

DISCLOSURES I have nothing to disclose concerning possible financial relationships with ineligible companies that may have a direct or indirect interest in the subject matter of this presentation.

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LEARNING OBJECTIVES Review the pharmacokinetics, efficacy, and safety considerations of diabetes medications, including glucagon-like peptide-1 (GLP-1) receptor agonists and sodium glucose cotransporter 2 (SGLT2) inhibitors. Explain the fundamental concepts of continuous glucose monitor (CGM) technology and interpretation of data. Assess the evidence supporting CGM use in diabetes. Discuss the role of pharmacists providing diabetes care within a multidisciplinary

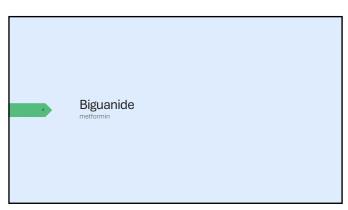
THE EVOLUTION OF DIABETES PHARMACOTHERAPY Concern for adverse events
 Required CV trials Cardiovascular Benefits

Major Adverse Cardiovascular Events (MACE)

Heart Failure Chronic Kidney Disease (CKD) Obesity Metabolic Dysfunction-Associated Steatotic Liver Disease (MASLD) **Liver Benefits** Metabolic Dysfunction-Associated Steatohepatitis (MASH)

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WHAT WE WILL NOT COVER TODAY: Insulin Sulfonylureas Dipeptidyl Thiazolidinediones peptidase-4 inhibitors (DDP-4 (TZDs) inhibitors)



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METFORMIN

- Mechanism: decreases hepatic glucose production, improves insulin sensitivity, thought to provide appetite regulation
- FDA indications: diabetes type 2
- Off label: prevention of diabetes type 2; gestational diabetes; polycystic ovarian syndrome, antipsychotic induced weight gain

exicomp. Metformin. Lexi-Drugs. Hudson, OH: Wolters Kluwer Clinical Drug Information, Inc.; (updated 2025 Mar 03; cited 2025 Mar 03). Available from

METFORMIN ADVERSE EFFECTS

- · Common side effects:
 - GI: nausea, vomiting, diarrhea, flatulence
- · Severe side effects (SAE):
 - · Lactic acidosis
 - Vitamin B12 deficiency (monitor every 2-3 years)



Lexicomp. Metformin. Lexi-Drugs. Hudson, OH: Wolters Kluwer Clinical Drug Information, Inc.; [updated 2025 Mar 03; cited 2025 Mar 03]. Available from:

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METFORMIN DOSING CONSIDERATIONS

Starting dose 500 mg daily; increase by **500 mg** per day every week as tolerated

Maximum: 2000-2500 mg total daily dose

Extended release better tolerated \rightarrow improved adherence

Counsel to take with food to minimize side effects

Avoid extended release if history of bariatric surgery; immediate release tablets can be cut/crushed

oxicomp. Metformin. Lexi-Drugs. Hudson, OH: Wolters Kluwer Clinical Drug Information, Inc.; [updated 2025 Mar 03; cited 2025 Mar 03]. Available from:

ADDITIONAL CLINICAL CONSIDERATIONS: METFORMIN

Contraindications:

- GFR < 30 ml/min (can use if GFR 30-44 ml/min if already taking but requires 50% dose reduction)
- Metabolic acidosis including lactic acidosis and diabetic ketoacidosis and associated risk factors (excessive alcohol intake)
- Severe liver disease or heart disease (caution)

Safe in pregnancy

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Lexicomp. Metformin. Lexi-Drugs. Hudson, OH: Wolters Kluwer Clinical Drug Information, Inc.; [updated 2025 Mar 03; cited 2025 Mar 03]. Available from:

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Sodium glucose cotransporter –2 inhibitors (SGLT2 inhibitors)

bexagliflozin, canagliflozin, dapagliflozin, empagliflozin, ertugliflozin, sotagliflozin

SGLT-2 INHIBITORS

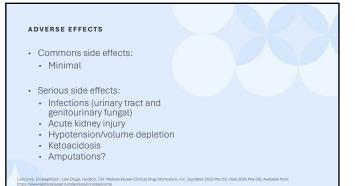
- Agents: bexagliflozin, canagliflozin, dapagliflozin, empagliflozin, ertugliflozin, sotagliflozin
- Mechanism: decreases reabsorption of glucose in the proximal tubules by inhibiting sodiumglucose cotransporter 2 → increased glucose excretion

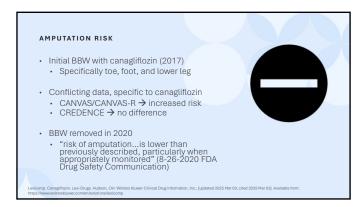


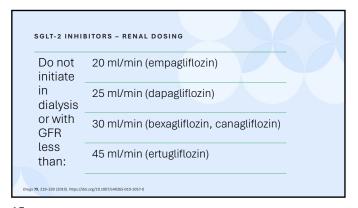
FDA indications: Diabetes type 2, chronic kidney disease, heart failure

Lexicomp. Empagliflozin. Lexi-Drugs. Hudson, OH: Wolters Kluwer Clinical Drug Information, Inc.; [updated 2025 Mar 03; cited 2025 Mar 03]. Available from:

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CLINICAL CONSIDERATIONS FOR SGLT-2
INHIBITORS

Contraindications:

DM type 1
Any history of ketoacidosis
Pregnancy

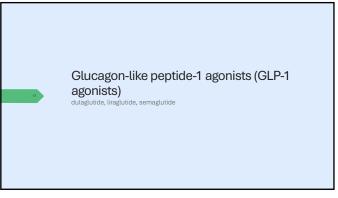
Euglycemia

Must be able to maintain adequate PO intake

Weight loss
Variable, average 1-3 kg

\$\$\$ brand only, expensive

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Mechanism: GLP-1 is an incretin hormone
 → glucose dependent insulin secretion,
 decreases inappropriate glucagon secretion,
 increases B-cell growth, slows gastric
 emptying, decreases food intake, improves
 insulin sensitivity
 FDA indications: Diabetes type 2, weight
 management – chronic

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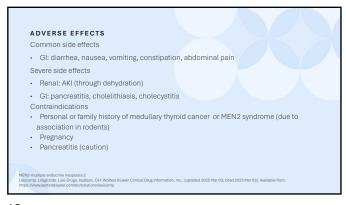
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DPP-4 inhibitors

Pancreatitis risk
No increased efficacy
AVOID combination

Insulins/secretagogues
Hypoglycemia risk
Use with caution
Monitor and adjust doses

Lexicomp. Lingluidie. Levi-Drugs. Huddon, ON: Wilhers Klaver Clinical Drug Information, Inc.; [updated 2025 Mar OI], Available from: https://www.woothersklaver.com/en/beldicinor/lexicomp

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Liraglutide (Victoza®, Saxenda®)
^{2010, 2014}

LIRAGLUTIDE AND CARDIOVASCULAR
OUTCOMES IN TYPE 2 DIABETES
(LEADER)

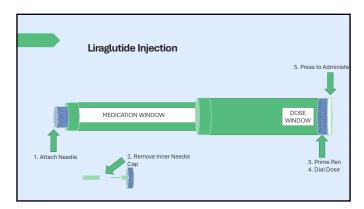
- Study Design: Multicenter, double-blind, randomized, placebo-controlled trial
- Participants: 9,340 adults with type 2 diabetes and established cardiovascular disease or high cardiovascular risk
- Duration: Median follow-up 3.8 years
- Intervention: Liragllutide (up to 1.8 mg daily) vs. placebo
- Primary Endpoint: Composite major adverse cardiovascular events (MACE):
- Cardiovascular death
- Nonfatal trnyocardial infarction (MI)
- Nonfatal stroke
- Key Findings:
- Significant reduction in MACE (HR: 0.87, 95% CI: 0.78–0.97, p=0.01)
- Lower cardiovascular mortality (HR: 0.78, p=0.007)
- No increased risk of severe hypoglycemia or pancreatic adverse events

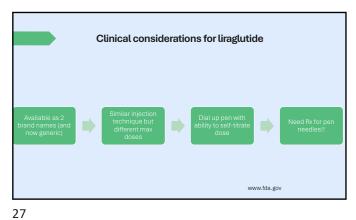
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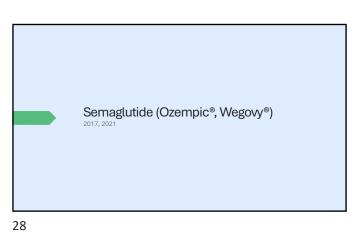
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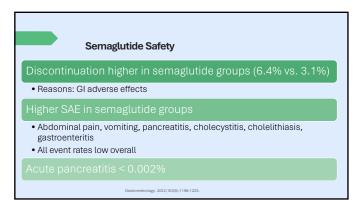


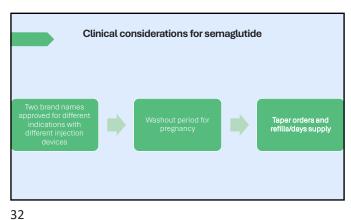


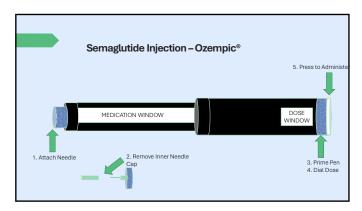


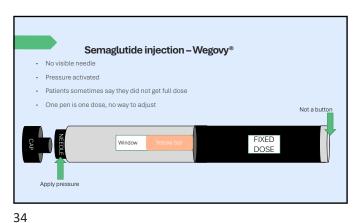
Semaglutide Literature – Weight Loss Trials included: 8 RCTs, n=2658 in treatment groups Mean age: 46-59.5 years Predominantly female Lifestyle: most included 500 kcal/d deficit						
Mean Starting Weight	Mean Starting BMI	Mean Starting Waist Circumference	Comorbidities Included	Follow Up Duration		
86.9-113.2 kg	32-39.9kg/m ²	103.8-119 cm	4 excluded DM, 3 mixed, 1 required DM	52-72 weeks		
	Gestroenterolog	(): 2022;163(5):1198-1225.				

% TBWL*	Weight loss*	Percent achieving 5% TBWL*	Percent achieving 10% TBWL*	Percent achieving 15% TBWL*
10.76%	10.8 kg	51.7%	52.6%	40.7%
	nean difference o	compared to plad	ceno group	

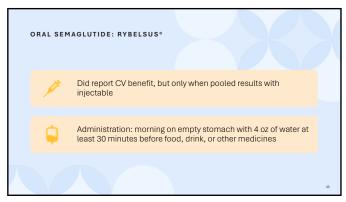






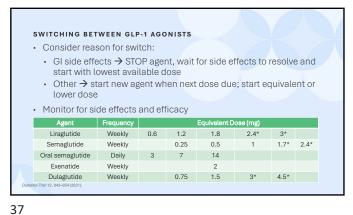


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GLP-1 and GIP Dual **Agonist** TIRZEPATIDE (MOUNJARO®, ZEPBOUND®)

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TIRZEPATIDE · Mechanism: • Dual glucose-dependent insulinotropic polypeptide (GIP) and GLP-1 agonist GIP – intestinal hormone, stimulates glucose dependent insulin secretion postprandially, stimulated glucagon secretion in periods of eu- and hypoglycemia • FDA indication: Diabetes type 2; weight management, chronic · Contraindications, cautions, side effects: same as GLP-1 agonists

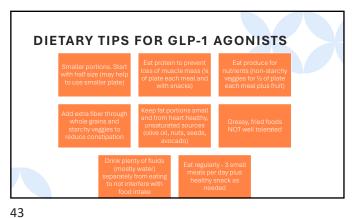
Tirzepatide Literature - weight loss • n=3964 in treatment group · Mean age: 44.9 years Predominantly female · Lifestyle: calorie deficit Mean Starting | Mean Starting 10.8.8 kg 38 kg/m² 114.1 cm Excluded DM 72 weeks

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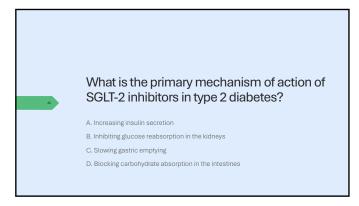
Tirzepatide Outcomes Dose 5 mg 11.9% 50% 49.7% 10 mg 16.4% 54% 59.3% 47% 17.8% 56% 54% *Reported as mean difference compared to placebo group · Improvements in all prespecified cardiometabolic measures with tirzepatide

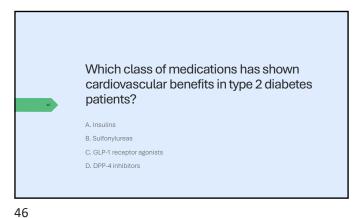
Tirzepatide Safety • Reasons: mild-moderate GI adverse effects No difference in SAF

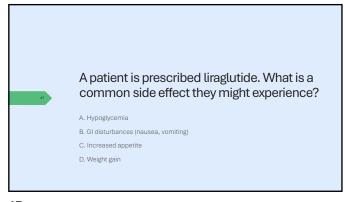
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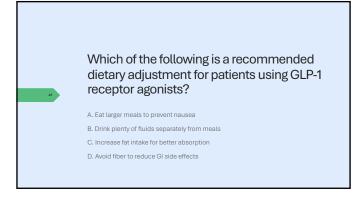


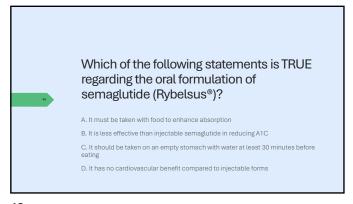


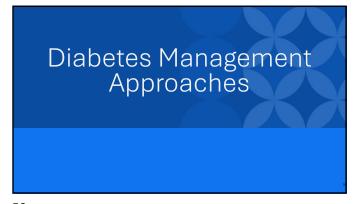


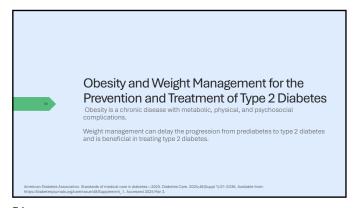


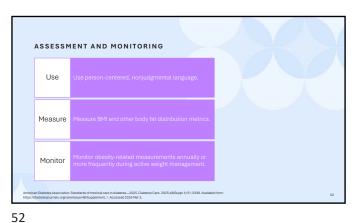






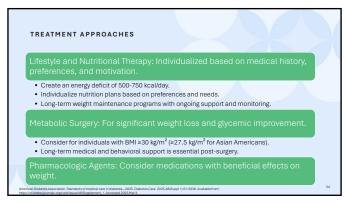




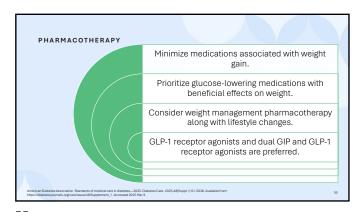


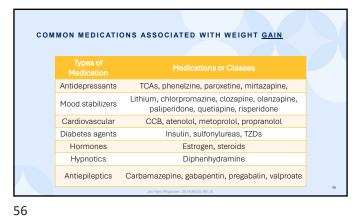
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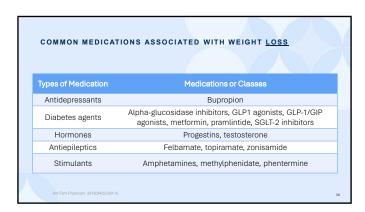


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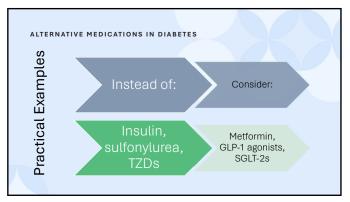


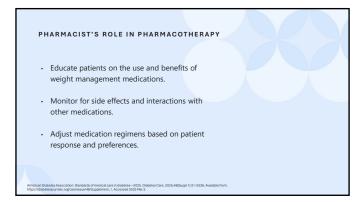


COMMON MEDICAT	IONS THAT ARE WEIGHT <u>NEUTRAL</u>	
Types of Medication	Medications or Classes	
Antidepressants	Citalopram, desvenlafaxine, duloxetine, escitalopram, fluoxetine, sertraline, venlafaxine	
Antipsychotics	Aripiprazole, haloperidol, ziprasidone	
Cardiovascular	ACE inhibitors	
Diabetes agents	DPP4 inhibitors	
Hypnotics	Benzodiazepines, trazodone	
Mood stabilizers/ antiepileptics	Oxcarbazepine, lamotrigine, levetiracetam, phenytoin	
	Am Fam Physician. 2016;94(5):361-8.	

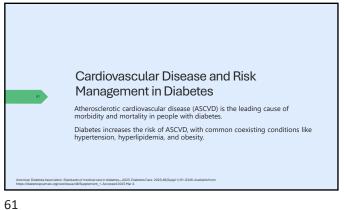


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LIPID MANAGEMENT · High-intensity statin therapy is recommended for individuals aged 40-75 years with diabetes and high cardiovascular risk. - Target LDL cholesterol reduction by \geq 50% of baseline and aim for LDL <70 mg/dL (<1.8

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ANTIPLATELET THERAPY

- · Aspirin (75-162 mg/day) is recommended for individuals with diabetes and a history of ASCVD. (SECONDARY PREVENTION)
- · Consider aspirin therapy for primary prevention in individuals at high cardiovascular risk after discussing potential benefits and risks.

HEART FAILURE MANAGEMENT

- · Heart failure is a major cause of morbidity and mortality in people with diabetes.
- · Use SGLT2 inhibitors to reduce the risk of heart failure hospitalization.

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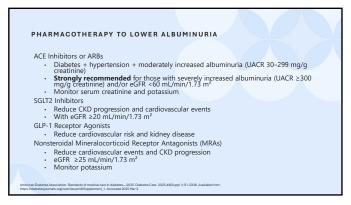
SCREENING RECOMMENDATIONS

- Assess kidney function (urine albumin creatinine ratio (UACR) and eGFR) in type 1 diabetes (duration ≥5 years) and all type 2 diabetes patients.
- · Monitor urinary albumin and eGFR 1-4 times per year in established CKD.

Chronic Kidney Disease and Risk Management in Diabetes Chronic Kidney Disease (CKD) is a common complication in diabetes, affecting 20-40% of people with diabetes.

CKD increases the risk of cardiovascular disease and healthcare costs

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ADDITIONAL CONSIDERATIONS TO LOWER ALBUMINURIA

Effective glucose management

Optimize Blood Pressure Management

• Blood pressure target of <130/80 mmHg

Dietary Protein Intake

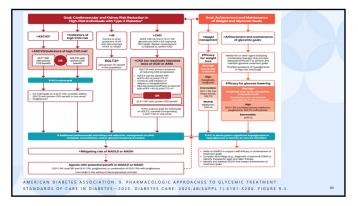
- non-dialysis stage G3 or higher CKD → 0.8 g/kg body weight/day
- Dialysis →1.0–1.2 g/kg/day

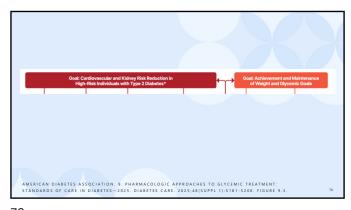
Lifestyle Modifications Smoking cessation

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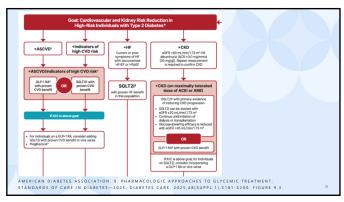
- Weight loss
- Changes in eating patterns (e.g., decreased salt and protein intake)

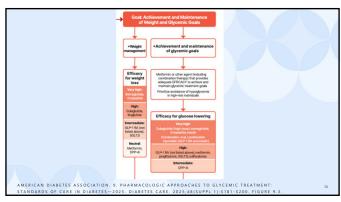
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What is a key potential benefit of GLP-1 receptor agonists in patients with type 2 diabetes beyond glycemic control? в. Weight gain c. Cardiovascular risk reduction D. Increased hepatic glucose production

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Which class of diabetes medications has been shown to provide cardiovascular benefits, including reduced risk of major adverse cardiovascular events in patients with type 2 diabetes and established cardiovascular disease? B. Dipeptidyl peptidase-4 (DPP-4) inhibitors c. Sodium-glucose cotransporter-2 (SGLT2) inhibitors D. Insulin

What class of medication has NOT been shown to help reduce albuminuria? Nonsteroidal Mineralocorticoid Receptor Antagonists (MRAs) B. ACE Inhibitors c. Sulfonvlureas D. GLP-1 Receptor Agonists E. GLP-1 Receptor Agonists

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WHAT ARE CGMS?

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- · Thin filament inserted under skin that connects to a small sensor and wirelessly transmits to a reader/smartphone
- · Personal (patient monitored)
 - · Can upload device to computer
 - · Can remotely transmit data to provider practice
- · Professional (provider monitored)
- · Real time or intermittent scanning

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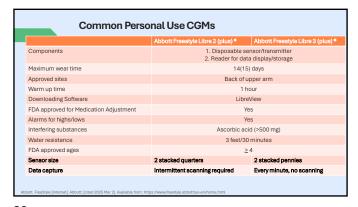
Select CGM brands

Abbott Freestyle Libre®

Dexcom®

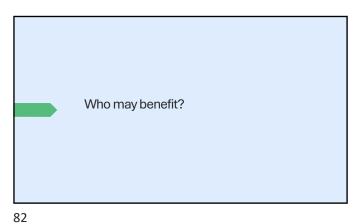
Eversense®

Guardian®



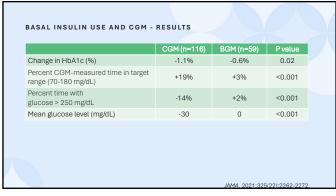
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	Dexcom G6 •	Dexcom G7 •		
Components	Disposable sensor Reusable transmitter Reader for data display/sto	Disposable sensor/transmitt Reader for data display/stora rage		
Maximum wear time	10 days	10 days*> 15 days		
Approved sites	Lower abdomen	Back of upper arm Upper buttocks (children)		
Warm up time	2 hours	30 minutes		
Downloading Software	Dexcom Clarity, Glooko, Tidepool			
FDA approved for Medication Adjustment	Yes			
Alarms for highs/lows	Yes			
Interfering substances	Hydroxyurea, acetaminophen (≥ 1 gram)			
Water resistance	8 feet/24 hours ≥ 2 years			
FDA approved ages				
Sensor size	oval size of 2 quarters	2 stacked pennies		
Data capture	Every 5 minutes (no scanning	g)		



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Design	Multicenter, randomized, parallel group
Intervention	CGM or traditional blood glucose meter (BGM)
Primary Outcome	HbA1c
Key Secondary Outcomes	*CGM-measured time in target glucose range (70-180 mg/dL) *Time with glucose > 250 mg/dL *Mean glucose level
Population	Adults with type 2 diabetes Receiving care from PCP 1-2 daily injections of long or intermediate acting basal insulin without prandial insulin With or without noninsulin medications



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How much do CGMs cost?

CASH

- Abbott Freestyle Libre 3: \$140-170/month with GoodRx coupon (cost for 2 boxes, one sensor each)
- Dexcom G7: \$170-190/month with GoodRx coupon (cost for 3 boxes, one sensor each)

85 86

INSURANCE COVERAGE: MEDICARE

- Requires diagnosis of diabetes
- As of 2021, no longer requires patients to check blood glucose 4 times per day
- MUST have a daily insulin regimen (1 injection per day) OR have a history of documented problematic hypoglycemia
- Have been trained (or had caregiver trained) to use a continuous glucose monitor
- Provider visits every 6 months
- Covered as durable medical equipment in many plans

Centers for Medicare & Medicaid Services. Local Coverage Determination (LCD): Stucose Monitors (L33822) [Internet]. Baltimore (MD): CMS; [cited 2025 Mar 2]. Available from: https://www.cms.snu/medicare.coverage.cds/absset/viewlicd.ssar/2/cride/33892

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INSURANCE COVERAGE: WV MEDICAID

REQUIRES 3+ daily insulin administrations

6 months of adherence based on fill history

INSURANCE COVERAGE: OH MEDICAID

All patients with diagnosis of diabetes (2023)

INSURANCE COVERAGE: COMMERCIAL

Variable

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- May or may not require prior authorization
- WVU Employee Health Plan and Highmark covered with prior authorization

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OTC Options

- Stelo* (Dexcom)

- Subscriptions available for discount

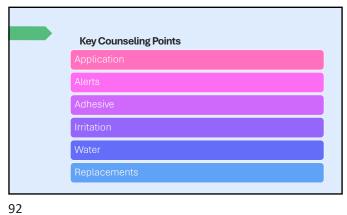
- Lingo* (Abbott)

- Subscription based

- Sensors + coaching

- FSA/HSA eligible

- Targeted for patients without diabetes



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 Key Standardized Metrics

 Time in Range
 • Fasting AND post-prandial while avoiding hypoglycemia (70-180)

 Number of Days Worn
 • 14+ recommended

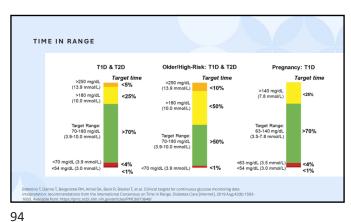
 Percent of Time Active
 • >70% recommended

 Mean Glucose
 • based on selected date range

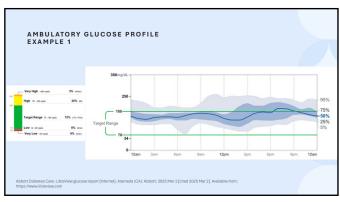
 Glucose Management Indicator (GMI)
 • Estimate of HbA1c

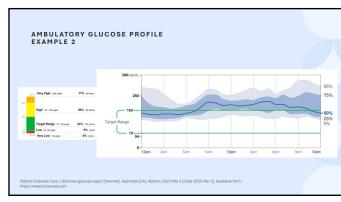
 Coefficient of Variation
 • ≤ 36% recommended

 Ambulatory Glucose Profile (AGP)
 • Summary of glucose values with mean and percentiles as one day



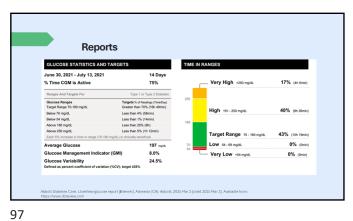
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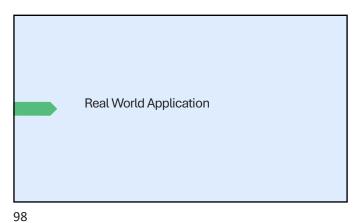


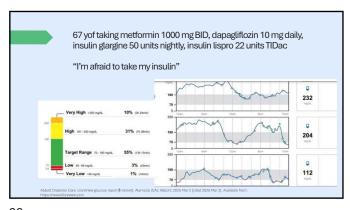


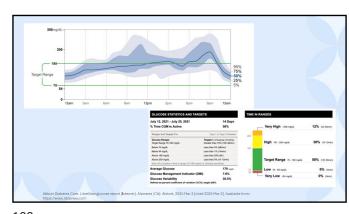
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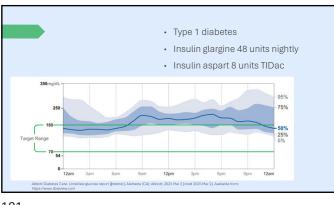
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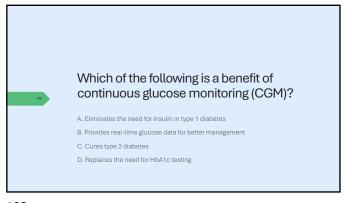












What is a key requirement for Medicare coverage of CGM?

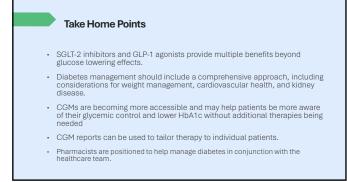
A. The patient must be using insulin or have hypoglycemia

B. The patient must check blood glucose 4 times daily

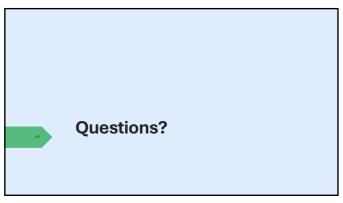
C. The patient must have type 1 diabetes

D. The patient must be over 65 years old

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CE Evaluation Access Code

Capital Letters, No spaces, complete by March 16, 2025

Note: CE credit will be reported to NABP CPE Monitor within 4-6 weeks

107 108

