

Predictors of long-term prognosis of depression

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See related commentary by Kurdyak and Cairney on page 1955 and at www.cmaj.ca/lookup/doi/10.1503/cmaj.111664

ABSTRACT

Background: Many people with depression experience repeated episodes. Previous research into the predictors of chronic depression has focused primarily on the clinical features of the disease; however, little is known about the broader spectrum of sociodemographic and health factors inherent in its development. Our aim was to identify factors associated with a long-term negative prognosis of depression.

Methods: We included 585 people aged 16 years and older who participated in the 2000/01 cycle of the National Population Health Survey and who reported experiencing a major depressive episode in 2000/01. The primary outcome was the course of depression until 2006/07. We grouped individuals into trajectories of depression using growth trajectory models. We included demographic, mental and physical health factors as predictors in the

multivariable regression model to compare people with different trajectories.

Results: Participants fell into two main depression trajectories: those whose depression resolved and did not recur (44.7%) and those who experienced repeated episodes (55.3%). In the multivariable model, daily smoking (OR 2.68, 95% CI 1.54–4.67), low mastery (i.e., feeling that life circumstances are beyond one's control) (OR 1.10, 95% CI 1.03–1.18) and history of depression (OR 3.5, 95% CI 1.95–6.27) were significant predictors ($p < 0.05$) of repeated episodes of depression.

Interpretation: People with major depression who were current smokers or had low levels of mastery were at an increased risk of repeated episodes of depression. Future studies are needed to confirm the predictive value of these variables and to evaluate their accuracy for diagnosis and as a guide to treatment.

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Depression is a common and often recurrent disorder that compromises daily functioning and is associated with a decrease in quality of life.¹⁻³ Guidelines for the treatment of depression, such as those published by the Canadian Network for Mood and Anxiety Treatments (CANMAT)⁵ and the National Institute for Health and Clinical Excellence (NICE) in the United Kingdom,⁴ often recommend antidepressant treatment in patients with severe symptoms and outline specific risk factors supporting long-term treatment maintenance.^{4,5} However, for patients who do not meet the criteria for treatment of depression, the damaging sequelae of depression are frequently compounded without treatment.⁵ In such cases, early treatment for depression may result in an improved long-term prognosis.⁶⁻⁸

A small but growing number of studies have begun to characterize the long-term course of depression in terms of severity,⁹ life-time prevalence¹⁰ and patterns of recurrence.¹¹ However, a recent systematic review of the risk factors of

chronic depression highlighted a need for longitudinal studies to better identify prognostic factors.¹² The capacity to distinguish long-term patterns of recurrence of depression in relation to the wide range of established clinical and non-clinical factors for depression could be highly beneficial. Our objective was to use a population-based cohort to identify and understand the baseline factors associated with a long-term negative prognosis of depression.

Methods

Study sample

The National Population Health Survey is a nationwide longitudinal study conducted by Statistics Canada. The survey began in 1994/95 and collects health information as well as other health-related information such as economic, social, demographic and occupation data. At study inception, 17 276 people were randomly selected by use of a stratified two-stage sample design. The cohort is representative of the Cana-

dian population and has been followed every two years. The first interview cycle in 1994/95 had a response rate of 83.6%; the follow-up rate for the fourth cycle (2000/01) was 78.7% of the original cohort.¹³ In the present study, we included the 585 survey respondents who were 16 years or older and reported having a major depressive episode in that year.

Definition of depression

Major depression in the study cohort was measured by use of the Composite International Diagnostic Interview Short Form. This form, completed during a 10-minute interview, has 90% sensitivity and 94% specificity for identifying major depressive episodes compared with the full hour-long Composite International Diagnostic Interview,¹⁴ which can identify depressive episodes consistent with the Diagnostic and Statistical Manual of Mental Disorders (DSM-IV).¹⁵

For each individual, the Composite International Diagnostic Interview Short Form produces a predictive probability of a major depressive episode in the past 12 months.¹⁴ An individual with a predictive probability of 90% or higher was considered to have major depression (a score of 5 or higher on a 0–8 scale).¹⁴ This corresponds to the DSM-IV criteria for a major depressive episode (five of nine depressive symptoms in a two-week period during the past year, including either loss of interest or depressed mood).¹⁵

Trajectory of depression

Of specific interest in this study was the recurrence of major depression in the biannual cycles of data collection from 2000/01 (cycle 4) to 2006/07 (cycle 7). Although occurrence at single points was reported, the primary outcome for this study was the trajectory of depression over these four periods. All individuals included in the sample had experienced a major depressive episode in 2000/01; there were therefore eight possible patterns of occurrence (i.e., [1,0,0,0], [1,1,0,0], ... , [1,1,1,1], where 0 indicates the absence of depression and 1 indicates the presence of depression). We used growth trajectory models to group individuals according to similar experiences of depression over the six-year follow-up period.¹⁶

Prognostic factors

Individuals in each cycle of the survey provided a detailed record of many health-related factors. In this study, we explored the demographic, physical health and mental health factors that have previously been reported to be associated with depression. We sought to determine which

factors were significantly associated with a negative trajectory of depressive symptoms during the follow-up period. Data about sexually transmitted infections were collected from 1996/97 (not reported in 2000/01). All other data were collected at baseline in 2000/01.

The following demographic factors were explored: age, sex, marital status, social support, education level, ethnic background, immigration status, income level and current employment status. We explored the following physical health factors: self-reported physical health status, sexually transmitted infection, obesity, chronic medical condition, chronic pain, smoking status and physical activity level. We collected data on the following mental health factors: severity of present symptoms, history of depression, heavy drinking, use of antidepressants, body image, low self-esteem, low mastery (i.e., feeling that life circumstances are beyond one's control), frequent thoughts about death, work stress, childhood trauma, stressful life events, self-perceived stress and chronic stress.

Statistical analysis

Trajectory analysis uses a semiparametric, group-based modelling strategy to identify homogeneous latent trajectory classes based on longitudinal data.¹⁶ We used growth trajectory models (PROC TRAJ in SAS 9.2) to group individuals based on episodes of major depression from 2000/01 to 2006/07. We used the Bayesian information criteria to identify the best-fitting model with the least number of trajectories. Missing data during the follow-up period were handled by PROC TRAJ under the missing-at-random assumption, which allows patterns with missing values to share information with patterns with more data points through the latent variable.¹⁷ Individuals with missing data were assigned to their most likely group.

We investigated the possible predictive effects of factors at baseline on trajectory patterns by use of logistic regression adjusted for age and sex. We used purposeful selection for multivariable logistic regression models that considered all prognostic factors. Income was included as a covariate. We performed two stratified analyses. The first analysis investigated predictive factors among those with mild symptoms (e.g., depressed individuals whose distress score was below the median) separately from those with more severe symptoms. The second analysis investigated predictive factors among those with a history of major depression separately from those with no history of major depression.

The National Population Health Survey used a multistage sampling design with unequal selec-

Table 1: Characteristics of respondents in the 2000/2001 cycle of the National Population Health Survey

Factor	% of respondents*					
	History of depression n = 236	No history of depression n = 277	p value	All respondents with depression n = 585	Excluded cohort (no depression) n = 10 908	p value
Age, yr, mean (SD)	38.8 (1.2)	41.6 (1.1)	0.10	38.5 (0.8)	44.6 (0.1)	< 0.001
Sex (female)	77.4	57.5	< 0.001	64.5	51.4	< 0.001
Income (low; Low Income Cut-Off ¹⁹)	25.1	12.9	0.007	18.5	9.8	< 0.001
Employment status (employed)	55.2	64.0	0.12	59.9	66.8	0.008
Any postsecondary education	67.3	64.0	0.52	63.6	63.6	0.99
Marital status			0.17			< 0.001
Single	35.2	28.7		36.5	25.4	
Married, partner or common-law	38.3	48.6		40.6	60.2	
Divorced, separated or widowed	26.5	22.8		22.9	14.4	
Social support (Medical Outcomes Study social support scale ¹⁹ ; out of 5), mean (SD)	3.7 (0.1)	4.1 (0.1)	0.006	3.9 (0.1)	4.5 (0.1)	< 0.001
Ethnic background (white)	93.0	91.7	0.69	91.7	89.9	0.39
Immigration status	11.6	9.0	0.44	9.1	17.5	< 0.001
Self-reported health scale (out of 5; higher score indicates better health), mean (SD)	3.0 (0.1)	3.2 (0.1)	0.08	3.1 (0.1)	3.7 (0.1)	< 0.001
Chronic health condition†	83.8	77.7	0.17	78.6	61.9	< 0.001
Chronic pain (reported as "not usually free of pain")	38.3	27.9	0.08	29.9	12.2	< 0.001
Obese or overweight‡	55.1	49.8	0.36	50.8	52.4	0.60
Physical inactivity (Ontario Health Survey ¹⁹)	38.2	42.2	0.51	43.9	44.3	0.88
Daily smoking	66.0	53.6	0.027	38.0	21.4	< 0.001
Sexually transmitted infection	2.2	6.7	0.06	4.1	2.0	0.024
Severity of symptoms scale (out of 24; Kessler K6 Psychological Distress Score ¹⁹), mean (SD)	9.9 (0.5)	6.3 (0.4)	< 0.001	8.0 (0.3)	2.2 (0.1)	< 0.001
Duration of depression in the past year, weeks, mean (SD)	14.6 (1.4)	10.7 (0.9)	0.018	0.2 (0.1)	12.2 (0.8)	< 0.001
History of depression§	NA	NA		47.0	10.9	< 0.001
Frequent thoughts about death, self-reported	57.0	49.1	0.18	54.0	0.3	< 0.001
Heavy drinking¶	15.1	6.0	0.52	13.3	7.0	0.020
Self-reported antidepressant use in the past month	31.7	36.7	0.33	31.4	3.7	< 0.001
Self-reported body image			0.006			0.36
Overweight	51.2†	38.8		43.5	39.1	
Just about right	7.5	3.1		4.7	4.7	
Underweight	41.3	58.1		52.2	56.2	
Self-esteem scale (out of 24; Rosenberg Scale), ²⁰ mean (SD)	7.2 (0.4)	5.3 (0.3)	< 0.001	6.4 (0.3)	4.4 (0.1)	< 0.001
Mastery scale (out of 28; Pearlin & Schooler ²¹), mean (SD)	12.7 (0.4)	10.6 (0.4)	< 0.001	11.6 (0.3)	7.9 (0.1)	< 0.001
Childhood traumatt			0.18			< 0.001
0 events	23.4	30.4		29.0	51.9	
1 events	24.2	28.0		26.2	26.6	
≥ 2 events	52.1	41.7		44.8	21.5	
High work stress‡‡	35.0	26.0	0.19	30.1	12.1	< 0.001
Stressful life events (≥ 1 of 10 negative events in previous year§§)	71.0	57.6	0.014	64.0	26.8	< 0.001
Self-perceived stress (out of 5; higher score indicates greater stress), mean (SD)	3.7 (0.1)	3.4 (0.1)	0.003	3.5 (0.1)	2.8 (0.1)	< 0.001
Chronic stress¶¶	26.1	41.6	0.005	33.2	6.8	< 0.001

Note: NA = not available, SD = standard deviation.

*Unless stated otherwise

†Presence of 1 of 22 common chronic conditions (complete list available from corresponding author).

‡Body mass index > 25 for people ≥ 18 years. Cole cut-off points used for those aged 16–17 years.¹⁹

§Composite International Diagnostic Interview Short Form assessment from previous three survey cycles.

¶Men: > 15 drinks/wk; women: > 10 drinks/wk.

††Childhood and Adult Stress Index¹⁹ — seven possible events.

‡‡75th percentile or above of the National Population Health Survey Work Stress Index.²⁰

§§A complete list is available from the authors.

¶¶75th percentile or above of the General Chronic Stress Index.²⁰

tion probabilities, and normalized sampling weights for the survey are available in the survey data set. To account for the complex survey design, we included normalized sampling weights in the descriptive data analysis, trajectory modelling and logistic regression. We used Taylor linearization for the variance estimation of regression coefficients. The standard errors for all estimates were calculated using the bootstrap method.¹⁸ We used the SURVEYLOGISTIC procedure for the analysis of complex surveys in SAS 9.2 for all statistical analyses. We considered *p* values of less than 0.05 to indicate statistically significant associations.

This study was approved by the Health Research Ethics Board of the University of Alberta.

Results

Of the 585 people in the survey aged 16 or older who reported major depression in 2000/01, 64.5% were women, and the average age was 38.5 (Table 1). Most earned a middle to high income (81.5%), were white (91.7%) and had achieved greater than high school education (63.6%). There were significant differences between the 585 participants with depression and those who reported no depressive episodes in 2000/01 for many factors, including social support, immigration status and self-reported health (Table 1). It was not possible to ascertain a history of depression for 72 individuals who had missing data from the first three waves of the study (between 1994 and 1999). We excluded 10 908 respondents who reported not having depression during 2000/01. Among participants

with depression, we found significant differences in many factors, most notably self-esteem, daily smoking and body image, between those with and without a history of depression (Table 1).

The trajectory analysis identified a two-group model as the best-fitting model: those who experienced subsequent episodes of depression in the follow-up period (55.3%) and those who did not (44.7%). Among those with repeated episodes of depression, the probability of being depressed at any point during the follow-up period was between 35% and 50% (Figure 1).

The adjusted odds ratios (ORs) for a negative depression trajectory in association with demographic, physical health and mental health factors are shown in Tables 2 and 3. Of the variables that were significantly predictive of repeat episodes of depression in the age- and sex-adjusted analyses, only daily smoking (OR 2.68, 95% CI 1.54–4.67), a history of depression (OR 3.50, 95% CI 1.95–6.27) and low mastery (OR 1.10, 95% CI 1.03–1.18) remained significant in the multivariable model (Table 4). We inspected the correlation coefficients for all predictor variables using a correlation matrix, and the coefficients were determined to be sufficiently low such that issues of multicollinearity within the model were not indicated. The highest correlation between predictor variables was below 0.6.

We conducted stratified analyses separately for individuals with or without a history of major depression (Tables 2 and 3). Age- and sex-adjusted analyses revealed significant associations between repeat episodes of depression and both high severity of symptoms and low mastery in both groups. A number of other significant associations were observed in either group. In the multivariable analysis, age, sex and income were not significantly associated with repeated episodes of depression (Table 4). Repeat depression remained significantly associated only with daily smoking (OR 4.79, 95% CI 2.03–11.32) and low mastery (OR 1.14, 95% CI 1.03–1.25) among those with a history of depression. Among people with no history of depression, high severity of symptoms (OR 1.09, 95% CI 1.00–1.20) and migraine headaches (OR 2.75, 95% CI 1.00–7.55) were predictive of repeat depression in the multivariable analysis.

We performed a final stratified analysis based on the severity of depression symptoms. The age- and sex-adjusted analyses revealed multiple significant predictors of repeat episodes of depression among those with mild symptoms or those with severe symptoms (Tables 2 and 3). At both levels of severity in the multivariable analysis, a negative trajectory remained significantly associated with daily smoking (mild:

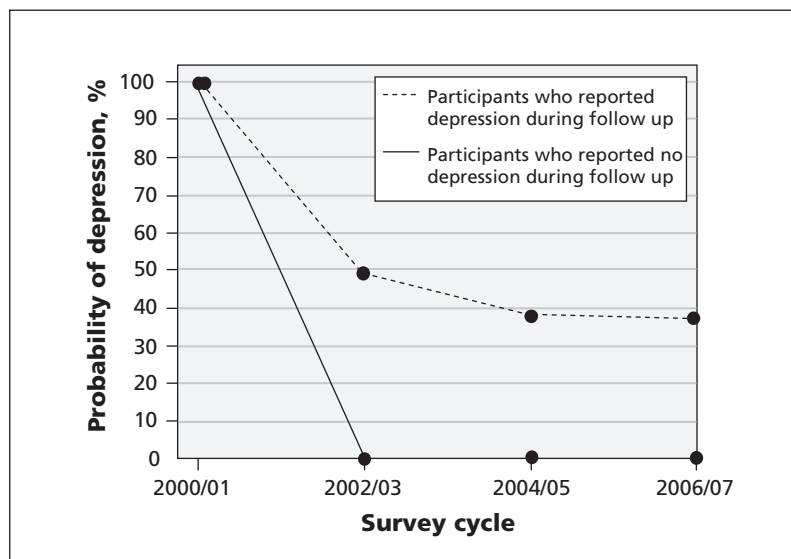


Figure 1: Trajectories of depression over a six-year follow-up period among 585 people who reported experiencing an episode of major depression in 2000/01.

OR 2.59, 95% CI 1.16–5.79; severe: OR 2.79, 95% CI 1.17–6.65) and history of depression (mild: OR 4.34, 95% CI 1.66–11.35; severe: OR 3.15; 95% CI 1.31–7.58). In the multivariable model, being an immigrant remained as a significant protective effect among people with severe symptoms (OR 0.23, 95% CI 0.06–0.92) (Table 4).

Interpretation

In this population-based survey of 585 Canadians who reported an episode of major depression in 2000/01, we found that more than half experi-

enced one or more repeated episodes in the following six years. We found that a history of depression, daily smoking and low mastery were significant predictors of a negative trajectory of depression.

A history of depression is well-documented in the medical literature to be a risk factor for future depressive episodes.^{6,22} The association between smoking and depression is more complex, however, with evidence accumulating in the neurobiological,²³ genetic²⁴ and psychosocial realms,²⁵ and there is a lack of clarity about the direction of effect and possible causal mechanisms. In our study, the strongest predictor of

Table 2: Association between demographic and physical health factors and a negative trajectory of depression among 585 participants who experienced depression in 2000/01, with stratification by severity of symptoms and history of depression

Factor	Group of participants; adjusted odds ratio* (95% CI)				
	All participants <i>n</i> = 585	Mild symptoms <i>n</i> = 290	Severe symptoms <i>n</i> = 295	History of depression <i>n</i> = 236	No history of depression <i>n</i> = 277
Age	1.00 (0.98–1.01)	0.98 (0.96–1.01)	1.01 (0.99–1.03)	0.99 (0.96–1.02)	0.99 (0.96–1.01)
Sex (female)	0.84 (0.54–1.29)	0.90 (0.47–1.72)	0.77 (0.39–1.52)	1.25 (0.51–3.06)	0.89 (0.46–1.72)
Income (middle or high)	0.51 (0.28–0.94)	0.81 (0.26–2.48)	0.53 (0.24–1.17)	0.61 (0.25–1.48)	0.98 (0.31–3.07)
Employment status (employed)	0.73 (0.44–1.21)	0.72 (0.34–1.52)	0.97 (0.46–2.03)	1.00 (0.44–2.27)	0.97 (0.46–2.04)
Education (> high school)	0.74 (0.48–1.13)	0.79 (0.37–1.67)	0.80 (0.42–1.51)	0.99 (0.46–2.13)	0.58 (0.30–1.13)
Marital status					
Single (ref)	1.00	1.00	1.00	1.00	1.00
Married, partner or common-law	0.91 (0.52–1.57)	0.81 (0.36–1.82)	1.14 (0.49–2.66)	1.21 (0.44–2.84)	0.58 (0.26–1.30)
Divorced, separated or widowed	1.31 (0.68–2.52)	0.54 (0.20–1.45)	2.94 (1.04–8.32)	1.81 (0.62–5.27)	0.60 (0.23–1.55)
Social support	0.66 (0.52–0.83)	0.82 (0.56–1.20)	0.74 (0.53–1.04)	0.62 (0.40–0.96)	0.75 (0.53–1.05)
Ethnic background (white)	0.91 (0.37–2.20)	0.57 (0.03–12.78)	1.60 (0.47–5.37)	1.55 (0.05–47.96)	0.63 (0.02–22.95)
Immigrant	0.60 (0.25–1.42)	1.54 (0.06–41.66)	0.25 (0.07–0.87)	0.33 (0.09–1.22)	0.78 (0.13–4.77)
Health status†	0.73 (0.58–0.90)	1.05 (0.73–1.51)	0.71 (0.52–0.97)	0.70 (0.48–1.03)	0.83 (0.60–1.15)
Chronic health condition	1.66 (0.97–2.87)	1.39 (0.62–3.14)	1.57 (0.65–3.80)	1.82 (0.68–4.90)	1.46 (0.69–3.61)
Chronic pain	2.03 (1.28–3.23)	1.05 (0.44–2.53)	2.56 (1.26–5.22)	1.52 (0.70–4.48)	1.51 (0.75–3.03)
Overweight or obese	1.25 (0.79–1.96)	1.30 (0.65–2.61)	1.39 (0.70–2.78)	0.94 (0.45–2.02)	1.06 (0.54–2.10)
Physical activity (active or moderate level)	0.86 (0.54–1.39)	1.02 (0.56–1.87)	1.07 (0.51–2.24)	0.76 (0.29–1.59)	1.07 (0.53–2.17)
Daily smoking	3.82 (2.45–5.95)	2.75 (1.40–5.42)	3.81 (1.96–7.44)	6.45 (2.92–14.29)	2.35 (1.18–4.67)
Sexually transmitted infection	1.81 (0.42–7.77)	‡	‡	‡	‡

Note: CI = confidence interval, ref = reference category.

*Adjusted for age and sex.

†Measured on a five-point scale, with higher scores indicating better self-reported health.

‡Estimates are unstable because of the small numbers reporting.

repeated episodes of depression was smoking among participants with a history of depression. Among those with no history of depression, smoking did not increase the likelihood of future episodes. This apparent potentiating influence of one factor on the other may, in fact, fit a model in which a lag exists between a primary depressive episode and subsequent changes (e.g., neurochemical or behavioural alterations), which could increase a smoker's vulnerability to further depressive relapses.

Mastery is a psychological concept that measures whether an individual has an internal locus of control and a strong belief that they have command over his or her life circumstances. The mastery scale used in this survey has shown high validity¹⁹ and reliability²⁶ in studies that have linked low mastery to less frequent health

behaviours, such as physical activity and vegetable consumption,²⁷ and to outcomes such as higher metabolic risk²⁸ and cardiovascular-related death.²⁹ In our study, high levels of mastery at baseline appeared to provide protection against a negative trajectory of depression, an observation supported by existing cross-sectional research.³⁰ Again, this association was significant only among participants with a history of depression.

Interventions designed to increase mastery and promote smoking cessation carry the additional benefit of positive health outcomes in other areas, notably a substantial reduction in the risk of smoking-related diseases and, by increasing mastery, improvements in weight and dietary control, pain management, adherence to preventive health programs and successful postoperative recovery.

Table 3: Association between mental health factors and a negative trajectory of depression among 585 participants who experienced depression in 2000/01, with stratification by severity of symptoms and history of depression

Factor	Group of participants; adjusted odds ratio* (95% CI)				
	All participants <i>n</i> = 585	Mild symptoms <i>n</i> = 290	Severe symptoms <i>n</i> = 295	History of depression <i>n</i> = 236	No history of depression <i>n</i> = 277
Severity of symptoms	1.12 (1.07–1.17)	NA	NA	1.09 (1.00–1.18)	1.11 (1.02–1.20)
Duration of depression	1.02 (1.01–1.04)	1.03 (0.99–1.06)	1.01 (0.98–1.03)	1.02 (0.99–1.05)	0.74 (0.33–1.66)
History of depression	4.00 (2.46–6.50)	3.37 (1.45–7.79)	2.79 (1.35–5.79)	NA	NA
Frequent thoughts about death	1.17 (0.73–1.88)	1.26 (0.66–2.43)	0.91 (0.45–1.83)	1.23 (0.56–2.72)	1.05 (0.56–1.95)
Heavy drinking	1.72 (0.71–4.17)	0.84 (0.22–3.13)	–†	–†	1.51 (0.13–17.79)
Antidepressant use	1.69 (1.04–2.77)	1.38 (0.63–3.00)	1.68 (0.84–3.38)	1.93 (0.87–4.27)	1.00 (0.48–2.06)
Body image					
Overweight	1.29 (0.80–2.09)	1.22 (0.58–2.53)	1.28 (0.62–2.63)	1.20 (0.55–2.59)	0.62 (0.32–1.19)
Just about right (ref)	1.00	1.00	1.00	1.00	1.00
Underweight	2.40 (0.69–8.37)	1.49 (0.01–196.41)	2.35 (0.36–15.31)	2.21 (0.02–309.92)	0.62 (0.01–143.79)
Social support	0.66 (0.52–0.83)	0.82 (0.56–1.20)	0.74 (0.53–1.04)	0.62 (0.40–0.96)	0.75 (0.53–1.05)
Low self-esteem	1.09 (1.03–1.14)	1.02 (0.93–1.14)	1.04 (0.97–1.14)	1.10 (1.00–1.21)	1.03 (0.94–1.12)
Low mastery	1.15 (1.09–1.20)	1.08 (0.99–1.18)	1.11 (1.03–1.22)	1.19 (1.09–1.30)	1.11 (1.03–1.20)
Childhood trauma					
0 events (ref)	1.00	1.00	1.00	1.00	1.00
1 events	1.35 (0.72–2.51)	1.95 (0.77–4.95)	0.89 (0.35–2.31)	1.01 (0.34–2.98)	1.69 (0.72–4.01)
2 events	2.87 (1.63–5.05)	3.45 (1.51–7.91)	2.12 (0.87–5.17)	3.28 (1.11–9.74)	1.97 (0.91–4.26)
High work stress	2.13 (1.13–4.04)	1.60 (0.59–4.34)	1.86 (0.66–5.22)	4.39 (0.71–27.34)	1.54 (0.53–4.44)
Stressful life events	1.65 (1.06–2.55)	1.18 (0.60–2.31)	1.56 (0.77–3.18)	0.79 (0.33–1.86)	1.85 (0.92–3.73)
Self-perceived stress	1.63 (1.27–2.10)	1.62 (1.10–2.38)	1.30 (0.83–2.02)	1.34 (0.80–2.24)	1.62 (1.06–2.48)
High chronic stress	2.48 (1.52–4.03)	1.31 (0.58–2.96)	1.97 (1.01–3.82)	1.84 (0.82–4.12)	2.62 (1.19–5.80)

Note: NA = not applicable, ref = reference category.
*Adjusted for age and sex.
†Estimates are unstable because of the small numbers reporting.

Limitations

As is expected with long-term studies, losses to follow-up were observed. We used a missing-at-random assumption in our analyses to minimize selection bias when dealing with incomplete or missing data. We were also unable to distinguish between participants whose depression remitted before experiencing a repeat episode (e.g., between those with chronic or recurrent depression), which may have important clinical implications.

Lastly, we looked at smoking status only at baseline, and we did not consider changes in smoking behaviour at subsequent times (in par-

ticular, we did not take into account smoking cessation, which has known associations with recurrences of major depression³¹). This evidence does not conflict with our finding that current smokers were at an increased risk of repeated depressive episodes. The purpose of this paper was not to disentangle shared etiologies between these two factors, but simply to provide clinicians with practical prognostic information. This was best achieved by the use of baseline measures that are most readily available to clinicians.

Conclusion

History of depression is a well-known clinical indicator of future depressive episodes;^{6,22} however, smoking and mastery are more novel prognostic factors that are not well accounted for in current clinical practice. Assessment of these two factors can be easily achieved by use of a single question about smoking and a simple seven-item questionnaire about mastery.¹⁹ Our results will be strengthened by future studies that follow a cohort of patients with newly diagnosed depression more frequently and collect additional data about the short-term course of depressive symptoms. Future research should evaluate the benefits of including smoking cessation and mastery in existing clinical guidelines for the treatment of depression.

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Table 4: Demographic, physical and mental health predictors of a negative trajectory of depression in the multivariable models

Model; factor	Odds ratio* (95% confidence interval)
All individuals, n = 585	
Age	0.99 (0.96–1.01)
Sex (female)	0.91 (0.49–1.66)
Income (high and middle)	0.93 (0.48–1.82)
Daily smoking	2.68 (1.54–4.67)
Had depression history	3.50 (1.95–6.27)
Low mastery	1.10 (1.03–1.18)
History of depression, n = 236	
Age	0.99 (0.95–1.02)
Sex (female)	0.85 (0.30–2.38)
Income (high and middle)	0.94 (0.35–2.53)
Daily smoking	4.79 (2.03–11.32)
Low mastery	1.14 (1.03–1.25)
No history of depression, n = 277	
Age	0.98 (0.95–1.02)
Sex (female)	1.27 (0.57–2.82)
Income (high and middle)	1.13 (0.33–3.88)
Severity of symptoms	1.09 (1.00–1.20)
Has migraine headaches	2.75 (1.00–7.55)
Mild symptoms, n = 290	
Age	0.98 (0.95–1.02)
Sex (female)	0.85 (0.34–2.12)
Income (high and middle)	0.87 (0.27–2.85)
Daily smoking	2.59 (1.16–5.79)
History of depression	4.34 (1.66–11.35)
Severe symptoms, n = 295	
Age	0.98 (0.95–1.02)
Sex (female)	0.84 (0.32–2.24)
Income (high and middle)	0.92 (0.34–2.52)
Daily smoking	2.79 (1.17–6.65)
History of depression	3.15 (1.31–7.58)
Immigration	0.23 (0.06–0.92)

*Adjusted for age and sex.

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