## CONTINUOUS GLUCOSE MONITORING FOR THE INTERNIST

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Adult Endocrinology

#### Disclosures:

No conflict of interest

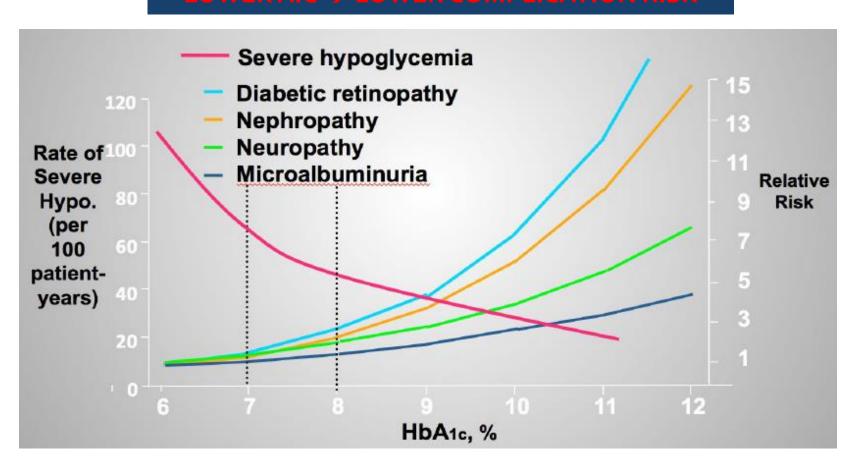
## Objectives

- Is it meaningful to use diabetes technology in 2020?
- Explain how continuous glucose monitoring (CGM) works
- Explain how to interpret main aspects of AGP
- Most common and accessible alternatives in Puerto Rico
- Main differences among them
- Cases for discussion

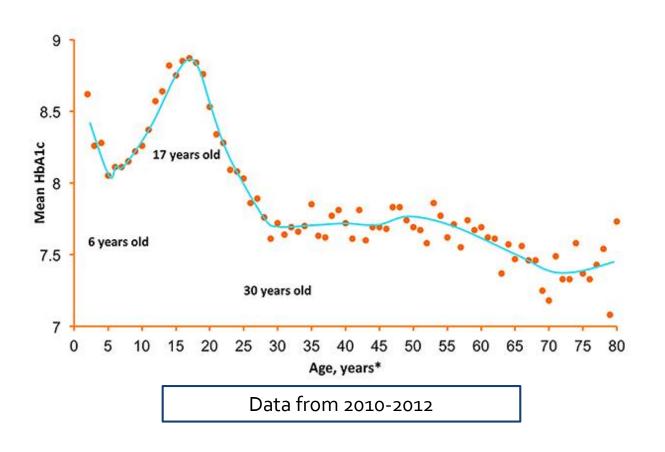
# IS IT MEANINGFUL TO USE DIABETES TECHNOLOGY IN 2020?

## Glucose Control and Complications (Diabetes Control and Complication Trial, DCCT)

#### I OWER AIC → I OWER COMPLICATION RISK



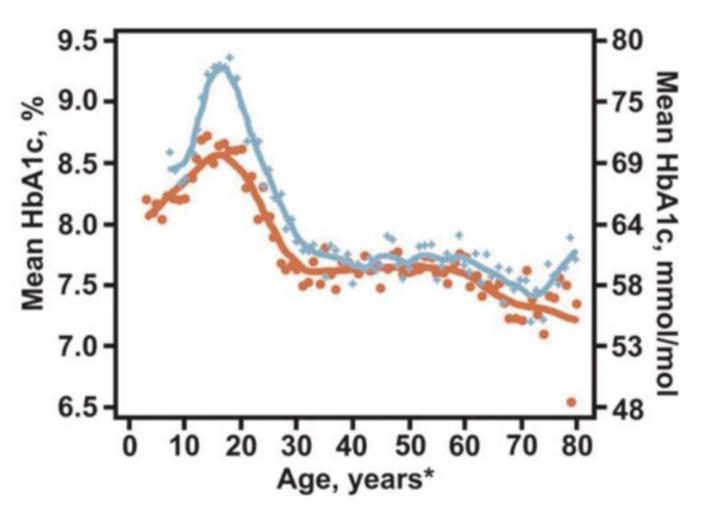
#### How controlled are our patients?



• The majority of young adults in their 20's do not fully achieve glycemic control until they are 30.

 Only 14 % of registrants between 18 and 25-years old met the recommended 7 % A1c level compared with 30 % of older adults.

#### More recent data (2016-2018 cohort)

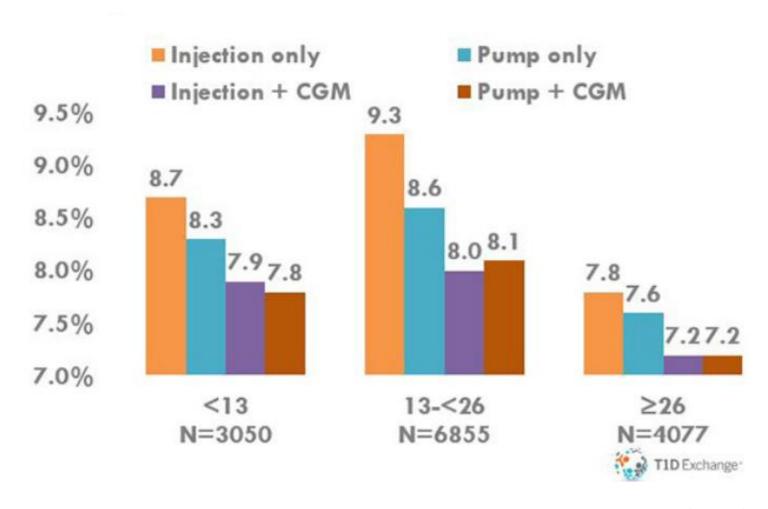


• Orange: 2010-2012

• Blue: 2016-2018

 No significant improvement in A1C despite CGM/insulin pump/diabetes technology

#### A1C comparison regardless of insulin delivery method

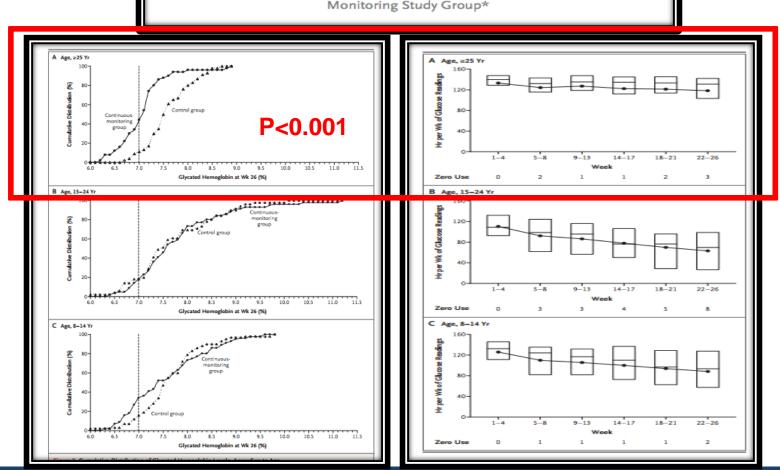


The NEW ENGLAND JOURNAL of MEDICINE

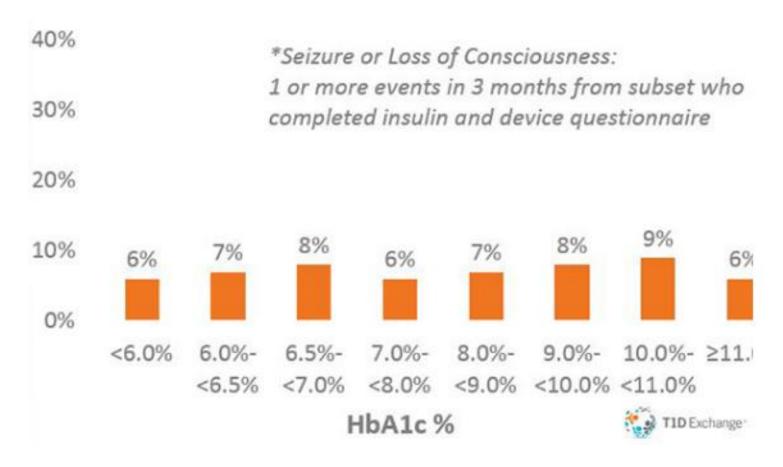
#### ORIGINAL ARTICLE

#### Continuous Glucose Monitoring and Intensive Treatment of Type 1 Diabetes

The Juvenile Diabetes Research Foundation Continuous Glucose Monitoring Study Group\*



# 3 Month frequency of severe hypoglycemia according to A1C



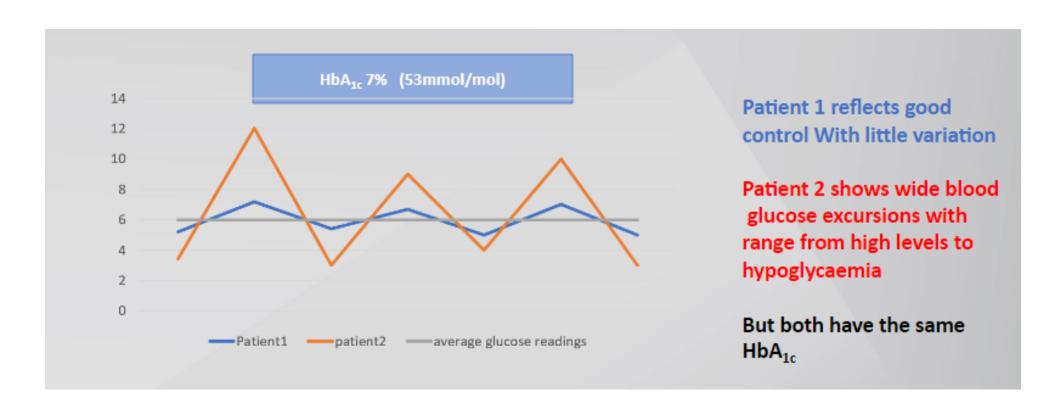
#### HBA1C pitfalls and interpretation

Table 6.1—Estimated average glucose (eAG)							
A1C (%)	mg/dL*	mmol/L					
5	97 (76–120)	5.4 (4.2–6.7)					
6	126 (100–152)	7.0 (5.5–8.5)					
7	154 (123–185)	8.6 (6.8–10.3)					
8	183 (147–217)	ABG of 173 mg/dl					
9	212 (170–249)	11.8 (9.4–13.9)					
10	240 (193–282)	13.4 (10.7–15.7)					
11	269 (217–314)	14.9 (12.0–17.5)					
12	298 (240–347)	16.5 (13.3–19.3)					

Data in parentheses are 95% CI. A calculator for converting A1C results into eAG, in either mg/dL or mmol/L, is available at professional.diabetes.org/eAG. \*These estimates are based on ADAG data of  $\sim$ 2,700 glucose measurements over 3 months per A1C measurement in 507 adults with type 1, type 2, or no diabetes. The correlation between A1C and average glucose was 0.92 (6,7). Adapted from Nathan et al. (6).

Inappropriately Low HbA1c	Inappropriately High HbA1c	Variable Effect on HbA1c+
<ul> <li>Hemolysis</li> <li>Certain hemoglobinopathies</li> <li>Recent blood transfusion</li> <li>Acute blood loss</li> <li>Hypertriglyceridemia</li> <li>Drugs*</li> <li>Chronic liver disease</li> </ul>	<ul> <li>Iron deficiency</li> <li>Vitamin B 12 deficiency</li> <li>Alcoholism</li> <li>Uremia</li> <li>Hyperbilirubinemia</li> <li>Drugs*</li> </ul>	<ul> <li>Fetal hemoglobin</li> <li>Methemoglobin</li> <li>Certain hemoglobinopathies</li> </ul>

#### A1C by itself does not tell much...



Laboratory measured A1C lacks: actual Average Blood Glucose, Variability and interference with other conditions

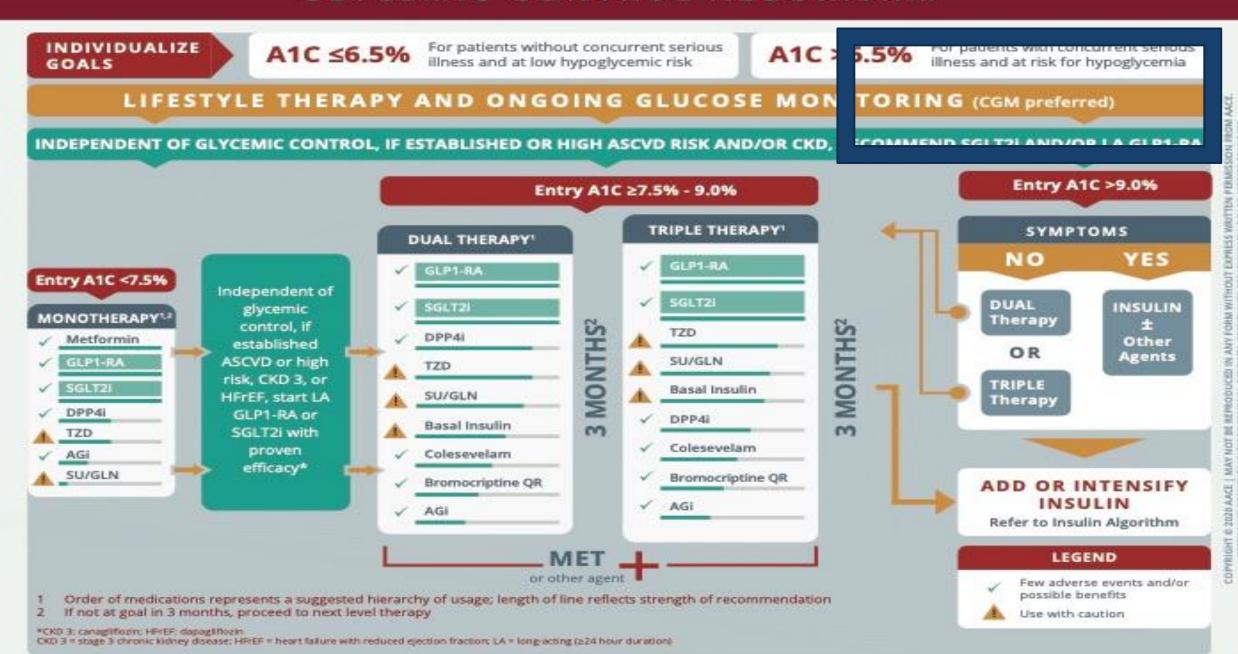
## Candidates for CGM technology

- Patient is on MDI of insulin therapy
- Frequent hypoglycemia, nocturnal hypoglycemias and/or unawareness
- Increased glucose variability (discrepancy between FBG, SMBG, A1C)
- Variations in physical activity
- Willing to use CGM in a daily basis
- Willing to assist to a certified diabetes educator for training
- Children
- Seeking insulin pump therapy or already on it
- To document the need for intensifying therapy

#### 2020 AACE Recommends

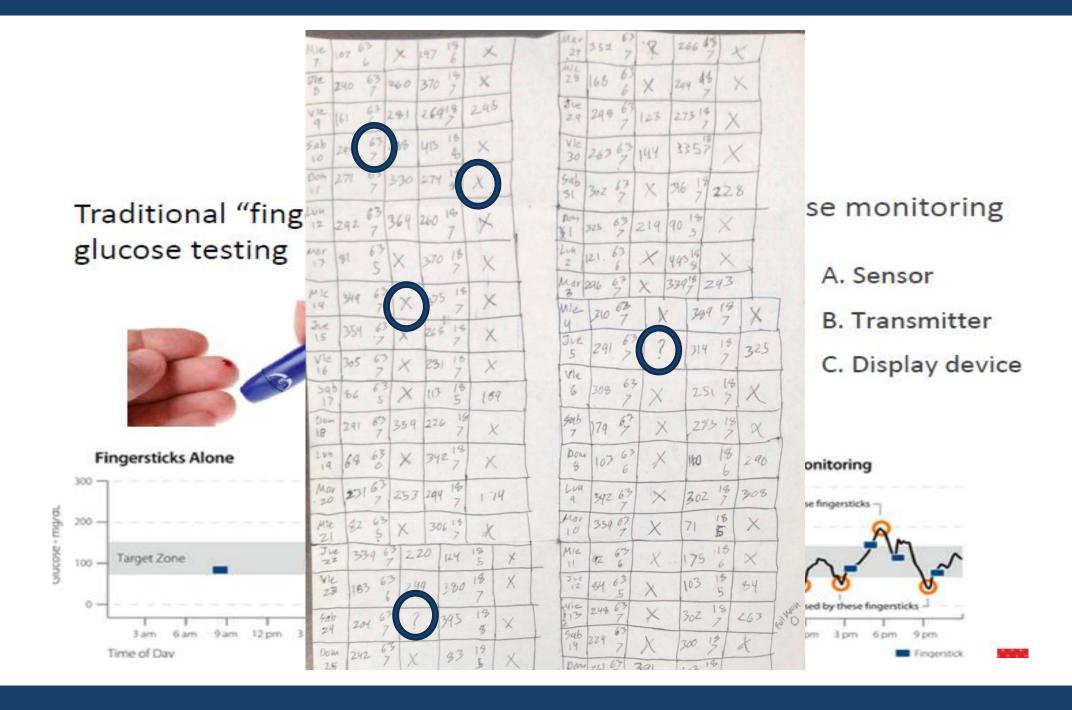
- Professional CGM devices (those owned by the clinician's practice) should be considered in patients:
  - who have not reached their glycemic target after 3 months of the initial antihyperglycemic therapy
  - those who require therapy that is associated with risks of hypoglycemia (SU, insulin)
- Personal CGM devices (those owned by the patient) should be considered for:
  - patients who are on intensive insulin therapy (3 to 4 injections/day or on insulin pump)
  - history of hypoglycemia unawareness
  - recurrent hypoglycemia

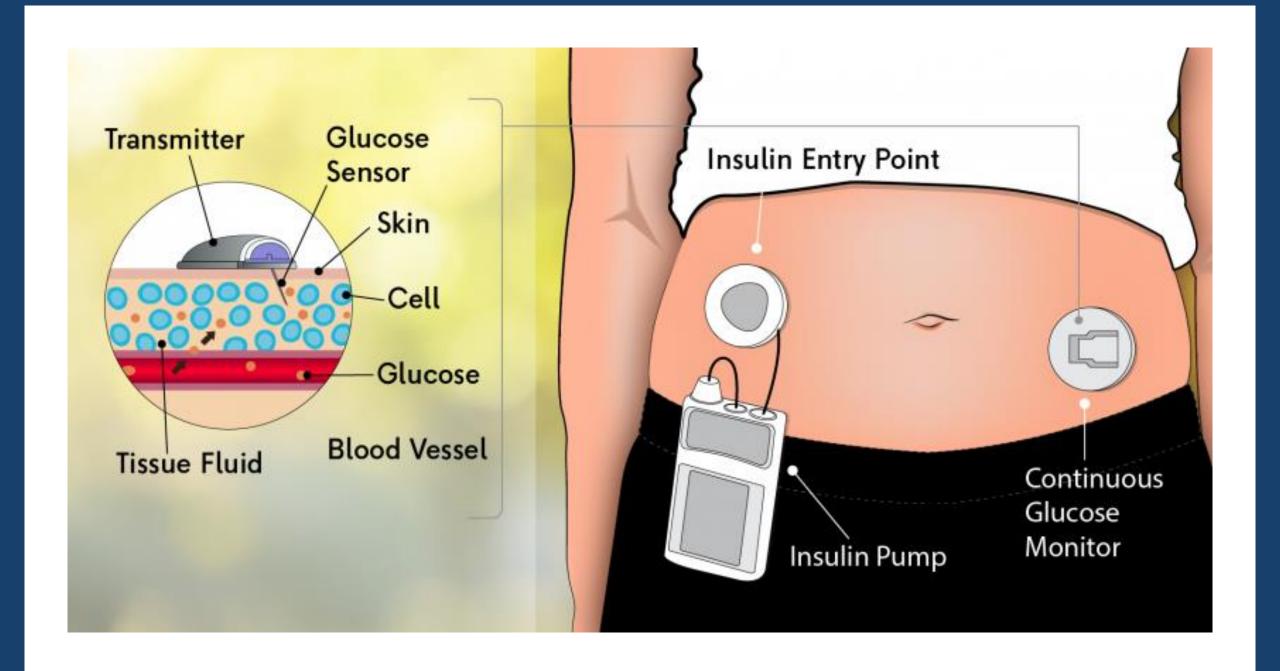
#### **GLYCEMIC CONTROL ALGORITHM**



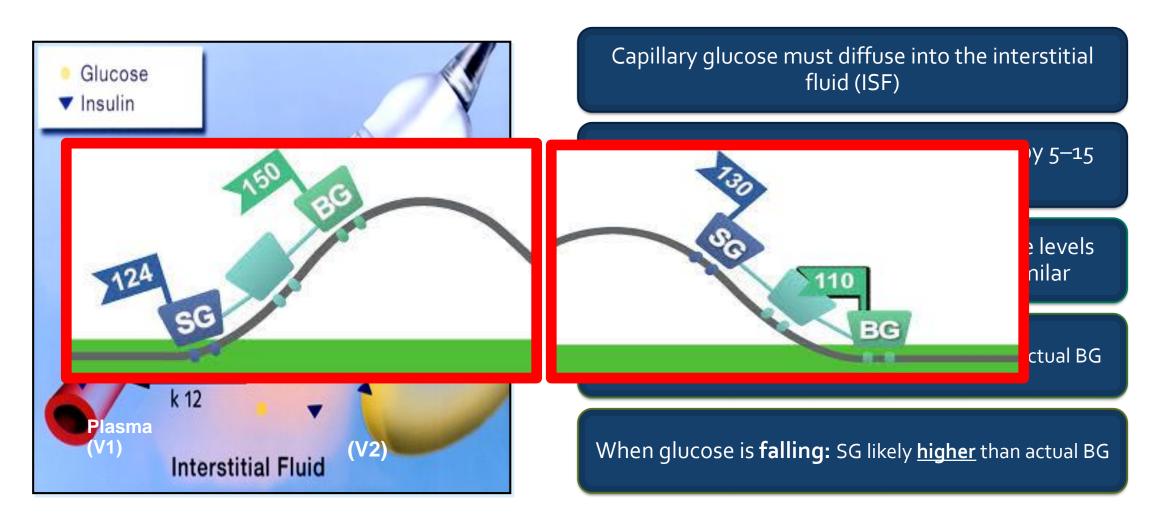
PROGRESSION OF DISEASE

#### HOW CONTINUOUS GLUCOSE MONITORING (CGM) WORKS





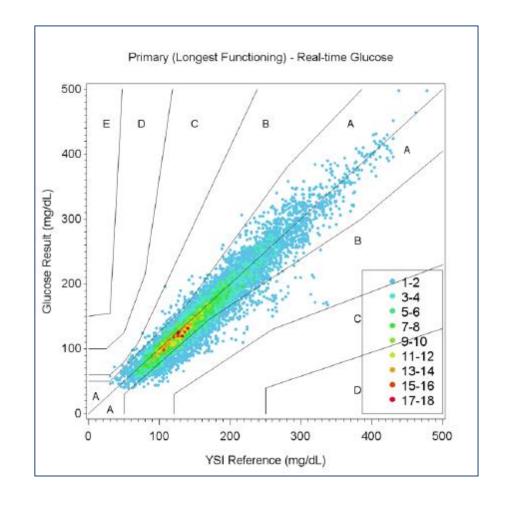
## Interstitial Fluid and "Lag Time"



#### Sensor Accuracy – improving over time

MARD: average difference between sensor glucose and YSI

	MARD
MiniMed iPro retrospective CGM	26%
GlucoWatch	22%
Original Dexcom	21.2%
Medtronic Soft Sensor	18.4%
Dexcom 7 Plus	16%
Medtronic Enlite	15.3%
Dexcom G4	13%
FreeStyle Libre	11.4%
Medtronic Guardian 3	10%
Dexcom G5	9%
Senseonics Eversense	9.4%
Dexcom G6*	9%





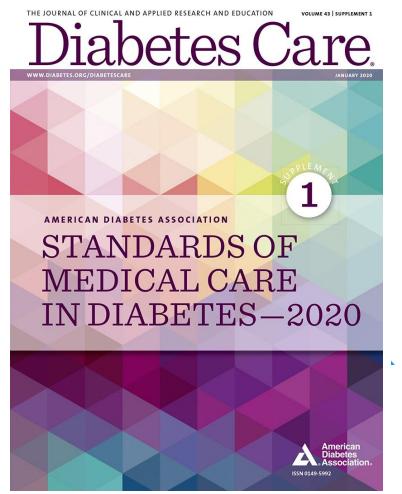
Connected for Life





Clinical Targets for Continuous Glucose Monitoring Data Interpretation: Recommendations From the International Consensus on Time in Range

Diabetes Care 2019;42:1593-1603 | https://doi.org/10.2337/dci19-0028









#### Device-Driven Diabetes Management

## International consensus on *TIME IN RANGE* standardized CGM metrics More Actionable

## Guiding medical nutrition therapy and physical activity Prevent Hypoglycemia

- 1. Number of days CGM device is worn (recommend 14 days)
- 2. Percentage of time CGM device is active (recommend 70% of data from 14 days)
- 3. Mean glucose
- 4. Glucose management indicator (GMI)
- Glycemic variability (%CV) target ≤36%\*
- 6. Time above range (TAR): % of readings and time >250 mg/dL (>13.9 mmol/L)
- 7. Time above range (TAR): % of readings and time 181-250 mg/dL (10.1-13.9 mmol/L)
- 8. Time in range (TIR): % of readings and time 70-180 mg/dL (3.9-10.0 mmol/L)
- 9. Time below range (TBR): % of readings and time 54-69 mg/dL (3.0-3.8 mmol/L)
- 10. Time below range (TBR): % of readings and time <54 mg/dL (<3.0 mmol/L)

CGM, continuous glucose monitoring; CV, coefficient of variation. \*Some studies suggest that lower %CV targets (<33%) provide additional protection against hypoglycemia for those receiving insulin or sulfonylureas. Adapted from Battelino et al. (17).

Level 2

Level 1

Level 1

Level 2

In range

#### AGP (Ambulatory Glucose Profile)

#### **AGP** Report

#### **GLUCOSE STATISTICS AND TARGETS**

26 Feb 2019–10 Mar 2019 13 days % Time CGM is Active 99.9%

Targets [% of Readings (Time/Day)]
Target Range 70–180 mg/dL .......Greater than 70% (16h 48min)
Below 70 mg/dL ......Less than 4% (58min)
Below 54 mg/dL .....Less than 1% (14min)
Above 180 mg/dL .....Less than 25% (6h)
Above 250 mg/dL .....Less than 5% (1h 12min)

Each 5% increase in time in range (70-180 mg/dL) is clinically beneficial.

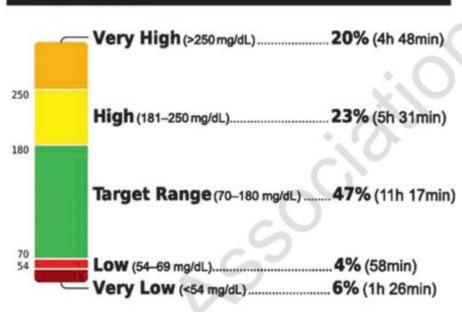
Average Glucose 173 mg/dL Glucose Management Indicator (GMI) 7.6% 49.5%

Defined as percent coefficient of variation (%CV); target ≤36%

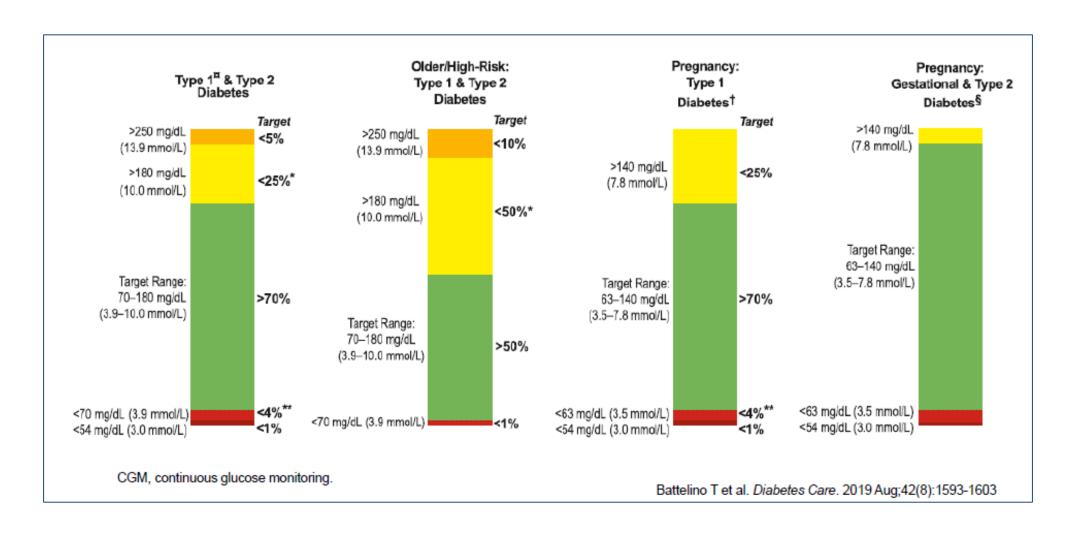
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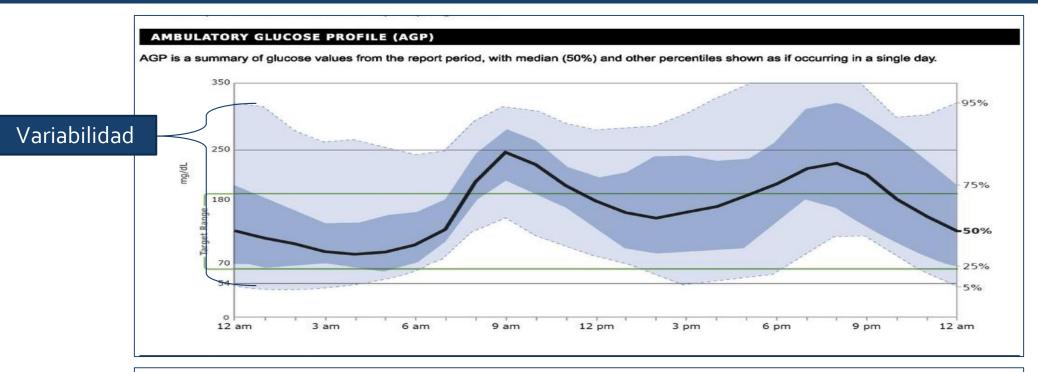
MRN

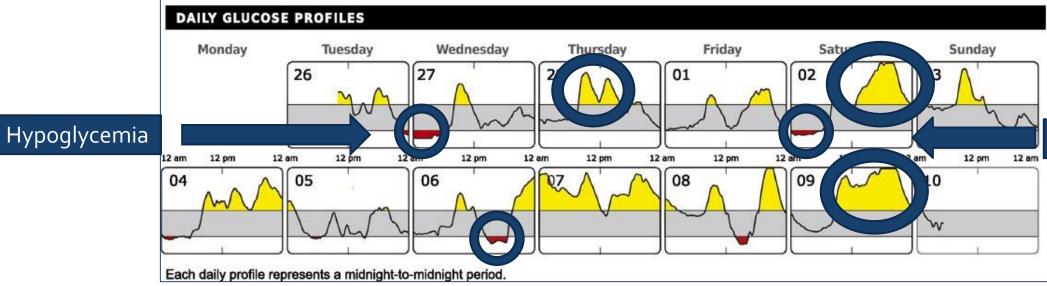
#### TIME IN RANGES



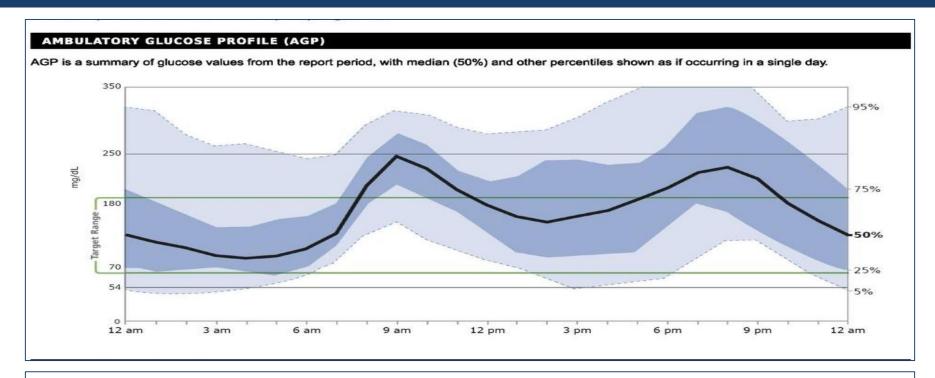
#### We may personalize individual glycemic targets

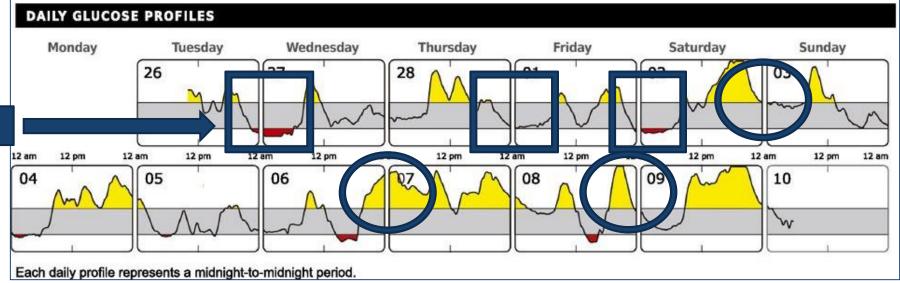






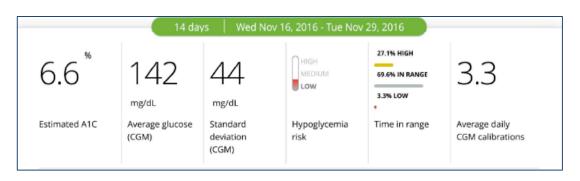
Hyperglycemia

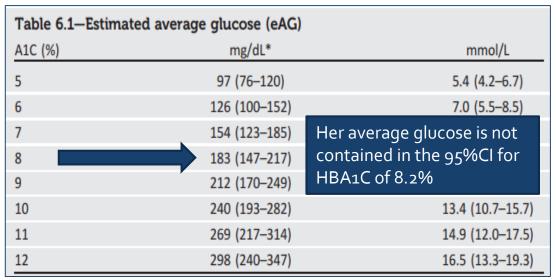




Trends

#### Glucose Management Indicator (GMI) aka Real time A1C





- 25 y/o using Dexcom CGM with Hba1c of 8.2%, Hb of 10 g/dl, Hct 31%
- Patient note with glycation gap due to iron deficiency anemia
- Up to 15% of glycated hemoglobin may be misleading in clinical practice
- GMI lower that lab A1C, means glucose is lower that reported by lab and caution is needed in order to avoid hypoglycemia
- Plan: iron supplements to correct anemia, NO INSULIN optimization

## Benefits of Time in Range

Table 5—Estimate of A1C for a given TIR level based on type 1 diabetes and type 2 diabetes studies

Beck et al. (26) (n = 545 participants with type 1 diabetes)			participants with type 1 or type 2 diabetes)	
TIR 70–180 mg/dL (3.9–10.0 mmol/L)	A1C, % (mmol/mol)	95% CI for predicted A1C values, %	TIR 70-180 mg/dL (3.9-10.0 mmol/L)	A1C, % (mmol/mol)
20%	9.4 (79)	(8.0, 10.7)	20%	10.6 (92)
30%	8.9 (74)	(7.6, 10.2)	30%	9.8 (84)
40%	8.4 (68)	(7.1, 9.7)	40%	9.0 (75)
50%	7.9 (63)	(6.6, 9.2)	50%	8.3 (67)
60%	7.4 (57)	(6.1, 8.8)	60%	7.5 (59)
70%	7.0 (53)	(5.6, 8.3)	70%	6.7 (50)
80%	6.5 (48)	(5.2, 7.8)	80%	5.9 (42)
00%	6.0 (42)	(47.72)	00%	E 1 /22\

Every 10% increase in TIR =  $\sim$ 0.5% (5.5 mmol/mol) A1C reduction

Every 10% increase in TIR =  $\sim$ 0.8% (8.7 mmol/mol) A1C reduction

The difference between findings from the two studies likely stems from differences in number of studies analyzed and subjects included (RCTs with subjects with type 1 diabetes vs. RCTs with subjects with type 2 diabetes with CGM and SMBG).

Vigorely, and McNahan (27) /n - 1 127

## Ambulatory Glucose Profile works if:

- Data can be effectively downloaded for each patient visit
  - Limitations in software, hardware, TIME! or even having the device present
- Data can be effectively interpreted
  - Assess patent routine, wake and sleep times, feedings, medication intake
  - Ask the patient to remember exactly what he was doing for a specific hypoglycemia event in that 14 day log
  - Discuss times of the day where more glucose fluctuations (variability) exist
- Physician and Patient design a course of action (simple solutions are always better!)
  - Correct hypo/hyperglycemia management
  - Teach the patient to carb counting
  - Teach the patient to calculate Sensitivity Factors for corrections, Carbs Ratio, Active insulin time, ect
  - Exercise activity and glucose changes

#### Reimbursement

- 95249 CGM patient provided equipment, sensor placement, hook-up, calibration of monitor, patient training, and printout
- 95250 CGM HCP (office) provided equipment, sensor placement, hook-up, calibration of monitor, patient training, removal of sensor, and printout
- 95251 CGM analysis, interpretation and report
  - Can be billed monthly on ongoing basis
- General comments:
  - All codes require a minimum of 72 hours of data
  - Use -25 modifier for CGM codes if billing same day as a Problem Visit code (99212-99215) if significant an separately identifiable service took place
    - le 99212-99215: Pre-CGM evaluation (+) -25 95250: CGM start-up and instruction

# CGM'S CURRENTLY AVAILABLE IN PR

#### Types of CGM

- Real Time CGM
  - CGM systems that measure glucose levels continuously and provide the user automated alarms and alerts at specific glucose levels and/or for changing glucose levels.
- Intermittent Scanning CGM
  - CGM systems that measure glucose levels continuously but only display glucose values when swiped by a reader or a smart phone that reveals the glucose levels
- Blinded (professional) CGM
  - CGM devices that measure glucose levels that are not displayed to the patient in real time. These devices are generally initiated in a clinic, using a reader that is owned by the clinic. They are removed after a period of time (generally 10–14 days) and analyzed by the patient and provider to assess glycemic patterns and trends

## Abbot FreeStyle Libre

- Intermittent Scanning CGM
- Professional (Blinded) CGM available
- Factory Calibrated
- 14-day wear following 12-hr warm-up
- For best results need to read at least q8 hours
- No acetaminophen interference



#### **Daily Patterns**

a graph showing the pattern and variability of your Sensor glucose over a typical day.



#### Time in Target

displays percentage of time your sensor glucose readings were within, above or below your Target Glucose Range



#### Low Glucose Events

shows readings lower than 70 mg/dL (for a minimum of 15 minutes) in four different 6-hour periods of the day









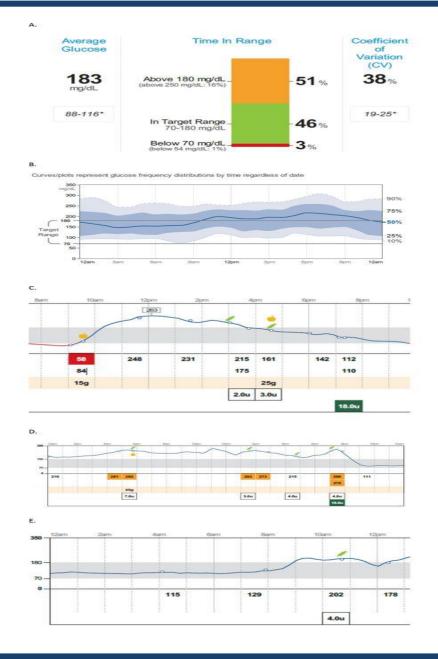




User only gets a glucose reading when they wave the receiver device over the sensor which is unique to this CGM system. 10:23pm 🖭 © Ends in 13 days The reader is 95mm x 60mm x 16mm and store No calibration needed; preup to 90 days of glucose calibrated at the factory. readings. 250 150 Can be used with or without and insulin pump. FreeStyle Libre 14 day Lowest cost of any CGM on the market. No automatic alerts for high/low 14 day sensor, the longest lasting glucose levels when user doesn't wave sensor in the CGM industry. the receiver over the sensor.

#### Freestyle Libre Report:





☐ Gluc

MEDIAN G

Settings

Legend



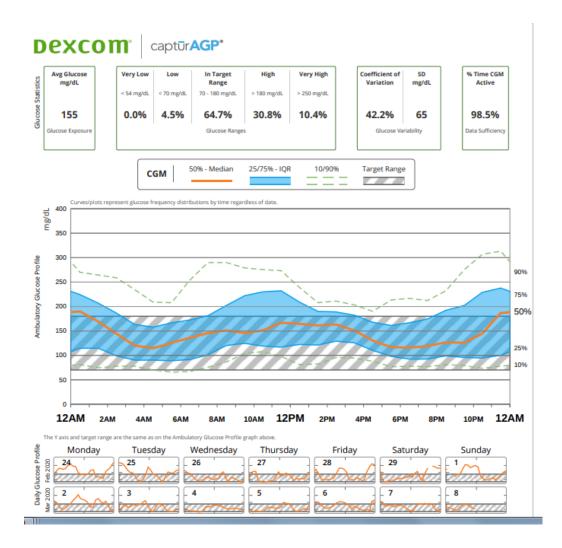
#### Dexcom G6

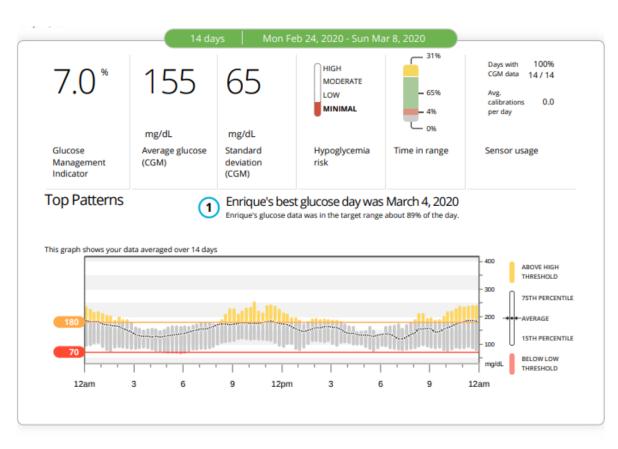
- Real Time, Stand Alone CGM
- No confirmatory fingersticks unless
  - Discrepancy with symptoms and readings
  - Device malfunction
- No Calibrations Needed
- Remote monitoring
- Alerts available
- 10 days sensor wear





# **Dexcom Clarity**







#### Medtronic 670G



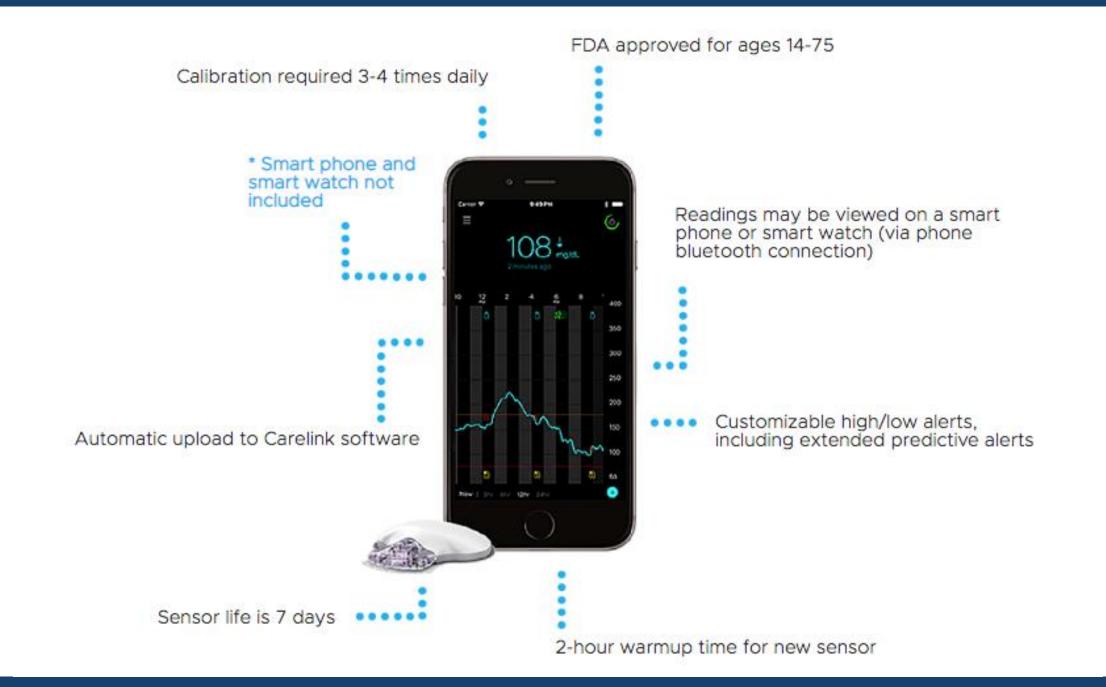
## Medtronic



1081

- Key unique features
  - First hybrid closed loop system
    - Modulates insulin infusion based on sensor glucose information
  - Predictive Low Glucose Suspend and Low Glucose Suspend
  - Calibration at least once every 12 hours (3-4x/day recommended)
  - 7 day sensor use
  - Acetaminophen sensitive
  - Approved for 14 years and older

- Stand-alone CGM system
- Key unique features
  - No receiver; display device is iOS phone only
  - Predictive Alert Schedules
    - 10-60 minute prediction of hypo-/ hyperglycemia based on threshold settings
  - Calibration at least once every 12 hours (4x/day recommended)
  - 7 day sensor
  - Acetaminophen sensitive
  - Approved for 18 and older



## Medtronic Carelink



	Guardian Connect/640G SmartGuard	Dexcom G6	FreeStyle Libre
	108 =		6.22
Sensor life (days)	7	10	14
Alarms	Multiple	High, low and trend	None
Predictive	Yes	Yes	n/a
Trends	Yes	Yes	n/a
Rate change	Yes	Yes	n/a
Calibration	12-hourly	None	None
MARD	9.64%	9.0%	9.7%

Data taken from manufacturers' specifications

<sup>\*</sup>The manufacturer claims a sensor life up to 14 days; however, the device is only licesnsed for up to 7 days. MARD=mean absolute relative difference.

# CLINICAL EXAMPLES

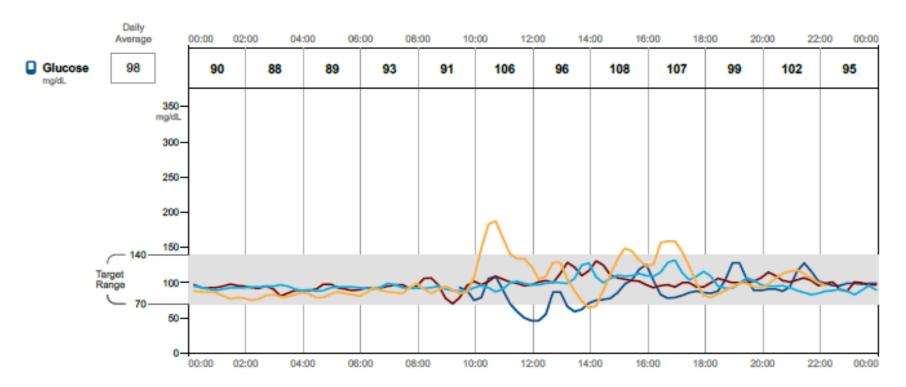
Patients with DM 1-2 and Insulin Pump Therapy

# CGM on non-diabetic patient

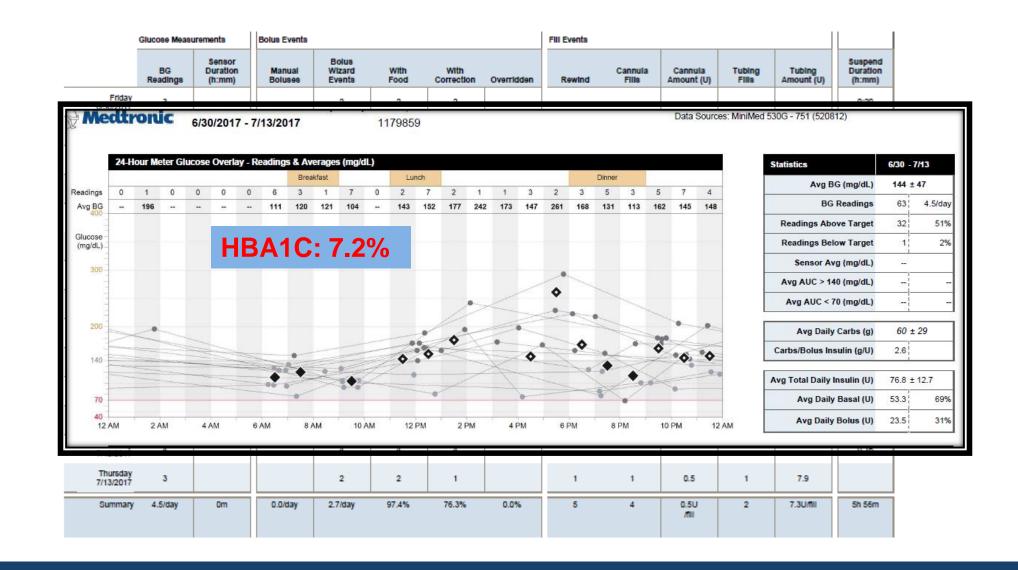
#### Daily Patterns (with glucose readings)

FreeStyle Libre Pro

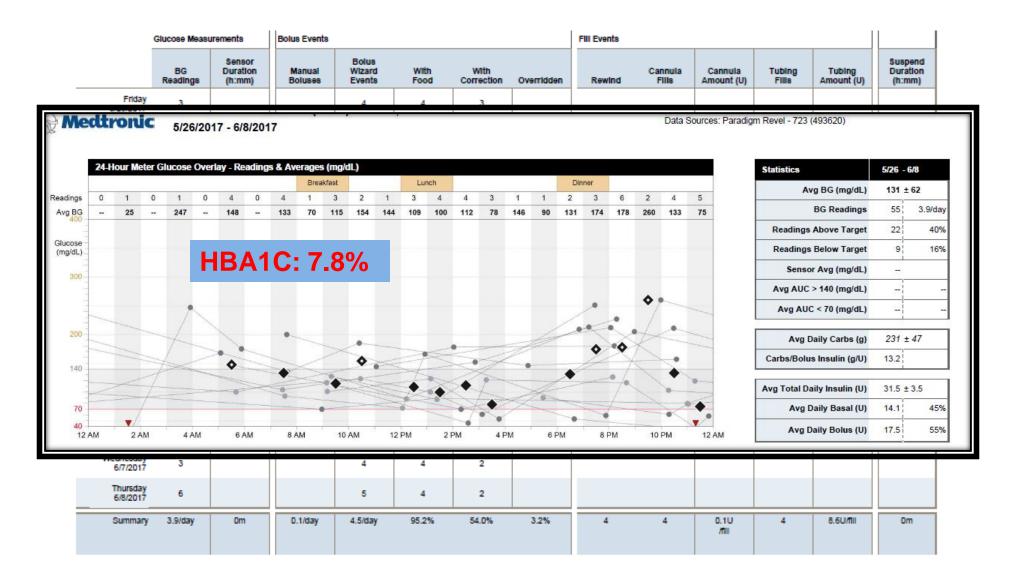
24 July 2017 - 27 July 2017 (4 days)



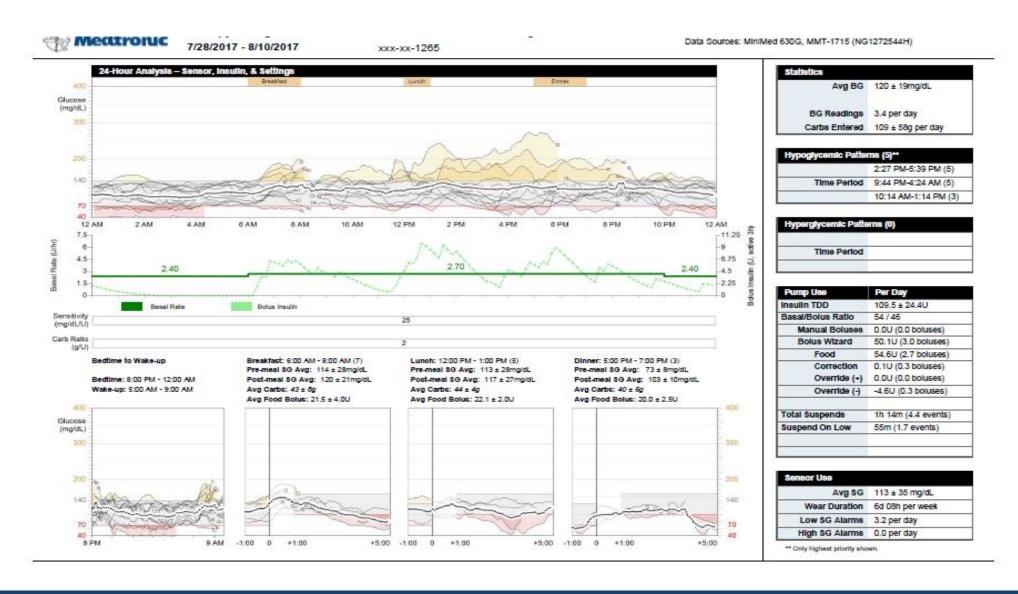
### 59 y/o F, BMI: 29 kg/m2, HBP, no CGM

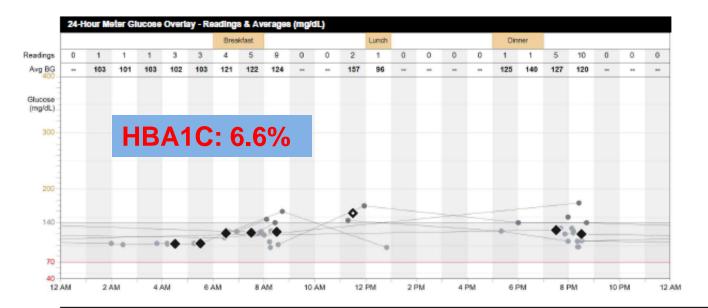


### 55 y/o F, BMI: 23.5 kg/m2, HBP, no CGM



### 61 y/o F, BMI: 38 kg/m2, HBP, Sensor Augmented Tx





Statistics	7/28	-8/10
Avg BG (mg/dL)	120	± 19
BG Readings	47	3.4/day
Readings Above Target	6	13%
Readings Below Target	-	0%
Sensor Avg (mg/dL)	113	±35
Avg AUC > 140 (mg/dL)	4.8	12d 17h
Avg AUC < 70 (mg/dL)	1.2	12d 17h

Avg Dally Carbs (g)	109 ± 58
Carbs/Bolus Insulin (g/U)	2.2

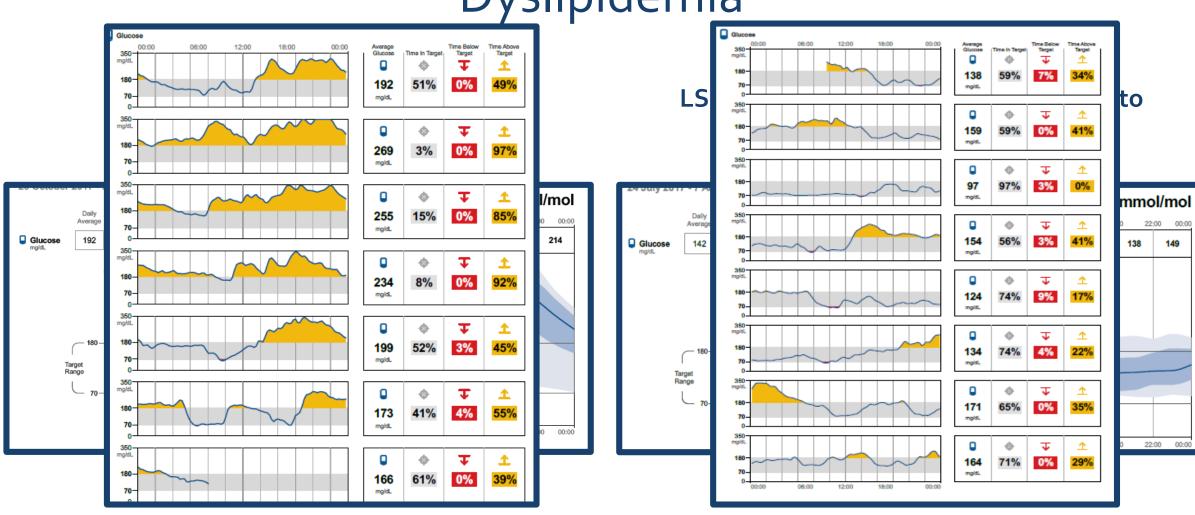
Avg Total Dally Insulin (U)	109.45	± 24.4
Avg Daily Basal (U)	59.33	54%
Avg Dally Bolus (U)	50.13	46%

Basal					
Maximu	ım Basal R	ate 4.00 U/Hr			
Basal 1 (a	active)	Day Off		Workday	
24-Hour Total	62.400 U	24-Hour Total		24-Hour Total	-
Time	U/Hr	Time	U/Hr	Time	U/Hr
0:00	2.40	-	-	-	-
6:00	2.70				
22:00	2.40				

Dolu	•								
В	olus Wizard	On		Π [		Eas	y Bolus	Off	$\neg \neg$
	Units		g, mg/dL		Bolus Increment		crement	0.025 U	
Active	Active Insulin Time (h:mm)		3:00		Bolus Speed		Standard		
Max	lmum Bolus	25.0	25.0 U Dual/Square		Off/Off				
Carbohydrate Rat (g/U)		io	Insulin (mg/dL				Blood ( Target	Glucos (mg/dL	_
Time	Ratio		Time	Sen	sitivity		Time	Low	High
0:00	2.0		0:00		25		0:00	80	120

		Sensor	On	
High A	lerts		On (	Snooze 1:00)
Start Time	High (mg/dL)	Alert On High	Alert Before High	Rise Alert Limit (mg/dL)
0:00	250			
				<del>                                     </del>

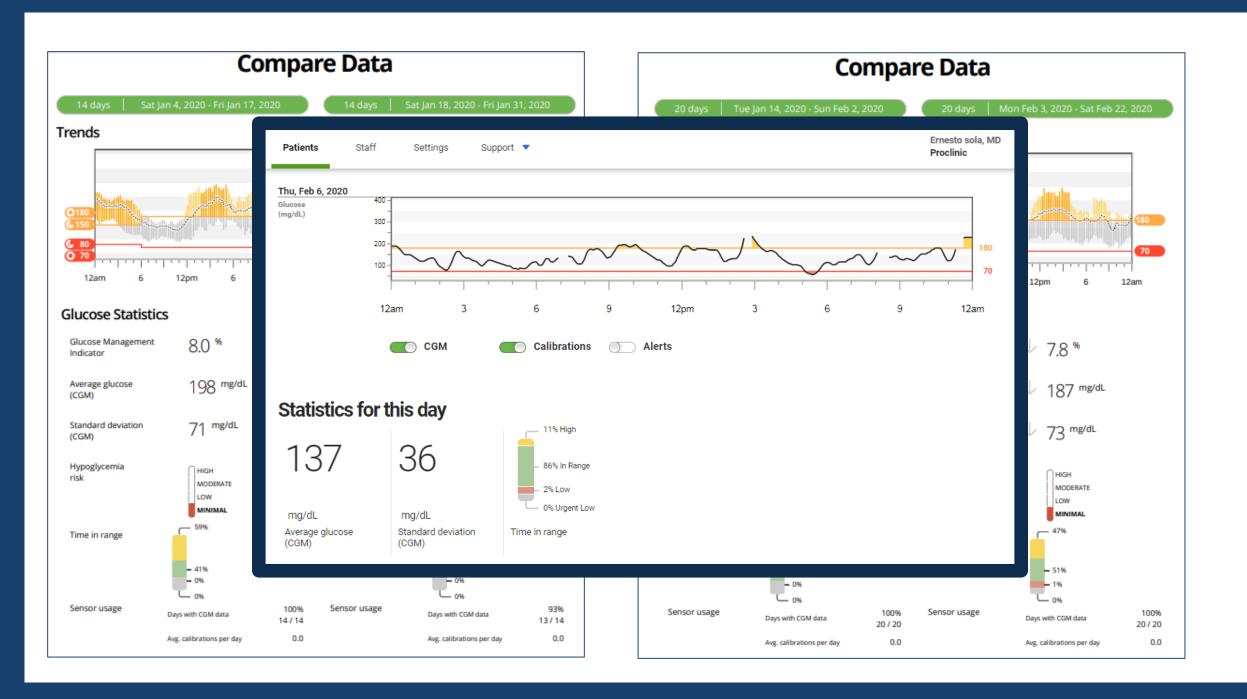
56 y/o DM2 A1C 8.9% CAD, obesity, HBP, Dyslipidemia



61 F DM1 x 45 years: HbA1c: 11% – Happy to learn that will not perform finger sticks anymore!







# There is no age limit for technology!!

- 82 y/o male
- Long standing DM2, low C-peptide
- On basal bolus
- Hx of recurrent severe hypoglycemia with 1-2 glucagon emergency kits per MONTH!
- rtCGM use, eliminated 100% hypoglycemia, A1C unchanged



# THANKS!!