

Depression in Mexican Americans with diagnosed and undiagnosed diabetes

R. L. Olvera^{1*}, S. P. Fisher-Hoch², D. E. Williamson¹, K. P. Vatcheva² and J. B. McCormick²

¹Department of Psychiatry, Division of Genetic Epidemiology, The University of Texas Health Science Center at San Antonio, San Antonio, TX, USA

²Division of Epidemiology Human Genetics and Environmental Health, The University of Texas School of Public Health, Brownsville, Campus, Brownsville TX, USA

Background. Depression and diabetes commonly co-occur; however, the strength of the physiological effects of diabetes as mediating factors towards depression is uncertain.

Method. We analyzed extensive clinical, epidemiological and laboratory data from $n=2081$ Mexican Americans aged 35–64 years, recruited from the community as part of the Cameron County Hispanic Cohort (CCHC) divided into three groups: Diagnosed (self-reported) diabetes (DD, $n=335$), Undiagnosed diabetes (UD, $n=227$) and No diabetes (ND, $n=1519$). UD participants denied being diagnosed with diabetes, but on testing met the 2010 American Diabetes Association and World Health Organization definitions of diabetes. Depression was measured using the Center for Epidemiological Studies – Depression (CES-D) scale. Weighted data were analyzed using dimensional and categorical outcomes using univariate and multivariate models.

Results. The DD group had significantly higher CES-D scores than both the ND and UD ($p \leq 0.001$) groups, whereas the ND and UD groups did not significantly differ from each other. The DD subjects were more likely to meet the CES-D cut-off score for depression compared to both the ND and UD groups ($p=0.001$), respectively. The UD group was also less likely to meet the cut-off score for depression than the ND group ($p=0.003$). Our main findings remained significant in models that controlled for socio-demographic and clinical confounders.

Conclusions. Meeting clinical criteria for diabetes was not sufficient for increased depressive symptoms. Our findings suggest that the ‘knowing that one is ill’ is associated with depressive symptoms in diabetic subjects.

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Introduction

The co-occurrence of diabetes and depression has been well established with the odds of depression in those with diabetes being approximately twice that of patients without diabetes (Anderson *et al.* 2001). The increased prevalence of depression has been reported in both type 1 (Gendelman *et al.* 2009) and type 2 diabetes (Ali *et al.* 2006) and depression has been linked with poor glycemic control (Lustman *et al.* 2000; Kendzor *et al.* 2014) and diabetes complications (de Groot *et al.* 2001). Potential explanatory models underlying the link between depression and diabetes have included lifestyle changes and stress associated with having diabetes (Dziemidok *et al.* 2011). Other potential contributors to this link include hypothalamic pituitary axis abnormalities (Gragnoli, 2012), and

mechanisms suggesting the effect of stress on insulin resistance through inflammation, stress hormones, the rennin-angiotensin system, endothelial cells, adipocytes and the liver (Black, 2006).

Recent studies, however, draw into question the strength of the physiological effects of diabetes as mediating factors towards depression. A longitudinal 3-year study of patients with type 2 diabetes, found the incidence of depressive symptoms was elevated only in subjects undergoing treatment for diabetes compared to subjects with impaired fasting glucose, those with normal fasting glucose, and those with untreated type 2 diabetes (Golden *et al.* 2008). Moreover subjects with impaired fasting glucose actually had a lower risk of depression compared to subjects with normal fasting glucose, and those with untreated type 2 diabetes had similar risk compared to those with normal fasting glucose (Golden *et al.* 2008). Along these lines, a recent meta-analysis noted the risk for depression was increased in individuals previously diagnosed with type 2 diabetes compared to subjects with undiagnosed diabetes and impaired glucose

* Address for correspondence: R. L. Olvera MD, MPH, Department of Psychiatry, The University of Texas Health Science Center at San Antonio, 7703 Floyd Curl Drive, San Antonio, TX 78229, USA.
(Email: olverar@UTHSCSA.edu)

metabolism (Nouwen *et al.* 2011), furthermore the risk for depression did not differ in subjects with impaired glucose metabolism compared to those with undiagnosed diabetes (Nouwen *et al.* 2011). Similarly data from the National Health and Nutrition Examination Study (NHANES) revealed clinically identified type 2 diabetes was associated with an increase odds ratio of depression, but undiagnosed diabetes was not (Mezuk *et al.* 2013). These studies suggest that 'knowing that one is ill' and being in treatment may be keys for becoming depressed in those with diabetes.

The consequences of depression and diabetes may have major public health implications for Mexican Americans. Existing studies note that diabetes is highly prevalent in Mexican American populations with approximately 25% meeting the World Health Organization's (WHO) definition for diabetes (Quinones *et al.* 2013) and depression is the most common mental illness in Mexican American subjects (Alegria *et al.* 2007). Our own work has found that 29% of Mexican Americans in South Texas suffer from depression (Olvera *et al.* *in press*) and about one-third also suffer from diabetes (Fisher-Hoch *et al.* 2015). Herein we examine the prevalence of depression in subjects with diagnosed diabetes as well as subjects with undiagnosed diabetes and subjects without evidence of diabetes from a randomly selected population-based cohort of Mexican Americans living on the US–Mexico border.

Method

Sample

Participants in this study were recruited between the years 2004–2013, into the Cameron County Hispanic Cohort (CCHC; Fisher-Hoch *et al.* 2010). Households were randomly selected based on the 2000 census tract data in the city of Brownsville, Texas, situated on the US–Mexico border. All selected households were visited, and all occupants over the age of 18 years invited to participate. This cohort is predominantly Mexican American (>98%). Willing participants completed comprehensive questionnaires regarding basic demographic information, medical history, medication use, and social and family history as described previously (Fisher-Hoch *et al.* 2010). All participants provided written informed consent and this study has been approved by the Institutional Review Board of the University of Texas Health Science Center at Houston.

Measures

Based on self-reported medical history, we categorized our subjects ($n = 2081$) into three groups: (1) 'Diagnosed

diabetes' (DD, $n = 335$) based on the subject being previously informed by a health professional that they had diabetes and meeting the 2010 American Diabetes Association (ADA) and WHO definitions of diabetes. (2) 'Undiagnosed diabetes' (UD, $n = 227$) were those who denied being diagnosed with diabetes and were not on appropriate treatment, but who on testing met the 2010 ADA/WHO definitions of diabetes. (3) 'No diabetes' (ND, $n = 1519$) were those who denied having received a diagnosis of diabetes, were not on appropriate treatment, and did not meet the ADA/WHO criteria for the diagnosis at the time of the visit. The 2010 ADA/WHO definitions of diabetes is: a mean fasting blood glucose (FBG) >126 mg/dl on two consecutive visits, and/or a glycosylated hemoglobin (HbA_{1c}) of >6.5% (ADA, 2010).

Depression was measured using the Center for Epidemiological Studies – Depression (CES-D) a 20-item scale developed for epidemiologic studies of depressive symptoms in the general population (Radloff, 1977) with extensive use in epidemiological samples (Sayetta & Johnson, 1980). A Spanish version has been used with Mexican Americans, with good reliability with Cronbach's $\alpha = 0.88$ and Spearman–Brown split-half = 0.91 (Roberts, 1980). Further studies with Spanish versions of the CES-D have also noted good validity (Ring & Marquis, 1991; Ruiz-Grosso *et al.* 2012) and diagnostic accuracy in Spanish-speaking populations (Reuland *et al.* 2009). Our own sample is consistent with other studies as the CES-D for the Spanish respondents ($n = 1530$) has a Cronbach's $\alpha = 0.89$ with a Spearman–Brown split-half = 0.83, in the English respondents ($n = 551$) Cronbach's $\alpha = 0.91$ with Spearman–Brown split-half = 0.88 and for the total sample Cronbach's $\alpha = 0.90$ with Spearman–Brown split-half = 0.89. Consistent with prior studies (Zich *et al.* 1990), we classified individuals as non-depressed if their CES-D score was <16, and depressed if their score was ≥ 16 . Although an acculturation scale was not used we measured preferred language, country of birth, time lived in the United States, and scores on language ability in Spanish measured by the Word Accentuation Test (Del Ser *et al.* 1997) and English measured by the reading portion of the Wide Range Achievement Test 3 (WRAT-3; Wilkinson, 1993).

Anthropometric measures were taken as described previously (Fisher-Hoch *et al.* 2010). Blood specimens were taken and aliquots immediately stored at -70°C for a range of clinical and experimental assays. Blood glucose measurement was performed on site, HbA_{1c} was measured by high-performance liquid chromatography and stored specimens were sent in batches to a Clinical Laboratory Improvement Amendments (CLIA)-approved clinical laboratory for clinical chemistries.

Statistical analysis

This is a nested cohort of $n=2081$ with complete data taken from the CCHC ($n=3500$). These 2081 subjects did not differ from the entire cohort in terms of age and gender status. We report results at the participant level. Our sample is 67% female therefore we incorporated sampling weights into our analysis as fully described previously to enhance generalizability (Fisher-Hoch *et al.* 2010). In the survey data analysis, taking into consideration of the complex sampling design, we also accounted for the potential clustering effect among participants from the same household. All analyses were performed using SAS v. 9.1 (SAS Institute Inc., USA) and Stata 10 SE (StataCorp LP, USA). For descriptive purposes, categorical variables for demographic and clinical characteristics were summarized in unweighted frequencies and weighted percentages. The Rao–Scott design-adjusted χ^2 test was used to test for equality of proportions across the diabetes status groups. Continuous variables for demographic and clinical characteristics were summarized using weighted means and their standard errors. To assess independent effects of the multiple factors on the CES-D score, a multivariable weighted linear regression model for depression was performed with Wald's F tests to assess interactions. We assessed the overall effect of diabetes status on depression score in bivariate regression using design-based Wald F tests. *Post-hoc* pairwise comparisons of the means were assessed for significance using a Tukey–Kramer adjustment to correct for the multiple comparisons. In both linear and bivariate regression analyses a step wise strategy was performed to test for potential confounders. We only retained variables whose presence altered the estimated coefficients of other variables in the model by more than 10%. A variance inflation factor (VIF) indicated that there was not problematic multicollinearity among the independent variables included in the regression models ($VIF < 1.5$). *Post-hoc* analyses using t tests were performed comparing depression scores within subjects diagnosed with diabetes divided into those with and without reported medical complications. A *post-hoc* analysis of variance (ANOVA) was used to compare depression scores between the three groups after removing subjects with skin ulcers.

Results

We found the DD participants had significantly higher depression scores than both the UD and ND groups ($p \leq 0.001$) and the UD and ND groups did not significantly differ from each other on CES-D scores (see Table 1, Fig. 1). The DD group was significantly

older than the UD and ND groups ($p < 0.0001$) and both the DD and UD groups had significantly higher body mass index (BMI) than the ND group ($p < 0.001$) (see Table 1). Repeating the analyses including age, gender and BMI in the model, the difference between groups on CES-D depression scores remained significant ($p < 0.001$) with the DD subjects having significantly higher scores than the both the UD and ND groups, respectively, on pairwise comparisons ($p < 0.001$), with no significant difference between the UD and ND subjects. This model revealed a significant main effect for gender ($p < 0.001$) with females having significantly higher depression scores than males ($p < 0.001$) across all groups, without an interaction effect for gender.

Using the CES-D established cut-off score of ≥ 16 as suggestive of depression 41% of DD subjects qualified as depressed whereas 26% of ND subjects and only 17% of UD subjects were depressed ($\chi^2 = 19.57$, df_2 , $p < 0.001$). The increased percentage of DD subjects meeting the cut-off for depression was significant compared to UD subjects ($\chi^2 = 21.03$, df_1 , $p < 0.001$) and the ND group ($\chi^2 = 10.52$, df_1 , $p = 0.001$), respectively. In addition there was a significant difference between the ND and UD groups ($\chi^2 = 4.65$, df_1 , $p = 0.003$).

On laboratory measures the DD and UD subjects did not differ from each other in terms of HbA_{1c}; however, the DD subjects did have significantly higher FBG levels than UD subjects ($p < 0.001$). As anticipated both the DD and UD subjects had significantly higher HbA_{1c} and FBG levels than the ND group ($p < 0.001$), respectively. HbA_{1c} and FBG were highly correlated ($r = 0.66$, $p < 0.0001$) and since these variables are used to define the presence of diabetes, we did not attempt to covary for their effects on depression. Within the total sample only FBG (not HbA_{1c}) was modestly correlated ($r = 0.074$, $p = 0.001$) with depression scores; however, within each group (DD, UD, ND) neither HbA_{1c} nor FBG were significantly correlated with depression scores.

Examining socio-demographic variables, revealed the DD subjects scored lowest on both Spanish- and English-language tests compared to the ND group, and were least likely to have finished high school and had the highest levels of unemployment compared to both of the other groups (Table 1). The DD subjects also had lived the longest in the United States but there was no difference between groups on the percent born in Mexico or Spanish preference (Table 1).

We then explored the effects of potential confounding variables such as gender, marital status, employment and education in both linear (Table 2) and logistic (Table 3) regression models to examine depression scores as dimensional and categorical outcomes respectively. In the linear model the socio-demographic

Table 1. Demographic and clinical variables: weighted means and percentages

| | Diagnosed with diabetes (<i>n</i> = 335) Mean (s.e.) | Undiagnosed with diabetes (<i>n</i> = 227) Mean (s.e.) | No diabetes (<i>n</i> = 1519) Mean (s.e.) | <i>p</i> |
|-----------------------------------------------|----------------------------------------------------------|------------------------------------------------------------|-----------------------------------------------|----------|
| Age | 58.00 (1.7) ^a | 45.93 (1.9) ^b | 43.11 (0.9) ^b | <0.0001 |
| Body mass index | 33.28 (0.7) ^a | 33.18 (0.7) ^a | 30.04 (0.3) ^b | <0.0001 |
| CES-D score | 15.65 (0.9) ^a | 9.73 (1.0) ^b | 10.86 (0.4) ^b | <0.0001 |
| HbA _{1c} | 7.33 (0.2) ^a | 7.40 (0.2) ^a | 4.78 (0.0) ^b | <0.0001 |
| HbA _{1c} mmol/mol (IFCC units) | 57 | 57 | 29 | |
| FBG, mg/dl | 171.34 (5.1) ^a | 134.07 (6.8) ^b | 96.11 (0.4) ^c | <0.0001 |
| Spanish ability (<i>n</i> = 2031) | 35.60 (0.86) ^a | 34.66 (1.39) ^a | 38.32 (0.37) ^b | 0.0014 |
| English ability (<i>n</i> = 1880) | 26.01 (1.43) ^a | 27.61 (1.67) ^{ab} | 31.42 (0.65) ^b | 0.0006 |
| Years lived in USA (<i>n</i> = 1879) | 30.55 (2.97) ^a | 25.32 (2.00) ^{ab} | 21.73 (0.91) ^b | 0.0006 |
| | <i>n</i> (%) | <i>n</i> (%) | <i>n</i> (%) | <i>p</i> |
| Depression (%) | 123 (41.3%) ^a | 46 (17.2%) ^b | 452 (25.8%) ^c | <0.0001 |
| Female (%) | 228 (62.7%) | 145 (51.4%) | 1013 (55.9%) | 0.19 |
| Insured (%) | 134 (46.0%) ^a | 63 (31.3%) ^b | 401 (33.1%) ^b | 0.03 |
| Born in Mexico (%) | 221 (61.46) | 150 (60.78) | 959 (58.99) | 0.8467 |
| Spanish as a preferred language (%) | 271 (77.13) | 173 (69.85) | 1125 (69.41) | 0.1750 |
| Married (%) | 206 (62.3%) | 147 (66.9%) | 975 (63.6%) | 0.77 |
| High school education | 64 (19.1) ^a | 94 (41.4) ^b | 756 (49.8) ^b | <0.0001 |
| Employed full time | 69 (18.3) ^a | 70 (32.5) ^b | 535 (36.9) ^b | |
| Employed part time | 51 (16.7) ^a | 50 (22.5) ^b | 270 (16.4) ^a | <0.0001 |
| Unemployed | 215 (65.0) ^a | 107 (45.0) ^b | 714 (46.7) ^b | |
| Cardiovascular disease, high lipids or cancer | 298 (91.2) ^a | 132 (55.9) ^b | 849 (58.0) ^b | <0.0001 |

CES-D, Center for Epidemiological Studies – Depression; s.e., standard error; *n*, frequency IFCC, International Federation of Clinical Chemistry; FBG, fasting blood glucose.

Different superscript letters (a, b, c) denote significantly different pairwise comparisons $p < 0.05$.

Frequencies are unweighted. Percentages use weighted data. Means and standard errors (s.e.) are weighted.

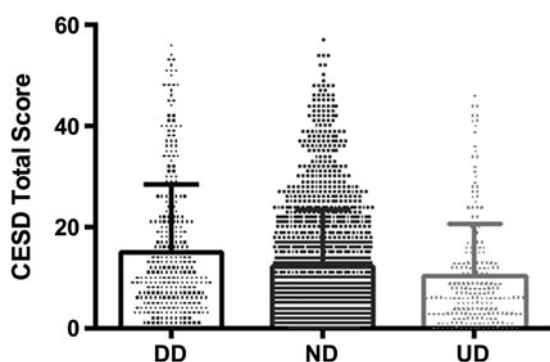


Fig. 1. Center for Epidemiological Studies – Depression (CES-D) scores by diagnostic groups. DD, Diagnosed with diabetes; ND, no diabetes; UD, undiagnosed diabetes.

variable regression estimates were in the anticipated direction as female gender, and lacking a high-school diploma were predictive of higher depression scores whereas being married and in full-time employment were significant for predicting lower depression scores. Notably, even with these socio-demographic variables in the model the DD group still was significantly higher

on CES-D depression scores compared to ND subjects whereas the UD and ND groups did not significantly differ from each other (Table 2). Likewise, using the CES-D cut-off score in a logistic regression model, the DD group had a significantly higher odds ratio for depression compared to the ND group, whereas being UD was protective for depression even with other variables in the model (see Table 3). Similar to the linear model, being female and lacking a high-school degree increased the risk for depression and being married and having full-time employment were protective (See Table 3). Potential acculturation factors: English and Spanish ability, preferred language, country of birth and years lived in United States did not significantly contribute to these models nor were they significant for depression.

When we included the presence of additional medical conditions (cardiovascular disease, high blood pressure, high lipids, and cancer) in a logistic regression model with other significant socio-demographic variables, our main findings remained unchanged as the DD group remained at higher risk of depression and the UD subjects were at lower risk compared to the ND

Table 2. Linear regression model for CES-D depression score controlling for socio-demographic variables^a

| Parameter | Estimate | S.E. | t | p |
|-----------------------------------|----------|------|-------|---------|
| Estimated regression coefficients | | | | |
| Intercept | 10.80 | 1.14 | 9.47 | <0.0001 |
| Diagnosed diabetes | 3.77 | 1.01 | 3.72 | 0.0002 |
| Undiagnosed diabetes | -1.00 | 1.05 | -0.95 | 0.3418 |
| No diabetes | 0.00 | 0.00 | | |
| Female | 3.50 | 0.74 | 4.75 | <0.0001 |
| Male | 0.00 | 0.00 | | |
| Married | -2.83 | 0.69 | -4.08 | <0.0001 |
| Not married | 0.00 | 0.00 | | |
| Full-time employed | -1.96 | 0.82 | -2.39 | 0.0170 |
| Part-time employed | -0.63 | 0.90 | -0.70 | 0.4860 |
| Not employed | 0.00 | 0.00 | | |
| No high school | 1.66 | 0.69 | 2.42 | 0.0155 |
| High school | 0.00 | 0.00 | | |

CES-D, Center for Epidemiological Studies – Depression.

^a All listed variables included in the model.

Table 3. Logistic regression model for categorical (CES-D ≥ 16) depression by socio-demographic variables^a

| Effect | OR | 95% CI |
|----------------------------------------------|------|-----------|
| OR estimates | | |
| Diagnosed diabetes <i>v.</i> no diabetes | 1.70 | 1.12–2.58 |
| Undiagnosed diabetes <i>v.</i> no diabetes | 0.58 | 0.36–0.94 |
| Female <i>v.</i> male | 2.12 | 1.51–2.97 |
| Married <i>v.</i> not married | 0.62 | 0.47–0.82 |
| No high-school ed. <i>v.</i> high-school ed. | 1.38 | 1.04–1.83 |
| Employed full time <i>v.</i> (not employed) | 0.59 | 0.41–0.84 |
| Employed part time <i>v.</i> (not employed) | 0.91 | 0.62–1.33 |

CES-D, Center for Epidemiological Studies – Depression;
OR, odds ratio; CI, confidence interval.

^a All listed variables included in the model.

group (Table 4). In a *post-hoc* analysis we attempted to examine potential diabetes-related medical complications within the DD group (the other groups denied such complications). Within the DD group $n=37$ (11%) reported retinopathy, $n=19$ (6%) reported ketoacidosis, $n=4$ (1%) reported needing dialyses and 24 (7%) reported having skin ulcers. Within this group only the presence of a skin ulcer was predictive of increased depression scores, as those with ulcers had a mean CES-D of 22.21 (S.E. = 3.68) compared to those without (14.44, S.E. = 0.73, $p=0.003$). Removing the 24 subjects with skin ulcers did not alter our main findings as the DD group still had significantly higher CES-D scores compared to the UD ($p<0.001$) and ND

Table 4. Logistic regression model for categorical (CES-D ≥ 16) by socio-demographic variables^a and medical illness

| Effect | OR | 95% CI |
|-----------------------------------------------|------|-----------|
| Odds ratio estimates | | |
| Diagnosed diabetes <i>v.</i> no diabetes | 1.90 | 1.25–2.90 |
| Undiagnosed diabetes <i>v.</i> no diabetes | 0.62 | 0.38–0.99 |
| Female <i>v.</i> male | 2.52 | 1.85–3.44 |
| Married <i>v.</i> not married | 0.64 | 0.48–0.84 |
| Cardiovascular disease, high lipids or cancer | 1.11 | 0.82–1.50 |

CES-D, Center for Epidemiological Studies – Depression;
OR, odds ratio; CI, confidence interval.

^a All listed variables included in the model.

($p=0.002$) subjects. Within the DD group we had data on duration of diabetes in a subset ($n=164$) with a mean duration of 10.82 years (S.E. = 0.63). We did not find a significant difference in the mean duration of diabetes in depressed (10.00 years, S.E. = 1.06) compared to non-depressed (11.29 years, S.E. = 0.79) ($p=0.33$) subjects. Along these lines there was not a significant correlation between depression scores and illness duration ($r=-0.09$, $p=0.27$).

Discussion

Our main findings were that subjects with DD had higher depression scores and a higher prevalence of depression compared to UD and those with ND. What is noteworthy is that participants with UD despite having obesity, and elevated HbA_{1c} and FPG, reported the lowest depressive symptoms and when using dichotomized depression cut-offs, even had a lower odds ratio for depression relative to those in the ND group. As in other studies (Fisher *et al.* 2001; Egede & Zheng, 2003) we found significant effects for potential socio-demographic confounders such as female gender, low education and unemployment for depression. However, even when we included these variables in our models, the main effect of diabetes diagnostic status remained significant.

Our findings suggest that 'knowing that one is ill' and being in treatment could be major contributors to depression in persons with diabetes. A systematic review of the responses to being diagnosed with diabetes revealed up to one half of newly diagnosed patients reporting negative emotions; however, there was great variability in the emotional cognitive and behavioral responses (Thoolen *et al.* 2006). Many newly diagnosed patients initially downplay the seriousness of their illness (Thoolen *et al.* 2006) and full

understanding of the implications of illness may take years (Lawson *et al.* 2008). Individual factors that influence the emotional response of diabetic patients to receiving the diagnosis include personality traits (Lyness *et al.* 1998), perceptions of illness, coping mechanisms (Duangdao & Roesch, 2008; Bazzazian & Besharat, 2012) and severity of symptoms (Thoolen *et al.* 2006). An example of the complexity of this issue is the personality trait of 'neuroticism' that is defined by subdomains of worry and self-consciousness (Lane *et al.* 2000). In some instances neuroticism may be protective in diabetes; providing the vigilance needed for good glycemic control (Lane *et al.* 2000). However, subjects with high neuroticism may also be at greater risk for depression (Fanous *et al.* 2002) especially when exposed to increased illness burden (Lyness *et al.* 1998). Furthermore, the 'Burden of Illness' i.e. worries about complications has been associated with depression in patients with diabetes (Karlson & Agardh, 1997) suggesting there may be a tipping point where the propensity to worry combined with diabetes becomes deleterious. These concepts may account for our findings where within those diagnosed with diabetes, the presence of ulcerations, an obvious physical symptom, was significantly associated with increased CES-D scores. This suggests that there may be a certain threshold (i.e. a notable clinical manifestation of illness) that cannot be readily ignored that leads to the emergence of mood symptoms. Once the threshold is reached then a vicious cycle may occur as depression can hamper self-care and the ability to follow healthy diet and exercise (Katon *et al.* 2010b). In Mexican Americans with diabetes, this synergy between depression and diabetes has been documented where the presence of depression and diabetes predicted earlier mortality, and multiple complications that affected daily living (Black *et al.* 2003).

Given the negative impact from having both depression and diabetes there has been added attention to addressing the emotional response to living with this serious chronic illness as part of the overall treatment. In addition the importance of considering family dynamics has been demonstrated as families with high conflict have poorer diabetic control (Miller-Johnson *et al.* 1994; Fisher *et al.* 2000) and depression (Fisher *et al.* 2001). With such issues in mind psychosocial support is recommended as a standard of care by the American Diabetes Association (ADA, 2014); however, it is only as a category 'C' as the findings are considered relatively weak with conflicting empirical evidence. The evidence does note benefits from psychological and pharmacological interventions in terms of depressive symptoms but there have been mixed results for glycemic control (Markowitz *et al.* 2011; Baumeister *et al.* 2012, 2014). A collaborative

care model that included pharmacotherapy, individualized goals, medication adherence monitoring, motivational coaching and self-care guide resulted in improvement across multiple domains including depression scores, glycemic control, and reports of quality of life and satisfaction compared to non-intervention controls (Katon *et al.* 2010a). The collaborative care intervention models' benefits for depressive symptoms and adherence have been replicated, however, the beneficial effects of this intervention on glycemic control have not been consistent (Huang *et al.* 2013).

In regards to Mexican American populations, numerous individual and cultural factors have been found to impact the management of diabetes (Brown & Hanis, 2014). For example, even though lifestyle changes such a healthy diet and increased physical activity are the accepted interventions for controlling type 2 diabetes (Tuomilehto *et al.* 2001), these dietary and behavioral changes, if perceived as 'restrictions', have actually been associated with increased depression in some subjects with diabetes (Karlson & Agardh, 1997). Focus groups with Mexican Americans with diabetes found they did not want to participate in weight loss focused outcomes in particular those that with an emphasis on 'diet' but were highly motivated by a concern for the welfare of their children and other family members (Brown & Hanis, 2014). Other issues that may be particularly salient to Mexican Americans include beliefs that being heavyset represents health (Stern *et al.* 1982; Diaz *et al.* 2007), food as a representation of love (Allan, 1998) or food security as a symbol of socio-economic status (Kumanyika, 2008) may magnify the negative perception to dietary restrictions. However culturally sensitive interventions that include family involvement and that incorporate cultural foods have been successful in better glycemic control in Mexican Americans from the Texas–Mexico border (Brown *et al.* 2002).

Even though we did not use an acculturation scale, we were able to examine English and Spanish ability, preferred language, country of birth and years lived in the United States, and such items have been reliable measures of acculturation (Marin *et al.* 1987; Wallen *et al.* 2002). As noted, we did not find a significant association between any of these measure and depression but it is interesting to note that the ND subjects had spent the fewest years living in the United States. Foreign-born immigrant status and longer duration in the United States have been associated with increased obesity (Kaplan *et al.* 2004) and greater acculturation may also lead to poorer dietary practices (Neuhouser *et al.* 2004; Montez & Eschbach, 2008). However, the ND group also scored the highest on tests of language, were the most likely to finish high school and had the highest levels of employment

which suggests the ND subjects have overall better function. Of note both of language tests used: the Word Accentuation Test and the reading portion of the WRAT-3 have been associated with levels of intelligence (Del Ser *et al.* 1997; Griffin *et al.* 2002). Higher intelligence may account for better function across multiple domains and better health outcomes (Gottfredson & Deary, 2004; Batty *et al.* 2007). As our study is cross sectional we cannot conclude if this better function is part of why they are free of diabetes or whether the differences between groups are due to the detrimental effects of diabetes and associated conditions. It might seem paradoxical that the ND group scored highest on both English and Spanish but many border residents are bilingual and bicultural. Given the proximity to Mexico, the ability to access services across the border, and the potential social and employment advantages to remaining bicultural on the border (Crouch, 2004), the effects of acculturation toward US cultural norms on our outcomes may have been diminished.

While there are numerous strengths of our population-based randomly recruited sample of Mexican Americans living on the US–Mexico border, there are several limitations to bear in mind when evaluating our results. First, this was a cross-sectional study and therefore cannot establish causality as we did not prospectively follow them in time after receiving their diabetes diagnosis. It is possible that those with depression were more likely to receive the diagnosis of diabetes as much as it is possible that having the diagnosis of diabetes increased the risk for depression. Only a longitudinal study could disentangle these two possibilities. Second, although the CES-D is a well-accepted measure of depressive symptoms in a population (Radloff, 1977; Lewinsohn *et al.* 1997), it does not follow strict DSM criteria, cannot establish chronicity and number of episodes and does not account for confounding or co-morbid or pre-existing psychiatric conditions which may play a role in depression in diabetic patients (Bot *et al.* 2010). Despite these limitations the breadth and sample size of this study allow for exploration based diabetic status, knowledge of illness and multiple established risk factors. In addition the randomized, population-based recruitment strategy used allows us to extrapolate to the community. Furthermore, even though our sample was Mexican American our findings are identical to a large ($n > 5000$) longitudinal study (Golden *et al.* 2008) a meta-analysis of 13 studies comprising $n = 1483$ cases (Nouwen *et al.* 2011) and to a nationally representative cross-sectional survey (NHANES) $n = 3183$ (Mezuk *et al.* 2013) confirming findings from samples of different racial and ethnic make-up. These findings dispute the notion that major depressive disorder results

directly from diabetes in favor of a multidimensional approach including consideration of biological and psychosocial factors (Talbot & Nouwen, 2000) that appear to accompany being diagnosed with diabetes. Hispanics now comprise the largest ethnic minority group residing in the United States accounting for 15% of the population; and Mexican Americans are the single largest Hispanic subgroup and number over 46 million people (Ennis *et al.* 2011). The consequences of depression and diabetes in this population have major public health implications with the need for individualized culturally sensitive personalized treatment that includes both medical and psycho-social considerations (Brown *et al.* 2002; Dziemidok *et al.* 2011).

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Declaration of Interest

None.

References

- American Diabetes Association (ADA) (2010). American diabetes association. Standards of medical care in diabetes – 2010. *Diabetes Care* **33**, S11–S61.
- Alegria M, Mulvaney-Day N, Torres M, Polo A, Cao Z, Canino G (2007). Prevalence of psychiatric disorders across Latino subgroups in the United States. *American Journal of Public Health* **97**, 68–75.
- Ali S, Stone MA, Peters JL, Davies MJ, Khunti K (2006). The prevalence of co-morbid depression in adults with Type 2 diabetes: a systematic review and meta-analysis. *Diabetic Medicine* **23**, 1165–1173.
- Allan JD (1998). Explanatory models of overweight among African American, Euro-American, and Mexican American women. *Western Journal of Nursing Research* **20**, 45–66.
- American Diabetes Association (2014). Standards of medical care in diabetes – 2014. *Diabetes Care* **37** (Suppl. 1), S14–S80.

- Anderson RJ, Freedland KE, Clouse RE, Lustman PJ** (2001). The prevalence of comorbid depression in adults with diabetes: a meta-analysis. *Diabetes Care* **24**, 1069–1078.
- Batty GD, Deary IJ, Gottfredson LS** (2007). Premorbid (early life) IQ and later mortality risk: systematic review. *Annals of Epidemiology* **17**, 278–288.
- Baumeister H, Hutter N, Bengel J** (2012). Psychological and pharmacological interventions for depression in patients with diabetes mellitus and depression. *Cochrane Database of Systemic Reviews* **12**, CD008381.
- Baumeister H, Hutter N, Bengel J** (2014). Psychological and pharmacological interventions for depression in patients with diabetes mellitus: an abridged Cochrane review. *Diabetic Medicine* **31**, 773–786.
- Bazzazian S, Besharat MA** (2012). An explanatory model of adjustment to type I diabetes based on attachment, coping, and self-regulation theories. *Psychology Health & Medicine* **17**, 47–58.
- Black PH** (2006). The inflammatory consequences of psychologic stress: relationship to insulin resistance, obesity, atherosclerosis and diabetes mellitus, type II. *Medical Hypotheses* **67**, 879–891.
- Black SA, Markides KS, Ray LA** (2003). Depression predicts increased incidence of adverse health outcomes in older Mexican Americans with type 2 diabetes. *Diabetes Care* **26**, 2822–2828.
- Bot M, Pouwer F, Ormel J, Slaets JP, de Jonge P** (2010). Predictors of incident major depression in diabetic outpatients with subthreshold depression. *Diabetic Medicine* **27**, 1295–1301.
- Brown SA, Garcia AA, Kouzekanani K, Hanis CL** (2002). Culturally competent diabetes self-management education for Mexican Americans: the Starr County border health initiative. *Diabetes Care* **25**, 259–268.
- Brown SA, Hanis CL** (2014). Lessons learned from 20 years of diabetes self-management research with Mexican Americans in Starr County, Texas. *The Diabetes Educator* **40**, 476–487.
- Crouch N** (2004). *Mexicans & Americans: Cracking The Cultural Code*. Nicholas Brealey: London.
- de Groot M, Anderson R, Freedland KE, Clouse RE, Lustman PJ** (2001). Association of depression and diabetes complications: a meta-analysis. *Psychosomatic Medicine* **63**, 619–630.
- Del Ser T, Gonzalez-Montalvo JI, Martinez-Espinosa S, Delgado-Villapalos C, Bermejo F** (1997). Estimation of premorbid intelligence in Spanish people with the Word Accentuation Test and its application to the diagnosis of dementia. *Brain and Cognition* **33**, 343–356.
- Diaz VA, Mainous AG III, Pope C** (2007). Cultural conflicts in the weight loss experience of overweight Latinos. *International Journal of Obesity* **31**, 328–333.
- Duangdao KM, Roesch SC** (2008). Coping with diabetes in adulthood: a meta-analysis. *Journal of Behavioral Medicine* **31**, 291–300.
- Dziemidok P, Makara-Studzinska M, Jarosz MJ** (2011). Diabetes and depression: a combination of civilization and life-style diseases is more than simple problem adding – literature review. *Annals of Agricultural and Environmental Medicine* **18**, 318–322.
- Egede LE, Zheng D** (2003). Independent factors associated with major depressive disorder in a national sample of individuals with diabetes. *Diabetes Care* **26**, 104–111.
- Ennis SR, Rios-Vargas M, Albert NG** (2011). The Hispanic Population: 2010. 2010 Census Briefs, United States Census Bureau 1–3.
- Fanous A, Gardner CO, Prescott CA, Cancro R, Kendler KS** (2002). Neuroticism, major depression and gender: a population-based twin study. *Psychological Medicine* **32**, 719–728.
- Fisher-Hoch SP, Rentfro AR, Salinas JJ, Perez A, Brown HS, Reininger BM, Restrepo BI, Wilson JG, Hossain MM, Rahbar MH, Hanis CM, McCormick JB** (2010). Socioeconomic status and prevalence of obesity and diabetes in a Mexican American community, Cameron County, Texas, 2004–2007. *Preventing Chronic Disease* **7**, A53.
- Fisher-Hoch SP, Vatcheva KP, Rahbar MH, McCormick JB** (2015). Undiagnosed diabetes and pre-diabetes in Health Disparities. *PLoS ONE* **10**, e0133135.
- Fisher L, Chesla CA, Mullan JT, Skaff MM, Kanter RA** (2001). Contributors to depression in Latino and European-American patients with type 2 diabetes. *Diabetes Care* **24**, 1751–1757.
- Fisher L, Chesla CA, Skaff MM, Gilliss C, Mullan JT, Bartz RJ, Kanter RA, Lutz CP** (2000). The family and disease management in Hispanic and European-American patients with type 2 diabetes. *Diabetes Care* **23**, 267–272.
- Gendelman N, Snell-Bergeon JK, McFann K, Kinney G, Paul Wadwa R, Bishop F, Rewers M, Maahs DM** (2009). Prevalence and correlates of depression in individuals with and without type 1 diabetes. *Diabetes Care* **32**, 575–579.
- Golden SH, Lazo M, Carnethon M, Bertoni AG, Schreiner PJ, Diez Roux AV, Lee HB, Lyketos C** (2008). Examining a bidirectional association between depressive symptoms and diabetes. *Journal of the American Medical Association* **299**, 2751–2759.
- Gottfredson LS, Deary IJ** (2004). Intelligence predicts health and longevity, but why? *Current Directions in Psychological Science* **13**, 1–4.
- Graglioli C** (2012). Depression and type 2 diabetes: cortisol pathway implication and investigational needs. *Journal of Cellular Physiology* **227**, 2318–2322.
- Griffin SL, Mindt MR, Rankin EJ, Ritchie AJ, Scott JG** (2002). Estimating premorbid intelligence: comparison of traditional and contemporary methods across the intelligence continuum. *Archives of Clinical Neuropsychology* **17**, 497–507.
- Huang Y, Wei X, Wu T, Chen R, Guo A** (2013). Collaborative care for patients with depression and diabetes mellitus: a systematic review and meta-analysis. *BMC Psychiatry* **13**, 260.
- Kaplan MS, Huguette N, Newsom JT, McFarland BH** (2004). The association between length of residence and obesity among Hispanic immigrants. *American Journal of Preventive Medicine* **27**, 323–326.
- Karlson B, Agardh CD** (1997). Burden of illness, metabolic control, and complications in relation to depressive symptoms in IDDM patients. *Diabetic Medicine* **14**, 1066–1072.

- Katon WJ, Lin EH, Von Korff M, Ciechanowski P, Ludman EJ, Young B, Peterson D, Rutter CM, McGregor M, McCulloch D (2010a). Collaborative care for patients with depression and chronic illnesses. *New England Journal of Medicine* **363**, 2611–2620.
- Katon WJ, Russo JE, Heckbert SR, Lin EH, Ciechanowski P, Ludman E, Young B, Von Korff M (2010b). The relationship between changes in depression symptoms and changes in health risk behaviors in patients with diabetes. *International Journal of Geriatric Psychiatry* **25**, 466–475.
- Kendzor DE, Chen M, Reininger BM, Businelle MS, Stewart DW, Fisher-Hoch SP, Rentfro AR, Wetter DW, McCormick JB (2014). The association of depression and anxiety with glycemic control among Mexican Americans with diabetes living near the U.S.-Mexico border. *BMC Public Health* **14**, 176.
- Kumanyika SK (2008). Environmental influences on childhood obesity: ethnic and cultural influences in context. *Physiology & Behavior* **94**, 61–70.
- Lane JD, McCaskill CC, Williams PG, Parekh PI, Feinglos MN, Surwit RS (2000). Personality correlates of glycemic control in type 2 diabetes. *Diabetes Care* **23**, 1321–1325.
- Lawson VL, Bundy C, Harvey JN (2008). The development of personal models of diabetes in the first 2 years after diagnosis: a prospective longitudinal study. *Diabetic Medicine* **25**, 482–490.
- Lewinsohn PM, Seeley JR, Roberts RE, Allen NB (1997). Center for Epidemiologic Studies Depression Scale (CES-D) as a screening instrument for depression among community-residing older adults. *Psychology and Aging* **12**, 277–287.
- Lustman PJ, Anderson RJ, Freedland KE, de Groot M, Carney RM, Clouse RE (2000). Depression and poor glycemic control: a meta-analytic review of the literature. *Diabetes Care* **23**, 934–942.
- Lyness JM, Duberstein PR, King DA, Cox C, Caine ED (1998). Medical illness burden, trait neuroticism, and depression in older primary care patients. *American Journal of Psychiatry* **155**, 969–971.
- Marin G, Sabogal F, Marin BV, Oterosabogal R, Perezstable EJ (1987). Development of a Short Acculturation Scale for Hispanics. *Hispanic Journal of Behavioral Sciences* **9**, 183–205.
- Markowitz SM, Gonzalez JS, Wilkinson JL, Safren SA (2011). A review of treating depression in diabetes: emerging findings. *Psychosomatics* **52**, 1–18.
- Mezuk B, Johnson-Lawrence V, Lee H, Rafferty JA, Abdou CM, Uzogara EE, Jackson JS (2013). Is ignorance bliss? Depression, antidepressants, and the diagnosis of prediabetes and type 2 diabetes. *Health Psychology* **32**, 254–263.
- Miller-Johnson S, Emery RE, Marvin RS, Clarke W, Lovinger R, Martin M (1994). Parent-child relationships and the management of insulin-dependent diabetes mellitus. *Journal of Consulting and Clinical Psychology* **62**, 603–610.
- Montez JK, Eschbach K (2008). Country of birth and language are uniquely associated with intakes of fat, fiber, and fruits and vegetables among Mexican-American women in the United States. *Journal of the American Dietetic Association* **108**, 473–480.
- Neuhouser ML, Thompson B, Solomon CC (2004). Higher fat intake and lower fruit and vegetables intakes are associated with greater acculturation among Mexicans living in Washington state. *Journal of the American Dietetic Association* **104**, 51–57.
- Nouwen A, Nefs G, Caramlau I, Connock M, Winkley K, Lloyd CE, Peyrot M, Pouwer FC (2011). Prevalence of depression in individuals with impaired glucose metabolism or undiagnosed diabetes: a systematic review and meta-analysis of the European Depression in Diabetes (EDID) Research Consortium. *Diabetes Care* **34**, 752–762.
- Olvera RL, Williamson DE, Fisher-Hoch SP, Vatcheva KP, McCormick JB (in press). Depression, obesity, and metabolic syndrome: prevalence and risks of comorbidity in a population-based representative sample of Mexican Americans. *Journal of Clinical Psychiatry*.
- Quinones AR, Liang J, Ye W (2013). Differences in diabetes mellitus onset for older Black, White, and Mexican Americans. *Ethnicity & Disease* **23**, 310–315.
- Radloff LS (1977). The CES-D Scale: a self-report depression scale for research in the general population. *Applied Psychological Measures* **1**, 385.
- Reuland DS, Cherrington A, Watkins GS, Bradford DW, Blanco RA, Gaynes BN (2009). Diagnostic accuracy of Spanish language depression-screening instruments. *Annals of Family Medicine* **7**, 455–462.
- Ring JM, Marquis P (1991). Depression in a Latino immigrant medical population: an exploratory screening and diagnosis. *American Journal of Orthopsychiatry* **61**, 298–302.
- Roberts RE (1980). Reliability of the CES-D Scale in different ethnic contexts. *Psychiatry Research* **2**, 125–134.
- Ruiz-Grosso P, Loret de Mola C, Vega-Dienstmaier JM, Arevalo JM, Chavez K, Vilela A, Lazo M, Huapaya J (2012). Validation of the Spanish Center for Epidemiological Studies Depression and Zung Self-Rating Depression Scales: a comparative validation study. *PLoS ONE* **7**, e45413.
- Sayetta RB, Johnson DP (1980). Basic data on depressive symptomatology. United States, 1974–75. *Vital and Health Statistics* **11**, i-v, 1–37.
- Stern MP, Pugh JA, Gaskill SP, Hazuda HP (1982). Knowledge, attitudes, and behavior related to obesity and dieting in Mexican Americans and Anglos: the San Antonio Heart Study. *American Journal of Epidemiology* **115**, 917–928.
- Talbot F, Nouwen A (2000). A review of the relationship between depression and diabetes in adults: is there a link? *Diabetes Care* **23**, 1556–1562.
- Thoolen BJ, de Ridder DT, Bensing JM, Gorter KJ, Rutten GE (2006). Psychological outcomes of patients with screen-detected type 2 diabetes: the influence of time since diagnosis and treatment intensity. *Diabetes Care* **29**, 2257–2262.
- Tuomilehto J, Lindstrom J, Eriksson JG, Valle TT, Hamalainen H, Ilanne-Parikka P, Keinanen-Kiukaanniemi S, Laakso M, Louheranta A, Rastas M, Salminen V, Uusitupa M (2001). Prevention of type 2 diabetes mellitus by changes in lifestyle among

subjects with impaired glucose tolerance. *New England Journal of Medicine* **344**, 1343–1350.

Wallen GR, Feldman RH, Anliker J (2002). Measuring acculturation among Central American women with the use of a brief language scale. *Journal of Immigrant and Minority Health* **4**, 95–102.

Wilkinson GS (1993). *Wide Range Achievement Test*. 3. PAR/ Psychological Assessment Resources, Inc: Lutz, FL.

Zich JM, Attkisson CC, Greenfield TK (1990). Screening for depression in primary care clinics: the CES-D and the BDI. *International Journal of Psychiatry in Medicine* **20**, 259–277.