The Association of the Prevalence of Depression in Type 2 Diabetes Mellitus with Visual-Related Quality of Life and Social Support

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Aim: To report the prevalence of depression and its association with vision-related quality of life and social support in a type 2 diabetes mellitus (T2DM) population.

Methods: Patients were recruited from a community-based study, Fushun Diabetic Retinopathy Cohort Study (FS-DIRECT), conducted between July 2012 and May 2013 in China. Depression was assessed using the Centre for Epidemiological Studies Depression Scale (CES-D). Vision-related quality of life was evaluated using the Visual Function Questionnaire-25 (VFQ-25). Social support was captured with the Social Support Rating Scale (SSRS). Generalized linear models were used to estimate the individual and joint association of VFQ-25 composite score (VFQCS) and SSRS score (SSRSS) with depression.

Results: A total of 1618 subjects (60.9% female) aged 61.69 ± 8.72 years in an urban district of Jiangjun Street, Fushun City, Liaoning province, Northeast China from July 2012 to May 2013 were recruited, of which, 23.36% (95% CI: 21.30–25.42%) were identified with depression. Every 14.1 increase in VFQ-25 composite score decreased the risk of depression by half (OR = 0.5; 95% CI: 0.4–0.6); with the elevation of 10.0 SSRS score the risk of depression decreased by 40% (OR = 0.6; 95% CI: 0.5–0.7). Patients with the VFQCS less than 91.3 and SSRSS less than 38.0 had 5.9 times more risk of depression (OR = 5.9; 95% CI: 3.6–9.7). Age (over 60 years) (OR = 0.6; 95% CI: 0.4–0.9) and medical history of cardiovascular disease (OR = 1.7; 95% CI: 1.1–2.5) were independently correlated with depression symptom.

Conclusion: The prevalence of depression is high among patients with T2DM in urban district in northeast China. Vision-related quality of life and social support scores are significantly associated with depression. Measures should be taken to screen depressive symptoms in patients with type 2 diabetes patients. These patients need to be intervened with appropriate and effective treatment as early as possible. Meanwhile, behavioral health specialists should guide the patient to get and use social support sources effectively.

Keywords: diabetic retinopathy, depression, quality of vision, Visual Function Questionnaire-25, Social Support Rating Scale

Introduction

The number of people with diabetes aged 20–79 years was predicted to rise to 642 million by 2040.¹ Diabetes has become a serious public health burden in China,² and its prevalence will likely increase due to population growth, aging, urbanization, and increasing number of obesity and physical inactivity.¹ Diabetes affects patient’s visual and overall well-being in many ways. Diabetic retinopathy (DR), a leading cause of preventable blindness in working-aged people in the US,³ has an age-standardized prevalence of 43.1%.⁴ Diabetes also increases the risk of cataract, glaucoma, strabismus, and dry eye.
Depression is also a public health problem in China. Many chronic or non-chronic systemic diseases are often considered to be associated with depression, such as diabetes, rheumatoid arthritis, and malignancies.\textsuperscript{5–7} The risk of developing depression in diabetes population is two times higher than that in the diabetes-free population.\textsuperscript{5} The occurrence of depression and diabetes leads to worse outcomes including reduced treatment adherence, which may lead to worse glycemic control, increased risk of complications, increased disability, reduced work productivity, increased treatment costs and reduced quality of life.\textsuperscript{8–11} In addition, previous studies show that the coexistence of these two diseases can increase the risk of morbidity and mortality with major negative implications on the individuals, families and society.\textsuperscript{12} International diabetes guidelines had recommended screening for psychosocial problems including depression.\textsuperscript{13}

Many demographic and clinical factors have been shown to be associated with the depression in patients with diabetes, while the results are still inconsistent. Öztürk ZA reported that depression scores (GDS) of diabetes patients with retinopathy and neuropathy were significantly higher than those of patients without complications,\textsuperscript{14} Lin et al showed there was a significant relationship between depression and diabetic microvascular complication in a prospective cohort study,\textsuperscript{11} while Hirai FE found no association between depression and severity of diabetic retinopathy or visual impairment in a cohort of persons with long-term type 1 diabetes mellitus (T1DM).\textsuperscript{15}

Several vision-related quality of life instruments have been used to assess the vision-related functional impairment such as Vision Function 14-item scale (VF-14),\textsuperscript{16} Activities of Daily Vision Scale (ADVS),\textsuperscript{17} the National Eye Institute Visual Function Questionnaire (NEI-VFQ).\textsuperscript{18} Of which, the National Eye Institute Visual Function Questionnaire-25 (VFQ-25), which was introduced by Mangione et al, had been used to evaluate visual function in patients with various chronic eye diseases, including glaucoma, age-related macular degeneration, cataract, diabetic retinopathy.\textsuperscript{19–22} The Chinese version of NEI-VFQ-25 (CHI-NEI-VFQ-25) developed by Chan and co-workers\textsuperscript{23} in Hong Kong was used to measure visual function in our study.

Although there were so many risk factors related to depression, there are some protective factors that can reduce the risk of depression, such as social support. According to “buffering hypothesis”,\textsuperscript{24} social support can protect a person from the potentially pathogenic influence of stressful events. Social support can be captured with the Social Support Rating Scale (SSRS).

Considering the above, the main objectives of this study were (1) to investigate the point prevalence of depression symptoms in T2DM; (2) to assess the association of VFQ-25 composite score, SSRS and their joint association with the risk of depression symptom in T2DM.

**Subjects and Methods**

**Ethics**

The study followed the tenets of the Declaration of Helsinki (2008). The study protocol was approved by the Ethics Committee of Fushun Eye Hospital. Informed consents were obtained from all participants.

**Subjects**

Fushun Diabetic Retinopathy Cohort Study (FS-DIRECT) was a community-based study in an urban district of Jiangjun Street, Fushun City, Liaoning province, Northeast China. The study subjects were residents with confirmed diagnosis of T2DM. The methodologies and baseline data were reported elsewhere.\textsuperscript{25} Briefly, all the study subjects met the following criteria: (1) who have documented history of diagnosis or treatment of T2DM; (2) residence in the Jiangjun Street for at least half year with an age of 30 years or older at the time of the survey. The subjects were recruited from the community healthcare center through phone calls or a home visit.

**Methods**

The comprehensive ophthalmic examination, body check and blood tests were reported elsewhere.\textsuperscript{25} Those who accepted the invitation were scheduled to have a clinical examination in the community healthcare center. Blood and urine samples were sent to the laboratory of Fushun Eye Hospital for analysis. At the same time, all subjects need to complete general
questionnaire mainly include demographic characteristics (sex, age, marital status, etc.) and income, religion, dwelling status, medical history during investigation, etc.

**Grading for Diabetic Retinopathy**

Six-field stereoscopic color fundus photographs were taken from each eye with a digital camera (Kowa, VK-2, Tokyo). All the color fundus photographs were graded for diabetes retinopathy severity, macular edema scores following the guidelines of the Early Treatment Diabetic Retinopathy Study classification (ETDRS).\(^{26}\) The severity of DR was categorized into no apparent diabetic retinopathy (ETDRS level 10–20), mild non-proliferative diabetic retinopathy (NPDR) (ETDRS level 31–37), moderate NPDR (ETDRS level 43–47), severe NPDR (ETDRS level 53), and proliferative diabetic retinopathy (PDR) (ETDRS level 60–85). The severity scores of the worse eye were used for the individual. If the images in one eye were ungradable, the scores for the fellow eye were used to define these outcomes.

**Depression Symptom**

Depressive symptoms were measured using the Center for Epidemiologic Studies Depression (CES-D) Scale. CES-D was a 20-item self-report scale developed by the National Institute of Mental Health in 1977 to assess depressive symptoms among both clinical and samples drawn from general populations.\(^{27}\) Briefly, CES-D consists of four components: dysthymic affect, the lack of positive affect, somatic and retarded activity difficulties, and interpersonal difficulties. CES-D score ranged between 0 and 60. A score of 16 or greater was an indicator of clinically significant sign of depression.\(^{27}\) The CES-D had satisfactory reliability and validity in studies conducted in Chinese populations.\(^{28,29}\)

**Vision-Related Quality of Life**

The NEI-VFQ 25 was designed to evaluate vision-related quality of life with excellent reliability and validity.\(^{30}\) It comprises 25 items regarding the level of difficulty of particular visual symptoms and day-to-day activities. Each item is assigned to 1 of the 12 subscales, namely, general health, general vision, ocular pain, near activities, distance activities, social functioning, mental health, role difficulties, dependency, driving, color vision, and peripheral vision. A Chinese version of the NEI-VFQ 25 was administered to all the enrolled subjects in this study, and the reliability and validity of the Chinese version of the NEI-VFQ 25 questionnaire have been assessed, and the questionnaire was proven to accurately measure the VR-QOL in Chinese individuals.\(^{23}\) The NEI-VFQ 25 composite score is calculated as the unweighted average response to all items, excluding the questions regarding general health.\(^{23}\)

**Social Support**

Social support was evaluated using the Social Support Rating Scale (SSRS).\(^{31}\) It had 10 items, measuring three dimensions of social support: subjective support (4 items), objective support (3 items), and support-seeking behavior (3 items). SSRS had a total score ranging from 12 to 66, where higher scores indicated stronger social support. The SSRS has been widely used in Chinese populations showing high reliability and validity,\(^{32}\) and exhibited strong internal consistency (Cronbach’s alpha = 0.954) in the former study.\(^{33}\)

**Statistical Analysis**

Continuous and normally distributed variables were presented as mean ± standard deviation (SD), continuous and abnormally distributed data were described using the median and interquartile range (IQR) and categorical variables were presented as frequency.

We evaluated the individual and joint association of VFQ-25 composite score (VFQCS) and SSRS score (SSRSS) with the prevalence of depression using generalized linear models. All models of the independent association of VFQCS or SSRSS were performed in 2 ways: exposure as a categorical variable (quartiles) and as a continuous variable (scaled to interquartile range [IQR]). For models of the joint association of VFQCS and SSRSS, VFQCS and SSRSS are averagely divided into 2 groups. We adjusted potential confounders known to be associated with depression, such as age, sex, body mass index, education, religion, dwelling state, marital status, employment status, family income status and so on. Odds ratios (OR) and their 95% confidence interval (CI) were estimated for each independent variable.
In addition, we examined the relationship between the odds of depression and VFQCS or SSRSS with restricted cubic splines. All tests were two-sided, and a $P \leq 0.05$ was considered significant. All statistical analyses were performed using SAS 9.4 (SAS Institute Inc. Cary, NC, USA).

**Results**

**Features of Study Population**

2006 subjects completed the examinations in the Fushun-DIRECT Study, among them, 1618 subjects aged 61.69 ± 8.72 years who completed CES-D, NEI-VFQ 25, and SSRS were included in the current analysis (respond rate was 80.66%). Individuals refused to accept depressive symptoms check were excluded (n = 388). The demographic, clinical characteristics of subjects included in the study are summarized in Table 1.

**Prevalence of Depressive Syndrome in Type 2 Diabetes and Factors Associated with Depression**

The score of CES-D in our study is 8.0 [11.0, median (IQR)]. The prevalence of depressive syndrome, defined as having a CES-D score of ≥16, was 23.36% (95% CI 21.30–25.42) in our study. The scores of VFQ composite score and SSRSS score in our study are 91.3 [14.1, median (IQR)] and 38.0 [11.0, median (IQR)], respectively.

The univariate logistic regression showed that binocular visual acuity, sex, education, religion, dwelling status, marital status, employment status, income were significantly correlated with depression ($P < 0.05$). Severity of diabetes retinopathy, diagnosed with cataract, glaucoma, nephropathy, cardiovascular disease, stroke, hyperlipidemia, hypertension, VFQ-25 composite score (VFQCS) and SSRSS score (SSRSS) were significantly correlated with depressive symptom ($P < 0.05$).

The results of multivariate logistic regression suggested that there was a significantly strong correlation between VFQCS and depression. Patients with the VFQCS > 81.4 had 30% potential risk of depression than patients with the VFQCS < 81.4 (OR = 0.3; 95% CI: 0.2–0.5). Every increase by 14.1 in VFQCS decreased the risk of depression by half. (OR = 0.5; 95% CI: 0.4–0.6). The results of multivariate logistic regression suggested that SSRSS is a significant protective factor for depression. With the elevation of SSRSS, the prevalence of depression in quartile decreases significantly. Patients with the SSRSS >32.0 had 0.6 times potential risk of depression than patients with the SSRSS <32.0 (OR = 0.6; 95% CI: 0.4–0.9). With the elevation of 10.0 in SSRSS, the risk of depression decreased by 40% (OR = 0.6; 95% CI: 0.5–0.7). The individual association of VFQ composite score (quartile) and SSRSS score (quartile) with the prevalence of depression in T2DM patients is listed in Table 2.

In addition, multivariate logistic regression suggested that age ≥60 years (OR = 0.6; 95% CI: 0.4–0.9), diagnosis of cardiovascular disease (OR = 1.7; 95% CI: 1.1–2.5) were independently correlated with depression. Diabetes retinopathy was not independent risk of depression.

Table 3 shows the combined correlation of VFQCS and SSRSS to the risk of depression in type 2 diabetes mellitus patients. Multivariate logistic regression analysis shows that the joint correlation of VFQCS and SSRSS contributes to the decrease of the risk of depression. The prevalence of depression increases in the sequence of patients with perfect VFQCS and SSRSS, perfect VFQCS with poor SSRSS, poor VFQCS with perfect SSRSS, and both poor VFQCS and SSRSS. Patients with the VFQCS less than 91.3 and SSRSS less than 38.0 had 5.9 times potential risk of depression than patients with the VFQCS larger than 91.3 and SSRSS larger than 38.0 (OR = 5.9; 95% CI: 3.6–9.7).

The non-linear relationship between the odds of depression and VFQCS or SSRSS is listed in Figure 1. The adjusted OR of depression symptom decreases sharply with the increase of VFQ-25 composite score or SSRSS score, respectively. All estimates are adjusted for age, gender, body mass index, binocular visual acuity, education, religion, dwelling state, marital status, employment status, family income status, cataract, glaucoma, severity of diabetes retinopathy, nephropathy, cardiovascular disease, stroke, hyperlipidemia, and hypertension.
Table 1: Demographic, Clinical Characteristics of Subjects Included in Study

| Subject                        | Included (n=1618) |
|-------------------------------|-------------------|
| Age (years)                   | 61.69±8.72        |
| Duration of diabetes (years)  | 7.65±5.93         |
| FPG (mmol/L)                  | 9.39±3.52         |
| HbA1c (%)                     | 7.79±2.00         |
| Cholesterol (mmol/L)          | 5.30±1.18         |
| BMI (kg/m²)                   | 26.47±3.47        |
| Binocular visual acuity (LogMar) | 0.14±0.22   |
| Sex (%)                       |                   |
| Male                          | 632(39.06)        |
| Female                        | 986(60.94)        |
| Education (%)                 |                   |
| Less than high school         | 997(61.66)        |
| High school and technical school | 387(23.93)   |
| More than high school         | 233(14.41)        |
| Religion (%)                  |                   |
| No                            | 1390(85.91)       |
| Yes                           | 228(14.09)        |
| Dwelling status (%)           |                   |
| Alone                         | 135(8.34)         |
| With partner                  | 1483(91.66)       |
| Marital status (%)            |                   |
| Married                       | 1350(83.44)       |
| Divorced                      | 39(2.41)          |
| Widowed                       | 225(13.91)        |
| Spinsterhood                  | 4(0.25)           |
| Employment status (%)         |                   |
| Employed                      | 191(11.80)        |
| Retired                       | 1306(80.72)       |
| Unemployed                    | 121(7.48)         |
| Smoking (%)                   |                   |
| No                            | 1276(78.86)       |
| Yes                           | 342(21.14)        |
| Income levels (%)             |                   |
| Low                           | 617(38.35)        |
| Middle                        | 915(56.87)        |
| High                          | 77(4.79)          |
| Treatment regimen of diabetes (%) |                 |
| Diet alone                    | 175(11.24)        |
| Oral hypoglycemic agents      | 1005(64.55)       |
| Insulin dependent or with oral hypoglycemic agents | 377(24.21) |
| Severity of DR (%)            |                   |
| No DR                         | 858(54.75)        |
| Mild NPDR                     | 488(31.14)        |
| Moderate NPDR                 | 92(5.87)          |
| Severe NPDR                   | 69(4.40)          |
| PDR                           | 60(3.83)          |
| AMD (%)                       |                   |
| No                            | 1418(97.66)       |
| Yes                           | 34(2.34)          |

(Continued)
Discussion

This study is the first community-based survey of prevalence of depression in patients with T2DM in urban district in northeast China. Using the CES-D cut off score 16, prevalence of depression symptom in this population of patients with T2DM is 23.36% (95% CI 21.30–25.42), which is consistent to the result found in Shandong (23%), though the latter used a different tool to identify depression (ZESD ≥ 40). A few studies were carried out on diverse populations using different methodologies and a wide variety of ascertainment methods for depression. The reported rates of depression in people with diabetes also varied greatly with prevalence rates ranging from a low rate of 5.0% to a high rate of 71.8%. In the mainland of China, one study found that about 35.1% of patients with type 2 diabetes mellitus in Shanghai experienced depressive symptoms (ZSDS ≥ 53). A general hospital-based study reported that the prevalence of comorbid depression (BDI scores ≥14) was 5.7% among Chinese subjects with type 2 diabetes mellitus in Beijing. Using the same method (CES-D score of ≥16), prevalence of depression symptom is slightly higher (31.0%) in Hong Kong Chinese patients with type 2 diabetes than the result of our study.

Our study confirmed our hypothesis that the visual function damage is the key factor of depression instead of diabetic retinopathy. The NEI VFQ-25 is a subjective assessment by patients of their perception of visual function. It can address the subject’s vision and subject’s vision-specific quality of life. People with poor visual function score have trouble dealing with daily life, frequent social activities, and support from the external world. These problems create a large gap between expectations and reality and lead to more physical and mental burden. Although mechanism of depression remains unclear, “psychological burden hypothesis” is likely the underlying mechanisms of depression. This hypothesis may decipher the relationship between VFQCS and depression. Many conditions can affect the visual function such as diabetic retinopathy, cataract, glaucoma, and ophthalmologist can improve VFQCS through different approaches.

Table 1 (Continued).

| Subjects Included (n=1618) |
|-----------------------------|
| Cataract (%)                |
| No                          | 1239 (84.80) |
| Yes                         | 222 (15.20)  |
| Glaucoma (%)                |
| No                          | 1432 (97.48) |
| Yes                         | 37 (2.52)    |
| Nephropathy (%)             |
| No                          | 1174 (74.30) |
| Yes                         | 406 (25.70)  |
| Cardiovascular disease (%)  |
| No                          | 721 (46.61)  |
| Yes                         | 826 (53.39)  |
| Stroke (%)                  |
| No                          | 1286 (81.34) |
| Yes                         | 295 (18.66)  |
| Hyperlipidemia (%)          |
| No                          | 583 (48.58)  |
| Yes                         | 617 (51.42)  |
| Hypertension (%)            |
| No                          | 912 (58.31)  |
| Yes                         | 652 (41.69)  |

Notes: Duration of diabetes is defined by the period from the date of diagnosis to the date of the study examination, Religion is defined by if having any kind of religious belief, Smoking is defined by if smoke at the time of study, family income status is defined accordance with the perceptions of the patients, other ocular diseases is defined by self-report history at the time of study.

Abbreviations: AMD, age-related macular degeneration; BCVA, best corrected visual acuity; BMI, Body Mass Index; DR, diabetic retinopathy; FPG, fasting plasma glucose; HbA1c, glycated hemoglobin; NPDR, non proliferative diabetic retinopathy; PDR, proliferative diabetic retinopathy; PRP, panretinal photocoagulation.
based on patients’ conditions, such as early cataract surgery, effective lowering intraocular pressure, and timely photocoagulation for diabetic retinopathy.

We also found that social support did serve as a powerful protective factor of depression. These results were consistent with the findings in previous studies.\textsuperscript{38,43} Social support is a perception that one is accepted, cared for, and provided with assistance from certain individuals or a specific group or the realization of actual support received from another. Social support can arise from different sources, including family members, friends, peers, healthcare professionals and organizations.\textsuperscript{44} SSRS questionnaire can comprehensively evaluate social support from three aspects including objective support, subjective support, and social support utilization. It was well known that emotional loss,
physical impairment, lifestyle interference, or threatened disability can bring diabetes patients much burden. The buffering hypothesis asserts that social support is a protective barrier from stressful events.\textsuperscript{45}

Bio-psycho-social medical model implements comprehensive therapy for patients with negative emotions (including antidepressant use), which can significantly improve the clinical therapeutic effect. Psychological and pharmacological treatments for depression in diabetes have been validated in majority and mixed samples.\textsuperscript{46,47} However, the assessment to such treatment may be difficult, because most people in the current study district still believe that mental illness is a shame. On the contrary, the social environment can be improved relatively easily. Doubtlessly, better social support and more adequate resources are effective tools to reduce the perception of stress. Behavioral health specialists should guide the diabetes patient and the family to get familiar with social support sources and use these sources effectively.

Our study also found the joint correlation of VFQCS and SSRSS with prevalence of depression. If the cut-off value was 91.3 for VFQCS and 38.0 for SSRSS, with a poor VFQCS and SSRSS, the risk of depression could rise to 5.9. In addition, age and cardiovascular disease were found to be independent risk factors of depression in type 2 diabetes mellitus in our study.

This study was conducted at the community level without the influence of the hospital environment. The high response rate of the target population greatly reduced the bias of our research. It was the first survey of depression symptom of T2DM patients in the old industrial city in northeast China. In addition, we found that VFQ-25 composite score is a strong protective factor of depression, which is not reported before. Improvement of visual function might reduce the occurrence of depression in diabetes patients. The diagnosis of depression may have been neglected in ophthalmology department as ophthalmologist always concentrates on the primary illness. Based on this study, physicians and ophthalmologists should be aware of the high prevalence of depression, especially in diabetes patients with poor VFQ-25 composite score. At the same time, social support should be improved to alleviate depression, this is of great reference value for health care personnel who provide care and guidance to patients in such situation.

Our study has its limitations. First, our study population, as a community of the old industrial city in northeast China, may not be the perfect representative of the whole population of China. Second, with these cross-sectional data, the nature of the relationship between VFQ –25 composite score, social support and depression cannot be fully characterized. Additionally, it is possible that self-reports of other medical diagnoses and their medications were subject to bias, recall problems, or incomplete knowledge leading to potentially inaccurate identification of patients’
conditions. Medical record did not include diagnosed depression and used antidepressant drugs, which will influence the nature of prevalence of depression symptom. In the current study, we did not include some variables such as obesity and renal disorders, which might induce controversial. Instead, we included BMI and history of retinopathy to control the bias.

**Conclusions**

This study found that about one quarter of patients with T2DM in urban district in northeast China were presence of depression symptoms. We also revealed that VFQ −25 composite score and social support were significantly associated with the occurrence of depression symptoms. The joint associations of VFQ −25 composite score and social support with the risk of depression are strong. Measures should be taken to control depressive symptoms in patients with T2DM. These patients need to be intervened on controlling variables such as blood sugar as early as possible. Meanwhile, behavioral health specialists should guide the patient to get and use social support sources effectively.

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**Disclosure**

The authors report no conflicts of interest in this work.

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