BMJ Open Diabetes Research & Care

Improved glucometrics in people with type 1 diabetes 1 year into the COVID-19 pandemic

Namam Ali ⁽ⁱ⁾, ¹ Soumia El Hamdaoui, ¹ Giesje Nefs,^{2,3,4} Cornelis J Tack ⁽ⁱ⁾, ¹ Bastiaan E De Galan ⁽ⁱ⁾, ^{1,5,6}

To cite: Ali N, El Hamdaoui S, Nefs G, *et al.* Improved glucometrics in people with type 1 diabetes 1 year into the COVID-19 pandemic. *BMJ Open Diab Res Care* 2022;**10**:e002789. doi:10.1136/ bmjdrc-2022-002789

Additional supplemental material is published online only. To view, please visit the journal online (http://dx.doi. org/10.1136/bmjdrc-2022-002789).

Received 25 January 2022 Accepted 1 May 2022



© Author(s) (or their employer(s)) 2022. Re-use permitted under CC BY-NC. No commercial re-use. See rights and permissions. Published by BMJ.

For numbered affiliations see end of article.

Correspondence to

Namam Ali; namam.ali@radboudumc.nl

ABSTRACT

Introduction Various studies have shown a number of glycemic parameters to improve over several weeks in people with type 1 diabetes during the first surge of the COVID-19 pandemic. Whether and to what extent such improvement is sustained during following COVID-19 surges remains unknown. Therefore, the aim of this study was to investigate glycemic parameters during the first year of the COVID-19 pandemic in people with type 1 diabetes and to determine factors associated with glycemic improvement.

Research design and methods This was an observational cohort study in people with type 1 diabetes, aged \geq 16 years. We compared glycated hemoglobin (HbA_{1c}) and flash glucose monitoring (FGM) downloads between the prelockdown period and approximately 1 year thereafter. Using logistic regression analysis, we assessed associations between an HbA_{1c} reduction of at least 0.5% (~5.5 mmol/mol) with baseline clinical characteristics and self-reported changes in psychological well-being and lifestyle behavior related to COVID-19.

Results A total of 437 participants were included. As compared with prepandemic data, 1 year after the start of the COVID-19 pandemic and associated lockdowns, HbA_{1c} had decreased from 7.9%±1.1% (63±12 mmol/mol) to 7.5%±1.0% (59±11 mmol/mol) (p<0.001), whereas time in range increased from 55.8%±16.7% to 58.6%±16.7% (p=0.004) and time below (<3.9 mmol/L) and above (>13.9 mmol/L) range and glucose variability all decreased (all p<0.05). FGM use, higher HbA_{1c} at baseline and current smoking were independently associated with an HbA_{1c} decrease of at least 0.5%, whereas self-reported changes in psychological well-being and lifestyle behavior related to the first surge of the COVID-19 pandemic and associated lockdowns were not.

Conclusions The COVID-19 pandemic and related lockdown measures were associated with improvement in glucometrics, including HbA_{1c} and FGM data, in individuals with type 1 diabetes, particularly in FGM users, those with higher HbA_{1c} at baseline or current smokers.

INTRODUCTION

As of January 1, 2022, worldwide >312 million people have been diagnosed with infection caused by SARS-CoV-2 and >5.5 million have died.¹ People with both type 1 and type 2 diabetes mellitus have a twofold increased risk for severe COVID-19 infection

WHAT IS ALREADY KNOWN ON THIS TOPIC

⇒ Short-term improvement in several glycemic parameters has been shown in people with type 1 diabetes during the first surge of the COVID-19 pandemic.

WHAT THIS STUDY ADDS

- ⇒ Glycated hemoglobin (HbA_{1c}) and flash glucose monitoring (FGM) data improve, over 1 year of follow-up after the start of the COVID-19 pandemic and implementation of various lockdown measures.
- ⇒ The improvement in glycemic parameters is largely consistent across subgroups.
- ⇒ Use of FGM, higher HbA_{1c} at baseline and current smoking are independently associated with an HbA_{1c} decrease of at least 0.5% (~5.5 mmol/mol).

HOW THIS STUDY MIGHT AFFECT RESEARCH, PRACTICE AND/OR POLICY

⇒ Pandemics and the associated lockdown measures are associated with improvement rather than with worsening of glycemic control in individuals with type 1 diabetes.

and mortality compared with people without diabetes.²⁻⁴ Both hyperglycemia at presentation and elevated glycated hemoglobin (HbA_{1c}) are independently associated with severity of COVID-19 infections and increased mortality.⁵⁻⁷

To control the COVID-19 outbreak, many countries implemented several (lockdown) measures over the past 1.5 years. In the Netherlands, such lockdown measures, first implemented in March 2020 and changed throughout the year based on rates of infection, included stay-at-home orders for people working in non-vital areas of society, social distancing, a curfew and closures of schools, restaurants, theatres and other public spaces. There were also changes in the healthcare system with downscaling of outpatient clinic visits, partly replaced by virtual meetings and postponement of certain (surgical) procedures.⁸ ⁹ Unique and widespread impact of these lockdown measures have been

described on (societal) behavior, including those related to lifestyle and self-care, which may have directly or indirectly affected glucose (self) management and glycemic parameters.¹⁰

Several studies have reported various glycemic parameters to improve over several weeks in people with type 1 diabetes during the first surge of the COVID-19 pandemic.^{11–16} Whether and to what extent such improvement is sustained during following COVID-19 surges remains unknown, as studies with longer follow-up are lacking. Furthermore, few studies have investigated factors that contribute to this glycemic improvement. Therefore, the aim of the present study was to investigate, in a contemporary cohort of people with type 1 diabetes, glycemic parameters prestart and 1 year after the start of the COVID-19 pandemic and associated lockdown measures and factors associated with HbA_{1c} improvement.

RESEARCH DESIGN AND METHODS Study design and population

This study was an observational cohort study. People with type 1 diabetes who attend the diabetes outpatient clinic of the Radboud University Medical Center were invited to participate in the study. Inclusion criteria for the study were type 1 diabetes, age ≥ 16 years, sufficient comprehension of the Dutch language and ability to provide informed consent. Type 1 diabetes diagnosis was based on clinical criteria. Participants were excluded when the medical history mentioned severe psychiatric comorbidity or other comorbidity interfering with completing the questionnaires. After providing consent, participants were either invited to the outpatient clinic of our hospital or a virtual appointment was made for data collection.

We assessed 702 people with type 1 diabetes for eligibility in the study, 184 of whom did not want to participate and 81 persons were excluded for various reasons. As such, a total of 437 people with type 1 diabetes agreed to participate in the study (online supplemental figure 1). Individuals who declined to participate or were excluded did not differ from those who were included with respect to age and sex. For 427 participants (97.7%), HbA_{1c} data were available for both time periods (ie, period before and 1 year after the start of the COVID-19 pandemic and associated lockdown measures), whereas 200 participants (45.8%) gave consent to access their flash glucose monitoring (FGM) data. A total of 190 (43.5%) participants had both HbA_{1c} and FGM data for these periods.

Demographics and clinical characteristics

Data collection took place from February 2020 to April 2021. Demographics and clinical characteristics, including age, sex, presence of microvascular and macrovascular diabetes complications, smoking, alcohol use, data on insulin treatment (mode and dose), mode of glucose monitoring, diabetes duration, number of (severe¹⁷) hypoglycemic events, hypoglycemia awareness status assessed with the Clarke questionnaire¹⁸ and frequency of hospitalizations in the past year, were obtained by questionnaires and verified from clinical records, wherever possible. Body mass index was measured at the outpatient clinic or retrieved from the clinical record if the appointment for data collection was virtual.

At the end of the first COVID-19 surge, we administered the Problem Areas in Diabetes (PAID-5) questionnaire, where a score ≥ 8 indicates possible diabetes-related emotional distress,¹⁹ and a questionnaire, containing questions about whether there was any emotional distress, worries and changes in diabetes self-management or behavior due to the COVID-19 pandemic and the associated lockdown during the first COVID-19 surge (online supplemental table 1). Emotional distress due to the COVID-19 pandemic could include any of the following: having little interest or pleasure in activities, feeling nervous, anxious or tense, unable to stop worrying, feeling lonely or isolated, feeling little control over life or feeling insecure about the future.

Glycemic parameters

HbA, levels were derived from the electronic medical record, whereas online data sharing platforms were used to gain access to FGM data. Both HbA_{1c} and FGM data before the onset of the COVID-19 pandemic and associated lockdown measures (ie, prelockdown data) were collected between September 2019 and March 2020, whereas 1-year follow-up data were collected between January and March 2021. The primary outcome in this study was the change in HbA_{1c} between these defined periods. Data from FGM were collected for the participants who were using FGM and had data for both defined periods. FGM data (representation of 28 days) used for the analysis were: time sensor is active (% of time), average number of scans per day (n), average glucose (mmol/L), glucose variability (%), time below target range (% of time glucose $\leq 3.9 \text{ mmol/L}$), time in target range (% of time glucose 3.9-10.0 mmol/L), time above target range (% of time glucose >10.0 mmol/L), number of low glucose events (glucose ≤3.9 mmol/L) and duration of low glucose events.

Statistical analyses

Continuous data are expressed as mean (±SD) if normally distributed or median (IQR) if not normally distributed. Categorical data are presented as number (percentage, %). Demographics and clinical characteristics were compared using one-way analysis of variance followed by post hoc analysis with least significant difference or Kruskal-Wallis followed by post hoc analysis Mann-Whitney U test for continuous variables, depending on their distribution, and χ^2 test was used for categorical data. Glycemic data from the prelockdown and the 1-year follow-up period were compared using the paired samples t-test or the Wilcoxon signed rank test or McNemar test, as appropriate. We performed subgroup analysis for the primary outcome (ie, change in HbA_{1c}) for subgroups defined by sex, age, baseline HbA_{1c}, diabetes duration, insulin administration strategy, hypoglycemia awareness status, mode of glucose monitoring, FGM start year, PAID-5 score ≥ 8 and any change in psychological well-being and lifestyle behavior due to the first surge of COVID-19 pandemic and associated lock-down. Mixed models were used to investigate differences between these subgroups.

We investigated demographics, clinical characteristics and COVID-19-related changes associated with an HbA_{1c} reduction of $\geq 0.5\%$ (~5.5 mmol/mol), using univariable and multivariable logistic regression analysis. For the multivariable regression analysis, we used different models to adjust for potential confounders. In model 1 of the analysis, we adjusted for age and sex; model 2 was additionally adjusted for impaired awareness of hypoglycemia (IAH) and hospitalization in the past year and model 3 was additionally adjusted for continuous subcutaneous insulin infusion (CSII) use and insulin dose. Self-monitoring of blood glucose (SMBG) was used as reference category for mode of glucose monitoring.

All statistical analyses were performed with IBM SPSS V.25 software (Armonk, New York, USA). Graphs were

made with GraphPad Prism V.8.0 (La Jolla, California, USA). Data are shown as mean±SD, median (IQR) or number (%), as appropriate. ORs with 95% CIs are reported for the logistic regression analysis and for mixed models. P values <0.05 were considered statistically significant.

RESULTS

Characteristics of the study sample

Demographic, clinical and behavioral characteristics stratified according to mode of glucose monitoring are shown in table 1. In the total study population there was an equal sex balance (50.3% male) and HbA_{1c} on average was $7.9\% \pm 1.1\%$ (63±12 mmol/mol). At the time of inclusion, 330 (75.5%) participants used the FGM sensor for a median duration of 1.0 year (IQR 0.0–2.0), 35 (8.0%) used real-time continuous glucose monitoring (rt-CGM) and 72 (16.5%) used SMBG. Eight individuals used a hybrid closed-loop system and one individual used a do-it-yourself closed-loop system. There were no differences between FGM, rt-CGM and SMBG users with respect to age, sex, diabetes duration, HbA_{1c} and most other clinical

Table 1 Characteristics of the study populations (n=437) stratified according to mode of glucose monitoring					
		FGM users (n=330)	rt-CGM users (n=35)	SMBG users (n=72)	
Age, years		48.9±15.7	47.9±13.5	53.6±17.4	
Males, n (%)		169 (51.2%)	11 (31.4%)	40 (55.6%)	
Microvascular complications,† n (%)		183 (55.5%)	18 (51.4%)	42 (59.2%)	
Macrovascular complications,‡ n (%)		40 (12.1%)	4 (11.4%)	11 (15.7%)	
Smoking, n (%)	Never	176 (54.2%)	20 (58.8%)	36 (50.7%)	
	Past	112 (34.5%)	11 (32.4%)	23 (32.4%)	
	Current	37 (11.4%)	3 (8.8%)	12 (16.9%)	
Alcohol use, n (%)		258 (79.4%)	23 (65.7%)	55 (77.5%)	
CSII, n (%)		162 (49.1%)	34 (97.1%)**	25 (34.7%)*	
Daily insulin use, IU/kg		0.6 (0.5–0.7)	0.5 (0.4–0.6)	0.6 (0.5–0.8)	
Diabetes duration, years		29.5 (15.0–40.0)	31.0 (21.0–44.0)	26.0 (15.0–41.5)	
Number of hypoglycemic events, n/week		3 (2–7)	5 ((3–7)*	2 (1–4)**	
Number of severe hypoglycemic events in past year, n/participant/year		1 (0–2)	1 (1–2)	1 (0–2)	
At least one severe hypoglycemic event in past year, n (%) $$		43 (13.1%)	5 (14.3%)	8 (11.6%)	
Impaired awareness of hypoglycemia (Clarke score ≥4), n (%)		46 (14.0%)	20 (57.1%)**	13 (18.3%)	
Hospitalization in past year, n (%)		11 (3.3%)	4 (11.4%)	4 (5.6%)	
HbA _{1c} , % (mmol/mol)		7.6±1.0 (60±11)	7.6±1.1 (59±12)	7.8±1.2 (62±13)	
Body mass index, kg/m ²		25.9±4.1	25.1±4.1	25.6±4.4	

Data are presented as mean±SD or median (IQR) or number (%).

*P<0.05, **p<0.01: vs FGM users.

†Microvascular complications: retinopathy, nephropathy or neuropathy.

#Macrovascular complications: coronary heart disease, cerebrovascular disease or peripheral artery disease.

CSII, continuous subcutaneous insulin infusion; FGM, flash glucose monitoring; HbA_{1c}, glycated hemoglobin; rt-CGM, real-time continuous glucose monitoring; SMBG, self-monitoring of blood glucose.

parameters (table 1). The prevalence of IAH, the rate of hypoglycemia and the rate of CSII use was higher in the rt-CGM users as compared with FGM users. The rate of CSII use and the number of hypoglycemic events was lower in people using SMBG as compared with those on FGM.

A PAID-5 score of at least 8 was present in 22.3% (n=86) of the individuals. Any emotional distress due to the COVID-19 pandemic was reported by 45.3% (n=177) of the participants and 8.9% (n=39) had worries about their diabetes and/or COVID-19 in this period. About 17% (n=63) of the individuals reported to aim for another (ie, higher or lower) blood glucose level and 22.7% (n=86) of participants expressed to be more involved in their diabetes care during the COVID-19 pandemic period compared with the period before. Changes in physical activity, diet and sleep duration after the first surge of the COVID-19 pandemic and associated lockdown measures were reported by 58.2% (n=221), 28.7% (n=109) and 23.2% (n=88) of the individuals, respectively. Any selfreported change in psychological well-being and lifestyle behavior due to the first surge of COVID-19 pandemic and associated lockdown was present in 75.3% (n=329) of the individuals.

Glycemic parameters

The average follow-up between the two periods of data collection was 376 ± 94 days. HbA_{1c} decreased from $7.9\%\pm1.1\%$ (63 ± 12 mmol/mol) before the lockdown to $7.5\%\pm1.0\%$ (59 ± 11 mmol/mol) 1 year later (mean

difference (95% CI): 0.4% (0.3 to 0.4) (4 mmol/mol (3 to 5)); p<0.001)). The fall in HbA_{1c} between the defined time periods occurred in all participants independent of the mode of glucose monitoring, but was slightly greater for participants using FGM compared with participants not using FGM, that is, rt-CGM users and SMBG (0.2% (0.1 to 0.4) vs 0.4% (0.3 to 0.5); p=0.044). There was no significant difference in HbA_{1c} change between participants using rt-CGM or SMBG (figure 1 and online supplemental figure 2). There was also no difference in HbA_{1c} decrease between participants using FGM with or without available FGM data (0.5% (0.3 to 0.6) vs 0.4% (0.3 to 0.6); p=0.923). The number of participants with HbA_{1c} <7% (53 mmol/mol) increased from 80 (18.4%) before the lockdown to 126 (29.5%) 1 year later (p<0.001).

With respect to FGM parameters, time in range increased over the 1 year of follow-up, whereas times below 3.9 and above 13.9 mmol/L and glucose variability all decreased (table 2). The number of low glucose events and the duration of these events did not change. Mean percentage of time during the day that FGM was active increased from $70\%\pm31\%$ in the prelockdown period to $87\%\pm22\%$ after 1-year follow-up (18% (95% CI 12.8 to 22.5); p<0.001)) and the median scan frequency per day (IQR) increased from 6 (4–10) to 10 (6–14)^{6–14} (median difference (IQR): 4 (2–4); p<0.001).

HbA_{1c} change in subgroups

Figure 2 shows the subgroup analysis for the primary outcome. The decrease in HbA_{1c} that we observed over



Figure 1 HbA_{1c} change in participants with available HbA_{1c} for both time periods (n=427), stratified according to mode of glucose monitoring. FGM, flash glucose monitoring; HbA_{1c}, glycated hemoglobin; rt-CGM, real-time continuous glucose monitoring; SMBG, self-monitoring of blood glucose.

Table 2 Flash glucose monitoring data before and 1 year after start of the pandemic (n=200)							
		Before the pandemic	One-year follow-up	P value			
Time sensor active, %		70±31	87±22	< 0.001			
Scan frequency per day		6 (4–10)	10 (6–14)	<0.001			
Time in target range, %		55.8±16.7	58.6±16.7	0.004			
Time below target range, %	<3.9 mmol/L	5.6±5.2	3.8±4.2	<0.001			
	3.0–3.9 mmol/L	4.0±3.1	3.3±3.2	0.001			
	<3.0 mmol/L	1.6±2.6	0.5±1.4	<0.001			
Time above target range, %	>10.0 mmol/L	38.6±18.7	37.6±18.0	0.336			
	10.1-13.9 mmol/L	24.2±8.4	25.3±8.9	0.050			
	>13.9 mmol/L	14.5±13.4	12.3±11.7	0.002			
Average glucose, mmol/L		9.4±1.9	9.3±1.7	0.405			
Glucose variability, %		39.2±6.4	36.5±5.9	<0.001			
Hypoglycemic events, n		12 (5–25)	11 (5–24)	0.223			
Duration of hypoglycemic events, min		90 (64–113)	86 (62–115)	0.105			
Data are shown as mean±SD or media	an (IQR).						

the 1-year follow-up period was consistent in subgroups defined by sex, age, baseline HbA_{1c} , diabetes duration, insulin administration strategy, hypoglycemia awareness status, mode of glucose monitoring, FGM start year, PAID-5 score and any self-reported change in

psychological well-being and lifestyle behavior due to the first surge of COVID-19 pandemic and associated lockdown. Although HbA_{1c} improved in both males and females, the fall in HbA_{1c} was greater in men compared with women (p=0.049). Furthermore, HbA_{1c} decrease



Figure 2 Subgroup analysis for change in HbA_{1c}. CSII, continuous subcutaneous insulin infusion; FGM, flash glucose monitoring; HbA_{1c}, glycated hemoglobin; IAH, impaired awareness of hypoglycemia; MDI, multiple daily injection; NAH, normal awareness of hypoglycemia; PAID-5, Problem Areas in Diabetes-5 questionnaire; rt-CGM, real-time continuous glucose monitoring; SMBG, self-monitoring of blood glucose.

minol/mol/(ii=+27) in different models					
	Model 1 OR (95% CI)	Model 2 OR (95% CI)	Model 3 OR (95% CI)		
FGM use	2.09 (1.21 to 3.63)	2.65 (1.40 to 5.01)	2.58 (1.35 to 4.93)		
Prelockdown HbA _{1c}	2.42 (1.91 to 3.07)	2.33 (1.83 to 2.98)	2.33 (1.81 to 2.99)		
Current smoking	2.37 (1.29 to 4.37)	2.06 (1.01 to 4.17)	2.11 (1.03 to 4.35)		

Table 3 Multivariable logistic regression analysis relating demographic and clinical factors to HbA_{1c} decrease of $\geq 0.5\%$ (~5.5 mmol/mol) (n=427) in different models

Model 1: adjusted for age and sex.

Model 2: additionally adjusted for IAH and hospitalization in past year.

Model 3: additionally adjusted for CSII and insulin dose.

CSII, continuous subcutaneous insulin infusion; FGM, flash glucose monitoring; HbA_{1c}, glycated hemoglobin; IAH, impaired awareness of hypoglycemia.

was greater in the participants with baseline HbA_{1c} \geq 7.5% compared with those with lower HbA_{1c} (p<0.001), in the participants using FGM compared with those using SMBG or rt-CGM (p=0.001) and in the individuals who started using FGM after January 1, 2020 compared with individuals who already used FGM before that date (p=0.001). A PAID-5 score \geq 8 or self-reported change in psychological well-being or lifestyle behavior during the first surge of the pandemic were not associated with a greater improvement in HbA_{1c}.

Factors associated with improved HbA₁₀

Univariable logistic regression analysis showed that use of FGM (OR (95% CI): 2.1 (1.2 to 3.6), higher HbA_{1c} level at baseline (2.4 (1.9 to 3.0) and current smoking (2.4 (1.3 to 4.3) were significantly associated with an HbA_{1c} decrease of at least 0.5% (online supplemental table 2). After correction for potential confounders, all these associations remained significant in all three models in the multivariable logistic regression analysis (table 3). No COVID-19-related change in psychological well-being or behavior were found to be related to an HbA_{1c} reduction of at least 0.5%.

DISCUSSION

This study shows a clinically significant improvement in a number of glycemic parameters in individuals with type 1 diabetes over 1 year follow-up after the start of the COVID-19 pandemic and implementation (and temporary lifting) of various lockdown measures. HbA_{1c} fell by 0.4%, which was mirrored by increases in time in range and decreases in times below and substantially above target range and in glucose variability. Although we found no relation between changes in psychological wellbeing or behavior and the change in HbA_{1c}, it is likely that the COVID-19 pandemic has played an important role in the improvement in glucometrics in people with type 1 diabetes 1 year after the onset of the pandemic.

Our data are in line with previous studies that examined the effect of the first COVID-19 wave on glucose outcomes^{11–16} and extend these to a longer period. Previous studies that examined FGM parameters over several weeks of the first COVID-19 wave reported improvements in time in range¹¹⁻¹⁶ and time above range¹¹⁻¹⁵ that were of about similar magnitude as what we observed. Only one study reported a decrease in time below range¹⁶ similar to our results, while other studies found no difference¹²⁻¹⁴ or even an increase in this outcome.¹⁵ An increase in the time that the sensor was active and number of daily scans was also reported by one study,¹¹ but not by other studies,^{12 14-16} whereas some,^{13 16} but not all studies^{11 12 14 15} reported a decrease in glucose variability. However, due to the much shorter follow-up periods, most of these studies were unable to observe an improvement in HbA_{1c}, at least not to the extent observed here.¹¹⁻¹⁶

Our data contrast with a study from India showing deterioration of glycemic parameters, including an increase in HbA_{1c}, in people with type 1 diabetes during a lockdown period in the first COVID-19 wave.²⁰ However, since this change was mainly due to non-availability of insulin or glucose strips during this period, the disparity between this study and our study are probably explained by the major differences between the Netherlands and India in healthcare system and resources. A study conducted in the UK described an ~80% fall in HbA_{1c} tests for diabetes diagnosis and management during the COVID-19 pandemic, which was associated with a rise in HbA1, among individuals with diabetes with pre-existing suboptimal glycemic outcomes.²¹ Also, the few studies that investigated glucose parameters during the lockdown period in people with type 2 diabetes have reported no improvement in HbA_{1c} or even an increase of HbA_{1c} in older people.^{11 22} Explanations for the discrepancy with people with type 1 diabetes include the usually much less frequent use of continuous or flash glucose monitoring and the potential educational gap with respect to handling of insulin therapy in the small proportion of people with type 2 diabetes using insulin.

In our study, we found that the use of FGM and higher HbA_{1c} at baseline were independent predictors of a larger HbA_{1c} decrease. This is in line with a meta-analysis supporting greater fall in HbA_{1c} in people using FGM and with higher initial HbA_{1c} values.²³ FGM enables people with diabetes to monitor glucose levels more closely, when compared with SMBG, and it has been shown that

Clinical care/Education/Nutrition

more daily scans are associated with a lower HbA_{1c} .²⁴ Our finding that scan frequency under FGM users increased during follow-up may support this notion. People using rt-CGM did not show a large increase, however, this was a relatively small group with a lower start HbA_{1c} , as well as avoiding (severe) hypoglycemia rather than lowering HbA_{1c} being the main indication for its use.

Surprisingly, our analysis also showed that current smoking was independently associated with greater HbA_{1c} improvement. This association remained significant after correction for multiple confounders. It could be argued that people who smoke are more inclined to improve glucose management since smoking is a risk factor for more severe COVID-19 infections. Whether there are other, (patho-) physiological, mechanisms underlying this association would require further study, yet this finding should not be regarded as a beneficial effect of smoking.

There could be several reasons for the improvement of glycemic parameters that we observed. First, fear of contracting the virus and the potential role of suboptimal glycemic control in becoming ill may have been incentives for people to increase efforts to improve glucose levels to lower the risk of COVID-19 infection.^{25 26} However, diabetes-related distress, emotional distress due to the first surge of the COVID-19 pandemic, worries about diabetes or COVID-19, more involvement in diabetes care and aiming for other blood glucose levels in the COVID-19 pandemic period, were not associated with improvement in glucose control. It should we acknowledged, however that the COVID-19 questionnaires were completed within the first months after the start of the pandemic and associated lockdown, of which most of the participants reported no detrimental effects. We cannot exclude an effect of changes in psychological well-being later during the pandemic on the change in HbA_{1c}.

The improvement in glycemic parameters could also have been due to changes in lifestyle or daily activities in the lockdown period with less fluctuation in glucose profiles and insulin needs. Indeed, a recent study showed that time in range increased from 54.4% to 65.2% (p=0.010) in people with type 1 diabetes who stopped working and stayed at home during the pandemic compared with those who continued working.¹² It could also be speculated that these conditions made it easier for people with diabetes to count carbohydrates, administer insulin in time, have more time to monitor their glucose profiles (eg, from sensors) and pay more attention to diet and exercise.

The improvement in glucometrics could also be independent of the pandemic, for example, with more people becoming increasingly familiar with FGM. HbA_{1c} decreased to a greater extent in people using FGM as compared with people not (yet) using FGM and the decrease was more pronounced in participants who recently started FGM. However, glycemic parameters also improved in people who used FGM longer, which argues against this option being the only explanation. In the Netherlands, the introduction of FGM had largely taken place before the COVID-19 pandemic, and this study shows paired data of those already on FGM. The magnitude of the improvement that we show is larger than the HbA_{1c} drop of 0.2% associated with flash monitor initiation, reported by a nationwide Scottish observational study.²⁷

A key strength of our study is the 1-year follow-up, with data on HbA_{1c} and FGM parameters enabling us to study the long-term impact of the COVID-19 pandemic on glucose management in people with type 1 diabetes. Our study also has limitations. First, we only included people with type 1 diabetes with HbA_{1c} and/or FGM data that could be evaluated after 1 year, which may introduce selection bias. Not everyone consented in providing us access to their FGM data, which resulted in missing FGM data. However, the fall in HbA_{1c} for participants with FGM data was comparable to that among participants without FGM data. Another limitation is that we did not investigate glycemic parameters from rt-CGM in the participants using rt-CGM. However, although such data would have helped to explain the apparent smaller change in HbA₁ in participants using rt-CGM, the number of people using rt-CGM was small. Due to data protection regulation, we could only make a group comparison between the participants and those who declined to participate or were excluded, with respect to age and sex. We can therefore not fully rule out selection bias. Furthermore, several aspects around changes in behavior because of the COVID-19 pandemic and associated lockdown measures (eg, change in work habit, change in insulin dose, etc) were not addressed in our questionnaire and the questionnaire was administered only once. Finally, this was a monocentre study, potentially limiting extrapolation to the wider diabetes population, and since the study was observational, we were unable to attribute causation to the results.

In conclusion, our study shows an association between COVID-19 pandemic and related lockdown measures with a clinically relevant improvement of various glycemic parameters in individuals with type 1 diabetes during 1-year follow-up. We found a greater fall in HbA_{1c} in participants who used FGM, had higher HbA_{1c} at baseline or were current smokers. The glycemic improvements were substantial and seemingly sustainable, but further studies are needed to see whether and to what extent these improvements can be sustained after the COVID-19 pandemic-associated lockdown measures are lifted.

Author affiliations

¹Department of Internal Medicine, Radboud University Nijmegen, Nijmegen, The Netherlands

²Department of Medical Psychology, Radboudumc Radboud Institute for Health Sciences, Nijmegen, The Netherlands

³Center of Research on Psychological disorders and Somatic diseases (CoRPS), Department of Medical and Clinical Psychology, Tilburg University, Tilburg, The Netherlands

⁴Diabeter, National Treatment and Research Center for Children, Adolescents and Adults with Type 1 Diabetes, Rotterdam, Netherlands

⁶CARIM School for Cardiovascular Disease, Maastricht University, Maastricht, Netherlands

Acknowledgements The authors thank the participants of the study and the contributing physicians.

Contributors NA and BEDG designed the study. NA and SEH recruited the participants and collected the clinical data. NA analyzed the data and wrote the first draft of the manuscript. NA, BEDG, GN, CJT discussed the results and provided input for and commented on the manuscript at all stages. NA, BEDG and SEH had full access to the data and all authors approved the final version, and accept the responsibility to submit the study for publication. The guarantors of the study are NA and BEDG.

Funding The authors have not declared a specific grant for this research from any funding agency in the public, commercial or not-for-profit sectors.

Competing interests None declared.

Patient consent for publication Not applicable.

Ethics approval Ethical approval for the study was obtained from the Institutional Review Board of the Radboud University Medical Center (NL-71207.091.19) and the study was conducted in accordance with the Declaration of Helsinki. All participants gave written informed consent before participation.

Provenance and peer review Not commissioned; externally peer reviewed.

Data availability statement All data relevant to the study are included in the article or uploaded as supplementary information.

Supplemental material This content has been supplied by the author(s). It has not been vetted by BMJ Publishing Group Limited (BMJ) and may not have been peer-reviewed. Any opinions or recommendations discussed are solely those of the author(s) and are not endorsed by BMJ. BMJ disclaims all liability and responsibility arising from any reliance placed on the content. Where the content includes any translated material, BMJ does not warrant the accuracy and reliability of the translations (including but not limited to local regulations, clinical guidelines, terminology, drug names and drug dosages), and is not responsible for any error and/or omissions arising from translation and adaptation or otherwise.

Open access This is an open access article distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited, appropriate credit is given, any changes made indicated, and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/.

ORCID iDs

Namam Ali http://orcid.org/0000-0003-4302-7820 Cornelis J Tack http://orcid.org/0000-0003-0322-1653 Bastiaan E De Galan http://orcid.org/0000-0002-1255-7741

REFERENCES

- 1 WHO. Who coronavirus (COVID-19) Dashboard, 2021. Available: https://covid19.who.int/ [Accessed 01 Jan 2022].
- 2 O'Malley G, Ebekozien Ö, Desimone M, *et al*. CÓVID-19 hospitalization in adults with type 1 diabetes: results from the T1D exchange multicenter surveillance study. *J Clin Endocrinol Metab* 2021;106:e936–42.
- 3 Nassar M, Nso N, Baraka B, *et al*. The association between COVID-19 and type 1 diabetes mellitus: a systematic review. *Diabetes Metab Syndr* 2021;15:447–54.
- 4 Yang J, Zheng Y, Gou X, et al. Prevalence of comorbidities and its effects in patients infected with SARS-CoV-2: a systematic review and meta-analysis. Int J Infect Dis 2020;94:91–5.
- 5 Lim S, Bae JH, Kwon H-S, *et al.* COVID-19 and diabetes mellitus: from pathophysiology to clinical management. *Nat Rev Endocrinol* 2021;17:11–30.

- 6 Bhandari S, Rankawat G, Singh A, et al. Impact of glycemic control in diabetes mellitus on management of COVID-19 infection. Int J Diabetes Dev Ctries 2020:340–5.
- 7 Ruan Y, Ryder REJ, De P, *et al*. A UK nationwide study of people with type 1 diabetes admitted to hospital with COVID-19 infection. *Diabetologia* 2021;64:1717–24.
- 8 Tack CCJ. Would you believe? A virus changes diabetes care. J Diabetes Sci Technol 2020;14:795–6.
- 9 de Galan BE. Relative limited impact of COVID-19 on diabetes: a personal view. *J Diabetes Sci Technol* 2020;14:727–8.
- 10 Scott ES, Jenkins AJ, Fulcher GR. Challenges of diabetes management during the COVID-19 pandemic. *Med J Aust* 2020;213:56–7.
- 11 Ruissen MM, Regeer H, Landstra CP, et al. Increased stress, weight gain and less exercise in relation to glycemic control in people with type 1 and type 2 diabetes during the COVID-19 pandemic. BMJ Open Diabetes Res Care 2021;9:e002035.
- 12 Bonora BM, Boscari F, Avogaro A, *et al.* Glycaemic control among people with type 1 diabetes during Lockdown for the SARS-CoV-2 outbreak in Italy. *Diabetes Ther* 2020;11:1369–79.
- 13 Prabhu Navis J, Leelarathna L, Mubita W, et al. Impact of COVID-19 lockdown on flash and real-time glucose sensor users with type 1 diabetes in England. Acta Diabetol 2021;58:231–7.
- 14 Mesa A, Viñals C, Pueyo I, et al. The impact of strict COVID-19 lockdown in Spain on glycemic profiles in patients with type 1 diabetes prone to hypoglycemia using standalone continuous glucose monitoring. *Diabetes Res Clin Pract* 2020;167:108354.
- 15 Dover AR, Ritchie SA, McKnight JA, *et al.* Assessment of the effect of the COVID-19 lockdown on glycaemic control in people with type 1 diabetes using flash glucose monitoring. *Diabet Med* 2021;38:e14374.
- 16 Pla B, Arranz A, Knott C, *et al*. Impact of COVID-19 Lockdown on glycemic control in adults with type 1 diabetes mellitus. *J Endocr* Soc 2020;4:bvaa149.
- 17 Östenson CG, Geelhoed-Duijvestijn P, Lahtela J, et al. Self-Reported non-severe hypoglycaemic events in Europe. *Diabet Med* 2014;31:92–101.
- 18 Clarke WL, Cox DJ, Gonder-Frederick LA, et al. Reduced awareness of hypoglycemia in adults with IDDM. A prospective study of hypoglycemic frequency and associated symptoms. *Diabetes Care* 1995;18:517–22.
- 19 McGuire BE, Morrison TG, Hermanns N, et al. Short-form measures of diabetes-related emotional distress: the Problem Areas in Diabetes Scale (PAID)-5 and PAID-1. *Diabetologia* 2010;53:66–9.
- 20 Verma A, Rajput R, Verma S, *et al.* Impact of lockdown in COVID 19 on glycemic control in patients with type 1 diabetes mellitus. *Diabetes Metab Syndr* 2020;14:1213–6.
- 21 Holland D, Heald AH, Stedman M, et al. Impact of the UK COVID-19 pandemic on HbA1c testing and its implications for diabetes diagnosis and management. Int J Clin Pract 2021;75:e13980.
- 22 Falcetta P, Aragona M, Ciccarone A, *et al.* Impact of COVID-19 lockdown on glucose control of elderly people with type 2 diabetes in Italy. *Diabetes Res Clin Pract* 2021;174:108750.
- 23 Evans M, Welsh Z, Ells S, et al. The impact of flash glucose monitoring on glycaemic control as measured by HbA1c: a metaanalysis of clinical trials and real-world observational studies. *Diabetes Ther* 2020;11:83–95.
- 24 Dunn TC, Xu Y, Hayter G, *et al.* Real-world flash glucose monitoring patterns and associations between self-monitoring frequency and glycaemic measures: a European analysis of over 60 million glucose tests. *Diabetes Res Clin Pract* 2018;137:37–46.
- 25 Joensen LE, Madsen KP, Holm L, et al. Diabetes and COVID-19: psychosocial consequences of the COVID-19 pandemic in people with diabetes in Denmark-what characterizes people with high levels of COVID-19-related worries? *Diabet Med* 2020;37:1146–54.
- 26 Joensen LE, Steenberg JL, Madsen KP, et al. What people with diabetes in Denmark worry about during the COVID-19 pandemic: a longitudinal study of the first 3 months of the COVID-19 pandemic. *Diabet Med* 2021;38:e14665.
- 27 Jeyam A, Gibb FW, McKnight JA, et al. Flash monitor initiation is associated with improvements in HbA₁ levels and DKA rates among people with type 1 diabetes in Scotland: a retrospective nationwide observational study. *Diabetologia* 2022;65:159–72.