[ORIGINAL ARTICLE]

Medium-term Influence of the Coronavirus Disease 2019 Pandemic on Patients with Diabetes: A Single-center Cross-sectional Study

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Abstract:

Objective This study evaluated the lifestyle changes in patients with diabetes and their independent associations with glycemic and body weight control. In addition, the correlation between changes in mental health and lifestyles was evaluated.

Methods This single-center cross-sectional study included 340 patients with diabetes who periodically visited our department. Changes in dietary habits, activities of daily living, and mental health before and during approximately six months after the onset of the coronavirus disease 2019 (COVID-19) pandemic were evaluated using a questionnaire, including the International Physical Activity Questionnaire-Short Form.

Results Approximately 20%, 30%, and over 50% of patients had worsened dietary habits, decreased activities of daily living, and deteriorated mental health, respectively. A multiple regression analysis showed that irregular meal timing was significantly associated with change in HbA1c ($\beta=0.328$, $p=0.001$), and decreased walking time was significantly associated with changes in body weight ($\beta=-0.245$, $p=0.025$). The change in fear and anxiety was positively associated with changes in meal timing regularity ($r=0.129$, $p=0.019$) and carbohydrate consumption ($r=0.127$, $p=0.021$). Subsequently, the change in depressed mood was positively associated with changes in carbohydrate ($r=0.142$, $p=0.010$) and alcohol ($r=0.161$, $p=0.037$) consumption, and the change in psychological stress was positively associated with changes in carbohydrates ($r=0.183$, $p=0.001$) and snack ($r=0.151$, $p=0.008$) consumption as well as sedentary time ($r=0.158$, $p=0.004$).

Conclusion The COVID-19 pandemic has had a considerable medium-term impact on the lifestyle and mental health of patients with diabetes. Lifestyle changes were associated with glycemic and body weight control, and mental health changes were associated with lifestyle changes. These findings may provide important information on diabetes care during the pandemic.

Key words: COVID-19, diabetes, lifestyle, mental health

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Introduction

The novel coronavirus disease 2019 (COVID-19) has caused a global health emergency; governments of pandemic-hit countries have adopted different measures to prevent the spread of the disease and the collapse of the healthcare system. Depending on the rate of transmission and the robustness of the medical system, these measures

range from requests to promote social distancing to forced lockdowns of cities, with penalties for violations (1).

In Japan, the government declared a state of emergency over the COVID-19 outbreak, requesting that residents refrain from nonessential outings and restricting certain business operations without penalties for violations in seven prefectures, including Osaka, on April 7, 2020. This declaration was expanded nationwide on April 16, 2020, and phased out due to a reduction in the first wave of transmission on May

14, 2020 (2). This situation temporarily affected Japanese people's lifestyle and mental health, including their dietary habits and activities of daily living (3, 4). However, the second and third waves of infection have since begun, and as of December 31, 2020, Japan had recorded 233,785 confirmed COVID-19 cases and 3,459 deaths (2).

For patients with diabetes, dietary habits, activities of daily living, and mental health are particularly important in disease management, and any disruptive change is likely to adversely affect patients' glycemic and body weight control (5, 6). Indeed, recent studies have evaluated the impact of the COVID-19 pandemic on the changes in lifestyles and/or mental health and their impact on glycemic and body weight control (7-14). However, these reports evaluated the early phase, approximately two months after the beginning of the COVID-19 pandemic. The medium-term impact of the COVID-19 pandemic on changes in lifestyles and mental health and their independent impact on glycemic and body weight control in patients with diabetes thus remain unclear. Furthermore, the correlation between changes in mental health and lifestyle also remains unclear.

This single-center, cross-sectional study evaluated the lifestyle and mental health changes and their independent association with glycemic and body weight control in patients with diabetes during the medium-term period after the beginning of the COVID-19 pandemic. In addition, the correlation between mental health and lifestyle changes was also evaluated.

Materials and Methods

Study design and participants

This single-center, cross-sectional study was conducted at the Department of Diabetes, Metabolism, and Endocrinology of the Osaka Police Hospital in Japan. Patients with diabetes over 20 years old who periodically visited our department and provided their written consent to participate in the study were included. Individuals with malignant tumors, mental illnesses, or dementia and those with serious illnesses or conditions that significantly affected their daily lives were excluded or deemed ineligible for participation by an attending physician. Demographic data and anthropometric measurements were obtained from medical records at the time the questionnaire survey was conducted, from September 1 to October 30, 2020.

Questionnaire

The questionnaire was developed by our department and consisted of three sections. The first section contained questions on changes in dietary habits (regularity of meal timings, amount of total diet, and consumption of carbohydrates, snacks, fruits, and alcohol) during the pandemic (from April 2020 to the time the questionnaire was completed) compared to before the pandemic. The responses to the question on meal timing regularity included "became

regular," "no change," or "became irregular." The responses to the question on dietary habits included "decreased," "no change," or "increased."

The second section contained questions regarding activities of daily living, including physical activity and sedentary time before and during the pandemic, which were measured using the International Physical Activity Questionnaire-Short Form (IPAQ-SF) (15). This questionnaire comprises seven questions assessing the total time (in minutes) spent on vigorous-/moderate-intensity activities and walking per week and being sedentary per day. The responses regarding the vigorous-/moderate-intensity activities and walking were converted to metabolic equivalent task minutes per week (MET.min/week), using the IPAQ scoring protocol. The total weekly physical activity level was estimated by adding the scores for each activity. The last section contained questions regarding changes in mental health, including fear and anxiety, depressed mood, and psychological stress during the pandemic compared to before the pandemic. The three-item responses included "decreased," "no change," or "increased." The respondents completed the questionnaire at regular visits during the pandemic (September 1 to October 30, 2020).

Changes in HbA1c, body weight, and hypoglycemic agents

The change in HbA1c was calculated as the difference in this indicator at the time of questionnaire completion during the pandemic from that at the regular visits within nearly two months before the state of emergency over the COVID-19 outbreak (April 7, 2020). The same calculation applied to the change in body weight. The change in hypoglycemic agents was evaluated as "strengthened," "unchanged/modified," or "attenuated" during the pandemic (from April 7, 2020, to the time of questionnaire completion).

Ethics statement

The protocol was approved by the Osaka Police Hospital Clinical Research Review Committee in compliance with the Declaration of Helsinki and the current legal regulations in Japan. Written informed consent was obtained from all participants before participation in the study.

Statistical analyses

Statistical analyses were performed using the IBM SPSS Statistics for Windows, version 21.0, software program (IBM, Armonk, USA). Continuous variables are presented as means and standard deviations, while categorical variables are presented as valid percentages. The changes in dietary habits, mental health, and hypoglycemic agents were coded as follows: regularity of meal timing was -1= became regular, 0= no change, and 1= became irregular. Other dietary habits were coded as -1= decreased, 0= no change, and 1= increased. Mental health was coded as -1= decreased, 0= no change, and 1= increased. Hypoglycemic agents were as follows: -1= strengthened, 0= unchanged/modified, and 1= attenuated. Differences in physical activity and sedentary time

before and during the pandemic were analyzed using Wilcoxon's signed-rank test. A Pearson's correlation analysis and multiple regression analysis were used to evaluate if and how lifestyle changes were associated with those in HbA1c and body weight. A Pearson's correlation analysis was also used to examine the correlation between mental health and lifestyle changes. Statistical significance was set at $p < 0.05$.

Table 1. Demographic Characteristics and Anthropometric Measurements.

Variables	Value
N (Male/female)	340 (226/114)
Age (years)	67.2±11.2
Type of diabetes (type 2/type 1/others)	320/15/5
Duration of diabetes (years)	16.3±11.6
Body weight (kg)	68.8±14.5
BMI (kg/m ²)	25.5±4.3
Alcohol (yes/no) (%)	40.0/60.0
Smoking (yes/no) (%)	17.1/82.9
Established cardiovascular disease (yes/no) (%)	32.9/67.1
Employment (yes/no) (%)	48.4/51.6
HbA1c (%)	7.0±0.8
Systolic blood pressure (mmHg)	131.6±15.3
Diastolic blood pressure (mmHg)	77.4±11.6
eGFR (mL/min/1.73m ²)	67.7±20.7
Diabetes treatment	
Diet only (%)	2.1
Oral hypoglycemic agents (%)	90.6
GLP-1RAs (%)	25.0
Insulin (%)	25.0

Data are presented as means±standard deviation (SD) or proportion. BMI: body mass index, eGFR: estimated glomerular filtration ratio, GLP-1RAs: glucagon-like peptide-1 receptor agonists, HbA1c: glycated hemoglobin A1c

Results

Demographic characteristics and anthropometric measurements

A total of 360 patients completed the questionnaire, with HbA1c and body weight changes evaluated in 340. Overall, 226 (66.5%) were men, and the majority (94.1%) had type 2 diabetes. Their mean age was 67.2±11.2 years old; most patients (63.8%) were between 60 and 79 years old, followed by those between 40 and 59 years old (22.6%). Patients' mean body mass index (BMI) was 25.5±4.3 kg/m². The mean duration of diabetes and HbA1c levels was 16.3±11.6 years and 7.0% ±0.8%, respectively. Established cardiovascular diseases were present in 32.9% of patients. The summarized demographic data and anthropometric measurements of all patients are presented in Table 1.

Changes in dietary habits during the pandemic

The changes in dietary habits before and during the pandemic are shown in Fig. 1. Changes in the regularity of meal timing showed that 15.2% of patients reported regular meal habits, whereas 7.5% reported irregular meal timings. Total diet decreased in 11.8% of patients but increased in 13.0% during the pandemic. The proportions of patients who reported reduced carbohydrate, snack, and fruit consumption were 6.0%, 9.1%, and 12.9%, while 22.2%, 15.3%, and 21.8% reported increased consumption, respectively. Alcohol consumption decreased and increased in 24.9% and 10.1% of patients, respectively.

Activities of daily living before and during the pandemic

Table 2 presents the results of the IPAQ-SF, which evalu-

Figure 1. The changes in dietary habits during the pandemic.

Table 2. Activities of Daily Living before and during the Pandemic.

	Before the pandemic	During the pandemic	p value
Vigorous intensity (METs.min/week)	1,220.1±2,582.9	759.3±2,050.6	<0.001
Moderate intensity (METs.min/week)	466.9±1,468.8	297.8±1,150.4	<0.001
Walking (METs.min/week)	959.8±1,363.9	769.6±1,405.1	<0.001
All physical activity (METs.min/week)	2,592.6±4,333.5	1,803.5±3,339.0	<0.001
Sedentary time (min/day)	305.1±218.5	353.6±230.7	<0.001

Physical activity and sedentary time were measured using the International Physical Activity Questionnaire-Short Form (IPAQ-SF).

Figure 2. The changes in mental health, including fear and anxiety, depressed mood, and psychological stress during the pandemic.

ated patients' activities of daily living, including physical activity and sedentary time, before and during the pandemic. The proportions of patients with decreased and increased physical activity were 32.6% and 4.1%, respectively. Vigorous-/moderate-intensity activities and walking during the pandemic were significantly decreased compared to those before the pandemic (vigorous: 759.3±2,050.6 MET.min/week vs. 1,220.1±2,582.9 MET.min/week, $p<0.001$; moderate: 297.8±1,150.4 MET.min/week vs. 466.9±1,468.8 MET.min/week, $p<0.001$, walking; 769.6±1,405.1 MET.min/week vs. 959.8±1,363.9 MET.min/week, $p<0.001$, respectively). Furthermore, total physical activity during the pandemic was significantly decreased compared to before the pandemic (1,803.5±3,339.0 MET.min/week vs. 2,592.6±4,333.5 MET.min/week, $p<0.001$).

The sedentary time increased and decreased in 33.8% and 1.5% of patients, respectively. The sedentary time per day during the pandemic was significantly increased compared to before the pandemic (353.6±230.7 min/day vs. 305.1±218.5 min/day, $p<0.001$).

Changes in mental health during the pandemic

Regarding changes in mental health, increased fear and anxiety, depressed mood, and psychological stress were re-

ported in 50.0%, 27.7%, and 37.2% of patients, respectively during the pandemic (Fig. 2).

Hypoglycemic agent changes during the pandemic

During the pandemic, the usage of hypoglycemic agents was strengthened, unchanged/modified, and attenuated in 10.0%, 82.4%, and 7.6%, respectively.

Association of lifestyle changes and mental health with changes in HbA1c and body weight

The mean HbA1c value slightly but significantly decreased during the pandemic (7.09% ±0.73% to 6.94% ±0.73%, $p<0.001$). In contrast, the body weight did not significantly vary during the pandemic compared with before the pandemic (69.0±14.4 kg vs. 68.8±14.5 kg, $p=0.267$). Tables 3 and 4 describe how lifestyle changes were related to HbA1c and body weight changes. Irregular meal timing was positively associated, but changes in walking time were negatively associated with changes in HbA1c ($\beta=0.202$, $p<0.001$, and $r=-0.116$, $p=0.048$, respectively).

A multiple regression analysis included all items from the questionnaire and patient attributes (age, sex, BMI, duration of diabetes, and changes in hypoglycemic agents) as explanatory variables and the change in the HbA1c as an ob-

Table 3. The Association of Changes in Lifestyles and Mental Health with the Changes in HbA1c (n=340).

	Univariate		Regression coefficient in ordinal variables		Multivariate	
	r	p	β	p	β	p
Age (years)	0.016	0.774			-0.039	0.714
Sex (0=Female, 1=Male)	-0.067	0.224			-0.144	0.109
Duration of diabetes (years)	-0.079	0.149			0.091	0.350
BMI (kg/m ²)	0.085	0.122			-0.035	0.748
Employment (0=No, 1=Yes)	-0.067	0.223				
Change in hypoglycemic agents						
Unchanged/modified vs. strengthened			0.014	0.801	0.019	0.842
Attenuated vs. unchanged/modified			-0.034	0.543	0.182	0.068
Changes in dietary habits						
Regularity of meal timing						
Unchanged vs. became regular			-0.035	0.52	0.054	0.602
Became irregular vs. unchanged			0.202	<0.001	0.328	0.001
Amount of total diet						
Unchanged vs. decreased			0.052	0.349	-0.104	0.323
Increased vs. unchanged			0.107	0.056	0.053	0.656
Consumption of carbohydrates						
Unchanged vs. decreased			-0.049	0.383	0.074	0.495
Increased vs. unchanged			0.039	0.486	-0.151	0.156
Consumption of snacks						
Unchanged vs. decreased			-0.076	0.182	0.165	0.111
Increased vs. unchanged			-0.021	0.713	-0.063	0.506
Consumption of fruits						
Unchanged vs. decreased			-0.103	0.067	-0.155	0.122
Increased vs. unchanged			0.023	0.685	-0.101	0.294
Consumption of alcohol						
Unchanged vs. decreased			-0.041	0.611	-0.052	0.614
Increased vs. unchanged			0.058	0.469	-0.020	0.836
Changes in activities of daily living						
Vigorous intensive activity	-0.009	0.869			-0.029	0.825
Moderate intensive activity	0.011	0.854			0.143	0.267
Walking	-0.116	0.048			-0.146	0.166
Total activity	-0.028	0.644				
Sedentary time	0.077	0.173			-0.015	0.885
Changes in mental health						
Fear and anxiety						
Unchanged vs. decreased			0.038	0.498	-0.114	0.307
Increased vs. unchanged			0.033	0.552	0.064	0.563
Depressive mood						
Unchanged vs. decreased			0.106	0.054	0.228	0.207
Increased vs. unchanged			0.053	0.339	0.056	0.661
Psychological stress						
Unchanged vs. decreased			0.064	0.248	-0.117	0.489
Increased vs. unchanged			0.053	0.338	0.032	0.799

HbA1c: glycated hemoglobin A1c, BMI: body mass index

Change in hypoglycemic agents was coded as “strengthened” (-1 point), “unchanged/modified” (0 point), and “attenuated” (1 point).

Each lifestyle change was coded as “decreased/became regular” (-1 point), “unchanged” (0 point), and “increased/became irregular” (1 point).

jective variable. Irregular meal timing was identified as a factor significantly related to the change in HbA1c ($\beta=0.328$, $p=0.001$) (Table 3). In contrast, some factors, such as the BMI ($r=0.187$, $p=0.001$), irregular meal timing ($\beta=0.190$, $p=0.001$), changes in total diet ($r=0.158$, $p=0.004$), carbohydrate consumption ($r=0.120$, $p=0.029$), fear and anxiety ($r=$

0.162, $p=0.003$), and psychological stress ($r=0.122$, $p=0.026$), were positively associated with body weight change. However, only the change in walking time showed an inverse association with body weight change ($r=-0.154$, $p=0.008$).

With all items of the questionnaire and patient attributes (age, sex, BMI, duration of diabetes, and changes in hypoglycemic agents) included as explanatory variables and the change in body weight as an objective variable, we performed a multiple regression analysis. This analysis revealed that only the change in walking time significantly contributed to body weight change ($\beta=-0.245$, $p=0.025$) (Table 4).

Correlation between mental health and lifestyle changes

The correlations between mental health and lifestyle changes are shown in Table 5. The change in fear and anxiety was positively associated with changes in meal timing regularity ($r=0.129$, $p=0.019$) and carbohydrate consumption ($r=0.127$, $p=0.021$). The change in depressed mood was positively associated with changes in carbohydrate consumption ($r=0.142$, $p=0.010$) and alcohol ($r=0.161$, $p=0.037$). In addition, changes in psychological stress were positively associated with changes in consumption of carbohydrates ($r=0.183$, $p=0.001$), snacks ($r=0.151$, $p=0.008$), and sedentary time ($r=0.158$, $p=0.004$).

Discussion

In this study, we demonstrated considerable lifestyle and mental health changes and the independent impact of the lifestyle changes on glycemic and body weight control during the medium-term period of the COVID-19 pandemic in Japan. In addition, we also demonstrated an association between mental health and lifestyle changes.

Based on the study's findings, we confirmed that dietary habits worsened and activities of daily living decreased in approximately 20% and 30% of patients, respectively. In addition, the study demonstrated that irregular meal timing was positively associated with changes in HbA1c levels during the COVID-19 pandemic. Furthermore, decreased walking time was associated with changes in body weight. As is well known, dietary habits and activities of daily living are quite important factors for glycemic and body weight control in patients with diabetes (5, 6). Recent studies have revealed that lifestyle changes during the early phase of the COVID-19 pandemic influenced glycemic and body weight control in patients with diabetes in core pandemic areas (7-9). Kishimoto et al. investigated lifestyle changes, including dietary habits and physical activity, body weight, and HbA1c levels in 168 patients with diabetes (7). They revealed that physical activity levels (coded as increased, no change, and decreased) and dietary habits (coded as improved, no change, and deteriorated) were significant determinants of group categorization (patients with elevated HbA1c levels $>0.2\%$ and decreased HbA1c levels $>0.2\%$) in the

multiple logistic regression analyses. However, their analysis did not include individual components of dietary habits and physical activities, unlike our report. In addition, changes in body weight have not been analyzed. Munekawa et al. evaluated the association of changes in dietary habits quantitatively, including total diet, snacks, prepared food, and exercise, with changes in HbA1c and body weight in 203 patients with diabetes, using a visual analogue scale (8). They revealed that the total diet intake was positively associated with changes in HbA1c and that total diet intake and snack consumption were positively associated while exercise was negatively associated with changes in body weight, findings that are partially consistent with those of our study. However, these associations were unadjusted, and whether or not these factors are independently related to HbA1c and body weight changes is unclear.

In contrast, Takahara et al. evaluated the association of detailed lifestyle changes with HbA1c and body weight changes in 1,402 patients with diabetes using a linear regression model (9). They reported that the change in leisure time for physical activities was inversely associated with HbA1c and weight changes. In contrast, the quantitative change in meals, with a decline in meals eaten out and snacks, was positively associated with HbA1c and weight changes. However, in their reports, physical activity was not analyzed using the quantified indicators used in our study.

These previous studies were performed during the early stage of the COVID-19 pandemic. However, the present study was performed during the medium-term period of about half a year after the COVID-19 pandemic onset, a major difference from previous studies. We observed worsening of dietary habits and a decrease in activities of daily living in a considerable number of patients. Such changes may have led to a deterioration in glycemic control and weight gain, which are generally consistent with previous studies conducted during the early stage of the pandemic. Conversely, the mean HbA1c during the pandemic was significantly lower than before the pandemic. This improvement may have been due in part to patients' medical guidance from attending physicians and/or their self-care behavior in glycemic control during the pandemic. Thus, considering these findings, including our own, physicians should pay close attention to lifestyle changes leading to worsening glycemic control and body weight gain and provide careful dietary and exercise guidance to diabetic patients while the COVID-19 pandemic persists.

Regarding mental health changes, we confirmed that over 50% of patients had deteriorated mental health. A recent study reported that as many as 87% of people with diabetes were affected by psychological stress during the initial stage of the COVID-19 pandemic (11). It is well known that patients with diabetes tend to have more psychological problems, including anxiety, depression, and stress, than the general population (16). In addition, diabetes has been associated with higher severity and mortality rates due to COVID-19 (17, 18). Furthermore, because such warnings were

Table 4. The Association of Changes in Lifestyles and Mental Health with the Changes in Body Weight (n=340).

	Univariate		Regression coefficient in ordinal variables		Multivariate	
	r	p	β	p	β	p
Age (years)	0.025	0.651			0.043	0.692
Sex (0=Female, 1=Male)	0.034	0.224			0.092	0.316
Duration of diabetes (years)	-0.064	0.239			-0.003	0.973
BMI (kg/m ²)	0.187	0.001			0.085	0.449
Employment (0=No, 1=Yes)	-0.025	0.645				
Change in hypoglycemic agents						
Unchanged/modified vs. strengthened			0.034	0.533	-0.087	0.387
Attenuated vs. unchanged/modified			0.025	0.350	-0.001	0.994
Changes in dietary habits						
Regularity of meal timing						
Unchanged vs. became regular			-0.028	0.613	-0.072	0.498
Became irregular vs. unchanged			0.190	0.001	0.110	0.274
Amount of total diet						
Unchanged vs. decreased			0.089	0.104	-0.040	0.709
Increased vs. unchanged			0.158	0.004	0.028	0.818
Consumption of carbohydrates						
Unchanged vs. decreased			0.064	0.247	0.184	0.104
Increased vs. unchanged			0.120	0.029	0.053	0.626
Consumption of snacks						
Unchanged vs. decreased			-0.068	0.227	0.004	0.969
Increased vs. unchanged			-0.038	0.498	-0.025	0.797
Consumption of fruits						
Unchanged vs. decreased			-0.029	0.610	-0.064	0.533
Increased vs. unchanged			-0.022	0.691	-0.104	0.290
Consumption of alcohol						
Unchanged vs. decreased			-0.098	0.213	-0.102	0.338
Increased vs. unchanged			-0.017	0.828	-0.067	0.499
Changes in activities of daily living						
Vigorous intensive activity	-0.017	0.764			-0.045	0.739
Moderate intensive activity	-0.045	0.427			0.068	0.605
Walking	-0.154	0.008			-0.245	0.025
Total activity	-0.062	0.297				
Sedentary time	0.080	0.151			-0.045	0.677
Changes in mental health						
Fear and anxiety						
Unchanged vs. decreased			-0.035	0.517	-0.035	0.756
Increased vs. unchanged			0.162	0.003	0.052	0.651
Depressive mood						
Unchanged vs. decreased			-0.036	0.509	-0.176	0.344
Increased vs. unchanged			0.103	0.061	-0.041	0.753
Psychological stress						
Unchanged vs. decreased			0.020	0.712	0.186	0.288
Increased vs. unchanged			0.122	0.026	0.115	0.356

HbA1c: glycated hemoglobin A1c, BMI: body mass index

Change in hypoglycemic agents was coded as “strengthened” (-1 point), “unchanged/modified” (0 point), and “attenuated” (1 point).

Each lifestyle change was coded as “decreased/became regular” (-1 point), “unchanged” (0 point), and “increased/became irregular” (1 point).

widely disseminated through the mass media in Japan, fear of transmission may have deteriorated the mental health of patients with diabetes. Indeed, a recent meta-analysis re-

vealed that patients with noninfectious chronic diseases, including diabetes, had a higher risk of depression and anxiety than others during the COVID-19 pandemic (19).

Table 5. The Correlation between Mental Health and Lifestyle Changes.

	Regularity of meal timing	Amount of total diet	Consumption of carbohydrates	Consumption of snacks	Consumption of fruits	Consumption of alcohol	Total physical activity	Sedentary time
Fear and anxiety	r=0.129 p=0.019	r=0.036 p=0.514	r=0.127 p=0.021	r=-0.098 p=0.077	r=0.054 p=0.332	r=0.115 p=0.138	r=-0.051 p=0.391	r=0.103 p=0.064
Depressed mood	r=-0.004 p=0.943	r=0.106 p=0.055	r=0.142 p=0.010	r=-0.086 p=0.118	r=0.086 p=0.118	r=0.161 p=0.037	r=-0.058 p=0.331	r=0.106 p=0.057
Psychological stress	r=0.078 p=0.156	r=0.080 p=0.148	r=0.183 p=0.001	r=0.151 p=0.008	r=0.038 p=0.492	r=0.160 p=0.172	r=-0.024 p=0.682	r=0.158 p=0.004

HbA1c: glycated hemoglobin A1c

Although there was no significant association between changes in mental health and glycemic control in this study, mental health deterioration was associated with worsening of dietary habits, such as increased carbohydrates and snacks. Recent studies have reported an association between changes in mental health and dietary habits in patients with diabetes during the COVID-19 pandemic (8, 12). Sanker et al. reported that 15.5% of patients had increased mental stress during the lockdown. The majority had an unhealthy dietary pattern, including higher consumption of snacks, which was consistent with our study findings (12). Munekawa et al. also reported that 41.9% of patients experienced increased stress during the early phase of the COVID-19 pandemic, and this was positively associated with changes in prepared food intake (8). The causal relationship between mental health deterioration and increased consumption of carbohydrates and snacks may be attributed to the finding that negative emotion is likely to cause emotional hunger and eating. In addition, carbohydrate ingestion stimulates serotonin production, which enhances mood and alleviates stress (20, 21).

Regarding the association between mental health and activities of daily living, we found an association between increased psychological stress and sedentary time. Stanton et al. reported that higher depression, anxiety, and stress symptoms were associated with decreased physical activity during the COVID-19 pandemic, based on a study of 1,491 Australian adults (22). Although the association between mental health and activities of daily living is bidirectional (23), deterioration of mental health may be more likely to lead to a decrease in activities of daily living during the pandemic.

Given these findings, the deterioration of mental health may lead indirectly to worsened glycemic control through the worsening of self-care, including dietary habits and activities of daily living during the pandemic. Unfortunately, we found no significant association between changes in mental health and glycemic control in this study; however, negative psychological factors are directly associated with the worsening of glycemic control through increased generation of stress hormones, including cortisol (24, 25). However, a systematic review and meta-analysis showed that psychosocial interventions modestly but significantly improved glycemic control and mental health outcomes (26). Therefore, we believe that physicians should pay close attention to changes in mental health and lifestyle changes while

the COVID-19 pandemic persists.

Several limitations associated with the present study warrant mention. First, the responses to the questions were dependent on memory and the patients' subjective opinions, which may have affected the outcome validity. Second, this study was performed in a single diabetes center in Japan, with a relatively small sample size and no control. Third, we used the IPAQ-SF, a self-report questionnaire that assesses physical activity and has been accurately validated in people 15-69 years old. However, our study population included elderly people over 70 years old. This data collection tool has only recently been used in clinical practice to evaluate this age group (27, 28). Fourth, the COVID-19 infection rate was lower and measures employed in the study area less stringent than those in Western countries and other regions with government-enforced lockdowns and quarantines. Therefore, these results may not be representative of other populations. Furthermore, this study did not include patients whose visits to our department had been suspended due to the pandemic. Those patients may have been experiencing more lifestyle changes related to dietary habits, activities of daily living, and mental health. Therefore, the present results may have been underestimated. Finally, although we detected an association between lifestyle changes and glycemic and body weight control, and changes in mental health and dietary habits, we were unable to clarify any causal relationship because of the cross-sectional design of this study. Therefore, further prospective, large-scale clinical studies are needed to clarify the causality. However, we believe that this study's findings will provide important information on diabetes care during the COVID-19 pandemic.

In conclusion, the persistence of the COVID-19 pandemic has had a considerable impact on the lifestyles and mental health of patients with diabetes. In addition, lifestyle changes were shown to be associated with glycemic and body weight control, and mental health changes were associated with lifestyle changes. Thus, physicians need to provide more careful diet and exercise guidance and mental health-care to patients with diabetes during the COVID-19 pandemic.

Author's disclosure of potential Conflicts of Interest (COI).

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