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Joint trajectories of cigarette smoking and depressive symptoms from the mid-twenties to the mid-thirties predicting generalized anxiety disorder

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Abstract

The current study examines the longitudinal patterns of both cigarette smoking and depressive symptoms as predictors of generalized anxiety disorder (GAD) using data from the Harlem Longitudinal Development Study. There were 674 African American (53%) and Puerto Rican (47%) participants. Among the 674 participants, 60% were females. In the logistic regression analyses, the indicator variables of membership in each of the joint trajectories of cigarette smoking and depressive symptoms from the mid 20s to the mid 30s were used as the independent variables, and the diagnosis of GAD in the mid 30s was used as the dependent variable. The high cigarette smoking with high depressive symptoms group and the low cigarette smoking with high depressive symptoms group and the low cigarette smoking with high depressive symptoms group. The findings shed light on the prevention and treatment of GAD.

Keywords

generalized anxiety disorder; cigarette smoking; depressive symptoms; trajectory analysis; longitudinal study

Generalized anxiety disorder (GAD) has a number of negative consequences as noted below.^{1, 2} GAD has a long average duration of symptoms^{3, 4} and is associated with a significant impairment of quality of life or disability.⁵ The Christchurch Health and Development Study, a 25-year longitudinal study of over 1,000 participants, found that GAD increased the odds of suicidal ideation by a factor of 8.0 times and increased the rate of suicide attempts by a factor of 5.9 times.¹ In addition, exaggerated amygdala activation in

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response to fearful⁶ and masked angry facial expressions⁷ and during the anticipation of aversive photographs⁸ has been reported in patients with GAD. Also, a study using a clinical sample of 31 women (16 patients with GAD and 15 healthy control participants) reported that patients with GAD had a larger volume of the amygdala and dorsomedial prefrontal cortex,² both of which have been correlated with psychological symptoms.⁹

One of the important and common risk factors for GAD is cigarette smoking.^{10–12} About 15% of the ever-smokers¹³ compared with about 3% among the general population¹⁴ in the United States have had at least one anxiety disorder in their lifetime. The symptoms of general anxiety were more severe in nicotine-dependent smokers than in never-smokers, former smokers, and non-dependent smokers.¹⁵ Self-administration of nicotine via smoking may induce anxiety, as dysphoria is one of the observed pharmacologic effects of nicotine.¹⁶ The National Comorbidity Survey Replication research found that GAD is significantly associated with smoking behaviors including daily smoking, heavy smoking, and nicotine dependence.¹⁷

Comorbidity of anxiety symptoms and depressive symptoms exists.^{18, 19} The Dunedin birth cohort (N=1,037) study reported 12% of adults experience comorbidity between GAD and major depressive disorders.¹⁸ Numerous studies have reported a positive relationship between depressive symptoms and anxiety symptoms.^{20, 21} Norton²¹ reported that the correlation between depression and anxiety was higher than 0.7 in a sample of college students. **This may be that pathological worry also occurs in depression**²² **and that active and passive rumination is involved in anxiety**.²³ Hermans and Evenhuis²⁴ found that increased depressive symptoms were positively associated with increased anxiety symptoms using a cohort living in the Netherlands.

The present study seeks possible precursors of GAD given a number of negative consequences.^{1–5} Both smoking cigarettes¹⁰ and depressive symptoms¹⁸ can be the important possible risk factors for GAD. Furthermore, the two possible risk factors (cigarette smoking and depressive symptoms) commonly co-occur.^{25–28} However, only a few studies have focused on the joint trajectories of cigarette smoking and depressive symptoms as a risk factor for adverse conditions.²⁹ Leventhal and Zvolensky (2015) documented that cigarette smoking, depressive symptoms, and GAD are linked together.³⁰ The current study examines the earlier patterns of cigarette smoking and depressive symptoms simultaneously and their association with later GAD.

Our study is unique in three ways. First, we identify the joint trajectories of cigarette smoking and depressive symptoms occurring simultaneously among the relatively understudied ethnic groups of African Americans and Puerto Ricans living in an urban area. Second, we investigated the association between the earlier joint trajectories of cigarette smoking and depressive symptoms and later GAD. Third, we followed the young adult sample from mean age 14 to mean age 36.³¹

We hypothesize that there will be at least four joint trajectory groups **based on all possible combinations with two levels (high and low) in each** of cigarette smoking and depressive symptoms: a) a high level of both cigarette smoking and depressive symptoms trajectory

group, b) a high level of cigarette smoking and a low level of depressive symptoms trajectory group, c) a low level of cigarette smoking and a high level of depressive symptoms trajectory group, and d) a low level of both cigarette smoking and depressive symptoms trajectory group. We also hypothesize that **the trajectory group with higher levels of both cigarette smoking and depressive symptoms as well as the trajectory groups with a higher level of either cigarette smoking or depressive symptoms** will be associated with an increased likelihood of having GAD as compared with a low level of both cigarette smoking and depressive symptoms trajectory group.

Methods

Study Design

Data from the Harlem Longitudinal Development Study were first collected in 1990 (time 1; T1, N=1,332) when the participants were students (grades 7 to 10) attending schools in the East Harlem area of New York City. Data were collected in person or by phone by the National Opinion Research Center at time 2 (T2; 1994–1996; N=1,190). **The attrition rate at T2 as compared to T1 was 89%**. The Survey Research Center of the University of Michigan collected the data at time 3 (T3; 2000–2001; N=662). **At T3, a sub-sample was taken from the T2 sample due to budgetary limitations. The attrition rate at T3 as compared to T2 was 56%**. The data were collected by our research group: in person, by phone, or by mailed questionnaire at time 4 (T4; 2004–2006; N=838) and by mailed questionnaire at time 5 (T5; 2011–2013; N=674). **The attrition rates at T4 as compared to T3 and at T5 as compared to T4 were 127% and 80%, respectively**. For the interview in person or by phone, each participant was interviewed by an interviewer of the same sex and ethnic background. For the sections regarding drug use and possible involvement with the police, the participant marked his or her own responses directly on the questionnaire.

Participants

The present study (N=674; 53% African Americans, 47% Puerto Ricans) is based on time waves 1–5 of the Harlem Longitudinal Development Study. Sixty percent were females (n=405). The mean age of the participants at T1 was 14.1 years (Standard Deviation; SD=1.4 years). The mean age of the participants at this wave was 19.2 years (SD=1.5 years). At T3, the mean age of the participants was 24.4 years (SD=1.3 years). At T4, the mean age of the participants was 29.2 years (SD=1.4 years). At T5, the mean age was 35.9 years (SD=1.4 years). At T4, and 57% at T5 were non-smokers.

The Institutional Review Board (IRB) of the New York University School of Medicine approved the study for T4 and T5, and the IRBs of the Mount Sinai School of Medicine and New York Medical College (our former affiliations) approved the study's procedures for earlier waves of data collection. A Certificate of Confidentiality was obtained from the National Institutes of Health for each wave of data collection. Informed consent was obtained from all participants (18 years or older) at each time wave. At T1 and T2, passive consent was obtained from the parents of participants who were minors (< 18 years old).

Measures

Independent Variables—Cigarette use and depressive symptoms were measured from T3–T5 (mean ages 24 to 36 years) since we have only two items for depressive symptoms at T1 and T2. *Cigarette smoking (T3–T5)* was a single item, i.e., "In the past 5 years, how many cigarettes did you smoke?" The answer options were as follows: none (0), a few cigarettes or less a week (1), 1–5 cigarettes a day (2), about half pack a day (3), about one pack a day (4), about one and a half packs a day (5), and more than one and a half packs a day (6). *Depressive symptoms*³² (*T3–T5*) consisted of a 5 item scale, e.g., "Do you sometimes feel unhappy, sad, or, depressed?" The answer options were as follows: completely false (0), false (1), true (2), and completely true (3). The Cronbach's alphas were 0.74 (T3), 0.77 (T4), and 0.85 (T5).

Control Variables

The participants were asked about their *gender* (female=1, male=2) and *ethnicity* (African American=1, Puerto Rican=2). *Anxiety symptoms*³² (T3) was a 3 item scale, e.g., "Over the past few years, how much were you bothered by feeling nervous?" Answer options were as follows: not at all (0), a little (1), somewhat (2), quite a bit (3), and extremely (4). The Cronbach's alpha was 0.78. The participants provided their *age* in years, marital status (live alone=0, married and live together including cohabiting=1), annual income (less than or equal to 10,000=0, between 10,001 and 35,000=1, between 35,001 and 75,000=2, and greater than 75,000=3), and employment status (unemployed=0, employed=1) at T5. The measure *socioeconomic status* (T5) was a summed score of the marital status, annual income, and employment status which ranged from 0 to 5. *Physical disease* (T5) was a summed score of diabetes (no=0, yes=1), hypertension (no=0, yes=1), asthma (no=0, yes=1), and obesity (no=0, yes=1). The score ranged from 0 to 4.

Generalized Anxiety Disorder at T5—Generalized Anxiety Disorder (GAD) was assessed using an adaptation of the DSM-5 Generalized Anxiety Disorder measure.³³ The participants were asked nine questions: (1) Within the last 5 years, have you had a period of at least 6 months when you worried excessively or were anxious about several things? During this period of 6 months or more: (2) Were these worries present most days? (3) Was it difficult to control the worries or did they interfere with your ability to focus on what you were doing? (4) Did you feel restless, keyed up or on edge? (5) Did you feel tense? (6) Did you feel tired or weak, or were you easily exhausted? (7) Did you have difficulty concentrating or find your mind going blank? (8) Did you feel irritable? and (9) Did you have sleep problems (e.g., difficulty falling asleep, waking up in the middle of the night, early morning awakening, or sleeping excessively)? The internal reliability of this measure was satisfactory (Cronbach's alpha = 0.94).

If the participants answered "yes" to the first three questions and "yes" to 3 or more of the last six questions, the participant received a score of 1 on the measure of GAD; otherwise, the participant received a score of 0.

Analytic procedure

We used Mplus³⁴ to obtain the joint trajectories of cigarette smoking and depressive symptoms. Cigarette smoking and depressive symptoms at each time point were treated as censored normal variables. We used the Bayesian Information Criterion (BIC) to determine the number of trajectory groups.^{34, 35} The observed trajectories for each of the groups consisted of the averages of cigarette smoking and depressive symptoms, respectively, at each point in time when the participants were assigned to the group with the largest Bayesian posterior probability (BPP). We applied the full information maximum likelihood approach for missing data.³⁴

Logistic regression analyses were then conducted to examine whether the indicator of the trajectory group membership in either high use of cigarette smoking or a high level of depressive symptoms, compared with the indicator of membership in each of the other trajectory groups from T3 to T5, were associated with GAD at T5. In these analyses, the covariates included gender, ethnicity, anxiety symptoms at T3, age, SES, and physical disease at T5. That is, all pair wise comparisons among the joint trajectory groups were performed.

Results

A four-group model was selected, based on the BIC. The BICs were 8496, 8363, 8224, and 8241 for a 2, 3, 4, and 5-group model, respectively. Since the 4-group model had the smallest BIC score, we selected it. Figure 1 presents the observed trajectories and the percentages of the sample who were members of each of the four trajectory groups. The mean BPP in each trajectory group ranged from 0.86 to 0.96, which indicated an adequate classification.

The four trajectory groups were named: 1) high cigarette smoking and high depressive symptoms (prevalence 8%, mean BPP=0.91); 2) intermediate cigarette smoking and low depressive symptoms (prevalence 33%, mean BPP=0.96); 3) low cigarette smoking and high depressive symptoms (prevalence 12%, mean BPP=0.94); and 4) no cigarette smoking and low depressive symptoms (prevalence 47%, mean BPP=0.86). Table 1 presents summary statistics for each of the four trajectory groups. **As presented in** Table 2, the prevalence of GAD for the whole sample was 12.3%. The high cigarette smoking and high depressive symptoms group had the highest prevalence (50.0%) on the measure of GAD. On the other hand, the no cigarette smoking and low depressive symptoms group had the lowest prevalence (4.7%) on the measure of GAD.

Table 3 shows the adjusted odds ratios (AOR) for GAD in the logistic regression analysis. **Memberships in** the high cigarette smoking and high depressive symptoms group (AOR=11.2, p <.001) and **in** the low cigarette smoking and high depressive symptoms group (AOR=5.8, p<.001) were associated with an increased likelihood of having GAD at T5 as compared to **membership in** the no cigarette smoking and low depressive symptoms group. Additionally, **memberships in** the high cigarette smoking and high depressive symptoms group (AOR=5.7, p<.001) and **in** the low cigarette smoking and high depressive symptoms group (AOR=5.7, p<.001) and **in** the low cigarette smoking and high depressive symptoms group (AOR=3.8, p<.001) were associated with an increased likelihood of having GAD at

Page 6

T5 as compared with **membership in** the intermediate cigarette smoking and low depressive symptoms group. Lastly, **membership in** the high cigarette smoking and high depressive symptoms group was associated with an increased likelihood of having GAD at T5 as compared with **membership in** the low cigarette smoking and high depressive symptoms group (AOR=2.1, *p*<.05).

Discussion

The GAD rate of 12% in our study sample consisting of African Americans and Puerto Ricans is similar to the findings reporting 11% GAD among Blacks and Hispanics obtained in a nationally representative survey.³⁶ The hypotheses were partially supported with one exception. Our results are consistent with the findings of other investigators that cigarette smoking and depressive symptoms are associated with greater symptoms of anxiety or GAD.^{15, 24, 37} Biological pathways linking cigarette smoking to the risk of GAD may help to explain these findings. Anxiety symptoms may play a role in mitochondrial dysfunction which is also affected by exposure to cigarette smoking.^{38, 39} Mitochondria can be important sources of oxidative stress, and many abnormalities in mitochondrial function have been found in psychiatric disorders such as GAD.⁴⁰ Patients exhibiting mitochondrial disorders commonly demonstrate increased anxiety symptoms.^{41, 42}

The association between depressive symptoms and GAD may also be due in part to genetic influences. In adults, the symptoms of depression and anxiety have been linked to the personality trait of neuroticism.⁴³ The role of neuroticism, a known risk factor for common mental disorders including depressive disorder and anxiety disorder,⁴⁴ may partially explain the linkage between depressive symptoms and GAD. For example, psychological distress (e.g., sadness), core to the neuroticism domain, includes a proneness to irrational ideas, poor impulse control and poor stress management,⁴⁵ and thus may contribute to GAD.

Our findings regarding the association between joint trajectories of cigarette smoking and depressive symptoms and GAD can also be explained in part from a psychological perspective. Since the comorbidity of chronic/heavy cigarette smoking and chronic/high depressive mood was associated with low self-control, low self-esteem, and low coping,²⁹ individuals who smoke and have depressive symptoms may have narrower social networks and/or poorer relationships with co-workers, friends, and family members.^{46, 47} This might have an adverse effect on a smooth transition to adulthood including maintaining strong friendships, establishing financial independence, starting a family, and obtaining job security, which may ultimately predict GAD.

An interesting finding is that individuals who smoke very few cigarettes (i.e., less than 1–5 cigarettes a day) but are highly depressed have a 5.8 times greater risk for having GAD as compared with the individuals who do not smoke cigarettes and have low depressive symptoms. Also, individuals who smoke a few cigarettes but are highly depressed have a 3.8 times greater risk for having GAD, as compared with individuals who smoke cigarettes less than a half a pack a day and have low depressive symptoms. Depressive symptoms seem to have a stronger association with GAD than smoking. On the other hand, there is also a significant association between the high cigarette smoking with high depressive symptoms

group and GAD as compared with the low cigarette smoking with high depressive symptoms group. Cigarette smoking has some impact on GAD. We also examined the associations between single trajectory groups for cigarette smoking as well as for depressive symptoms and GAD without any other covariates; **memberships in the** higher level of cigarette smoking **trajectory group** (p<0.05) and **in the higher level of** depressive symptoms (p<0.001) trajectory group were associated with GAD (data not shown). These findings indicate that the interaction of cigarette smoking and depressive symptoms increases the

As noted earlier, one of our hypotheses was not supported since there was no significant difference between the intermediate cigarette smoking and low depressive symptoms trajectory group and the no cigarette smoking and low depressive symptoms trajectory group with respect to GAD. The trajectory group smoking less than a half a pack a day with low depressive symptoms, statistically speaking, has the same likelihood of GAD as the trajectory group not smoking with low depressive symptoms.

Limitations-Strengths

likelihood of GAD.

Our data are based on self-reports rather than on official records such as medical records. However, studies have shown that use of self-report data yields reliable results.⁴⁸ We also did not include genetic variables or specific ethnic variables for African Americans and Puerto Ricans. Future studies should include these assessments. **In this study, depressive symptoms were used so it is difficult to draw inferences about depressive disorders**. Lastly, the sample in this study consisted of African American and Puerto Rican adults residing in an urban area. Consequently, the results may not apply to the general population.

Despite these limitations, the study supports and adds to the literature in a number of important ways. First, unlike most research that focuses on one point in time, we assess cigarette smoking and depressive symptoms simultaneously over a span of up to 12 years. The prospective nature of the data allows us to go beyond a cross-sectional analysis and to consider the temporal sequencing of variables. Second, this community sample varying in SES is unique as it consisted of African American and Puerto Rican inner city adults studied until the mid 30s. Third, a significant contribution of the paper is a set of findings relating different joint trajectories of cigarette smoking and depressive symptoms beginning in the mid 20s to the later occurrence of GAD in the mid 30s. Fourth, the joint trajectory analysis based on a person-centered approach enables one to examine the magnitude, length of time, and the starting point of cigarette smoking and depressive symptoms simultaneously and their associations with later GAD. Therefore, the current study would have important implications for prevention and treatment of smoking and depression among African Americans and Puerto Ricans.

Conclusions

The findings of this study have implications for prevention and treatment, especially, for African American and Puerto Rican adults. Cigarette smoking encourages the brain to switch off its own mechanism for making dopamine, so in the long term the dopamine level decreases, which in turn prompts individuals to smoke more. Prevention strategy may focus

on decreasing the frequency of cigarette smoking among individuals who have depressive symptoms. Prevention and treatment programs for GAD may assess the extent of smoking among individuals who are depressed, and may offer treatment for depression as well as for smoking cessation. Cognitive behavioral therapy⁴⁹ is effective in both the treatment of cigarette smoking and of depression. Acceptance commitment therapy^{50, 51} and combined pharmacotherapy/psychotherapy⁵² could be other approaches to consider.

From a public health perspective, our longitudinal study designed to examine the association between earlier joint trajectories of cigarette smoking and depressive symptoms and later GAD suggests that interventions to reduce cigarette smoking and depressive symptoms may lower the rates of GAD in the mid 30s. Federal, state, and/or city governments may consider funding to support programs for treatment including smoking cessation programs focused on reducing the use of cigarettes as well as on the treatment of depressive symptoms.

Future research is necessary to examine the association between the earlier joint trajectories of cigarette smoking and depressive symptoms and later GAD within larger and more diverse samples of individuals at different developmental stages.

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Lee et al.



Figure 1. Joint trajectories of cigarette smoking and depressive symptoms among African American and Puerto Rican adults from the mid-twenties to the mid-thirties (N=674) Note. The x-axis and y-axis represent age in years and the level of cigarette smoking and depressive symptoms, respectively.

Answer options for cigarette smoking: none (0); a few cigarettes or less a week (1); 1-5 cigarettes a day (2); about half pack a day (3); about one pack a day (4); about one and half packs a day (5); more than one and half packs a day (6)

Answer options for depressive symptoms: completely false (0); false (1); true (2); completely true (3)

Table 1

Model fit indexes for the number of joint trajectory groups of cigarette smoking and depressive symptoms among African American and Puerto Rican adults from the mid twenties to the mid thirties (N=674)

Number of joint trajectory groups	Bayesian Information Criteria	Akaike Information Criteria	p-value for LRT comparing the k class to the k-1 class model
2	8496	8411	<0.0001
3	8363	8246	0.0593
4	8224	8076	<0.0001
5	8241	8061	0.4770

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Table 2

Summary statistics by the joint trajectory groups of cigarette smoking and depressive symptoms from the mid twenties to the mid thirties among African American and Puerto Rican adults (N=674)

Lee et al.

			Joint trajectory groups			Whole sample (N=674)
	High cigarette smoking and high depressive symptoms (n=54, 8%)	Intermediate cigarette smoking and low depressive symptoms (n=222, 33%)	Low cigarette smoking and high depressive symptoms (n=78, 12%)	No cigarette smoking and low depressive symptoms (n=320, 47%)	F or chi-square test statistics	
Independent variables						
Cigarette smoking (T3–T5)	T3: 3.14 (1.55)	T3: 2.33 (1.56)	T3: 0.20 (0.52)	T3: 0.10 (0.48)	$F\!\!=\!\!188.6^{**}$	T3: 1.20 (1.63)
	T4: 3.27 (1.18)	T4: 2.60 (1.34)	T4: 0.12 (0.40)	T4: 0.04 (0.20)	F=509.9 **	T4: 1.14 (1.58)
	T5: 2.91 (1.56)	T5: 2.63 (1.54)	T5: 0.37 (0.83)	T5: 0.03 (0.18)	F=330.7 **	T5: 1.18 (1.65)
Depressive symptoms (T3–T5)	T3: 2.67 (0.64)	T3: 1.42 (0.56)	T3: 2.18 (0.64)	T3: 1.23 (0.51)	$F_{=110.9}^{**}$	T3: 1.55 (0.72)
	T4: 2.54 (0.68)	T4: 1.29 (0.54)	T4: 2.26 (0.52)	T4: 1.16 (0.46)	$F_{=179.9}^{**}$	T4: 1.44 (0.70)
	T5: 2.66 (0.81)	T5: 1.23 (0.58)	T5: 2.25 (0.73)	T5: 1.06 (0.49)	$F\!\!=\!\!179.0^{**}$	T5: 1.38 (0.78)
Control variables						
Female	n=30, 55.6%	n=107, 48.2%	n=55, 70.5%	n=213, 66.6%	$x^{2}(3)=22.7^{**}$	n=405, 60.1%
African American	n=22, 40.7%	n=109, 49.1%	n=36, 46.2%	n=189, 59.1%	$x^2(3)=10.8^*$	n=256, 52.8%
Anxiety symptoms (T3)	1.93 (0.96)	0.84 (0.72)	1.41 (0.89)	0.75~(0.63)	F=52.1	0.95 (0.80)
Age (T5)	35.83 (1.29)	35.93 (1.39)	35.61 (1.36)	35.99 (1.43)	F=1.6	35.91 (1.40)
SES (T5)	1.56 (1.13)	2.07 (1.32)	2.24 (1.31)	2.78 (1.26)	$F=22.6^{**}$	2.39 (1.33)
Physical disease (T5)	1.39 (1.06)	0.98 (0.72)	1.41 (0.83)	1.11 (0.83)	F=7.4 **	1.12 (0.83)
Dependent variable						
Generalized anxiety disorder (T5)	n=27, 50.0%	n=18, 8.1%	n=23, 29.5%	n=15, 4.7%	$x^{2}(3) = **$	n=83, 12.3%
Notes. SES=Socioeconomic status						
* P<.05,						
** <i>p</i> <.001						

JAddict Dis. Author manuscript; available in PMC 2017 July 10.

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Answer options for cigarette smoking: none (0), a few cigarettes or less a week (1), 1–5 cigarettes a day (2), about half pack a day (3), about one pack a day (4), about one and half packs a day (5), more than one and half packs a day

Answer options for depressive symptoms: completely false (0), false (1), true (2), completely true (3)

Answer options for anxiety symptoms: not at all (0) to extremely (4)

Answer options for SES: lowest (0) to highest (5)

Answer options for physical disease: no disease (0) to 4 diseases (4)

Table 3

Adjusted odds ratios (95% Confidence Interval): the association of the membership in the joint trajectories of cigarette smoking and depressive symptoms with generalized anxiety disorder (GAD) among African American and Puerto Rican adults (N=674)

Comparisons of membership in each of the joint trajectory groups		
¹ High cigarette smoking and high depressive symptoms vs. No cigarette smoking and low depressive symptoms	11.2 (4.6, 27.0) ***	
² Intermediate cigarette smoking and low depressive symptoms vs. No cigarette smoking and low depressive symptoms	1.7 (0.8, 3.7)	
³ Low cigarette smoking and high depressive symptoms vs. No cigarette smoking and low depressive symptoms	5.8 (2.7, 12.5)***	
⁴ High cigarette smoking and high depressive symptoms vs. Intermediate cigarette smoking and low depressive symptoms	5.7 (2.4, 13.7)***	
⁵ Low cigarette smoking and high depressive symptoms vs. Intermediate cigarette smoking and low depressive symptoms	3.8 (1.7, 8.8)***	
⁶ High cigarette smoking and high depressive symptoms vs. Low cigarette smoking and high depressive symptoms	2.1 (1.0, 4.7)*	

Notes. Two-tailed test:

* p<.05,

*** p<.001

Gender, ethnicity, anxiety symptoms at T3, age at T5, socioeconomic status at T5, and physical disease at T5 were statistically controlled.

In models 1, 2, and 3, female gender and higher anxiety symptoms at T3 were associated with GAD (p<.05).

In model 4, higher anxiety symptoms at T3 were associated with GAD (p<.05).

In model 5, none of the other predictors was associated with GAD.

In model 6, female gender and lower socioeconomic status at T5 were associated with GAD (p<.05).