

# UC San Diego

## UC San Diego Previously Published Works

### Title

Accuracy and Glycemic Efficacy of Continuous Glucose Monitors in Critically Ill COVID-19 Patients: A Retrospective Study.

### Permalink

<https://escholarship.org/uc/item/1jv7q741>

### Journal

Journal of diabetes science and technology, 17(3)

### ISSN

1932-2968

### Authors

Boeder, Schafer  
Kobayashi, Emily  
Ramesh, Gautam  
[et al.](#)

### Publication Date

2023-05-01

### DOI

10.1177/19322968221113865

Peer reviewed

# Accuracy and Glycemic Efficacy of Continuous Glucose Monitors in Critically Ill COVID-19 Patients: A Retrospective Study

Journal of Diabetes Science and Technology  
2023, Vol. 17(3) 642–648

© 2022 Diabetes Technology Society



Article reuse guidelines:

sagepub.com/journals-permissions

DOI: 10.1177/19322968221113865

journals.sagepub.com/home/dst



Schafer Boeder, MD<sup>1,\*</sup>, Emily Kobayashi, BS<sup>2,\*</sup>,  
Gautam Ramesh, BS<sup>3</sup>, Brittany Serences, MSN, FNP, BC-ADM<sup>4</sup>,  
Kristen Kulasa, MD<sup>1</sup>, and Amit R. Majithia, MD<sup>1</sup>

## Abstract

**Background:** Continuous glucose monitoring (CGM) is approved for insulin dosing decisions in the ambulatory setting, but not currently for inpatients. CGM has the capacity to reduce patient-provider contact in inpatients with coronavirus disease 2019 (COVID-19), thus potentially reducing in hospital virus transmission. However, there are sparse data on the accuracy and efficacy of CGM to titrate insulin doses in inpatients.

**Methods:** Under an emergency use protocol, CGM (Dexcom G6) was used alongside standard point-of-care (POC) glucose measurements in patients critically ill from complications of COVID-19 requiring intravenous (IV) insulin. Glycemic control during IV insulin therapy was retrospectively assessed comparing periods with and without adjunctive CGM use. Accuracy metrics were computed and Clarke Error Grid analysis performed comparing CGM glucose values with POC measurements.

**Results:** Twenty-four critically ill patients who met criteria for emergency use of CGM resulted in 47333 CGM and 5677 POC glucose values. During IV insulin therapy, individuals' glycemic control improved when CGM was used (mean difference  $-30.7$  mg/dL). Among 2194 matched CGM: POC glucose pairs, a high degree of concordance was observed with a mean absolute relative difference of 14.8% and 99.5% of CGM: POC pairs falling in Zones A and B of the Clarke Error Grid.

**Conclusions:** Continuous glucose monitoring use in critically ill COVID-19 patients improved glycemic control during IV insulin therapy. Continuous glucose monitoring glucose data were highly concordant with POC glucose during IV insulin therapy in critically ill patients suggesting that CGM could substitute for POC measurements in inpatients thus reducing patient-provider contact and mitigating infection transmission.

## Keywords

accuracy, continuous glucose monitoring, coronavirus disease 2019, glycemic control, ICU, insulin titration

## Introduction

Uncontrolled hyperglycemia is associated with increased length of hospital stay and mortality.<sup>1</sup> Accurate blood glucose measurements are essential for safe and effective titration of insulin particularly in critically ill patients on intravenous (IV) insulin to achieve optimal blood glucose targets.<sup>2</sup> The current standard of care for measuring inpatient blood glucose for insulin dosing decision is to use point-of-care (POC) glucose meter devices which require hospital staff to manually sample patients at frequencies ranging from once every 4 to 6 hours for those receiving subcutaneous insulin injections, to once every 0.5 to 2 hours for those on IV insulin.<sup>2</sup> While POC is the standard of care, clinically significant glycemic events can be missed between POC tests even at the highest sampling frequency of 0.5 hours. Simply

increasing the frequency of POC testing would increase burden on the hospital staff who carry out the tests and increase

<sup>1</sup>Division of Endocrinology and Metabolism, Department of Medicine, University of California, San Diego, La Jolla, CA, USA

<sup>2</sup>Bioinformatics and Systems Biology Graduate Program, University of California, San Diego, La Jolla, CA, USA

<sup>3</sup>School of Medicine, University of California, San Diego, La Jolla, CA, USA

<sup>4</sup>Department of Nursing Education, Development and Research, University of California, San Diego, La Jolla, CA, USA

\*Schafer Boeder and Emily Kobayashi contributed equally.

## Corresponding Author:

Amit R. Majithia, MD, Division of Endocrinology and Metabolism, Department of Medicine, University of California, San Diego, 9500 Gilman Drive, La Jolla, CA 92093-0688, USA.  
Email: amajithia@health.ucsd.edu

their exposure to coronavirus disease 2019 (COVID-19) and other transmissible infections.

Continuous glucose monitoring (CGM), which requires one sensor insertion every 10 to 14 days and samples interstitial glucose every 1 to 15 minutes, has the potential to address the shortcomings of POC testing.<sup>3</sup> While non-adjunctive CGM has been approved by the US Food and Drug Administration (FDA) for use in the ambulatory setting since 2016,<sup>4,5</sup> their use in hospitals has not been approved and remains experimental despite mounting evidence that CGM improves glycemic control in inpatients.<sup>6-10</sup> The COVID-19 pandemic accelerated the adoption of CGM technology within hospitals,<sup>11,12</sup> reflecting the need to reduce health care provider exposure to severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) while improving glycemic control in patients with diabetes who are much more likely to be hospitalized after SARS-CoV-2 infection than their counterparts without diabetes.<sup>13,14</sup> In April 2020, the FDA announced that it would not object to expanded utilization of remote monitoring devices, including the use of CGM in hospitals.<sup>15</sup> More recently, the FDA provided the Dexcom G6 with a breakthrough device designation.<sup>16</sup>

Following the original FDA announcement, our institution developed an emergency protocol for the use of CGM in the intensive care unit (ICU) to assist with glycemic control in patients with COVID-19-related critical illness requiring IV insulin. Here, we report on data collected from 24 individuals relating glycemic outcomes and CGM accuracy to POC measurements.

## Methods

Continuous glucose monitoring sensors (Dexcom G6, Dexcom, San Diego, CA) were placed on hospitalized patients under an Emergency Operational Need Protocol approved by UC San Diego Health in 2020 allowing use in critically ill patients with COVID-19 requiring continuous IV insulin infusion for glycemic control. The emergency use protocol was designed to allow the frequency of POC glucose testing (and associated adjustment to the insulin infusion rate) to be extended to a maximum of every two hours if the CGM glucose and trend arrow indicated the patient was not in severe hyperglycemia (glucose >250 mg/dL) or at risk of developing hypoglycemia. Under this protocol, a trained physician, nurse practitioner, physician assistant, or diabetes nurse specialist placed the CGM sensor on the posterior upper arm of the patient and initiated the system. The posterior upper arm was chosen as the location for sensor placement to minimize the risk of device dislodgement, sensor compression during prone positioning, or skin injury. The CGM receiver (smartphone) was placed in the patient room within 20 feet of the patient, and data were shared in real time to a central iPad (iPad 5, iPadOS 14.7.1, Apple, Cupertino, CA) located at the nursing station and monitored by trained nursing staff. Use of the system for glucose monitoring began

after the initial two-hour sensor warm-up period. A Clinical Laboratory Improvement Amendments (CLIA)–waived hospital glucose meter, the Accu-Chek Inform II (Roche Diagnostics, Indianapolis, IN) Blood Glucose Monitoring System, was used to collect POC values.

Retrospective analysis of data obtained from the Emergency Protocol was approved by the UC San Diego Institutional Review Board. Data were collected from 23 patients who wore CGM in the hospital between July 2020 and February 2021 and met the following criteria: (1) age  $\geq 18$  years, (2) admitted with confirmed COVID-19 infection to a medical intensive care unit (MICU) at either UC San Diego Medical Center in Hillcrest or UC San Diego Jacobs Medical Center, and (3) required continuous IV insulin infusion therapy. One additional patient did not receive insulin infusion (IV insulin was planned but never implemented) but met all other inclusion criteria. The comparison of glycemic control during insulin infusion achieved with and without CGM excluded data from this patient, but the analysis of CGM accuracy included this patient. Continuous glucose monitoring data were downloaded using Dexcom CLARITY software. The POC glucose data and patient information, including demographics and medical history, were extracted from the electronic health record (EHR) system. Non-numeric glucose value entries were filtered, and POC data points with glucose values of less than 20 mg/dL were removed as they were considered artifacts.

Data analysis and plotting were implemented in Python 3.9.7 using the standard library, NumPy 1.20.3, SciPy 1.7.1, pandas 1.3.4, Matplotlib 3.4.3, python-dateutil 2.8.2, and statsmodels 0.12.2 packages. Glycemic control during periods “on” and “off” CGM was assessed by comparing POC glucose values during periods of IV insulin infusion with or without CGM use. A Kolmogorov-Smirnov (KS) test, a two-sided student *t*-test, and a  $\chi^2$  test were used to determine statistical significance. The CGM and POC pairs were matched based on timepoints, where each POC datapoint was matched with a CGM datapoint within five minutes. Clarke Error Grid analysis was computed using a publicly available Python script (<https://github.com/suetAndTie/ClarkeErrorGrid>) modified to allow for shading zones in different colors. Mean absolute relative difference (MARD) was calculated as previously described.<sup>17</sup> Univariate linear models were used to assess the effect of patient and clinical conditions on MARD.

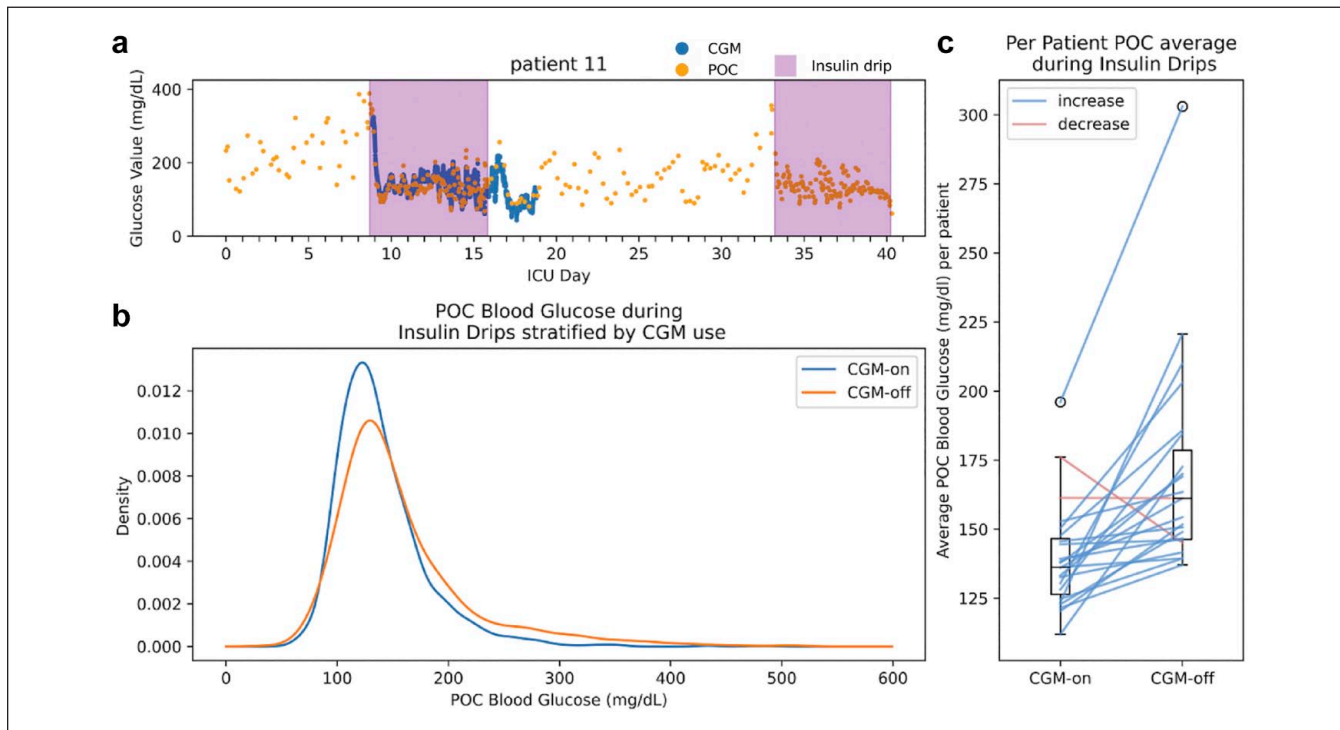
## Results

Characteristics of the patients included in the study ( $n = 24$ ) are shown in Table 1. Participants were predominantly men (71%) with poorly controlled diabetes (mean A1c 8.9%) and diagnosed with COVID-19 pneumonia/respiratory failure. Almost all participants required ventilatory ( $n = 23$ ) and hemodynamic ( $n = 22$ ) support with an eventual mortality rate of 54%. For each participant, periods of IV insulin

**Table 1.** Demographics Table.

Patient demographics	<i>n</i>	% or range
Sex		
Male	17	71%
Female	7	29%
Anthropometric measurements and lab values		
Age (median)	61	50-84
BMI (median)	31.0	20.5-45.0
A1c on admission (median)	8.9%	6.2%-15.8%
Race/ethnicity		
Hispanic (non-White)	11	46%
White	9	38%
Black or African American	2	8%
Other, mixed or unknown race	2	8%
Interventions and clinical status		
Mortality	13	54%
COVID-19 PNA/ARDS	24	100%
Intubation	23	96%
Vasopressors	22	92%
Glucocorticoids	22	92%

Abbreviations: BMI, body mass index; PNA, pneumonia; ARDS, acute respiratory distress syndrome.



**Figure 1.** (a) CGM and POC glucose values plotted by time for a representative patient. Insulin infusion times are highlighted. (b) Density plot of POC blood glucose values during insulin infusion over all patients. The POC values are stratified by concurrent CGM use ( $n = 2101$ ) or no CGM use ( $n = 1739$ ). (c) Boxplot of average POC values for each patient ( $n = 23$ ) during insulin infusion when on CGM compared with off CGM. Lines connect the means of individual patients. Blue lines indicate an increase in average blood glucose when off CGM. Orange lines denote a decrease in average blood glucose when off CGM. Abbreviations: CGM, continuous glucose monitoring; POC, point-of-care.

therapy were extracted from the EHR and intersected with CGM and POC glucose values. Figure 1a demonstrates the

POC glucose values during insulin infusion—with and without CGM use—for a representative patient.

**Table 2.** MARD and Median ARD by POC Categories.

POC cutoffs	MARD	Median ARD	n
<70	22.2%	16.2%	11
70-180	15.0%	12.8%	1806
180-250	13.6%	11.0%	284
≥250	12.9%	12.6%	93

Abbreviations: MARD, mean absolute relative difference; ARD, absolute relative difference; POC, point-of-care.

**Table 3.** Comparison of Patient Characteristics on MARD Using a Univariate Linear Model.

Patient characteristic	Coefficient	Standard error	P value
Sex	0.725	0.223	.748
Age	0.056	0.099	.581
BMI	-0.317	-0.154	.052
A1c on admission	0.795	0.382	.049
Interventions and clinical status			
Mortality	-1.698	2.006	.407
Intubation	3.827	5.018	.454
Vasopressors	3.418	3.602	.353
Glucocorticoids	1.088	3.668	.770

The Bonferroni threshold of significance for *P* values for this analysis is .05/8.

Abbreviations: MARD, mean absolute relative difference; BMI, body mass index.

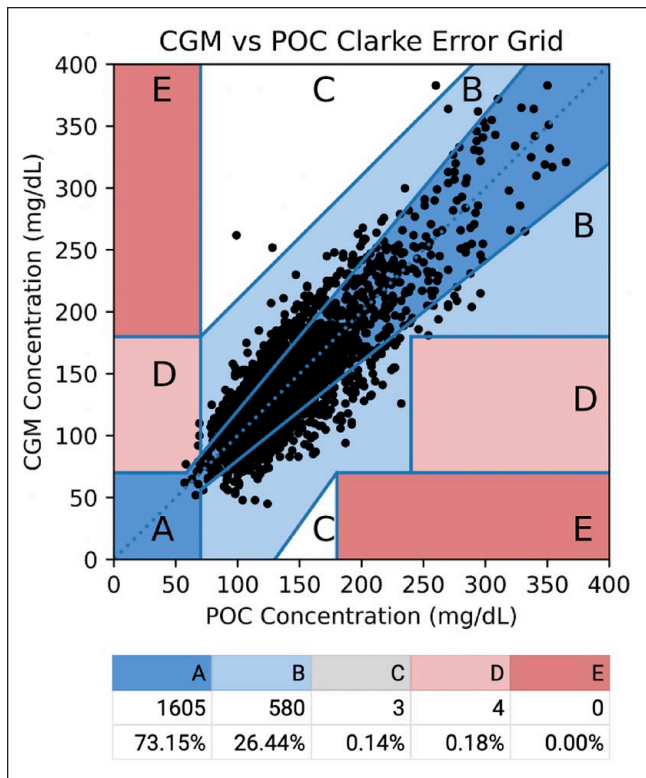
Stratifying the data by CGM use (ie, “CGM-on” and “CGM-off”), 2101 POC glucose measurements coincided with CGM-on and 1739 POC values coincided with CGM-off. The distributions of POC blood glucose levels stratified by CGM use are shown in Figure 1b, demonstrating a decrease in mean glucose (−18.5 mg/dL CGM-on vs CGM-off) and fewer extremely high glucose values in the CGM-on distribution. Statistically, these distributions were different under a two-sample KS test with test statistic 0.12 and *P* value 6.98e-13. We also performed paired analysis to evaluate intraindividual CGM-on and CGM-off periods (Figure 1c), finding a similarly decreased mean POC glucose during CGM-on of 139.5 mg/dL compared with 170.2 mg/dL CGM-off (mean decrease −30.7 mg/dL, *P* < .001 paired *t*-test). At the individual level, average POC blood glucose during CGM-on versus CGM-off periods decreased in all but two participants. The rate of hypoglycemia (<70 mg/dL) in CGM-on versus CGM-off (11 occurrences of hypoglycemia out of 2101 data points for CGM-on compared with 5 out of 1739 for CGM-off) was not significantly different ( $\chi^2$  *P* < .97). There were no instances of severe hypoglycemia (level 2 hypoglycemia; <55 mg/dL) in the CGM-on group and two (of five) instances of severe hypoglycemia in the CGM-off group.

To evaluate concordance between CGM and POC glucose, POC values were matched to the closest CGM value less than five minutes apart. From our dataset (*n* = 47333 CGM values and *n* = 5677 POC values), 2194 matched CGM: POC pairs were identified. The MARD across all matched pairs was 14.8% ± 0.5%. The median ARD was

12.7% (interquartile range [IQR]: 6.4%-20.9%). The MARD in the target range of 70 to 180 mg/dL was 15.0% (*n* = 1806). MARD for hypoglycemia (*n* = 11) and hyperglycemia (*n* = 93) ranges are listed in Table 2. To account for the unequal number of CGM: POC pairs per patient (median 70.5 pairs, interquartile range of 85), MARD was first computed on a per patient level and then averaged across all patients resulting in a similar value of 14.7%. To assess the effect of concurrent patient factors and clinical conditions on MARD, a univariate linear model regressed each condition's impact on per patient MARD (Table 3). None of the conditions tested significantly affected the MARD although A1c on admission met the nominal threshold of statistical significance (*P* = .049) for being positively associated with MARD.

To quantify clinical accuracy of CGM in this setting, we performed Clarke Error Grid analysis with the matched CGM: POC data pairs (Figure 2). Values over 400 mg/dL were excluded from analysis (two data points removed) as this exceeds the upper limit of measurement of the Dexcom G6. Of the remaining 2192 matched pairs, 73.15% fell into Zone A (clinically accurate), 26.44% fell into Zone B (benign errors that would not lead to inappropriate treatment), 0.14% fell into Zone C (overcorrection errors, harmless corrections), and 0.18% fell into Zone D (dangerous failure to detect hypo- or hyperglycemia). There were no data points in Zone E (erroneous treatment error). In summary, 99.5% of CGM: POC pairs fell into Zones A and B of the Clarke Error Grid indicating a high level of clinical accuracy for CGM in critically ill patients on IV insulin.





**Figure 2.** Clarke Error Grid of CGM versus POC blood glucose values in patients with COVID-19 ( $n = 24$ ). Dashed line indicates the 45° line. Data are rounded and may not add up to 100%. Two data points fell outside the range of this graph (POC glucose above 400 mg/dL) but would have fallen in the A or B zones. Abbreviations: CGM, continuous glucose monitoring; POC, point-of-care.

## Discussion

In this retrospective study of adjunctive CGM use in critically ill COVID-19 patients on IV insulin, we found a strong improvement in glycemic control (mean decrease 30.7 mg/dL) during periods of CGM use. An almost perfect clinical concordance was observed between CGM glucose values and POC glucose meters (99.5% CGM: POC matched values falling in Zones A and B of the Clarke Error Grid) suggesting that CGM could safely and effectively substitute POC glucose meters for IV insulin titration in this population where minimizing patient-provider contacts is imperative to infection control.

A major strength of our study includes enrollment of participants with COVID-19-related critical illness. Almost all participants required ventilatory support and over half required hemodynamic support with vasopressors—conditions that could theoretically affect interstitial glucose levels and decrease the concordance between CGM and blood glucose. These factors suggest that the observed concordance of CGM and POC glucose meter values (99.5% Clarke Error Grid Zones A and B) is likely a conservative estimate and in

the general, non-critically ill inpatient population, clinical concordance is likely to be even higher. Continuous glucose monitoring accuracy as quantified by MARD was unaffected by patient characteristics or clinical status (Table 3). Furthermore, the improved glycemic control during CGM-on versus CGM-off times during IV insulin was observed in almost all individual participants (21/23). Taken together, these findings support the use of CGM in critically ill COVID-19 patients in place of POC glucose meters, a timely finding given the trend toward increased infectiousness of newly arising SARS-CoV-2 variants.<sup>18,19</sup>

Notably, while rates of hypoglycemia during CGM-on versus CGM-off periods were statistically not different, there were more POC glucose values in the hypoglycemic range (<70 mg/dL) during CGM-on (11 of 2101 data points) than CGM-off (5 of 1739 data points). However, the hypoglycemic values were less severe during CGM-on with no instances of level 2 hypoglycemia (<55 mg/dL) compared with two instances during CGM-off. These findings are consistent with an increased detection of mild hypoglycemia by the bedside nurses who were notified of low blood glucose values by the CGM system, prompting them to check confirmatory POC glucose and provide treatment. This likely reflects a protective effect of CGM on the progression to severe hypoglycemia, a conclusion supported by multiple recent inpatient CGM trials.<sup>10,20,21</sup>

Our study has several limitations including a retrospective study design, a relatively small sample size, and lack of a CGM-only group. As patients were not prospectively enrolled, a cryptic bias in the individuals enrolled under the emergency use protocol could confound our findings. An additional cofounder is a mix of reference POC glucose values, as POC samples were taken from capillary, venous, and arterial blood. Despite a relatively small number of individuals enrolled ( $n = 24$ ), our study is among the largest testing CGM use in critically ill patients to date<sup>7,11,22-26</sup> and the amount of analyzable CGM and POC glucose data obtained ( $n = 47\,333$  CGM values,  $n = 5\,677$  POC values) drives statistically robust inferences. The lack of a CGM-only group prevents us from being able to formally test non-adjunctive CGM-based insulin titration, but our study provides evidence for the safety and potential efficacy of a CGM-based insulin titration that needs prospective validation.

## Conclusions

Continuous glucose monitors have not been widely studied in the ICU and have not been approved for inpatient use. In our study of critically ill COVID-19-positive patients on IV insulin, we observe improved glycemic control with adjunctive CGM use compared with standard POC testing alone. This finding was driven by a reduction in hyperglycemia, which translated to an improvement in average blood glucose during CGM use in 91% of participants. In addition, CGM demonstrated high concordance with POC, suggesting

that it can substitute for POC glucose measurements during IV insulin titration. Continuous glucose monitoring use would reduce patient-provider contact, thereby reducing in-hospital transmission of infectious illnesses such as SARS-CoV-2.

### Abbreviations

CGM, continuous glucose monitoring; POC, point-of-care; ICU, intensive care unit; COVID-19, coronavirus disease 2019; MARD, mean absolute relative distance; ARD, absolute relative difference; SARS-CoV-2, severe acute respiratory syndrome coronavirus 2; BMI, body mass index; PNA, Pneumonia; ARDS, Acute Respiratory Distress Syndrome; US FDA, US Food and Drug Administration; IV, intravenous; KS, Kolmogorov-Smirnov; MICU, medical intensive care unit

### Declaration of Conflicting Interests


The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.


### Funding


The author(s) received no financial support for the research, and/or authorship. Open access publication charges for this article were supported by the UC Libraries.

### ORCID iDs

Emily Kobayashi  <https://orcid.org/0000-0002-9041-5167>

Gautam Ramesh  <https://orcid.org/0000-0003-4064-3353>

Kristen Kulasa  <https://orcid.org/0000-0002-3559-5046>

Amit R. Majithia  <https://orcid.org/0000-0001-9978-2644>

### References

- Bode B, Garrett V, Messler J, et al. Glycemic characteristics and clinical outcomes of COVID-19 patients hospitalized in the United States. *J Diabetes Sci Technol*. 2020;14:813-821.
- American Diabetes Association Professional Practice Committee 16. Diabetes care in the hospital: standards of medical care in diabetes—2022. *Diabetes Care*. 2021;45:S244-S253.
- Olczuk D, Priefer R. A history of continuous glucose monitors (CGMs) in self-monitoring of diabetes mellitus. *Diabetes Metab Syndr*. 2018;12(2):181-187.
- FDA approval of Dexcom's non-adjunctive indication triggers a new era in diabetes management. <https://www.businesswire.com/news/home/20161220005992/en/FDA-Approval-of-Dexcom%E2%80%99s-Non-Adjunctive-Indication-Triggers-a-New-Era-in-Diabetes-Management>. Published 2016. Accessed July 9, 2022.
- Chamberlain JJ. Continuous glucose monitoring systems: categories and features. In: *Role of continuous glucose monitoring in diabetes treatment*. Arlington, VA: American Diabetes Association; 2018. <https://www.ncbi.nlm.nih.gov/books/NBK538975/> Accessed July 9, 2022.
- Krinsley JS, Chase JG, Gunst J, et al. Continuous glucose monitoring in the ICU: clinical considerations and consensus. *Crit Care*. 2017;21:197.
- Lu M, Zuo Y, Guo J, Wen X, Kang Y. Continuous glucose monitoring system can improve the quality of glucose control and glucose variability compared with point-of-care measurement in critically ill patients. *Medicine (Baltimore)*. 2018;97(36):e12138.
- Wallia A, Umpierrez GE, Rushakoff RJ, Klonoff DC, Rubin DJ, Hill Golden S, et al. Consensus statement on inpatient use of continuous glucose monitoring. *J Diabetes Sci Technol*. 2017;11(5):1036-1044.
- Fortmann AL, Spierling Bagsic SR, Talavera L, et al. Glucose as the fifth vital sign: a randomized controlled trial of continuous glucose monitoring in a non-ICU hospital setting. *Diabetes Care*. 2020;43(11):2873-2877.
- Singh LG, Satyarengga M, Marcano I, et al. Reducing inpatient hypoglycemia in the general wards using real-time continuous glucose monitoring: the Glucose Telemetry System, a Randomized Clinical Trial. *Diabetes Care*. 2020;43(11):2736-2743.
- Reutrakul S, Genco M, Salinas H, et al. Feasibility of inpatient continuous glucose monitoring during the COVID-19 pandemic: early experience. *Diabetes Care*. 2020;43(10):e137-e138.
- Galindo RJ, Aleppo G, Klonoff DC, et al. Implementation of continuous glucose monitoring in the hospital: emergent considerations for remote glucose monitoring during the COVID-19 pandemic. *J Diabetes Sci Technol*. 2020;14(4):822-832.
- de Almeida-Pititto B, Dualib PM, Zajdenverg L, Dantas JR, de Souza FD, Rodacki M, et al. Severity and mortality of COVID 19 in patients with diabetes, hypertension and cardiovascular disease: a meta-analysis. *Diabetol Metab Syndr*. 2020;12:75.
- Gregg EW, Sophiea MK, Weldegiorgis M. Diabetes and COVID-19: population impact 18 months into the pandemic. *Diabetes Care*. 2021;44(9):1916-1923.
- Coronavirus (COVID-19) update: FDA allows expanded use of devices to monitor patients' vital signs remotely. FDA. <https://www.fda.gov/news-events/press-announcements/coronavirus-covid-19-update-fda-allows-expanded-use-devices-monitor-patients-vital-signs-remotely>. Published 2020. Accessed July 9, 2022.
- FDA grants breakthrough device designation for Dexcom Hospital CGM system. DexCom, Inc. n.d. <https://investors.dexcom.com/news-releases/news-release-details/fda-grants-breakthrough-device-designation-dexcom-hospital-cgm>. Accessed July 9, 2022.
- Reiterer F, Polterauer P, Schoemaker M, et al. Significance and reliability of MARD for the accuracy of CGM systems. *J Diabetes Sci Technol*. 2016;11:59-67.
- Wang Y, Chen R, Hu F, Lan Y, Yang Z, Zhan C, et al. Transmission, viral kinetics and clinical characteristics of the emergent SARS-CoV-2 Delta VOC in Guangzhou, China. *EClinicalMedicine*. 2021;40:101129.
- Tian D, Sun Y, Xu H, Ye Q. The emergence and epidemic characteristics of the highly mutated SARS-CoV-2 Omicron variant. *J Med Virol*. 2022;94(6):2376-2383.
- Galindo RJ, Migdal AL, Davis GM, et al. Comparison of the FreeStyle Libre Pro flash continuous glucose monitoring (CGM) system and point-of-care capillary glucose testing in hospitalized patients with type 2 diabetes treated with basal-bolus insulin regimen. *Diabetes Care*. 2020;43(11):2730-2735.
- Singh LG, Levitt DL, Satyarengga M, Pinault L, Zhan M, Sorkin JD, et al. Continuous glucose monitoring in general wards for prevention of hypoglycemia: results from the

- Glucose Telemetry System Pilot Study. *J Diabetes Sci Technol*. 2019;14:783-790.
22. Longo RR, Elias H, Khan M, Seley JJ. Use and accuracy of inpatient CGM during the COVID-19 pandemic: an observational study of general medicine and ICU patients [published online ahead of print May 10, 2021]. *J Diabetes Sci Technol*. doi:10.1177/19322968211008446.
  23. De Block CE, Gios J, Verheyen N, et al. Randomized evaluation of glycemic control in the medical intensive care unit using real-time continuous glucose monitoring (REGIMEN Trial). *Diabetes Technol Ther*. 2015;17(12):889-898.
  24. Sadhu AR, Serrano IA, Xu J, et al. Continuous glucose monitoring in critically ill patients with COVID-19: results of an emergent pilot study. *J Diabetes Sci Technol*. 2020;14(6):1065-1073.
  25. Agarwal S, Mathew J, Davis GM, Shephardson A, Levine A, Louard R, et al. Continuous glucose monitoring in the intensive care unit during the COVID-19 pandemic. *Diabetes Care*. 2021;44(3):847-849.
  26. Preiser J-C, Lheureux O, Thoofit A, Brimioulle S, Goldstein J, Vincent J-L. Near-continuous glucose monitoring makes glycemic control safer in ICU patients. *Crit Care Med*. 2018;46(8):1224-1229.