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Hostility and depressive mood: results from the Whitehall II prospective cohort study

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Abstract

Background

The psychosocial vulnerability model of hostility posits that hostile individuals, given their oppositional attitudes and behaviours, are more likely to have increased interpersonal conflicts, lower social support, more stressful life events and higher likelihood of depression. However, little research has tested this hypothesis using large scale prospective samples. The present study aims to assess the predictive value of hostility for depressive mood.

Methods

Data are from 3,399 participants in the Whitehall II cohort study, aged 35–55 years at baseline (phase 1=1985–1988). Cynical hostility was measured at phase 1 using the Cook-Medley-Hostility-scale. Depressive mood was assessed at phase 7 (2002–2004) using the Center-for-Epidemiologic-Studies Depression-Scale (CES-D). Sociodemographic characteristics, health related-behaviours, common mental disorders and antidepressant medication intake were assessed at phase 1. Stressful life events (SL-E) and confiding/emotional support were measured at phases 1, 2 (1989–1990), and 5 (1997–1999).

Results

Compared to participants in the lowest quartile of cynical-hostility, those in the highest quartiles were more likely to have depressive mood (second quartile: OR=1.58, 95% CI; 1.14–2.20; third quartile: OR=2.78; 95% CI, 2.03–3.77; fourth quartile: OR=4.66; 95% CI, 3.41–6.36) in analysis adjusted for sociodemographic characteristics. This graded association was somewhat attenuated (up to 18%) but remained robust to adjustments for the covariates measured at baseline and follow-up. The association was also evident in participants free of mental health difficulties at baseline.

Conclusions

Cynical-hostility is a strong and robust predictor of depressive mood. Consideration of personality characteristics may be crucial to the understanding and management of depression.

Author Keywords hostility ; depressive mood ; psychosocial vulnerability ; stressful life events ; social support

INTRODUCTION

Depression is a major public health issue worldwide (Moussavi et al., 2007). Projections of the Global Burden of Disease (GBD) suggest that depression will account for 10% of the total disease burden in high-income countries by 2030 (Mathers and Loncar, 2006). The psychosocial vulnerability model of hostility posits that hostile individuals, given their oppositional attitudes and behaviours, are more likely to have increased interpersonal conflicts, lower social support, more stressful life events and higher likelihood of depression (Kivimäki et al., 2003, Miller et al., 1996). Research suggests that stressful life-events (SL-E) may be independent risk factors (Kendler et al., 1999) for depression with several studies showing SL-E to be associated with an increased risk of both the onset (Caspi et al., 2003) and recurrence of depression (Bifulco et al., 2000). Another well established factor in the aetiology of depression is social support. According to the “stress-buffering” hypothesis (Cohen and Wills, 1985), social support may protect from the negative effects of stressors such as SL-E, hence protecting against depression. Indeed, a large body of evidence has shown that a low level of social support predicts future depression and recovery from depressive episodes (Brown et al., 1994, Johnson et al., 1999).

We argue that cynical hostility, a personality trait characterized by general cynicism and interpersonal mistrust, may increase the risk of depressive disorders because hostility is related to both SL-E and social support (Hardy and Smith, 1988, Smith and Frohm, 1985). However, there is little research on the predictive value of hostility for depressive disorders using large scale prospective samples. A small scale cross-sectional study (Felsten, 1996) conducted among undergraduate students found cynical hostility to be strongly associated with

depressive mood. Another study (Heponiemi et al., 2006) examining the longitudinal effects of hostility on depressive tendencies among 1413 men and women found cynical hostility to be related to an increase in depressive tendencies after 5 years. Depressive mood may reinforce hostile feelings and behaviours toward others (Painuly et al., 2005), or influence the assessment of cynical hostility (Kendler et al., 2006); a longer time lag between assessment of hostility and the measurement of depression would allow the examination of whether the influence of cynical hostility on depressive mood persists over time. The aim of the present study is to examine the predictive value of cynical hostility measured in midlife age on depressive mood 19 years later by controlling for baseline common mental disorders, antidepressants medication intake as well as for SL-E and confiding/emotional support at the baseline and during the follow-up.

MATERIAL & METHODS

Data are drawn from the Whitehall II study, established in 1985 as a longitudinal study to examine the socioeconomic gradient in health and disease among 10,308 civil servants (6,895 men and 3,413 women). All civil servants aged 35–55 years in 20 London based departments were invited to participate by letter, and 73% agreed. Baseline screening (Phase 1) took place during 1985–1988, and involved a clinical examination and a self-administered questionnaire. Subsequent phases of data collection have alternated between postal questionnaire alone [Phases 2 (1989–1990), 4 (1995–1996), 6 (2001) and 8 (2006)] and postal questionnaire accompanied by a clinical examination [Phases 3 (1991–1993), 5 (1997–1999) and 7 (2002–2004)]. The University College London ethics committee approved the study.

Measures

Cynical hostility, defined as a personality trait characterized by general cynicism and interpersonal, was assessed using the Cook-Medley Hostility scale (Cook and Medley, 1954) at phase 1 (1985–1988). Internal consistency, test–retest reliability, and construct validity of this scale have been demonstrated (Smith, 1992). Participants completed an abridged 38-item version (Cronbach alpha = 0.83) of the original 50-item instrument. Item savings were necessary because of the extreme length of the original questionnaire; the Minnesota Multiphasic Personality Inventory (Hathaway and McKinley, 1943) (numbers of the omitted items are: 19, 183, 237, 253, 386, 394, 410, 455, 458, 485, 504, and 558). Cynical Hostility levels were determined based on the quartile distribution [lowest (0–6), middle lowest (7–10), middle highest (11–15) and highest (> 16)]. The lowest quartile was the reference category.

We also used the eight-item “Cynical Distrust Scale” (alpha = 0.72), an alternative short-form measure of cynical hostility, derived by factor analysis from the Cook-Medley Hostility Scale by Everson and colleagues. (Everson et al., 1997) Here again, cynical distrust levels were based on the quartile distribution [lowest (0–8), middle lowest (9), middle highest (10–11) and highest (> 11)]

Depressive mood at follow up was assessed using the Center for Epidemiologic Studies Depression Scale (CES-D, Cronbach alpha = 0.83) at phase 7 (2002–2004). The CES-D, a widely used and validated instrument, is a 20-item self-report questionnaire designed to measure depressive mood in community studies (Radloff, 1977). A score ≥ 16 from a total possible score of 60 reflects significant depressive mood and risk for presence of clinical depression (Radloff, 1977).

Covariates

Sociodemographic measures included age, sex, ethnicity, and socioeconomic position (SEP) assessed by British civil service grade of employment taken from the phase 1 questionnaire.

Health-related behaviours assessed at phase 1 included smoking status (never, ex, and current), exercise (≥ 1.5 or < 1.5 hours of moderate or vigorous exercise/week), heavy alcohol consumption in units of alcohol consumed per week (> 22 for men and > 15 for women), and body mass index (BMI) (< 20 , 20–24.9, 25–29.9, or ≥ 30 kg/m²).

Common mental disorder at baseline was assessed using the self-administered 30-item General Health Questionnaire at phase 1. In each GHQ item an enquiry is made about the presence or absence of a specific symptom. On the basis of receiver operating characteristics analysis and previous studies, we defined people with a GHQ sum score of 5 or more as cases and those scoring 0–4 as non-cases (Stansfeld and Marmot, 1992b). In the present study in which GHQ scores were validated against a Clinical Interview Schedule, the sensitivity (73%) and specificity (78%) using this measure of ‘caseness’ was acceptable (Stansfeld and Marmot, 1992b).

Antidepressant medication at phase 1 was assessed by asking participants whether in the last 14 days they had taken antidepressants prescribed by a doctor (yes/no).

Stressful life events at phases 1, 2, and 5 included the number stressful life events (0, 1, 2 and more) derived from an eight item self-reported question concerning experiences in the previous 12 months. The instruction “The following is a list of things that can happen to people. Try to think back over the past 12 months and remember if any of these things happened to you and, if so, how much you were upset or disturbed by it?” was followed by a list of events: 1) Personal serious illness, injury or operation; 2) Death of a close relative; 3) Serious illness, injury or operation of a close relative or friend; 4) Major financial difficulty; 5) Divorce, separation or break-up of personal

intimate relationship; 6) Other marital or family problem; 7) Any mugging, robbery, accident or similar event; 8) Change of job or residence.

Confiding/emotional support at phases 1, 2, and 5 was assessed using the Close Persons Questionnaire (Stansfeld and Marmot, 1992a) which included a 7-item scale measuring wanting to confide, confiding, sharing interests, boosting self-esteem and reciprocity relative to the first close relationship. Each item of the scale was evaluated using a Likert scale ranging from 1 to 4 with higher scores indicating more confiding/emotional support. The final confiding/emotional support scores were divided in three groups based on tertiles representing different levels of exposure to confiding/emotional support (low, middle, high).

Statistical analysis

Differences in cynical hostility score levels and depressive mood status as a function of the baseline covariates were assessed using a chi-square test.

The association between cynical hostility and the depressive mood at follow-up was assessed using logistic regressions in 5 serially adjusted models. In Model 1 no adjustment was made. Model 2 adjusted the likelihood of depressive mood for sex, age, ethnicity and socioeconomic position. In model 3, the analysis was additionally adjusted for smoking, BMI, alcohol consumption and physical exercise. Model 4 had two elements: model 4a was additionally adjusted for baseline stressful life events and confiding/emotional support score (phase 1) and model 4b for stressful life events and confiding/emotional support score at baseline and during the follow-up (phases 1, 2, and 5). Model 5 had further adjustments for antidepressant medication and common mental disorders at baseline. The same serial analyses were undertaken to examine the association between cynical distrust and depressive mood at follow-up. The interaction between cynical hostility with sex in relation to depressive mood was not statistically significant ($P > 0.05$), leading us to combine men and women in the analyses.

RESULTS

Only 75% of the 10308 participants were asked to complete the hostility scale at phase 1 due to this measure being introduced after the start of the baseline survey. 6484 participants responded to the hostility questions (84% of those asked). 6012 participants at Phase 7 responded to the CES-D Scale; 3639 of these had data cynical hostility. Finally, 3399 participants had complete data on cynical hostility, depressive mood and the 13 covariates. The mean age (SD) at baseline was 44 (5.9) years. The prevalence of depressive mood among these participants at phase 7 was 15.1%.

Table 1 shows the associations between covariates (Phase 1), cynical hostility (Phase 1) and depressive mood (Phase 7). Higher cynical hostility scores were associated with younger age, lower socioeconomic position, being non-white, higher BMI, antidepressant medication intake, having common mental disorders, higher number of SL-E, lower social network size, and higher social isolation (all $p \leq 0.007$). The presence of depressive mood at Phase 7 was associated with being female, younger age, lower socioeconomic position, being non-white, lower alcohol consumption, lower exercise, antidepressant medication, having common mental disorders, higher number of SL-E, and lower confiding/emotional support score at baseline (all $p < 0.001$).

Table 2 presents the association between cynical hostility at baseline (Phase 1) and depressive mood over 19 years later (Phase 7). In model 2, adjusting for sex, age, ethnicity and socioeconomic position, participants in the second quartile of cynical hostility had 1.58 times greater odds (95% CI; 1.14–2.20) of depressive mood compared to those in the first quartile. Those in the third (OR=2.78; 95% CI, 2.03–3.77) and fourth quartile (OR=4.66; 95% CI, 3.41–6.36) also had a greater likelihood of depressive mood when compared to those in the lowest quartile. Further adjustment for health-related behaviours in model 3 (BMI, alcohol consumption and exercise) did not much change these associations. In Model 4a, when further adjustment was made for baseline SL-E and confiding/emotional support score, the associations were attenuated, particularly for participants in the highest cynical hostility quartile (16% compared to model 2). In model 4b, when further adjustment was made for SL-E and confiding/emotional support score at the baseline and during the follow-up, a similar percentage of attenuation was observed. Finally, further adjustments (model 5) for antidepressant medication intake and common mental disorders at baseline, the odds of depressive mood at follow up was reduced, particularly for participants in the highest cynical hostility level (17% compared to model 3). However, the dose response association between cynical hostility levels and depressive mood was preserved even in the fully adjusted models. In the table we also present the association between cynical distrust and depressive mood. As with cynical hostility, we found evidence of a dose-response association between levels of cynical distrust and the likelihood of depressive mood at follow-up.

Sensitivity analyses

To test the robustness of the present findings, we examined the predictive value of hostility on depressive mood among participants with no mental health difficulties (common mental disorders or antidepressant medication) at study baseline (phase 1). After excluding participants who reported common mental disorders and antidepressant medication at baseline (phase 1), the number of participants with

depressive mood at follow decreased by 49% to 260. Nevertheless, the magnitude of the association between cynical hostility and depressive mood at follow-up was similar to that observed in the full sample. Participants in the second quartile of cynical hostility had 1.41 times greater odds (95% CI; 0.93–2.12) of depressive mood compared to those in the first quartile. Those in the third (OR=2.30; 95% CI, 1.57–3.37) and fourth quartile (OR=3.39; 95% CI, 2.27–5.07) also had greater likelihood of depressive mood, suggesting that cynical hostility is a strong predictor depressive mood even in individuals free of mental health difficulties at baseline.

Cynical distrust was also assessed at phase 5 of the study; analysis with this measure revealed that it also predicted depressive mood at follow-up, despite the shortened follow-up time (9 years instead of 19): participants in the second quartile of cynical distrust at phase 5 had 1.58 greater odds (95% CI; 2.21–2.05) of depressive mood compared to those in the first quartile. Those in the third (OR=2.03; 95% CI, 1.61–2.56) and fourth quartile (OR=4.06; 95% CI, 3.19–5.17) also had a greater likelihood of depressive mood, suggesting that cynical distrust is a strong and consistent predictor of depressive mood.

DISCUSSION

In this study we sought to examine the longitudinal association between cynical hostility assessed in midlife and depressive mood in early old age. The risk of depressive mood 19 years later increased in a dose-response relationship by level of cynical hostility. This graded association was preserved after controlling for sex, age, ethnicity, socioeconomic position, health-related behaviours (BMI, alcohol consumption, and exercise), common mental disorders, and antidepressant medication at baseline as well as SL-E, confiding/emotional support score at baseline and during the follow-up; all these factors were found to be associated with hostility or depressive mood or with both of them.

Comparison with previous studies

To the best of our knowledge, this is the first longitudinal cohort study to examine the predictive value of cynical hostility on depressive mood over a 19-year period. Both cynical hostility and depressive mood were assessed using standardized tools. We were able to control for a wide range of confounders that have been found to be important both for hostility and depressive mood. We were also able to control for common mental disorder at baseline. Previous studies have shown cross-sectional (Felsten, 1996) and prospective associations over a 5-year follow-up (Heponiemi et al., 2006) between neurotic or cynical hostility and depressive mood. Our findings show the effects of cynical hostility on depressive mood to persist over 19 years. Cynical hostility as personality trait is assumed to be relatively stable during adulthood.(McCrae and Costa Jr, 1994) In our sample the short-form 8-item cynical distrust scale showed moderate stability over 10 years (correlation coefficient = 0.53). The prospective association over the 19-year follow-up could imply that cynical hostility is relatively stable across the lifecourse and predicts depressive mood over time. It is also possible that the observed association is the product of a mutually reinforcing cycle between hostility and depression. Some evidence for the latter explanation comes from the stronger association of depression with short-form cynical distrust measured at Phase 5 compared to Phase 1. With either interpretation, our results clearly show cynical hostility to be risk factor for depressive mood.

Our results also show that the cynical distrust scale, a short-form measure of cynical hostility scale developed by Everson and colleagues, shown to be associated with mortality and myocardial infarction,(Everson et al., 1997) is also associated with depressive mood in a similar way to the longer version of the questionnaire. Thus, our results provide further validation of this shortened version of the cynical hostility scale; a finding that will be of interest to other researchers in the field.

The psychosocial vulnerability model of hostility (Kivimaki et al., 2003, Miller et al., 1996) suggests that hostile individuals may be at greater risk for depressive mood because they are more likely to have lower social support and experience more stressful life events. In the present study we found that participants who scored higher on cynical hostility scale were more likely to have a higher number of SL-E and reduced confiding/emotional support score. These factors have also been found to be related to the presence of depressive mood, making them potential mediators of the association between cynical hostility and depressive mood. However, statistical adjustment for baseline number of SL-E and confiding/emotional support score explained at best 16% of the association, providing only partial support for the psychosocial vulnerability model of hostility. Similar attenuation (at best 17%) was observed when the association between cynical hostility and depressive was adjusted for previous common mental disorders (depression and anxiety) and history of antidepressants intake. We were able to model potential mediators of the association between cynical hostility and depressive mood, particularly SL-E and confiding/emotional social support as time-dependent variables. However, controlling for the cumulative number of SL-E and exposure to confiding/emotional support did not strengthen their status as mediators between cynical hostility and depressive mood. The percentage of attenuation in the association was 16% at best.

Adjustment for socioeconomic position attenuated the association between hostility and depressive mood, suggesting that it is a possible confounder. On the other hand, we found no significant interactions between socioeconomic position and hostility in predicting depressive mood. However we cannot conclude that social context is of little importance for the development of hostility and ultimately the liability of depressive mood. Although personality is often seen as a relatively stable individual attribute, it is likely that socioeconomic circumstances affect personality, both in childhood and adulthood (McCrae and Costa, 1987). Previous studies have shown (Bifulco et al.,

1998, Brown et al., 1990a, Brown et al., 1990b, Brown et al., 1990c, Schwartz et al., 1995, Shaffer) that psychological attributes, personality characteristic and self-esteem for instance, are partially rooted in environmental conditions in childhood, (learning) experiences, and rearing styles and that the development of hostility is, in part, explained by factors such as parental behaviour that is overly strict, critical and demanding of conformity. It is also plausible that adult circumstances, such as work-related stressors, contribute to the development or promotion of personality traits, such as hostility. The parental behaviour pattern described above (i.e., overly strict, critical and demanding of conformity) may be viewed as a reflection of the parents' occupational and other life experiences, which are characterized, for example, by job-strain (Kivimaki et al., 2003).

There is evidence that personality characteristics are influenced by genetics factors (Heath et al., 1994). Similarly, genetic factors, such as serotonin transporter and receptor polymorphisms, are implicated in the aetiology of depressive disorders (Caspi et al., 2003, Hamet and Tremblay, 2005, Jokela et al., 2007). It is therefore impossible that genetics factors also influence or moderate the hostility-depressive mood link."

As research suggests that depression is also common in older adults (Jongenelis et al., 2004), we examined the effects of age on the strength of the association between cynical hostility and depressive mood. Results (not shown) revealed no significant interaction effects of age on this association, again supporting the finding that cynical hostility is a long-term vulnerability factor for depressive mood, irrespective of the effects of aging.

Study limitations

In interpreting the present results, it is important to note some limitations. First, our cohort of civil servants included neither blue collar workers, nor individuals who were unemployed or retired; thus it is not representative of the general population, which may limit the generalisability of our findings. Second, we assessed depressive mood instead of clinical depression. However, it has been suggested that significant depressive symptomatology could be a risk for clinical depression (Radloff, 1977). For example, findings from longitudinal data on 9,900 adults drawn from four sites in the United States showed depressive mood to be strongly associated with first-onset of major depression (Horwath et al., 1992). In that study, it was estimated that more than 50% of cases of first-onset of major depression were associated with prior depressive mood (Horwath et al., 1992). Thus, it is possible that cynical hostility is also associated with major depression although this needs to be confirmed in further studies. Third, only 3639 participants had data on cynical hostility (phase 1) and depressive mood (phase 7). As all analyses were based on complete data, only 3 399 (44%) participants were included in the present study. However, this did not compromise the statistical power of our analysis. In addition, compared to participants included in this study, those who did not respond to the CES-D and hostility scales were more likely to be: women (37.3% vs. 25.4%, $p < 0.001$), non-white (13.4% vs. 6.4%; $p < 0.001$), older (24% vs. 19.5% aged ≥ 50 years, $p < 0.001$), and from lower socioeconomic position (27.5% vs. 13.8%, $p < 0.001$). However, controlling for age, sex, ethnicity and socioeconomic position did not alter the graded association between cynical hostility and depressive mood as presented in table 2. We repeated our analyses modelling the association between hostility and depression mood stratified by sex, age groups, ethnicity and socioeconomic position. We found no significant interaction between these variables and hostility in relation to depressive mood, supporting therefore the validity of these findings.

Conclusions and implications

In summary, the present study based on a large occupational cohort, suggests that cynical hostility is a strong and robust predictor of depressive mood, even after a 19-year period. These findings emphasize the importance of considering individual-level psychological factors, alongside with social-cultural and biogenetic factors, in understanding the predictors of depressive mood or depression. If the relationship between cynical hostility and clinical depression is confirmed, it might have implications for the management of depression as understanding the role of hostility in the aetiology of depressive disorders might allow better assignment of a treatment.

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Footnotes:

Conflict of interest: None to declare

References:

- Bifulco A , Bernazzani O , Moran PM , Ball C 2000; Lifetime stressors and recurrent depression: preliminary findings of the Adult Life Phase Interview (ALPHI). *Soc Psychiatry Psychiatr Epidemiol.* 35: 264- 75
- Bifulco A , Brown GW , Moran P , Ball C , Campbell C 1998; Predicting depression in women: the role of past and present vulnerability. *Psychol Med.* 28: 39- 50

- Brown GW , Andrews B , Bifulco A , Veiel H 1990a; Self-esteem and depression. I. Measurement issues and prediction of onset. *Soc Psychiatry Psychiatr Epidemiol.* 25: 200-9
- Brown GW , Bifulco A , Andrews B 1990b; Self-esteem and depression. III. Aetiological issues. *Soc Psychiatry Psychiatr Epidemiol.* 25: 235- 43
- Brown GW , Bifulco A , Veiel HO , Andrews B 1990c; Self-esteem and depression. II. Social correlates of self-esteem. *Soc Psychiatry Psychiatr Epidemiol.* 25: 225- 34
- Brown GW , Harris TO , Hepworth C , Robinson R 1994; Clinical and psychosocial origins of chronic depressive episodes. II. A patient enquiry. *Br J Psychiatry.* 165: 457- 65
- Caspi A , Sugden K , Moffitt TE , Taylor A , Craig IW , Harrington H , McClay J , Mill J , Martin J , Braithwaite A , Poulton R 2003; Influence of life stress on depression: moderation by a polymorphism in the 5-HTT gene. *Science.* 301: 386- 9
- Cohen S , Wills TA 1985; Stress, social support, and the buffering hypothesis. *Psychol Bull.* 98: 310- 57
- Cook WW , Medley DM 1954; Proposed hostility and pharisaic-virtue scales for the MMPI. *Journal of Applied Psychology.* 38: 414- 418
- Everson SA , Kauhanen J , Kaplan GA , Goldberg DE , Julkunen J , Tuomilehto J , Salonen JT 1997; Hostility and increased risk of mortality and acute myocardial infarction: the mediating role of behavioral risk factors. *Am J Epidemiol.* 146: 142- 52
- Felsten G 1996; Hostility, stress and symptoms of depression. *Personality and Individual Differences.* 21: 461- 467
- Hamet P , Tremblay J 2005; Genetics and genomics of depression. *Metabolism.* 54: 10- 5
- Hardy JD , Smith TW 1988; Cynical hostility and vulnerability to disease: social support, life stress, and physiological response to conflict. *Health Psychol.* 7: 447- 59
- Hathaway SR , McKinley JC 1943; The Minnesota Multiphasic Personality Schedule. *The Minnesota Multiphasic Personality Schedule.* 22-
- Heath AC , Cloninger CR , Martin NG 1994; Testing a model for the genetic structure of personality: a comparison of the personality systems of Cloninger and Eysenck. *J Pers Soc Psychol.* 66: 762- 75
- Heponiemi T , Elovainio M , Kivimäki M , Pulkki L , Puttonen S , Keltikangas-Järvinen L 2006; The longitudinal effects of social support and hostility on depressive tendencies. *Social Science & Medicine.* 63: 1374- 1382
- Horwath E , Johnson J , Klerman GL , Weissman MM 1992; Depressive symptoms as relative and attributable risk factors for first-onset major depression. *Archives of General Psychiatry.* 49: 817- 823
- Johnson SL , Winett CA , Meyer B , Greenhouse WJ , Miller I 1999; Social support and the course of bipolar disorder. *J Abnorm Psychol.* 108: 558- 66
- Jokela M , Keltikangas-Jarvinen L , Kivimäki M , Puttonen S , Elovainio M , Rontu R , Lehtimäki T 2007; Serotonin receptor 2A gene and the influence of childhood maternal nurturance on adulthood depressive symptoms. *Arch Gen Psychiatry.* 64: 356- 60
- Jongenelis K , Pot AM , Eisses AMH , Beekman ATF , Kluiters H , Ribbe MW 2004; Prevalence and risk indicators of depression in elderly nursing home patients: the AGED study. *Journal of Affective Disorders.* 83: 135- 142
- Kendler KS , Gatz M , Gardner CO , Pedersen NL 2006; Personality and major depression: a Swedish longitudinal, population-based twin study. *Arch Gen Psychiatry.* 63: 1113- 20
- Kendler KS , Karkowski LM , Prescott CA 1999; Causal relationship between stressful life events and the onset of major depression. *Am J Psychiatry.* 156: 837- 41
- Kivimäki M , Elovainio M , Kokko K , Pulkkinen L , Kortteinen M , Tuomikoski H 2003; Hostility, unemployment and health status: testing three theoretical models. *Soc Sci Med.* 56: 2139- 52
- Mathers CD , Loncar D 2006; Projections of global mortality and burden of disease from 2002 to 2030. *PLoS Med.* 3: e442-
- McCrae RR , Costa PT Jr 1994; The Stability of Personality: Observations and Evaluations. *Current Directions in Psychological Science.* 3: 173- 175
- McCrae RR , Costa PT Jr 1987; Validation of the five-factor model of personality across instruments and observers. *J Pers Soc Psychol.* 52: 81- 90
- Miller TQ , Smith TW , Turner CW , Guizarro ML , Hallet AJ 1996; A meta-analytic review of research on hostility and physical health. *Psychol Bull.* 119: 322- 48
- Moussavi S , Chatterji S , Verdes E , Tandon A , Patel V , Ustun B 2007; Depression, chronic diseases, and decrements in health: results from the World Health Surveys. *Lancet.* 370: 851- 8
- Painuly N , Sharan P , Mattoo SK 2005; Relationship of anger and anger attacks with depression. *European Archives of Psychiatry and Clinical Neuroscience.* 255: 215- 222
- Radloff LS 1977; The CES-D Scale: A Self-Report Depression Scale for Research in the General Population. *Applied Psychological Measurement.* 1: 385-
- Schwartz JE , Friedman HS , Tucker JS , Tomlinson-Keasey C , Wingard DL , Criqui MH 1995; Sociodemographic and psychosocial factors in childhood as predictors of adult mortality. *Am J Public Health.* 85: 1237- 45
- Shaffer DR Social and personality development. Brooks/Cole; Monterey, Calif
- Smith TW 1992; Hostility and health: current status of a psychosomatic hypothesis. *Health Psychol.* 11: 139- 50
- Smith TW , Frohm KD 1985; What's so unhealthy about hostility? Construct validity and psychosocial correlates of the Cook and Medley Ho scale. *Health Psychol.* 4: 503- 20
- Stansfeld S , Marmot M 1992a; Deriving a survey measure of social support: the reliability and validity of the Close Persons Questionnaire. *Soc Sci Med.* 35: 1027- 35
- Stansfeld SA , Marmot MG 1992b; Social class and minor psychiatric disorder in British Civil Servants: a validated screening survey using the General Health Questionnaire. *Psychol Med.* 22: 739- 49

Table 1Bivariate associations of sample characteristics at baseline (phase 1) with cynical hostility score levels (phase 1) and depressive mood (phase 7), n=3399[†].

	Hostility score levels (quartiles)				p value	Depressive mood		p value
	1	2	3	4		No	Yes	
	Lowest			Highest				
Sex					0.218			< 0.001
Male	689(26.8)	682(26.5)	693(27.0)	506(19.7)		2222(86.5)	348(13.5)	
Female	221(26.7)	240(29.0)	231(27.9)	137(16.5)		664(80.1)	165(19.9)	
Age in years					< 0.001			< 0.001
35–40	303(25.0)	295(24.4)	365(30.2)	247(20.4)		988(81.7)	222(18.3)	
40–45	236(26.1)	243(27.5)	246(27.8)	160(18.1)		760(85.9)	125(14.1)	
45–50	167(25.7)	192(29.5)	168(25.8)	123(18.9)		569(87.5)	81(12.5)	
50–55	204(31.2)	192(29.4)	145(22.2)	113(17.3)		569(87.0)	85(13.0)	
SEP					< 0.001			< 0.001
High	357(30.4)	370(31.5)	286(24.4)	160(13.6)		1053(89.8)	120(10.2)	
Middle	452(25.2)	471(26.2)	510(28.4)	362(20.2)		1510(84.1)	285(15.9)	
Low	101(23.1)	81(18.8)	128(29.7)	121(28.1)		323(74.9)	108(25.1)	
Ethnicity					< 0.001			< 0.001
White	877(27.4)	894(27.9)	871(27.2)	562(17.5)		2750(85.8)	454(14.2)	
Other	33(16.9)	28(14.4)	53(27.2)	81(41.5)		136(69.7)	59(30.3)	
BMI					< 0.001			0.778
<19.9	59(29.9)	68(34.5)	42(21.3)	28(14.2)		157(79.7)	40(20.3)	
20–24.9	524(27.4)	520(27.2)	549(28.7)	318(16.6)		1631(85.3)	280(14.7)	
25–29.9	298(25.9)	302(26.3)	287(25.0)	262(22.8)		990(86.2)	159(13.8)	
>30	49(24.5)	42(21.0)	58(29.0)	51(25.5)		155(77.5)	45(22.5)	
Smoking status					0.176			0.296
Never smoker	462(27.0)	472(27.6)	459(26.8)	317(18.5)		1450(84.8)	260(15.2)	
Ex smoker	322(27.1)	329(27.6)	316(26.6)	223(18.7)		1030(86.6)	160(13.4)	
Current smoker	126(25.3)	121(24.2)	149(29.9)	103(20.6)		406(81.4)	93(18.6)	
Heavy alcohol use					0.112			< 0.001
No	131(27.2)	112(23.2)	128(26.6)	111(23.0)		382(79.3)	100(20.7)	
Yes	779(26.7)	810(27.8)	796(27.3)	532(18.2)		2504(85.8)	413(14.2)	
Exercise					0.508			< 0.001
≥1.5 h/week	620(26.9)	623(27.0)	645(28.0)	419(18.2)		1999(86.6)	308(13.4)	
< 1.5 h/week	290(26.6)	299(27.4)	279(25.5)	224(20.5)		887(81.2)	205(18.8)	
Antidepressants medication intake					0.007			< 0.001
No	902(26.9)	914(27.3)	909(27.1)	628(18.7)		2857(85.2)	496(14.8)	
Yes	8(17.4)	8(17.4)	15(32.6)	15(32.6)		29(63.0)	17(37.0)	
Common mental disorder					< 0.001			< 0.001

No	767(31.3)	694(28.2)	619(25.2)	378(15.4)		2193(89.2)	265(10.8)
Yes	14(15.2)	228(24.2)	305(32.4)	265(28.2)		693(73.6)	248(26.4)
Stressful life events					< 0.001		< 0.001
None	330(34.2)	247(25.6)	244(25.3)	145(15.0)		864(89.4)	102(10.6)
One	294(25.4)	349(30.2)	321(27.8)	192(16.6)		1006(87.0)	150(13.0)
Two & more	286(22.4)	326(25.5)	359(28.1)	306(24.0)		1016(79.6)	261(20.4)
Confiding/emotional support					< 0.001		< 0.001
Low	244(23.3)	269(25.7)	308(29.5)	224(21.4)		852(81.5)	193(18.5)
Middle	348(26.9)	329(25.4)	372(28.8)	244(18.9)		1105(85.5)	188(14.5)
High	318(30.0)	324(30.5)	244(23.0)	175(16.5)		929(87.6)	132(12.4)

† Data are number (%)

Table 2

Association of hostility (phase 1) with depressive mood (phase 7), n of depressive participants/n all participants = 513/3399.

	Depressive mood at phase 7 predicted by			
	Cynical hostility		Cynical distrust [±]	
	OR	95% CI	OR	95% CI
Model 1 (Unadjusted model)				
Lowest quartile	1	Ref	1	ref
Middle lowest	1.55	(1.11–2.15)**	1.06	(0.79–1.42)
Middle highest	2.91	(2.15–3.94)***	1.65	(1.28–2.12)***
Highest quartile	5.09	(3.75–6.91)***	3.14	(2.42–4.08)***
Model 2 (adjusted for sex, age, ethnicity, SEP)				
Lowest quartile	1	Ref	1	ref
Middle lowest	1.58	(1.14–2.20)**	1.04	(0.77–1.40)
Middle highest	2.78	(2.05–3.77)***	1.58	(1.22–2.05)***
Highest quartile	4.66	(3.41–6.36)***	2.80	(2.14–3.67)***
Model 3 (Model 2 + BMI, alcohol consumption, physical activity)				
Lowest quartile	1	ref	1	ref
Middle lowest	1.59	(1.14–2.20)**	1.06	(0.79–1.43)
Middle highest	2.84	(2.09–3.86)***	1.61	(1.24–2.08)***
Highest quartile	4.74	(3.47–6.48)***	2.85	(2.17–3.74)***
Model 4 a (Model 3 + stressful life events, confiding/emotional support at phase 1)				
Lowest quartile	1	ref	1	ref
Middle lowest	1.52	(1.09–2.12)*	0.99	(0.73–1.34)
Middle highest	2.59	(1.91–3.53)***	1.49	(1.15–1.94)**

Highest quartile	4.15	(3.03–5.69) ***	2.55	(1.93–3.35) ***
Model 4 b † (Model 3 + stressful life events and confiding/emotional support at phases 1, 2, and 5)				
Lowest quartile	1	ref	1	ref
Middle lowest	1.49	(1.07–2.08) *	1.00	(0.74–1.35)
Middle highest	2.57	(1.89–3.51) ***	1.50	(1.16–1.96) **
Highest quartile	4.13	(3.01–5.68) ***	2.55	(1.93–3.36) ***
Model 5 (Model 4 + antidepressants medication intake + common mental disorder at baseline)				
Lowest quartile	1	ref	1	ref
Middle lowest	1.45	(1.03–2.02) *	0.97	(0.72–1.33)
Middle highest	2.34	(1.72–3.19) ***	1.45	(1.11–1.89) **
Highest quartile	3.62	(2.63–4.98) ***	2.32	(1.76–3.07) ***

* p<0.05,

** p<0.01

*** p<0.001

† Role of cumulative stressful life events and confiding/emotional support (phases 1, 2 and 5) in the association between hostility (phase 1) and depressive mood (phase 7).

± Cynical distrust is a short-form 8-item subscale of the Cook-Medley Hostility Scale.