

In Brief

Patients with type 1 or type 2 diabetes are two times more likely to experience depression than their peers without diabetes. Comorbid depression results in deleterious effects on glycemic control, worsened diabetes complications, functional disability, and premature mortality. Once identified, depression can be effectively treated with antidepressant medications, psychotherapy, or a combination of both. Patients and providers should monitor depressive symptoms to identify their recurrence and work collaboratively to address barriers to care that exist in both urban and rural areas.

Depression Among Adults With Diabetes: Prevalence, Impact, and Treatment Options

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During the past 30 years, a mounting body of evidence has demonstrated that depression is a significant comorbid condition for patients with type 1 or type 2 diabetes. Patients with diabetes have been found to be two times more likely to experience depressive symptoms than their peers without diabetes.¹ Rates of elevated depressive symptoms have been found to range from 21 to 27% in type 1 diabetes and type 2 diabetes, respectively, with lower rates (11.4%) observed in studies using psychiatric interviews in which symptoms attributable to other causes can be evaluated.¹ In addition to higher depression rates among those already diagnosed with diabetes, multiple prospective studies have documented that a lifetime history of depression increases the risk of developing type 2 diabetes later in the life cycle.²⁻⁵

Impact of Depression and Diabetes

Depressive symptoms have been shown to be associated with worsened blood glucose levels⁶ and diabetes complications⁷ such as coronary heart disease.⁸ There is increasing evidence that significant additional functional, fiscal, and psychological costs are associated with depression

in patients with diabetes.⁹⁻¹² Several studies have documented decreased adherence to diet, exercise, and medication regimens associated with depression among adults with diabetes.⁹⁻¹¹ Medical costs associated with moderate to severe levels of depression have also been found to be 51-86% higher than among patients reporting low levels of depression.¹¹ Patients with diabetes and depression have been found to have 4.5 times higher medical expenditures than patients with diabetes alone. Patients with comorbid depression also have higher ambulatory care use and fill more prescriptions.¹²

Comorbid depression has also been shown to have a significant impact on functional disability. Data from the National Health Interview Study have shown that individuals with diabetes and comorbid depression are 7.15 times more likely to experience functional disability (i.e., impairment in work or social activities) compared to peers with either condition alone.¹³ Simon et al.¹⁴ found that > 50% of patients diagnosed with both conditions in a health maintenance organization population reported unemployment.

Finally, comorbid depression and diabetes have been found to increase the risk of early mortality 2.3 times compared to nondepressed patients with diabetes.¹⁵ Zhang et al.¹⁶ reported a 54% increased risk of early mortality among patients reporting elevated depression scores. As Lin et al.¹⁷ have recently documented, causes of mortality in this vulnerable population extend beyond cardiovascular disease to the full range of diseases and disorders.

Depression Treatment for People With Diabetes

Despite the significant costs of comorbid depression and diabetes, traditional treatment approaches such as psychotherapy and antidepressant medications have been found to be efficacious in treating depression in the short term. Lustman et al.¹⁸ conducted the benchmark randomized, controlled trial for cognitive behavioral therapy (CBT) in patients with comorbid depression and diabetes. In this study, 70.8% of patients randomized to the CBT treatment compared to 22.2% of patients in the control group were in depression remission at post-treatment assessment. At 6-month follow-up assessments, depression remission was evident in 66.6% of patients in the CBT group compared to 29.6% in the control group. Treatment responsiveness appeared to be associated with severity of depression and A1C at baseline.^{18,19} Improvements in glycemic control were observed among patients receiving CBT 6 months after completion of treatment.

Problem-solving therapy as an integrated treatment within the primary care setting has also been shown to be efficacious.²⁰ Participants in the Pathways study who were randomized to a stepped-care problem-solving therapy intervention reported higher levels of treatment exposure and satisfaction with care and improved depression outcomes compared to patients in the usual-care group. In this study, improvements in glycemic control were not observed immediately after care or at 6- or 12-month follow-up assessments.²⁰

Randomized, controlled trials have demonstrated the efficacy of antidepressant medications on depression outcomes in type 1 and type 2 diabetic patients. In a randomized trial, nortriptyline was found to improve depression compared to placebo with

hyperglycemic effects observed at post-treatment.²¹ A variety of studies have examined the efficacy of selective serotonin reuptake inhibitor medications, including fluoxetine, sertraline, paroxetine, and bupropion.²²⁻²⁶ All have been shown to be effective in reducing depressive symptoms with either hypoglycemic (e.g., fluoxetine, bupropion) or euglycemic (paroxetine) effects.

Challenges to Treatment Outcomes: Access to Care and Depression Relapse

Although treatment has been shown to be efficacious, challenges remain that contribute to health outcomes for patients with diabetes and depression. One challenge is access to mental health treatment, which includes significant delays in reaching administrative staff to discuss and schedule appointments in urban areas²⁷ and limited numbers of providers in rural areas.²⁸ Even when traditional treatment options are implemented that make use of the primary care medical system in underserved rural areas, barriers to adoption and implementation, including patient refusal of treatment, limited provider engagement, and difficulty training staff, pose threats to building treatment capacity in rural areas.²⁹ Among ethnic and racial groups, disparities in access and unique cultural barriers to quality care, including language translation, cultural understanding on the part of providers, and availability of a limited number of providers, pose even greater challenges to seeking and obtaining adequate mental health treatment.³⁰

Relapse of depressive symptoms also remains a challenge for patients and providers. In a longitudinal cohort of men and women with type 1 or type 2 diabetes, Lustman et al.³¹ found that 92% of individuals who had been successfully treated for major depression experienced a relapse of one or more episodes during a 5-year follow-up period. Depressive symptoms also appear to be persistent over time. Studies of type 2 diabetic patients have shown that between 70 and 77% of individuals reporting elevated depressive symptoms at a baseline survey continued to report depressive symptoms up to 18 months later.^{32,33} In contrast, epidemiological data of episode duration of major depressive disorder in the general population has noted an average duration of 8–12

weeks.³⁴ Additional data are needed to further characterize the long-term pattern of depression for people with diabetes.

Addressing the Challenges: Screening and Supporting Treatment

Taken together, these challenges highlight the need to work effectively and creatively within current health care settings to identify depression and support its treatment. The existence of significant discrepancies between observed rates of clinically significant depressive symptoms and diagnosis among primary care providers³⁵ points to the need for thoughtful and efficient screening of patients so that a dialogue about treatment may begin. In addition, studies documenting high rates of depression relapse suggest the need for routine monitoring and management of symptoms beyond depression treatment to effectively treat recurrent depression symptoms.

A variety of tools are available to screen patients for the presence and severity of depression, including self-report questionnaires and brief clinical interview questions. Tools such as the Patient Health Questionnaire–9³⁶ and the Beck Depression Inventory³⁷ may be incorporated into the beginning of a clinic visit or used interactively with a health care provider or staff member. Brief interpersonal questions such as those found on the two-item version of the Patient Health Questionnaire³⁸ can be used by providers to query patients about changes in mood during the past 2 weeks or since the last visit.

Such screening should be conducted in conjunction with staff-assisted diagnosis, treatment, and follow-up care as recommended by the U.S. Preventive Services Task Force.³⁹ For example, patient questionnaires should be reviewed before completion of the patient visit so that responses indicating severe levels of depressive symptoms or indications of suicidal intent or plan can be further assessed and addressed promptly and directly with the patient. Provider practices that engage in screening should train staff in notification and referral protocols so that patients' needs for immediate care can be met.

Once screening has taken place, providers have an opportunity to discuss with their patients the relationship between diabetes and depression and to review treatment options. This dialogue, however brief, serves multiple

purposes: to validate the importance of presenting mood symptoms to the medical encounter, educate patients about the impact of depression on diabetes self-management and outcomes, provide an opportunity for patients to express concerns about depression treatment and feelings of stigma, and empower patients to report changes in mood symptoms during future visits to facilitate follow-up assessment and treatment.

Finally, barriers to depression treatment can be addressed, in part, through integrated approaches to diabetes health care. The Pathways study²⁰ has provided one model for integrated care in which trained nursing staff provided problem-solving therapy to depressed patients within a primary care setting. Primary care psychology has emerged as a subfield that provides a variety of models of coordinated patient care for mental health and medical issues. Models range from curbside consultation to fully integrated care in which psychologists work side by side with medical providers to provide consultations and treatment on a part- or full-time basis.⁴⁰ Providers and health care systems can work creatively to lower barriers to care within the organizational, historical, and functional context of their practices.

Summary

During the past 30 years, findings from studies of the prevalence and impact of depression in people with diabetes have documented significant adverse effects of depression on morbidity and mortality when both conditions are present. Fortunately, conventional treatments for depression such as antidepressant medications and cognitive behavioral therapy have been shown to be effective in treating depression in people with diabetes. Challenges remain for providers and patients to be more aware of depressive symptoms. The inclusion of established depression screening protocols in diabetes clinical management pathways would increase provider awareness, screening, and psychological referral. These steps could result in earlier detection and initiation of depression treatment. These steps can also facilitate open dialogue between patients and their providers to overcome the attitudinal and logistical barriers to depression treatment and encourage careful monitoring of patients beyond treat-

ment to reduce the potential impact of depression relapse. Further research is needed to continue to develop effective and accessible treatment options for patients to manage persistent mood symptoms.

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References

- ¹Anderson RJ, Freedland K, Clouse R, Lustman PJ: The prevalence of comorbid depression in adults with diabetes: a meta-analysis. *Diabetes Care* 24:1069–1078, 2001
- ²Eaton WW: Epidemiologic evidence on the comorbidity of depression and diabetes. *J Psychosom Res* 53:903–906, 2002
- ³Kawakami N, Takatsuka N, Shimizu H, Ishibashi H: Depressive symptoms and occurrence of type 2 diabetes in Japanese men. *Diabetes Care* 22:1071–1076, 1999
- ⁴Arroyo C, Hu FB, Ryan LM, Kawachi I, Colditz GA, Speizer FE, Manson J: Depressive symptoms and risk of type 2 diabetes in women. *Diabetes Care* 27:129–133, 2004
- ⁵Knol MJ, Twisk JW, Beekman AT, Heine RJ, Snoek FJ, Pouwer F: Depression as a risk factor for the onset of type 2 diabetes mellitus: a meta-analysis. *Diabetologia* 49:837–845, 2006
- ⁶Lustman PJ, Anderson RJ, Freedland KE, de Groot M, Carney RM, Clouse RE: Depression and poor glycemic control: a meta-analytic review of the literature. *Diabetes Care* 23:934–942, 2000
- ⁷de Groot M, Anderson RJ, Freedland KE, Clouse RE, Lustman PJ: Association of depression and diabetes complications: a meta-analysis. *Psychosom Med* 63:619–630, 2001
- ⁸Clouse RE, Lustman PJ, Freedland KE, Griffith LS, McGill JB, Carney RM: Depression and coronary heart disease in women with diabetes. *Psychosom Med* 65:376–383, 2003
- ⁹Egede L, Ellis C, Grubaugh A: The effect of depression on self-care behaviors and quality of care in a national sample of adults with diabetes. *Gen Hosp Psychiatry* 31:422–427, 2009
- ¹⁰Katon WJ, Russo JE, Heckbert SR, Lin EHB, Ciechanowski P, Ludman E, Young B, von Korff M: The relationship between changes in depression symptoms and changes in health risk behaviors in patients with diabetes [article online]. *Int J Geriatr Psychiatry* Published electronically in 2009 (DOI: 10.1002/gps.2363)
- ¹¹Ciechanowski PS, Katon WJ, Russo JE: Depression and diabetes: impact of depressive symptoms on adherence, function, and costs. *Arch Intern Med* 160:3278–3285, 2000
- ¹²Egede LE, Zheng D, Simpson K: Comorbid depression is associated with increased health

care use and expenditures in individuals with diabetes. *Diabetes Care* 25:464–470, 2002

¹³Egede LE: Diabetes, major depression and functional disability among U.S. adults. *Diabetes Care* 27:421–428, 2004

¹⁴Simon GE, von Korff M, Lin E: Clinical and functional outcomes of depression treatment in patients with and without chronic medical illness. *Psychol Med* 35:271–279, 2005

¹⁵Katon WJ, Rutter C, Simon G, Lin EH, Ludman E, Ciechanowski P, Kinder L, Young B, Von Korff M: The association of comorbid depression with mortality in patients with type 2 diabetes. *Diabetes Care* 28:2668–2672, 2005

¹⁶Zhang X, Norris SL, Gregg EW, Cheng YJ, Beckles G, Kahn HS: Depressive symptoms and mortality among persons with and without diabetes. *Am J Epidemiol* 161:652–660, 2005

¹⁷Lin E, Heckbert S, Rutter C, Katon W, Ciechanowski P, Ludman E, Oliver M, Young B, McCulloch D, von Korff M: Depression and increased mortality in diabetes: unexpected causes of death. *Ann Fam Med* 7:414–421, 2009

¹⁸Lustman PJ, Griffith LS, Freedland KE, Kissel SS, Clouse RE: Cognitive behavior therapy for depression in type 2 diabetes mellitus: a randomized, controlled trial. *Ann Intern Med* 129:613–621, 1998

¹⁹Lustman PJ, Clouse RE: Treatment of depression in diabetes: impact of mood and medical outcome. *J Psychosom Res* 53:917–924, 2002

²⁰Katon WJ, von Korff M, Lin EH, Simon G, Ludman E, Russo J, Ciechanowski P, Walker E, Bush T: The Pathways study: a randomized trial of collaborative care in patients with diabetes and depression. *Arch Gen Psychiatry* 61:1042–1049, 2004

²¹Lustman PJ, Griffith LS, Clouse RE, Freedland KE, Eisen SA, Rubin EH, Carney RM, McGill JB: Effects of nortriptyline on depression and glycemic control in diabetes: results of a double-blind, placebo-controlled trial. *Psychosom Med* 59:241–250, 1997

²²Lustman PJ, Freedland KE, Griffith LS, Clouse RE: Fluoxetine for depression in diabetes: a randomized double-blind placebo-controlled trial. *Diabetes Care* 23:618–623, 2000

²³Goodnick PJ, Kumar A, Henry JH, Buki VM, Goldberg RB: Sertraline in coexisting major depression and diabetes mellitus. *Psychopharmacol Bull* 33:261–264, 1997

²⁴Gülseren L, Gülseren S, Hekimsoy Z, Mete L: Comparison of fluoxetine and paroxetine in type 2 diabetes patients. *Arch Med Res* 36:159–165, 2005

²⁵Williams MM, Clouse RE, Nix BD, Rubin EH, Sayuk GS, McGill JB, Gelenberg AJ, Ciechanowski PS, Hirsch IB, Lustman PJ: Efficacy of sertraline in prevention of depression recurrence in older adults with diabetes. *Diabetes Care* 30:801–806, 2007

²⁶Lustman PJ, Williams MM, Sayuk GS, Nix BD, Clouse RE: Factors influencing glycemic control in type 2 diabetes during acute- and maintenance-phase treatment of major

depressive disorder with bupropion. *Diabetes Care* 30:459–466, 2007

²⁷Rhodes KV, Vieth TL, Kushner H, Levy H, Asplin BR: Referral without access: for psychiatric services, wait for the beep. *Ann Emerg Med* 54:272–278, 2009

²⁸Hendryx M: Mental health professional shortage areas in rural Appalachia. *J Rural Health* 24:179–182, 2008

²⁹Luptak M, Kaas MJ, Artz M, McCarthy T: Project ADAPT: a program to assess depression and provide proactive treatment in rural areas. *Gerontologist* 48:542–548, 2008

³⁰Alegria M, Chatterji P, Wells K, Cao Z, Chen CN, Takeuchi D, Jackson J, Meng XL: Disparity in depression treatment among racial and ethnic minority populations in the United States. *Psychiatr Serv* 60:268–269, 2009

³¹Lustman PJ, Griffith L, Freedland K, Clouse R: The course of major depression in diabetes. *Gen Hosp Psychiatry* 19:138–143, 1997

³²Peyrot M, Rubin R: Persistence of depressive symptoms in diabetic adults. *Diabetes Care* 22:448–452, 1999

³³de Groot M, Risaliti C, Doyle T, Merrill J, Pinkerman B, Shubrook J, Gotfried R,

Schwartz F: Persistence of depression among type 2 diabetes Appalachians. *Ann Behav Med* 33 (Suppl.):S178, 2007

³⁴American Psychiatric Association: *Diagnostic and Statistical Manual of Mental Disorders*. 4th ed. Washington, D.C., American Psychiatric Press, 2000

³⁵Gary TL, Crum RM, Cooper-Patrick L, Ford D, Brancati FL: Depressive symptoms and metabolic control in African-Americans with type 2 diabetes. *Diabetes Care* 23:23–29, 2000

³⁶Kroenke K, Spitzer RL, Williams JB: The PHQ-9: validity of a brief depression severity measure. *J Gen Intern Med* 16:606–613, 2001

³⁷Beck AT, Steer RA, Brown GK: *BDI-II Beck Depression Inventory Manual*. 2nd ed. San Antonio, Texas, Harcourt Brace, 1996

³⁸Kroenke K, Spitzer RL, Williams JB: The Patient Health Questionnaire–2: validity of a two-item depression screener. *Med Care* 41:1284–1292, 2003

³⁹U.S. Preventive Services Task Force: Screening for depression in adults: U.S. Preventive Services Task Force Recommendation Statement. *Ann Intern Med* 151:784–792, 2009

⁴⁰Gatchel RJ, Oordt MS: *Clinical Health Psychology and Primary Care*. Washington, D.C., American Psychological Association, 2003

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