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Early childhood adversity and later hypertension: Data from the World Mental Health Survey

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Abstract

BACKGROUND—Although many studies have indicated that psychosocial factors contribute to hypertension, and that early childhood adversity is associated with long-term adverse mental and physical health sequelae, the association between early adversity and later hypertension is not well studied.

METHOD—Data from 10 countries participating in the World Health Organization (WHO) World Mental Health (WHM) Surveys (N = 18,630) were analyzed to assess the relationship between childhood adversity and adult-onset hypertension, as ascertained by self-report. The potentially mediating effect of early-onset depression-anxiety disorders, as assessed by the WHM Survey version of the International Diagnostic Interview (WMH-CIDI), on the relationship between early adversity and hypertension was also examined.

RESULTS—Two or more early childhood adversities, as well as early-onset depression-anxiety, were significantly associated with hypertension. A range of specific childhood adversities, as well as early-onset social phobia and panic/agoraphobia, were significantly associated with hypertension. In multivariate analyses, the presence of 3 or more childhood adversities was associated with hypertension, even when early-onset depression-anxiety or current depression-anxiety was included in the model.

CONCLUSIONS—Although caution is required in the interpretation of self-report data on adultonset hypertension, the results of this study further strengthen the evidence base regarding the role of psychosocial factors in the pathogenesis of hypertension.

INTRODUCTION

Research has found that psychosocial factors, in particular depression, anxiety, and anger, contribute to the etiology of hypertension. ^{1,2} Psychological trauma has been hypothesized to be an especially important contributor to hypertension. ³ Such a relationship would be consistent with research that has found chronic adverse psychobiological sequelae of early childhood adversity, ^{4,5} and with an increasingly sophisticated and integrative approach to understanding mental-physical comorbidity. ⁶

Nevertheless, studies on psychosocial contributors to hypertension have important limitations. ^{1,2} Most of these studies have been conducted in only a few countries. The relationship between childhood adversity and hypertension has received relatively little attention, with sparse data available on the contribution of different types of childhood adversity. Further, despite evidence that childhood adversity is a risk factor for depression and anxiety disorders, ⁷ there has been no research assessing whether the association of depression and anxiety with subsequent hypertension might be explained by exposure to childhood adversity.

Using data from cross-national population surveys, this paper reports on the relationships between childhood adversity, early-onset depression-anxiety disorders, and adult-onset

hypertension. Based on prior research, we hypothesized that exposure to significant childhood adversity is associated with adult-onset hypertension. To assess the potential mediation of early-onset depression-anxiety disorders on the association between childhood adversity and hypertension, we evaluated (1) whether such early-onset conditions are associated with an increased risk of adult-onset hypertension, and (2) whether childhood adversity and early-onset depression-anxiety disorders predict adult-onset hypertension independently of each other.

METHODS

Samples

Ten surveys were carried out in the Americas (Colombia, Mexico, and the United States), Europe (Belgium, France, Germany, Italy, the Netherlands, and Spain), and Asia (Japan). All surveys were based on multistage, clustered, area probability household samples. All interviews were carried out face-to-face by trained lay interviewers. Sample sizes ranged from 2372 (Netherlands) to 9282 (United States). Response rates ranged from 45.9% (France) to 87.7% (Colombia), with an average response rate of 66.7%.

Internal subsampling was used to reduce respondent burden and average interview time and cost by dividing the interview into 2 parts. Part 1 included the core diagnostic assessment of mental disorders. Part 2 included additional information relevant to a wide range of survey aims, including assessment of chronic physical conditions. All respondents completed part 1. All part 1 respondents who met criteria for any mental disorder as well as a probability sample of other respondents were administered part 2. Part 2 respondents were weighted by the inverse of their probability of selection for part 2 of the interview to adjust for differential sampling. Analyses in this article were based on the weighted part 2 subsample (n = 18,303). Additional weights were used to adjust for differential probabilities of selection within households, to adjust for nonresponse, and to match the samples to population sociodemographic distributions.

Training and field procedures

The central World Mental Health (WMH) staff trained bilingual supervisors in each country. The World Health Organization (WHO) translation protocol was used to translate instruments and training materials. Some surveys were conducted in bilingual form (Belgium). Other surveys were conducted exclusively in the country's official language. Persons who could not speak these languages were excluded. Quality control protocols, described in more detail elsewhere, 8 were standardized across countries to check on interviewer accuracy and to specify data cleaning and coding procedures. The institutional review board of the organization that coordinated the survey in each country approved and monitored compliance with procedures for obtaining informed consent and protecting human subjects.

Mental disorder status

All surveys used the WMH Survey version of the WHO Composite International Diagnostic Interview (WMH-CIDI, now CIDI 3.0),⁸ a fully structured diagnostic interview, to assess disorders. Clinical calibration studies have supported the reliability and validity of both the original CIDI⁹ and the WMH-CIDI.¹⁰ Disorders were assessed using the definitions and criteria of the DSM-IV.¹¹ CIDI organic exclusion rules were imposed (the diagnosis was not made if the respondent indicated that the episode of depressive or anxiety symptoms was due to physical illness or injury or the use of medication, drugs, or alcohol).

This article includes anxiety disorders (generalized anxiety disorder, panic disorder and/or agoraphobia, posttraumatic stress disorder, and social phobia) and depressive disorders (major depressive disorder). Anxiety and depressive disorders were aggregated into a single category, on the basis of prior WMH Survey findings that both anxiety disorders and depressive disorders have similar relationships with a wide range of chronic physical conditions. ¹² For each disorder present at any time in a respondent's lifetime, the age of onset was determined using standard CIDI questions. Persons were classified as having an early-onset depression/anxiety disorder if the earliest age of onset for any lifetime disorder was less than 21 years of age. In analyses for each mental disorder singly, early onset was also defined as age of onset prior to age 21.

Hypertension

In a series of questions about chronic conditions adapted from the US Health Interview Survey, respondents were asked about the lifetime presence of select chronic conditions. Respondents were asked, "Did a doctor or other health professional ever tell you that you had any of the following illnesses...hypertension?" Respondents were also asked how old they were when the condition began. Only those reporting hypertension onset after the age of 20 were included in the current analyses.

Childhood adversity

The following types of childhood adversity were included in the present report: physical abuse; sexual abuse; neglect; parental death, parent divorce, other parental loss; parental mental disorder; parental substance use; parental criminal behavior; family violence; and family economic adversity. The aim was to assess the occurrence of childhood adversity within the context of the family rather than all possible childhood adversities. Thus, natural disasters or exposure to war were not assessed in this section. Those respondents who reported that the experience occurred before age 18 and who met the criteria specified for a given adversity (described below) were coded as having experienced childhood family adversity.

Physical abuse—Respondents were classified as having experienced *physical abuse* when they indicated that, when they were growing up, their father or mother (includes biological, step, or adoptive parents) slapped, hit, pushed, grabbed, shoved, or threw something at them, or that they were beaten as a child by the persons who raised them.

Sexual abuse—For *sexual abuse*, the following questions were asked:

- "The next 2 questions are about sexual assault. The first is about rape. We define this as someone either having sexual intercourse with you or penetrating your body with a finger or object when you did not want them to, either by threatening you or using force, or when you were so young that you didn't know what was happening. Did this ever happen to you?"
- "Other than rape, were you ever sexually assaulted or molested?"

Sexual abuse was the only adversity for which information was not collected that would distinguish whether the perpetrator was a family member or someone else. However, previous research that used a similar measure but did allow such a distinction showed that a good indirect way to distinguish family vs nonfamily sexual abuse is to ask about the number of instances of victimization, with cases involving 1 or 2 instances typically perpetrated by a stranger and those involving 3 or more instances typically perpetrated by a family member. ¹³ In the WMH surveys, therefore, respondents who reported that they had

had any of these experiences 3 times or more were coded as having experienced sexual abuse (within the family context).

Neglect—For the assessment of *neglect*, 2 scales were created. These were based on responses to the neglect items:

- "How often were you made to do chores that were too difficult or dangerous for someone your age?"
- "How often were you left alone or unsupervised when you were too young to be alone?"
- "How often did you go without things you need like clothes, shoes, or school supplies because your parents or caregivers spent the money on themselves?"
- "How often did your parents or caregivers make you go hungry or not prepare regular meals?"
- "How often did your parents or caregivers ignore or fail to get you medical treatment when you were sick or hurt?"

The *serious* neglect scale was the sum of the number of neglect items in which the respondent replied "often" or "sometimes," plus 1 if the respondent rated either of his or her parents as having spent little or no effort in watching over them to ensure they had a good upbringing. The *severe* neglect scale is the sum of the number of neglect items where respondents replied "often" plus 1 if the respondent rated either of his or her parents as having spent no effort in watching over them to ensure they had a good upbringing. Both the serious and severe neglect scales ranged from 0 to 6. For the final definition of *neglect*, the respondent had to have a score of at least 1 on the *severe* neglect scale and at least 2 on the *serious* neglect scale. The coding of the *neglect* domain was determined empirically on the basis of frequency distributions to derive estimates in keeping with existing literature on the prevalence of these experiences in the general population.

Parental death, parental divorce, or other parental loss—Respondents were first asked whether they lived with both of their parents when they were brought up. If respondents replied in the negative, they were asked:

• "Did your biological mother or father die, were they separated or divorced, or was there some other reason?"

According to their answers to these questions, respondents were classified as having experienced parental death (when they indicated that one or both parents had died), parental divorce (when they indicated that their parents had divorced), and other parental loss (when respondents replied that they had either been adopted, had gone to boarding school, had been in foster care, or had left home before age 16).

Parental mental disorder—*Parental depression* was assessed by the following diagnostic items:

- "During the years you were growing up, did [woman/man who raised the respondent] ever have periods lasting 2 weeks or more where [she/ he] was sad or depressed most of the time?"
- "During the time when [her/his] depression was at its worst, did [she/he] also have other symptoms like low energy, changes in sleep or appetite, and problems with concentration?"

A positive response to item 1 was followed up with a frequency question:

• "Was this during all, most, some, or only a little of your childhood?"

Parental generalized anxiety disorder (GAD) was assessed as follows:

• "During the time you were growing up, did [woman/man who raised the respondent] ever have periods of a month or more when [she/he] was constantly nervous, edgy, or anxious?" and

• "During the time [her/his] nervousness was at its worst, did [she/he] also have other symptoms, like being restless, irritable, easily tired, and difficulty falling asleep?"

A positive response to GAD item 1 was followed up with a frequency question:

• "Was this during all, most, some, or only a little of your childhood?"

Parental panic disorder was assessed by the following item:

• "Did [woman/man who raised the respondent] ever complain about anxiety attacks where all of a sudden [she/he] felt frightened, anxious, or panicky?"

Respondents who replied positively to both diagnostic items for depression *and* who replied "all or most of the time" to the frequency item *and* who reported that their parents got professional help for depression *or* that depression interfered a lot with their parents' lives or activities were coded as respondents with parental depression. A similar logic applied to characterizing respondents whose parents had GAD. Respondents who responded positively to the single parental panic disorder item were coded as having parents with panic disorder.

Parental substance disorder—Similarly, *parental substance* disorder was assessed with the following items:

- (criterion a) "Did [woman/man who raised the respondent] ever have a problem with alcohol or drugs" and
- (criterion b) "Did [she/he] have this problem during all, most, some, or only a little
 of your childhood?"

Respondents who replied positively on the first and "all" and "most" on the second item were then asked whether the problem interfered a lot with life or activities of the woman or man who raised the respondent (criterion c), or whether they had sought professional help for this problem (criterion d). Those respondents who replied affirmatively on criteria a and b, and on either c or d were coded as having had parents with a substance disorder.

Parental criminal behavior—*Parental criminal behavior* was assessed by the following questions:

- "Was [woman/man who raised the respondent] ever involved in criminal activities like burglary or selling stolen property?" and
- "Was [woman/man who raised the respondent] ever arrested or sent to prison?"

Respondents who replied positively on either question were classified as having experienced criminal behavior in the family.

Family violence—Respondents were coded as having experienced *family violence* when they indicated that they

- "were often hit, shoved, pushed, grabbed, or slapped while growing up" or
- "witnessed physical fights at home, like when your father beat up your mother?"

Family economic adversity—Respondents were coded positive for *family economic adversity* if there was a positive response to either of the 2 following items:

- "During your childhood and adolescence, was there ever a period of 6 months or more when your family received money from a government assistance program like Welfare, Aid to Families with Dependent Children, General Assistance, or Temporary Assistance for Needy Families?" (This item was modified to be relevant to the welfare programs in each country where the survey was administered.) Or,
- If there was no male head of the family *and* the female head did NOT work all or most of the time during respondent's childhood; or if there was no female head of the family *and* the male head did NOT work all or most of respondent's childhood, or if there was no female head *and* no male head of the family.

Statistical analysis

The association of childhood adversity and early-onset mental disorders with adult-onset asthma was studied with discrete-time survival analyses, using retrospective self-report of age of onset of hypertension (reported in whole years). The start of the period at risk for adult-onset hypertension was set at age 20.

Cox proportional hazards models were constructed to estimate risk of adult-onset hypertension as a function of number of childhood adversities and early-onset depression-anxiety disorder status while adjusting for potential confounders (sex and current age). Country was included in the analyses as a stratifying variable, which allows each country to have a unique hazard function. This was necessary because of large differences in prevalence rates of childhood adversity, mental disorders, and chronic physical disorders between countries, and to allow for the slight differences in measurement of childhood adversities between countries (ie, measurement of neglect and parental divorce).

In the Cox proportional hazards regression analyses, time to age of onset of hypertension from age 20 was the dependent variable. Persons who had not developed hypertension were censored at their current age. The associations are expressed as hazard ratios (HRs) measuring relative risk (RR). Childhood adversities were analyzed in 4 categories of number of adversities (none, 1, 2, 3), with no adversities as the reference group. Childhood adversity and early-onset mental disorders were included in the models both separately and simultaneously. We screened for interaction of childhood adversity and early-onset depression-anxiety, but the interaction was nonsignificant; therefore, only main effects are reported in this article. To account for the possibility of differential recall of childhood adversity among those with a current mood or anxiety disorder, we performed an additional analysis that adjusted for current (12-month) anxiety or mood disorder. We performed yet another analysis, including educational attainment, to assess whether any associations between the risk factors of primary interest (childhood adversity and early-onset depression/anxiety) were explained by differences in educational level.

The assumption of proportional hazards was assessed by inspecting log-minus-log plots of the survival functions. Standard errors were calculated for all HRs. The statistical significance level was set at .05 for a 2-sided test. The analyses were performed using the SURVIVAL procedure in SUDAAN statistical software. ¹⁴ In all multivariate models, a complete-case approach was used, but the WMH survey used multiple imputation methods to impute missing values for the childhood adversities. The survival analyses employed sample weights to adjust for differences in probability of selection. Estimates were made using SUDAAN software to account for the complex sample design.

RESULTS

Characteristics of the sample are presented in TABLE 1, and prevalence of self-reported adult-onset hypertension, childhood adversity, and early-onset depression-anxiety disorders are presented in TABLE 2. Overall, 17.3% of the respondents reported hypertension, of whom almost half (48.6%) noted a history of childhood adversity, and about a fifth of whom (18.5%) were found to have early onset of depression-anxiety disorder. Prevalence of childhood adversity varied across countries, ranging from 30% in Spain to 65% in Colombia.

Kaplan-Meier curves for the cumulative percentage of hypertension by age for persons with 0, 1, or 2 childhood adversities (FIGURE 1) and for persons with vs without early-onset depression-anxiety disorders (FIGURE 2) were plotted; these curves indicate crude associations between hypertension and both childhood adversity and early-onset depression-anxiety disorders.

TABLE 3 provides HRs for the relationship between childhood adversity, early-onset depression-anxiety disorder, and adult-onset hypertension. The unadjusted HR for adult-onset hypertension for persons with early-onset depression-anxiety (relative to those without) was 1.34~(P<.001). After adjusting for number of childhood adversities, this HR was essentially unchanged, at 1.29~(P<.001). Unadjusted HRs for number of childhood adversities (relative to persons with no childhood adversity) were 1.03 for persons with 1 adversity (P= ns), 1.19 for persons with 2 adversities (P<.05), and 1.28 for persons with 3 or more adversities (P<.01), prior to adjusting for early-onset depression-anxiety disorder status. After adjusting for depression-anxiety disorder status, the HRs for number of childhood adversities were somewhat reduced but remained significantly greater than 1.0 for persons with 3 or more adversities (P<.05), as shown in the second column of TABLE 2. Adjusting for current depression-anxiety disorder status rather than early-onset depression-anxiety disorder status changed the HRs for number of childhood adversities only slightly, suggesting that effects of current mood on recall of childhood adversities or hypertension status does not explain their association.

HRs for each of the specific childhood adversities, adjusted for gender, age, and country are shown in TABLE 4. Of the 11 childhood adversities ascertained, 6 were significantly associated with adult-onset hypertension (with odds ratios [ORs] ranging from 1.17 to 1.53). Among the 5 specific DSM-IV mental disorders assessed, early-onset social phobia and panic disorder/agoraphobia were significantly associated with increased risk of adult-onset hypertension, whereas early-onset major depressive disorder, generalized anxiety disorder, and posttraumatic stress disorder were not (TABLE 4).

DISCUSSION

It is important to emphasize several limitations of this study. Blood pressure levels can be directly measured in epidemiological studies, but the current study relied on self-report and provided only a categorical measure of hypertension. At the same time, epidemiological studies of hypertension are rarely able to obtain detailed data on mental illness and psychological stressors. Partial reassurance is obtained from data that self-reported hypertension is moderately associated with objective data on hypertensive status. ^{15,16} Nevertheless, although estimates of sensitivity of self-report of hypertension range from 64% to 91% in several Western countries, it may be lower in other locations. ¹⁷ A second crucial limitation is that the focus of this study was childhood family adversity rather than a range of other possible early traumas, and the study relied on retrospective reports of childhood stressors. False-negative childhood reports may lead to underestimation of

associated effects. Some reassurance is obtained from a recent review of the validity of retrospective report of childhood adversities; this review found that although false-negative reports are common, false-positive reports are rare, and concluded that retrospective case-control studies of major, easily defined adversities are potentially valid. ¹⁸ Finally, retrospective reports of age of onset of both hypertension and mental disorders are subject to bias.

Given these limitations, we view the results reported in this article as exploratory findings that would need to be replicated with more rigorous ascertainment of hypertension and hypertension onset, along with prospective studies of subjects exposed to early adversity. The main findings of the current study were that (1) the presence of 2 or more early childhood adversities was significantly associated with adult-onset hypertension, (2) early-onset depression-anxiety disorder was also significantly associated with increased risk of hypertension, and (3) in multivariate analyses, 3 or more childhood adversities, as well as early-onset depression-anxiety disorder, continued to be significantly associated with increased risk of hypertension.

The observed HRs suggest a "dose-response" relationship between early childhood adversity and subsequent hypertension. The extent and number of childhood adversities may be more important than the specific type of adversity experienced. There was no increased risk of adult-onset hypertension among persons who reported only 1 childhood adversity, and the HRs for the relationship between early adversity and subsequent hypertension were relatively low. However, given that 22.8% of the respondents with adult-onset hypertension experienced 2 or more childhood adversities, the potential impact of these exposures on hypertension risk has potential importance.

Although childhood adversity is associated with an increased risk of early-onset depression-anxiety disorders, controlling for number of childhood adversities did not explain the association of early depression-anxiety disorders with adult-onset hypertension. Effect sizes for the association of psychosocial factors and hypertension have been modest to date,² and the findings here underscore previous recommendations that genetic, early environmental, and later environmental effects on hypertension and other cardiovascular conditions all require further delineation. ^{19,20} Effects sizes for behavioral treatments for hypertension are quite variable, ²¹ and this parallel area of research requires additional attention.

Additional work is needed to understand the specific psychobiological mechanisms that mediate the relationships between childhood adversity, early-onset depression and anxiety disorders, and subsequent adult-onset hypertension. Although the current data do not directly address these mechanisms, it is relevant to note the research documenting how both early trauma and depression-anxiety disorders alter the autonomic system, potentially contributing to increased allostatic load. 7.22–24 In addition to chronic sympathetic nervous system dysregulation, monoamine abnormalities and endothelial dysfunction might play a role. 2

In conclusion, the WMH Survey provides a unique opportunity to assess the relationship between childhood adversity, early-onset depression-anxiety disorders, and adult-onset hypertension in a large, multinational sample. The data are consistent with prior literature on the role of psychosocial factors in the pathogenesis of hypertension and call attention to the potential role of both early childhood adversity and early-onset depression-anxiety disorders as risk factors for hypertension. These exploratory results suggest the need for prospective studies assessing the independent and joint effects of childhood adversity and early-onset emotional disorders in influencing risk of developing hypertension as an adult.

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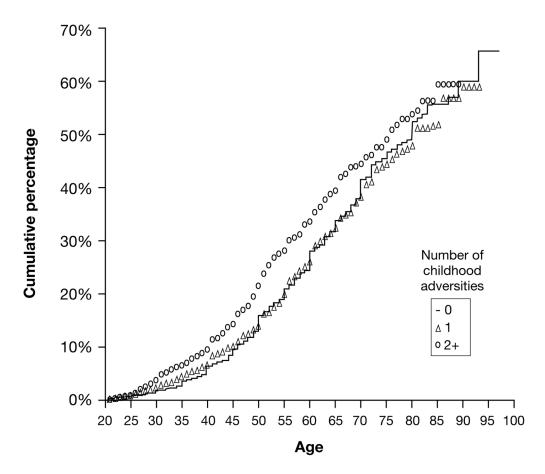


FIGURE 1. Kaplan-Meier curve for the cumulative percentage of onset of hypertension by age for persons with 0, 1, or 2+ childhood adversities

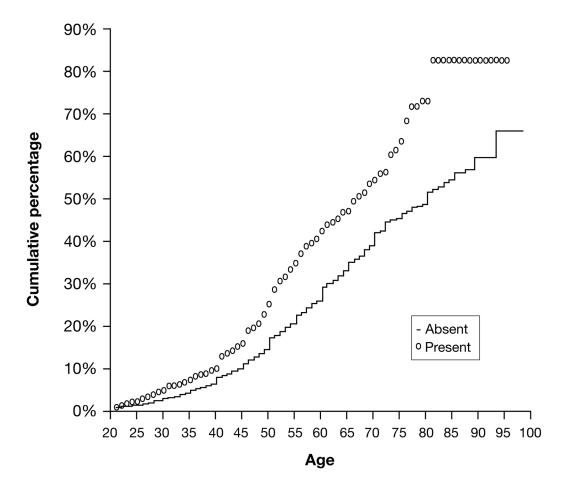


FIGURE 2.Kaplan-Meier curve for the cumulative percentage of hypertension onset by age for persons with vs without early-onset depression-anxiety disorders

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 TABLE 1

 Characteristics of total and adult onset hypertension samples

CHARACTERISTIC	TOTAL CROSS-NATIONAL SAMPLE (N = 18,600)		ADULT-ONSET HYPERTENSION (AGE 21) SAMPLE (N = 3215)	
	Unweighted, N	Weighted, %	Weighted, %	
Age (21 to 98)				
21 to 44	9827	52.5	18.0	
45	8803	47.5	82.0	
Female	11,092	52.8	55.6	
Adult onset of hypertension (21 years)	18,600	3215	17.3	
Country				
Colombia	2122	11.5	8.0	
Belgium	989	5.3	4.6	
France	1359	7.3	6.4	
Germany	1273	6.9	7.9	
Italy	1699	9.1	6.8	
Netherlands	1052	5.6	5.2	
Spain	2015	10.8	9.6	
Japan	873	4.7	4.4	
Mexico	2059	11.0	6.6	
United States	5189	27.8	23.6	

TABLE 2

 $Early-onset\ depressive/anxiety\ disorders\ and\ childhood\ adversities\ in\ total\ and\ adult-onset\ hypertension\ samples$

CHARACTERISTIC	TOTAL CROSS-NATIONAL SAMPLE (N = 18,600)	ADULT-ONSET HYPERTENSION (AGE 21) SAMPLE (N = 3215) Weighted, %	
	Weighted, %		
Early-onset (prior to age 21) mental disorders			
Any depressive/anxiety disorder	10.2	18.5	
Major depressive disorder	4.5	9.0	
Generalized anxiety disorder	1.3	2.5	
Social phobia	4.8	8.3	
Posttraumatic stress disorder	1.7	2.7	
Panic disorder/agoraphobia	1.8	3.3	
Number of childhood adversities			
0	56.1	51.4	
1	25.6	25.8	
2	10.0	11.6	
3	8.3	11.2	
Type of childhood adversity			
Physical abuse	9.6	12.9	
Sexual abuse	2.6	3.9	
Neglect	6.5	8.2	
Parent died	12.8	12.7	
Parents divorced	9.8	10.6	
Other loss of parent	5.3	6.2	
Parental mental disorder	7.2	10.0	
Parental substance use disorder	5.0	6.5	
Violence in family	9.4	11.9	
Parental criminal behavior	3.3	4.4	
Family economic adversity	5.8	6.3	

 TABLE 3

 Association between early-onset depression-anxiety disorders, childhood adversities, and hypertension

RISK FACTOR	HAZARD RATIOS (95% CONFIDENCE INTERVAL)			
	Main effect for hypertension	Adjusted for early-onset depression/anxiety	Adjusted for current depression/ anxiety	
Depression/anxiety onset before age 21	1.34 (1.20 to 1.48; <i>P</i> < .001)			
1 childhood adversity	1.03 (0.92 to 1.16; ns)	1.03 (0.91 to 1.15; ns)	1.02 (0.91 to 1.15, ns)	
2 childhood adversities	1.19 (1.01 to 1.41; P<.05)	1.17 (1.00 to 1.38; ns)	1.17 (1.00 to 1.38; <i>P</i> <.01)	
3 childhood adversities	1.28 (1.07 to 1.53; P<.01)	1.22 (1.02 to 1.46; P<.05)	1.21 (1.01 to 1.44; P< .001)	

TABLE 4

Effect of early-onset (prior to age 21) depressive/anxiety disorders and specific childhood adversities on adult-onset of hypertension (21 years and over)

Early-onset (prior to age 21) mental disorder	Hazard ratio (95% CI) for adult-onset hypertension (adjusted for age, sex, country)		
Major depressive disorder	1.15 (0.96 to 1.39; ns)		
Generalized anxiety disorder	1.16 (0.91 to 1.47; ns)		
Social phobia	1.36 (1.19 to 1.57; P< .001)		
Posttraumatic stress disorder	1.15 (0.89 to 1.48; ns)		
Panic disorder/agoraphobia	1.53 (1.19 to 1.97; P< .001)		
Childhood adversity			
Physical abuse	1.23 (1.04 to 1.46; P<.05)		
Sexual abuse	1.14 (0.84 to 1.54; ns)		
Neglect	1.10 (0.90 to 1.34; ns)		
Parent died	1.02 (0.89 to 1.18; ns)		
Parents divorced	0.96 (0.79 to 1.16; ns)		
Other loss of parent	1.26 (1.05 to 1.52; P<.05)		
Parental mental disorder	1.29 (1.11 to 1.50; P<.01)		
Parental substance use disorder	1.30 (1.07 to 1.56; P<.01)		
Violence in family	1.17 (1.02 to 1.34; P<.05)		
Criminal behavior in family	1.33 (1.09 to 1.62; P<.01)		
Family economic adversity	1.03 (0.83 to 1.27; ns)		

CI: confidence interval.