study limitations. First, our analysis focused on self-reported symptoms of anxiety and depression rather than diagnoses of anxiety disorders or major depression. Although the validity of the self-report measures used in this study is well-established (Timbremont et al. 2004; Wood et al. 2002), future research is needed to evaluate the relationships between onset, persistence, and recurrence of anxiety disorders and major depression during early adolescence. Second, adolescents with high levels of depressive symptoms at Time 1 were more likely to be lost to follow-up by the Time 3 assessment than adolescents with lower levels of symptoms. This likely contributed to our finding that the average level of depressive symptoms did not change over time in this sample. It is possible that we would have observed greater increases in depressive symptoms over time if we had been able to assess the entire Time 1 sample at both of the follow-up assessments. Moreover, our prospective models cover a relatively small age span, and only include two follow up time points. This precluded testing developmental hypotheses about sensitive periods for the association between depressive and specific forms of anxiety symptoms, including whether the association is stronger earlier or later in adolescence. The study design also limits the scope of what we may conclude about the associations among anxiety and depression symptoms to the period of early adolescence. Finally, we were unable to collect information about pubertal stage in this sample. Early

developmental timing of puberty has been shown to predict depression in females (Graber et al. 2004). The degree to which developmental timing of puberty influences both the trajectories of anxiety and depressive symptoms across adolescence, and their association with one another, is an important topic for future research.

Although early adolescence is a developmental period of increasing risk for major depression, most adolescents experience declines in anxiety symptomatology during this period. Study findings suggest that these declines over time are less marked for adolescents with high levels of depressive symptoms. Associations between symptoms of depression and anxiety over time were particularly strong when examined within individuals over time, such that adolescents with higher levels of anxiety or depressive symptoms at a particular time than would be expected given their overall symptom trajectory experienced subsequent elevations in other types of internalizing symptoms. These findings suggest that symptom elevations at one point in time can lead to a cascade of increases in other types of internalizing symptoms over time and highlight the importance of identifying these early shifts in symptom trajectories in order to prevent symptom acceleration and, ultimately, the onset of anxiety and mood disorders.

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Conflict of Interest The authors declare that they have no conflict of interest.

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