

OAAPN - Inpatient Diabetes Care: The Landscape for Management and Technologies in 2023

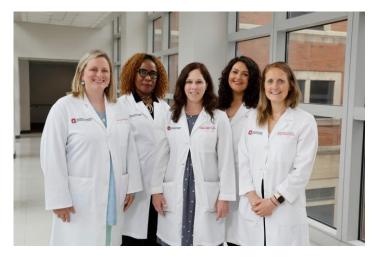
> Cara Harris, DNP, APRN-CNP, CDCES Certified Diabetes Care and Education Specialist



### **Objectives**

- 1. Discuss inpatient diabetes standards of care and best practices for glucose management
- 2. Describe recommended pharmacological therapies to treat hyperglycemia in the acute care setting
- 3. Review common challenges of Diabetes Care in the hospital
- 4. Compare and contrast between the types of diabetes technology

*Slides : many are transition and resource slides for reference* 







Acknowledge OSUWMC DM Team <u>inpt and outpt</u>

Some slides courtesy of team !!

QA join in and after...

#### Collaborative, resources included in this presentation:

CARDI-OH Oho Cardiovascular and Diabetes Health Collaborative

### About Cardi-OH

Founded in 2017, the mission of Cardi-OH is to improve cardiovascular and diabetes health outcomes and eliminate disparities in Ohio's Medicaid population.

**WHO WE ARE**: An initiative of health care professionals across Ohio's seven medical schools.

**WHAT WE DO:** Identify, produce and disseminate evidence-based cardiovascular and diabetes best practices to primary care teams.

**HOW WE DO IT:** Utilize monthly newsletters and an online repository of resources at Cardi-OH.org, podcasts available on <u>Cardi-OH</u> Radio, and the Project ECHO® virtual training model.

#### Learn more at cardi-oh.org





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# CARDI•OH

Ohio Cardiovascular and Diabetes Health Collaborative



### Interpretation of Continuous Glucose Monitoring in Primary Care: A Case-Based Approach

### Contributing Authors on Behalf of Team Best Practices:

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April 2023

Acknowledgement Director, Division of Endocrinology, Diabetes & Metabolism

### Cardi-OH ECHO

What's New in Cardiovascular Prevention? A Series of Case-Based Discussions

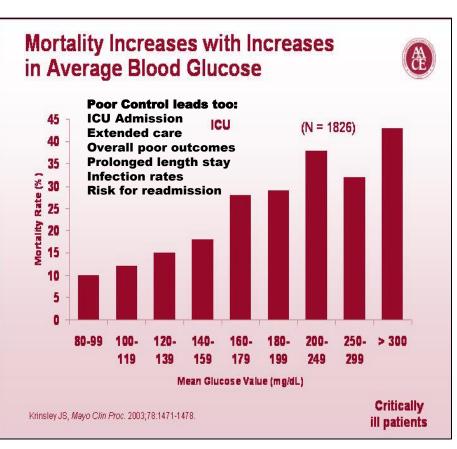
September 29, 2022

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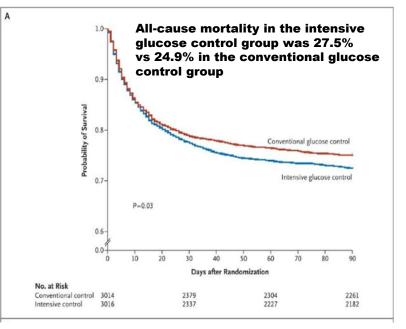
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#### Importance of Inpatient DM Care



### **NICE-Sugar**

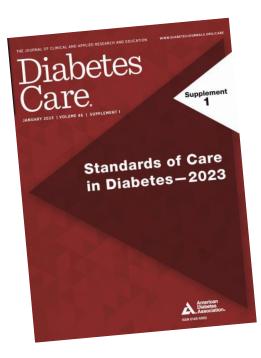


NICE-SUGAR (2009). Intensive versus Conventional Glucose Control in Critically III Patients. *New England Journal of Medicine*, *360*, 1283-1297.

ICU tight glycemic control increased mortality.

#### Inpatient Diabetes Standards of Care and Best Practice for Glucose Management

Diabetes Care Volume 46, Supplement 1, January 2023



# 16. Diabetes Care in the Hospital: Standards of Care in Diabetes—2023

Diabetes Care 2023;46(Suppl. 1):S267-S278 | https://doi.org/10.2337/dc23-S016

"High-quality hospital care for diabetes requires standards for care delivery, which are best implemented using a structured order sets and quality improvement strategies for process improvement."

> DM teams with specialist and educators in hospital can decrease LOS and improve outcomes





#### **Evidence Based Practice**



#### Topics

Search Guidelines Guidelines A-Z iTunes-U Course: Guidelines

**OSUWMC** Guidelines

About Evidence Based Practice Contact Us

OSUWMC Guideline Development <u>Future Guidelines</u> <u>New Guideline Request Process</u> <u>Guideline Feedback Form</u> <u>Guideline Template</u>

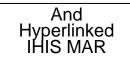
<u>Disclaimer</u>

- Diabetes: 4 Ross ONLY Type 2 Diabetes Mellitus and Other Non-Diabetes-Associated Hyperglycemia
- Diabetes: Foot Burn
- Diabetes: Hypoglycemia Treatment in Non-Pregnant Adults
- Diabetes: In Pregnancy Inpatient Management
- Diabetes: Non-Pregnant Adults Inpatient Management
- Diabetes: Outpatient Management
- Diabetes: Periop/Periprocedure Glucose Management
- , Diabetes: Type 1 Diabetes Mellitus (T1DM) and Diabetic Ketoacidosis (DKA)
- Diabetes: Type 2 Diabetes Mellitus (T2DM) and Other Non-Diabetes-Associated Hyperglycemia (i.e., Stress Induced)



#### Example:

Diabetes Guidelines Streamline Standardize Consistency



### Inpatient BG targets and recommended therapy to treat hyperglycemia – Insulin

#### Table 1.1—Summary of ADA/AACE Recommendations for Management of Hyperglycemia among Hospitalized Patients

|                                   | Critically ill  | Noncritically ill   |
|-----------------------------------|---|---|
| Blood glucose<br>target           | 140 to 180 mg/dL<br>(7.8 to 10.0 mmol/L)  | <ul> <li>Premeal: &lt;140 mg/dL (&lt;7.8 mmol/L)*</li> <li>Random: &lt;180 mg/dL (&lt;10.0 mmol/L)*</li> </ul>  |
| Preferred<br>treatment<br>regimen | <ul> <li>Intravenous insulin infusion of regular insulin</li> <li>Use validated insulin infusion protocol</li> <li>Frequently monitor blood glucose to minimize hypoglycemia</li> </ul> | <ul> <li>Scheduled subcutaneous administration of insulin, with basal, nutritional, and correction components</li> <li>Prolonged therapy with sliding-scale insulin as the sole regimen is discouraged</li> <li>Noninsulin antihyperglycemic agents are not appropriate for most hospitalized patients who require therapy for hyperglycemia</li> </ul> |

\*Provided these targets can be safely achieved. More stringent targets may be appropriate in stable patients with previous tight glycemic control; less stringent targets may be appropriate in terminally ill patients or those with severe comorbidities.

What would you do?

Admitted for CP and had Acute NSTEMI Blood sugar is 300 mg/dL.

- A. Restart home regimen of metformin and glipizide
- B. Start Insulin drip
- C. Start SQ Basal Bolus Insulin
- D. Hold Diabetes meds for now



#### Ex: Insulin Management Inpatient





### **Orals vs Insulin Inpatient**

| Non-Pharmacy<br>Standard of Care All | Pharmacologic agents     |
|--------------------------------------|--------------------------|
| <b>Diabetes Education</b>            | α-Glucosidase inhibitors |
| Lifestyle Change                     | Biguanides               |
| Exercise                             | Sulfonylureas            |
| Nutrition Therapy                    | Meglitinides             |
| Monitoring                           | Thiazolidinediones       |
| Stress Management                    | DDP-4 inhibitors         |
| Weight Management                    | GLP-1 injectable         |
| Physical Activity                    | SGLT – 2 Inhibitor       |
| Sleep                                | Bile acid sequestrates   |
| Smoking ETOH                         | Dopamine-2 agonists      |
| Mood Mental Health                   | Insulin                  |

- Insulin therapy recommended inpatient
- Resources and information on orals, included for your reference not focus this talk today
- Metformin not necessarily 1<sup>st</sup> line anymore now based on cardiorenal protection.



#### **Oral and Insulin Reference**



9. Pharmacologic Approaches to Glycemic Treatment: *Standards* of *Care in Diabetes*—2023

Diabetes Care 2023;46(Suppl. 1):S140-S157 | https://doi.org/10.2337/dc23-S009

10. Cardiovascular Disease and Risk Management: *Standards of Care in Diabetes—2023* 

Diabetes Care 2023;46(Suppl. 1):S158-S190 | https://doi.org/10.2337/dc23-S010



### **Oral Diabetes Medications – Mechanism of Action**

Multiple actions

GLP-1 based "incretin" therapies: <u>GLP-1 analogues</u> and <u>DPP4-inhibitors</u>

Increase insulin secretion (insulin secretagogues):

- Sulfonylureas and meglitinides
- Reduce endogenous glucose production:
  - Biguanides
  - Bile acid sequestrants (Colesevelam)

Increase insulin sensitivity:

Thiazolidinediones

Decrease GI glucose absorption:

- Amylin analogue
- Alpha-glucosidase inhibitors

Block renal glucose reabsorption:

SGLT-2 inhibitors









2008 FDA REQUIRE label indication of reducing CVD events



Composite endpoints !!

MACE Major Adverse CV Events 3 point CV Death, Nonfatal MI, Non-fatal Stroke 4 hospitalization unstable angina

Focus GLP SGLT2 DPP4 why why not ...

### **GLP-1** Receptor Agonists

GLP: Glucagon-like peptide

| Generic<br>Name | Brand<br>Name | Dose Forms       | Dosing<br>Interval | Cautions                                 |
|-----------------|---------------|------------------|--------------------|--|
| Exenatide BID   | Byetta        | 5, 10 µg         | BID                |  |
| Lixisenatide    | Lyxumia       | 10, 20 µg        | Daily              |  |
| Liraglutide*    | Victoza       | 1.6, 1.2, 1.8 mg | Daily              | Thyroid C-cell tumor                     |
| Exenatide QW    | Bydureon      | 2 mg             | Weekly             | warning, advanced<br>CKD, gastroparesis, |
| Semaglutide*    | Ozempic       | 0.5, 1.0 mg      | Weekly             | pancreatitis                             |
| Semagiulide     | Rybelsus      | 3, 7, 14 mg PO   | Daily              |  |
| Dulaglutide*    | Trulicity     | 0.75, 1.5 mg     | Weekly             |  |

CARDI**·OH** 

- No inherent hypoglycemia
- Modest weight 4-9 lbs. and BP reduction



\* Significantly reduce risk CV death, MI, CVA

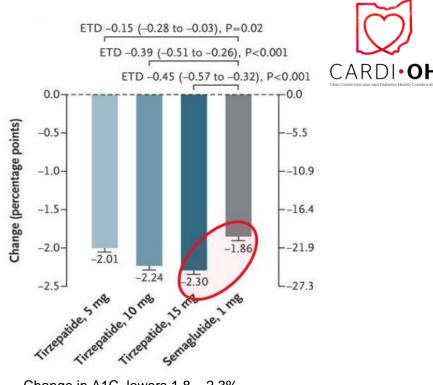
- Nausea/vomiting, usually self-limited
- Lower A1C 0.5 1.6%

# Tirzepatide - Mounjaro

- GLP-1/GIP analogue injection
- Superior A1C/weight loss/QOL vs. semaglutide 1.0 mg
- Similar tolerability
- No comparisons with semaglutide 2 mg or higher



GLP: Glucagon-like peptide GIP: Glucose dependent insulinotropic polypeptide



Change in A1C lowers 1.8 – 2.3% N=1878, 40 week RCT Additional 5.5 kg (12 lb.) weight loss vs. semaglutide

### SGLT2 Inhibitors

Sodium Glucose Transporter



| Name                           | Starting Dose | Max Dose     | Primary Effect          | Cautions                          |
|--------------------------------|---------------|--------------|-------------------------|-----------------------------------|
| Canagliflozin*<br>(Invokana®)  | 100 mg daily  | 300 mg daily |                         |                                   |
| Empagliflozin*<br>(Jardiance®) | 10 mg daily   | 25 mg daily  | Block renal             | UG infection<br>fluid/electrolyte |
| Dapagliflozin*<br>(Farxiga®)   | 5 mg daily    | 10 mg daily  | glucose<br>reabsorption | euglycemic DKA<br>Amputation? (C) |
| Ertugliflozin<br>(Steglatro®)  | 5 mg daily    | 15 mg daily  |                         |                                   |

- Modest blood pressure, weight reduction 3-4 lbs.
- No hypoglycemia
- Small rise in Cr early but long-term renoprotection

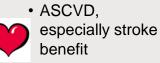


• 1<sup>st</sup> line HF, CKD, CVD before or with metformin Check GFR dosing, limited BG lower GFR<45

### GLP-1RA or SGLT2i ?

Weight loss in both No hypoglycemia in either \$\$

#### GLP1-RA



- Modest renal benefit
- Greater A1c
   reduction

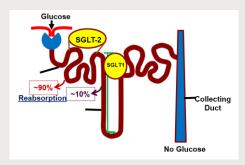
#### SGLT2i

 ASCVD, HF benefit

Renal benefit



Minimal A1c
 reduction at lower
 eGFR



### GLP- 1 agonists Patient education

- Most of these medicines are injectable.
- These medicines may cause some weight loss. The most common side effects are <u>nausea</u>, diarrhea, and <u>upset stomach</u>. These side effects often get better after the first few days.
- <u>Hold</u> day of or week prior to surgery, lower risk nausea, vomiting gastric contents.
- This medicine should be taken on an empty stomach at least 30 minutes before first food, drink, or other oral medicines with no more than 4 oz of plain water

Diabetes Education wexnermedical.osu.edu 23 Oral Diabetes Medicines DM ed book https://go.osu.edu/pted3577

#### SGLT2 Patient education

- Take1 time each day by mouth with or without food.
- These may cause some increased urination and weight loss and improve blood pressure.
- These may <u>reduce cardiovascular disease</u> and <u>chronic kidney disease</u>.
- The major risk is <u>yeast infections.genital infections</u>, and UTI.
- Practice good <u>personal hygiene</u>: wear cotton underwear; change out of wet clothes; do not wear tight clothing; always wipe from front to back; and change tampons, pads, and panty liners often.
- Hold 3 dy prior surgery lower risk ketoacidosis
- Inform patients and caregivers of the signs and symptoms of acidosis, such as rapid breathing, shortness of breath, abdominal pain, nausea, vomiting, feeling tired, or mental status changes, and seek medical attention immediately if they experience the signs or symptoms

| receptor agonists   |  |   |   |   |  |   |
|---|--|---|---|---|--|---|
|   | ELIXA (208)  | LEADER (203)  | SUSTAIN-6 (204)*  | EXSCEL (209)  | REWIND (207)   | PIONEER-6 (205)   |
|   | (n = 6,068)  | (n = 9,340)   | (n = 3,297)   | (n = 14,752)  | (n = 9,901)  | (n = 3,183)   |
| Intervention  | Lixisenatide/placebo                                 | Liraglutide/plaœbo  | Semaglutide s.c.<br>injection/placebo   | Exenatide QW/<br>plaœbo                               | Dulaglutide/<br>placebo  | Semaglutide oral/<br>placebo  |
| Main inclusion criteria   | Type 2 diabetes and<br>history of ACS<br>(<180 days) | Type 2 diabetes and<br>preexisting CVD,<br>CKD, or HF at<br>≥50 years of age<br>or CV risk at ≥60<br>years of age | Type 2 diabetes and<br>preexisting CVD,<br>HF, or CKD at<br>≥50 years of age<br>or CV risk at ≥60<br>years of age | Type 2 diabetes<br>with or without<br>preexisting CVD | Type 2 diabetes and<br>prior ASCVD<br>event or risk<br>factors for ASCVD | Type 2 diabetes and high<br>CV risk (age of ≥50<br>years with established<br>CVD or CKD, or age of<br>≥60 years with CV<br>risk factors only) |
| A1C inclusion criteria (%)  | 5.5-11.0   | ≥7.0  | ≥7.0  | 6.5-10.0  | ≤9.5   | None  |
| Age (years)†  | 60.3   | 64.3  | 64.6  | 62  | 66.2   | 66  |
| Race (% White)  | 75.2   | 77.5  | 83.0  | 75.8  | 75.7   | 72.3  |
| Sex (% male)  | 69.3   | 64.3  | 60.7  | 62  | 53.7   | 68.4  |
| Diabetes duration (years)†  | 9.3  | 12.8  | 13.9  | 12  | 10.5   | 14.9  |
| Median follow-up (years)  | 2.1  | 3.8   | 2.1   | 3.2   | 5.4  | 1.3   |
| Statin use (%)  | 93   | 72  | 73  | 74  | 66   | 85.2 (all lipid-lowering)   |
| Metformin use (%)   | 66   | 76  | 73  | 77  | 81   | 77.4  |
| Prior CVD/CHF (%)   | 100/22   | 81/18   | 60/24   | 73.1/16.2   | 32/9   | 84.7/12.2   |
| Mean baseline A1C (%)   | 7.7  | 8.7   | 8.7   | 8.0   | 7.4  | 8.2   |
| Mean difference in A1C<br>between groups at end of<br>treatment (%) | -0.3**   | -0.4‡   | -0.7 or -1.0^   | -0.53‡^   | -0.61‡   | -0.7  |
| Year started/reported   | 2010/2015  | 2010/2016   | 2013/2016   | 2010/2017   | 2011/2019  | 2017/2019   |
| Primary outcome§  | 4-point MACE 1.02<br>(0.89-1.17)                     | 3-point MACE 0.87<br>(0.78–0.97)  | 3-point MACE 0.74<br>(0.58-0.95)  | 3-point MACE 0.91<br>(0.83-1.00)                      | 3-point MACE 0.88<br>(0.79-0.99)   | 3-point MACE 0.79<br>(0.57-1.11)  |

Table 10.38-Cardiovascular and cardiorenal outcomes trials of available antihyperglycemic medications completed after the issuance of the FDA 2008 guidelines: GLP-1

recentor acconiste

Table 10.3B— Cardiovascular and cardiorenal outcomes trials of GLP-1 and SGLT2

UpToDate ....



Cardiovascular Disease and Risk Management: Standards of Care in Diabetes - 2023. Diabetes Care 2023;46(Suppl. 1):S158-S190 Table 9.3—Median monthly (30-day) AWP and NADAC of maximum approved daily dose of noninsulin glucose-lowering agents in the U.S.

| Class                           | Compound(s)  | Dosage strength/<br>product (if applicable)   | Median AWP<br>(min, max)†  | Median NADAC<br>(min, max)†                                | Maximum approved<br>daily dose*                                   |
|---------------------------------|--|---|--|--|---|
| Biguanides                      | Metformin  | 850 mg (IR)<br>1,000 mg (IR)<br>1,000 mg (ER)   | \$106 (\$5, \$189)<br>\$87 (\$3, \$144)<br>\$242 (\$242, \$7,214)                                      | \$2<br>\$2<br>\$32 (\$32, \$160)                           | 2,550 mg<br>2,000 mg<br>2,000 mg                                  |
| Sul fonylureas (2nd generation) | <ul> <li>Glimepiride</li> <li>Glipizide</li> <li>Glyburide</li> </ul>                                    | 4 mg<br>10 mg (IR)<br>10 mg (XL/ER)<br>6 mg (micronized)<br>5 mg  | \$74 (\$71, \$198)<br>\$70 (\$67, \$91)<br>\$48 (\$46, \$48)<br>\$52 (\$48, \$71)<br>\$79 (\$63, \$93) | \$3<br>\$6<br>\$11<br>\$12<br>\$9                          | 8 mg<br>40 mg<br>20 mg<br>12 mg<br>20 mg                          |
| Thiazolidinedione               | <ul> <li>Pioglitazone</li> </ul>   | 45 mg   | \$345 (\$7, \$349)   | \$4  | 45 mg   |
| α-Glucosidase inhibitors        | Acarbose     Miglit ol   | 100 mg<br>100 mg  | \$106 (\$104, \$106)<br>\$241 (\$241, \$346)   | \$29<br>NA   | 300 mg<br>300 mg  |
| Meglitinides                    | <ul> <li>Nateglinide</li> <li>Repaglinide</li> </ul>   | 120 mg<br>2 mg  | \$155<br>\$878 (\$58, \$897)   | \$27<br>\$31   | 360 mg<br>16 mg   |
| DPP-4 inhibitors                | <ul> <li>Alog liptin</li> <li>Saxagli pt in</li> <li>Linagliptin</li> <li>Sitagliptin</li> </ul>         | 25 mg<br>5 mg<br>5 mg<br>100 mg   | \$234<br>\$565<br>\$606<br>\$626   | \$154<br>\$452<br>\$485<br>\$500                           | 25 mg<br>5 mg<br>5 mg<br>100 mg                                   |
| SGLT2 inhibitors                | Ertugliflozin     Dapagliflozin     Canagliflozin     Empagliflozin                                      | 15 mg<br>10 mg<br>300 mg<br>25 mg   | \$390<br>\$659<br>\$684<br>\$685   | \$312<br>\$527<br>\$548<br>\$547                           | 15 mg<br>10 mg<br>300 mg<br>25 mg                                 |
| GLP-1 RAS                       | Exenatide<br>(extended release)<br>Exenatide<br>Dulaglutide<br>Semaglutide<br>Liraglutide<br>Liraglutide | 2 mg powder for<br>suspension or pen<br>10 µg pen<br>4.5 mg mL pen<br>1 mg pen<br>14 mg (tablet)<br>1.8 mg pen<br>20 µg pen | \$936<br>\$961<br>\$1,064<br>\$1,070<br>\$1,070<br>\$1,278<br>\$814                                    | \$726<br>\$770<br>\$852<br>\$858<br>\$858<br>\$1,022<br>NA | 2 mg**<br>20 μg<br>4.5 mg**<br>2 mg**<br>14 mg<br>1.8 mg<br>20 μg |
| GLP-1/GIP dual agonist          | Tirzepatide  | 15 mg pen   | \$1,169  | \$935  | 15 mg**   |
| Bile acid sequestrant           | Colesevelam  | 625 mg tabs<br>3.75 g suspension  | \$711 (\$674, \$712)<br>\$674 (\$673, \$675)   | \$83<br>\$177  | 3.75 g<br>3.75 g  |
| Dopamine-2 agonist              | <ul> <li>Bromocriptine</li> </ul>  | 0.8 mg  | \$1,118  | \$899  | 4.8 mg  |
| Amylin mimetic                  | <ul> <li>Pramlintide</li> </ul>  | 120 µg pen  | \$2,783  | NA   | 120 μg/injection+   |

AWP, average wholesale price; DPP4, dipeptidyl peptidase 4; ER and XL, extended release; GIP, glucose-dependent insulinotropic polypeptide; GLP-1 RA, glucagon-like peptide 1 receptor agonist; IR, immediate release; max, maximum; min, minimum; NA, data not available; NADAC, National Average Drug Acquisition Cost; SGLT2, sodium:glucose cotransporter 2. †Calculated for 30-day supply (AWP [72] or NADAC [73] unit price × number of doses required to provide maximum approved daily dose × 30 days); median AWP or NADAC listed alone when only one product and/or price. \* 'Utilized to calculate median AWP and NADAC (min, max); generic prices used, if available commercially. \*\*Administered once weekly. ††AWP and NADAC calculated based on 120 µg three times daily.

#### Review ADA section 9 Pharma Tx

# Summary of oral glucose-lowering agents.

Beyond scope this talk review orals but included reference.

Median monthly cost, options and doses.



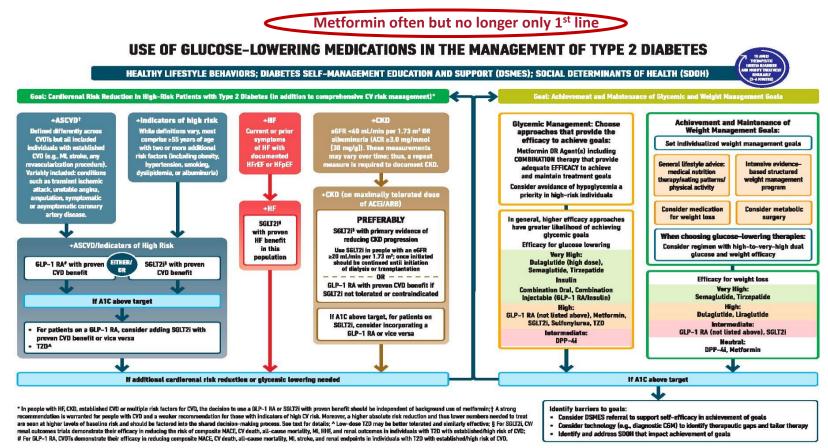
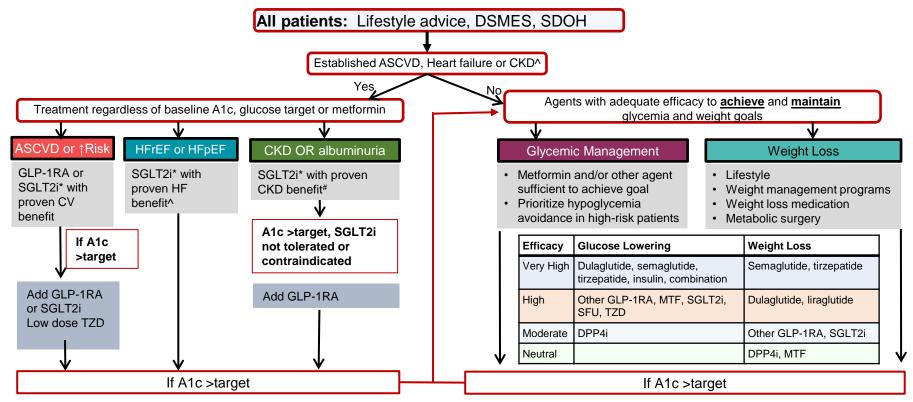


Figure 9.3—Use of glucose-lowering medications in the management of type 2 diabetes. ACEi, angiotensin-converting enzyme inhibitor; ACR, albumin-to-creatinine ratio; ARB, angiotensin receptor blocker; ASCVD, atherosclerotic cardiovascular disease; CGM, continuous glucose monitoring; CKD, chronic kidney disease; CV, cardiovascular disease; CVOT, cardiovascular outcomes trial; DPP-4i dipetidyl peptidase 4 inhibitor; eGFR, estimated glomerular filtration rate; GLP-1 RA, glucagon-like peptide 1 receptor agonist; HF, heart failure; HFpEF, heart failure; WACE, major adverse cardiovascular events; MJ, myocardial infarction; SDOH, social determinants of health; SGLT2i, sodium: glucose cotransporter 2 inhibitor; TZD, type 2 diabetes; TZD, thiaoidinedione. Adapted from Davies et al. (45).

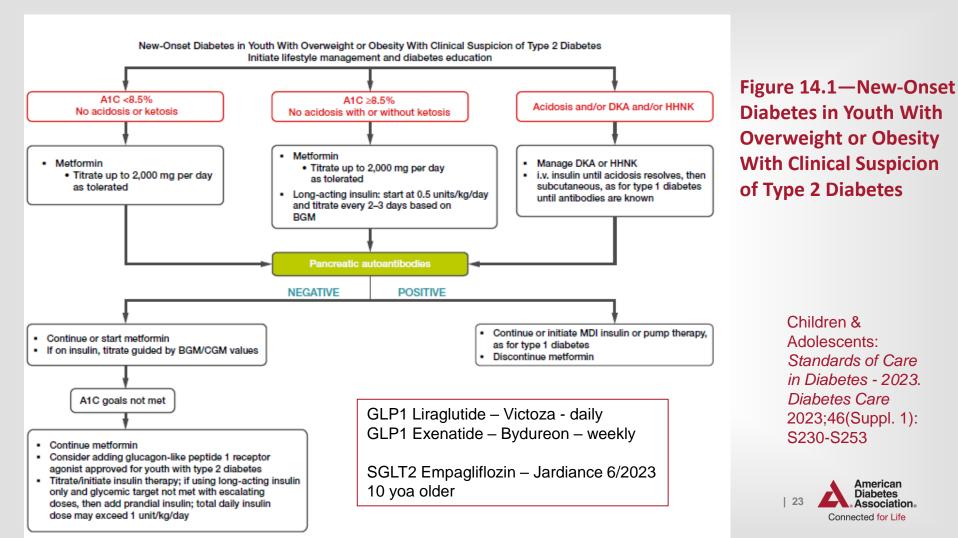
#### Standards of Medical Care in Diabetes 2023

514

#### Pharmacologic Management, ADA/EASD Consensus 2022







#### Final Oral Summary/Reference

Table 9.2—Medications for lowering glucose, summary of characteristics

|                        |                      |                         | Hypogly-   |   | CV ei   | fects  |  | Ranal effects   |             | A CONTRACTOR  |   |
|------------------------|----------------------|-------------------------|------------|---|---|--|--|---|-------------|---|---|
|                        |                      | Efficacy <sup>1</sup>   | cemia      | Weight change <sup>2</sup>                                      | Effect on MACE  | HF   | Progressien of DKD   | Dosing/use considerations*  | Oral/SQ     | Cost  | Clinical considerations   |
| Metformie              |                      | High                    | No         | Neutral (potential<br>for modest loss)                          | Potential<br>benefit                                  | Neutral  | Neutral  | <ul> <li>Contraindicated with eGFR &lt;30 mL/min<br/>per 1.73 m<sup>2</sup></li> </ul>  | Oral        | Low   | Bi side effects common; to mitigate Bi side effects, consider slow dose titration, extended retease formulations, and administration with food Potential for vitamin BI2 deficiency, monitor at regular intervals   |
| SGLT2 inhi             | ibitors              | Intermediate<br>to high | No         | Loss<br>(intermediate)  | Benefit:<br>canagliflozin,<br>empagliflozin           | Benefit:<br>canagliflozin,<br>dapagliflozin,<br>empagliflozin,<br>ertugliflozin                | Benefit:<br>canagliflozin,<br>dapagliflozin,<br>empagliflozin  | <ul> <li>See labels for renal dose considerations<br/>of individual agents</li> <li>Elucose-towering effect is lower for<br/>SGLT2 inhibitors at lower eGFR</li> </ul>  | Oral        | High  | DKA risk, rare in T2DM: discontinue, evoluate, and treat promptly if suspected; be aware of predisposing risk factors and clinical presentation (including eoglycamic DKA); discontinue before scheduled surgey (e.g., 3–4 days), during critical illness, or during prolonged fasting to mitigate potential risk.     Increase risk of genital mycotic infactions     Necrotizing fascilitis of the perineum (Fournier gangrene), rare reports: institute prompt treatment if suspected     Attention to volume status, blood pressure; adjust other volume-contracting agents as applicable   |
| GLP-1 RAs              | High to<br>very high | No                      | very high) | Benefit:<br>dulaglutide,<br>liraglutide,<br>semaglutide<br>(SQ) | Neutral   | Benefit for renal<br>endpoints in CVOTs,<br>driven by albuminuria<br>outcomes:<br>dulaglutide, | See labels for renal dose considerations<br>of individual agents     No dose adjustment for dulaglutide,<br>liarglutide, semaglutide     Monitor renal function when initiating or | SQ; oral<br>(semaglutide)   | High        | <ul> <li>Risk of thyroid C-cell tumors in rodents; human relevance not determined (liraglutide,<br/>dutaglutide, exonatide extended release, senaglutide)</li> <li>Coursel patients on potential for GI side effects and their typically tamporary nature; provide<br/>guidance on dietary modifications to mitigate GI side effects (moduction in meal size, mindful<br/>eating practices (e.g., store sitting once full, decreasing initiate of high-ratic or spicy food);</li> </ul> |   |
|                        |                      |                         |            |   | Neutral:<br>exenatide<br>once weekly,<br>lixisenatide |  | liraglutide,<br>semaglutide (SQ)   | escalating doses in patients with renal<br>impairment reporting severe adverse<br>GI reactions  |             |   | consider slower dose titration for patients experiencing 61 challenges Pancreatitis has been reported in clinical trials but causality has not been established. Disconfigue if pancreatitis is suspected Evaluate for gallbladder disease if cholelithiasis or cholecystitis is suspected  |
| 6IP and 6I             | LP-1 RA              | Very high               | No         | Loss (very high)  | Under<br>investigation                                | Under<br>investigation   | Under investigation  | <ul> <li>See label for renal dose considerations</li> <li>No dose adjustment</li> <li>Monitor renal function when initiating or<br/>escalating doses in patients with renal<br/>impairment reporting severe adverse<br/>61 reactions</li> </ul> | 50          | High  | Risk of thyroid C-cell tumors in rodents; human relevance not determined     Coursel patients on potential for Gi side effects and their typically temporary nature; provide     puidance on diretary modifications to mitigate G side effects finduction in meal size, mindful     sating practices (e.g., stop eating once full, decreasing intake of high-fat or spicy food);     consider solver dose titration for patients experimenting Gi challenges     Pancreatitis has been reported in clinical trials but causality has not been established.     Discontinue if pancreatitis is suspected     Evaluate for gallbadder disease if chaltelithasis or cholecystilts is suspected |
| DPP-4 inh              | nibitors             | Intermediate            | No         | Neutral   | Neutral   | Neutral<br>(potential risk,<br>saxagliptin)  | Neutral  | <ul> <li>Renal dose adjustment required<br/>(sitagliptin, saxagliptin, alogliptin); can<br/>be used in renal impairment</li> <li>No dose adjustment required for<br/>linagliptin</li> </ul>   | Oral        | Kigh  | Pancmatilis has been reported in clinical trials but causality has not been established. Discontinue if pancmatilis is suspected     Joint pain     Bulleus pernphigoid (postmarketing): discontinue if suspected   |
| Thiazolidi             | inediones            | High                    | No         | Gain  | Potential benefit:<br>pioglitazone                    | Increased risk   | Neutral  | <ul> <li>No dose adjustment required</li> <li>Generally not recommended in renal<br/>impairment due to potential for fluid<br/>retention</li> </ul>   | Oral        | Low   | Congestive HF (plogitizzone, rosigilizzone)     Fuid retardion (deema; heart failure)     Benefit in NASH     Risk of bone fractures     Weight gain: consider tower doses to mitigate weight gain and edema  |
| Sulfonylu<br>(2nd gene |                      | High                    | Yes        | 6ain  | Neutral   | Neutral  | Neutral  | Glyburide: generally not recommended<br>in chronic kidney disease     Glipizide and glimepiride: initiate<br>conservatively to avoid hypoglycemia   | Oral        | Low   | <ul> <li>FDA Special Warning on increased risk of CV mortality based on studies of an older sutionylurea<br/>(totbutanide); glimepiride shown to be CV safe (see text)</li> <li>Use with caution in persons at risk for hypoglycemia</li> </ul>   |
| Insulin                | Human                | High to                 | Yes        | 6ain  | Neutral   | Neutral  | Neutral  | <ul> <li>Lower insulin doses required with a<br/>decrease in eGFR; titrate per clinical</li> </ul>  | SQ; inhaled | Low (SQ)  | <ul> <li>Injection site reactions</li> <li>Higher risk of hypoplycemia with human insulin (NPH or premixed formulations) vs. analogs</li> </ul>   |
|                        | Analogs              | very high               |            |   |   |  |  | decrease in e644; titrate per cunical<br>response   | 50          | High  | <ul> <li>nighter risk or hypoglycemia with numan insulin (NPH of premixed formulations) vs. analogs</li> </ul>  |

CV, cardiovascular; CVOT, cardiovascular outcomes trial; DKA, diabetic ketoacidosis; DKD, diabetic kidney disease; DPP-4, dipeptidyl peptidase 4; eGFR, estimated glomerular filtration rate; FDA, U.S. Food and Drug Administration; GI, gastrointestinal; GIP, gastric inhibitory polypeptide; GLP-1 RA, glucagon-like peptide 1 receptor agonist; HF, heart failure; NASH, nonalcoholic steatohepatitis; MACE, major adverse cardiovascular events; SGLT2, sodium–glucose cotransporter 2; SQ, subcutaneous; T2DM, type 2 diabetes mellitus. \*For agent-specific dosing recommendations, please refer to manufacturers' prescribing information. <sup>3</sup>Tsapas et al. (62). <sup>3</sup>Tsapas et al. (14). Reprinted from Davies et al. (45).

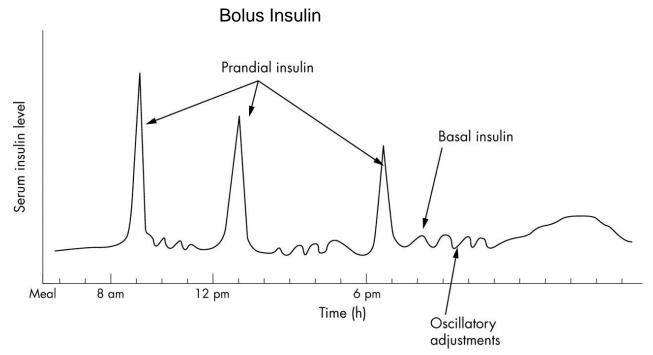
### Inpatient Management of Glucose Control

- Oral and non-insulin injectable therapy
  - No safety data available for use in hospital setting (oral intake varies, n/v, renal clearance)
  - Continuation of home therapy in select patients
    - Typically stop oral diabetes agents in hospital
    - Resuming if potential discharge and pt stable
- Insulin therapy preferred for majority of patients initiated for BG >180 mg/dL
  - IV infusion preferred for critical care patients
  - SubQ insulin (including basal, nutritional, and correction doses) preferred for general medicine patients – may need reduction in home total daily dose 20-50%.
- GLP1 injectables not used inpatient not on formulary and can cause nausea and vomiting and with limited oral intake.
- SGLT2 some evidence beneficial inpatient T2DM CHF Cardiology weigh risk / benefit

Slide adopted OSUWMC Med Safety 2012 Standards of Medical Care in Diabetes 2023

#### Insulin therapy preferred for majority of inpatients :

Representation of variation in serum insulin concentration during a 24 hour period to illustrate components of insulin secretion/delivery.



Hindmarsh P C Arch Dis Child 2005;90:1144-1147

### High Risk Medication - Insulin

- Definition
  - "High-risk drugs involved in a high percentage of medication errors and/or other adverse outcomes."
- Rationale/risk
  - Cause profound hypoglycemia accompanied by EKG changes and arrhythmias
  - Lethal if given in substantially excessive doses or in place of other medications
- Common causes of inpatient hypoglycemia:
  - Acute kidney injury, decreasing insulin clearance
  - Prescribing errors
  - Failure to respond to nutritional interruption
  - Failure to manage a prior hypoglycemic event
- Recommendations
  - Use electronic health record (EMR) protocols for insulin administration
  - Insulin order sets and guidelines
  - Insulin dosing algorithms in the EMR
  - Evaluate hypoglycemia trends and address systemic issues committees

Table 9.4-Median cost of insulin products in the U.S. calculated as AWP (72) and NADAC (73) per 1,000 units of specified dosage form/product

| Insulins                   | Compounds  | Dosage form/product                                     | Median AWP<br>(min, max)* | Median<br>NADAC*             |
|----------------------------|--|---|---------------------------|------------------------------|
| Rapid-acting               | <ul> <li>Lispro follow-on product</li> </ul>         | U-100 vial  | \$118 (\$118, \$157)      | \$94                         |
|                            |  | U-100 prefilled pen                                     | \$151                     | \$121                        |
|                            | Lispro   | U-100 vial  | \$99†                     | \$79†                        |
|                            |  | U-100 cartridge   | \$408                     | \$326                        |
|                            |  | U-100 prefilled pen                                     | \$127+                    | \$102†                       |
|                            |  | U-200 prefilled pen                                     | \$424                     | \$339                        |
|                            | <ul> <li>Lispro-aabc</li> </ul>                      | U-100 vial  | \$330                     | \$261                        |
|                            |  | U-100 prefilled pen                                     | \$424                     | \$339                        |
|                            |  | U-200 prefilled pen                                     | \$424                     | NA                           |
|                            | Glulisine  | U-100 vial  | \$341                     | \$272                        |
|                            |  | U-100 prefilled pen                                     | \$439                     | \$351                        |
|                            | Aspart   | U-100 vial  | \$174+                    | \$140+                       |
|                            |  | U-100 cartridge   | \$215+                    | \$172*                       |
|                            |  | U-100 prefilled pen                                     | \$224†                    | \$180+                       |
|                            | <ul> <li>Aspart ("faster acting product")</li> </ul> | U-100 vial  | \$347                     | \$277                        |
|                            |  | U-100 cartridge   | \$430                     | \$344                        |
|                            |  | U-100 prefilled pen                                     | \$447                     | \$357                        |
|                            | <ul> <li>Inhaled insulin</li> </ul>                  | Inhalation cartridges                                   | \$1,418                   | NA                           |
| Short-acting               | <ul> <li>Human regular</li> </ul>                    | U-100 vial  | \$165++                   | \$132++                      |
|                            |  | U-100 prefilled pen                                     | \$208                     | \$166                        |
| Intermediate-acting        | Human NPH  | U-100 vial<br>U-100 prefilled pen                       | \$165++<br>\$208          | \$132 <sup>++</sup><br>\$168 |
|                            |  |   |                           |                              |
| Concentrated human regular | <ul> <li>U-500 human regular insulin</li> </ul>      | U-500 vial  | \$178                     | \$142                        |
| insulin                    |  | U-500 prefilled pen                                     | \$230                     | \$184                        |
| Long-acting                | <ul> <li>Glargine follow-on products</li> </ul>      | U-100 prefilled pen                                     | \$261 (\$118, \$323)      | \$209 (\$209, \$258          |
|                            | -  | U-100 vial  | \$118 (\$118, \$323)      | \$95                         |
|                            | <ul> <li>Glargine</li> </ul>                         | U-100 vial; U-100 prefilled pen                         | \$136†                    | \$109+                       |
|                            | Partner to   | U-300 prefilled pen                                     | \$346                     | \$277                        |
|                            | Detemir  | U-100 vial; U-100 prefilled pen                         | \$370                     | \$296                        |
|                            | Degludec   | U-100 vial; U-100 prefilled pen;<br>U-200 prefilled pen | \$407                     | \$326                        |
| Premixed insulin products  | NPH/regular 70/30                                    | U-100 vial  | \$165++                   | \$133++                      |
|                            |  | U-100 prefilled pen                                     | \$208                     | \$167                        |
|                            | <ul> <li>Lispro 50/50</li> </ul>                     | U-100 vial  | \$342                     | \$274                        |
|                            |  | U-100 prefilled pen                                     | \$424                     | \$339                        |
|                            | <ul> <li>Lispro 75/25</li> </ul>                     | U-100 vial  | \$342                     | \$273                        |
|                            |  | U-100 prefilled pen                                     | \$127†                    | \$103+                       |
|                            | Aspart 70/30   | U-100 vial  | \$180†                    | \$146†                       |
|                            |  | U-100 prefilled pen                                     | \$224†                    | \$178†                       |
| Premixed insulin/GLP-1 RA  | <ul> <li>Glargine/Lixisenatide</li> </ul>            | 100/33 µg prefilled pen                                 | \$646                     | \$517                        |
| products                   | <ul> <li>Degludec/Liraglutide</li> </ul>             | 100/3.6 µg prefilled pen                                | \$944                     | \$760                        |

AWP, average wholesale price; GLP-1 RA, glucagon-like peptide 1 receptor agonist; NA, data not available; NADAC, National Average Drug Aquisition Cost. \*AWP or NADAC calculated as in Table 9.3. + Generic prices used when available. +TAWP and NADAC data presented do not include vials of regular human insultin and NPH available at Walmart for approximately 525/vial; median listed alone when only one product and/or price. Review ADA section 9 Pharma Tx

### Reference for all Insulin options

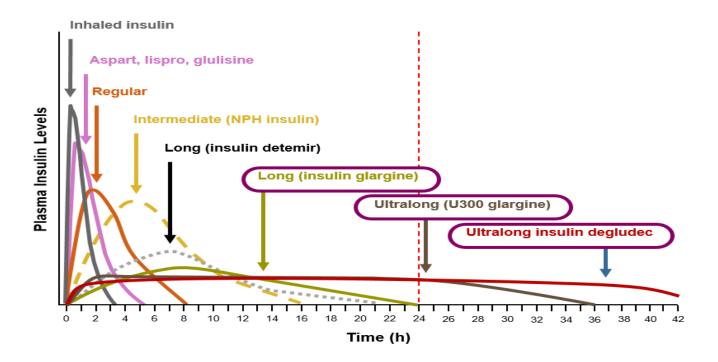
### Median monthly cost, and options to Rx.



### Insulin Types

| Types  | Formulations   |
|--|--|
| Bolus - Rapid Acting   | Aspart-Novolog, Glulisine-Apidra<br>Lispro-Humalog U100&U200, Admelog  |
| Ultra –Rapid Acting<br>Ultra –Rapid Acting- Inhaled  | Aspart-Fiasp; Lispro-Lyumjev<br>Afreeza<br>(no COPD,Asthma, Smoke, need PFTs)  |
| Bolus - Short Acting   | Regular  |
| Basal Insulin – Long Acting<br>Intermediate Acting   | Glargine U100-Lantus; Basaglar; Semglee<br>Glargine U300-Toujeo<br>Detemir-Levemir<br>Degludec –Tresiba U100&U200<br>NPH       |
| Pre – Mixed<br>NPH/Regular Based<br>NPL Based/Lispro/Humalog<br>NPH/Aspart/Novolog<br>Degludec/Novolog Ryzodeg | 70/30<br>75/25, 50/50<br>70/30<br>70/30  |
| Concentrations   | U-100; U-200, U-300, U-500 specialist<br>Use cautiously inpatient – follow a guideline<br>and specialist consult best practice |

# Insulin Pharmacology



#### PK = pharmacokinetic; NPH = neutral protamine Hagedorn.

Adapted from Hirsch IB. NEJM. 2005;352:174-183. Flood TM. J Fam Pract. 2007;56(suppl 1):S1-S12. Becker RH, et al. Diabetes Care. 2015;38:637-643. http://www.pdr.net/full-prescribing-information/afrezza?druglabelid=3540. Accessed April 5, 2015. Hompesch M, et al. Clin Ther. 2014;36(4):507-515. Adapted K.Wyne 2-2017

#### Table 1. IV Insulin Infusion

| 5 columns=                            |  | Change in Glucose from Prior Measure   |   |  |  |  |                                       |  |
|---------------------------------------|--|--|---|--|--|--|---------------------------------------|--|
| greater<br>precision                  | Current<br>Glucose   | Decreased<br>> 100 mg/dL <sup>1</sup>  | Decreased<br>50-100 mg/dL   | Decreased<br>25-50 mg/dL                               | Increased or<br>decreased<br>< 25 mg/dL      | Increased<br>25-50 mg/dL                               | Increased<br>> 50 mg/dL               |  |
| precision                             | > 400 mg/dL  | <ul> <li>&gt; 400 mg/dL</li> <li>Contact the prescriber.</li> <li>Increase infusion rate according to the row for 301-400 mg/dL.</li> <li>If glucose is &gt; 400 mg/dL and the decline in glucose is &lt; 25 mg/dL per hour for two consecutive checks, consider doubling the rate of infusion.</li> </ul> |   |  |  |  |                                       |  |
| Discrete                              | 301-400 mg/dL  | No Change  | Increase<br>infusion rate by<br>1 unit/hr                                       | Increase<br>infusion rate by<br>2 units/hr             | Increase<br>infusion rate by<br>2.5 units/hr | Increase<br>infusion rate by<br>3 units/hr             | Increase<br>infusion by 4<br>units/hr |  |
|                                       | 201-300 mg/dL  | Run infusion at<br>75% of current<br>rate <sup>3</sup>   | No Change   | Increase<br>infusion by 1<br>unit/hr                   | Increase<br>infusion rate by<br>1 unit/hr    | Increase<br>infusion by 2<br>units/hr                  | Increase<br>infusion by 3<br>units/hr |  |
|                                       | 151-200 mg/dL  | Run infusion at<br>50% of current<br>rate  | Decrease<br>infusion by 1<br>unit/hr  | No Change  | Increase<br>infusion<br>by 0.5 unit/hr       | Increase<br>infusion<br>by 1 unit/hr                   | Increase<br>infusion<br>by 2 unit/hr  |  |
| Determine Insulin                     | 120-150 mg/dL<br>OPTIMAL   | Run infusion at<br>25% of current<br>rate <sup>2</sup>   | Run infusion at<br>50% of current<br>rate                                       | Run infusion<br>at 75% of<br>current rate              | No Change                                    | No Change  | Increase<br>infusion by 1<br>unit/hr  |  |
| •Change in BG                         | 80-120 mg/dL   | Stop the<br>infusion,<br>contact the<br>prescriber and<br>recheck glucose<br>in 15 minutes   | Run infusion at<br>10% of current<br>rate; consider<br>contacting<br>prescriber | Run infusion at<br>25% of current<br>rate <sup>2</sup> | Run infusion at<br>50% of current<br>rate    | Run infusion at<br>75% of current<br>rate <sup>3</sup> | No Change                             |  |
| •Direction of change in BG            | Stop infusion of insulin and contact the prescriber.     Double current infusion rate of dextrose solution.     If not receiving dextrose IV infusion, start D5W at 50 ml/hr.     Consider giving D50% according to the <u>Hypoglycemia Treatment in Non-Pregnant Adults of</u>  |  |   |  |  |  | Its guideline.                        |  |
| •Current BG<br>•Current infusion rate | <ul> <li>S0 mg/dL</li> <li>Recheck glucose and treat according to the Hypoglycemia Treatment in Non-Pregnant Adults glucose and treat according to the Hypoglycemia Treatment in Non-Pregnant Adults gevery 15 minutes until glucose &gt; 80 mg/dL.</li> <li>Resume insulin at 25% of previous dose and reduce dextrose back to previous rate when glut 150 mg/dL in the absence of subcutaneous basal insulin (determir, glargine, NPH).</li> <li>This applies to patients with type 2 diabetes or other causes of hyperglycemia. Click here to access OSUWMC Type 1 Diabetes Mellitus (T1DM) and Diabetic Ketoacidosis (DKA) guideline.</li> </ul> |  |   |  |  | t Adults guideline<br>when glucose ><br>to access the  |                                       |  |

**Example** 

IV insulin Infusion Guideline

Streamline Standardize Consistency

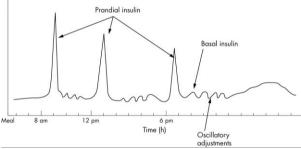
Can also include fluid replacement parameters

<sup>1</sup> Contact prescriber if rate of decline in glucose >100 mg/dL/hr. Patient may need a more rapid taper of the drip than indicated in the table above.



### Example: Guide to Inpatient Insulin Management

| Basal Insulin                       | <ul> <li>Long-acting insulin</li> <li>NPH</li> <li>Continuous SQ rapid acting insulin (pump)</li> <li>IV insulin drip</li> </ul>  |   |
|-------------------------------------|---|---|
| Prandial Insulin                    | <ul> <li>Rapid-acting insulin</li> <li>(High) 1 unit for every 5 g CHO</li> <li>(STD) 1 unit for every 10 g CHO</li> <li>(Low) 1 unit for every 20 g CHO</li> <li>Regular insulin (tube feeds)</li> </ul>                       | Meal 8 am   |
| Correction/Supplement<br>al Insulin | <ul> <li>Rapid-acting insulin</li> <li>(High) 1 unit for every 25 mg/dl BG &gt;150</li> <li>(STD) 1 unit for every 50 mg/dl BG &gt;150</li> <li>(Low) 1 unit for every 100 mg/dl BG &gt;150</li> <li>IV insulin drip</li> </ul> | Standard:<br>150-200 = 1 unit<br>201-250 = 2 units<br>251-300 = 3 units<br>301 -350 = 4 units |





### Summary of Inpatient Insulin Pitfalls:

- Stopping the infusion without basal insulin coverage
- Failure to restart the infusion following recovery from hypoglycemia
- Stopping the infusion before stable infusion rate is achieved
- T1DM patients should *never* go without basal insulin leads to <u>iatrogenic</u> DKA
- In general, DO NOT hold basal insulin even with procedures
- Avoid Regular sliding scale monotherapy in most patients, unless NPO.
- Insulin pens "For Single patient use only"





What would you do?

Admitted for CP and had Acute NSTEMI Blood sugar is 300 mg/dL.

- A. Restart home regimen of metformin and glipizide
- B. Start Insulin drip
- C. Start SQ Basal Bolus Insulin
- D. Hold Diabetes meds for now

Answer:

B. Start Insulin drip acute ill, insulin tx of choice inpatient

D/C plan depends on hospital course and A1C. Likely will need insulin at discharge.

### **Case 1- the plan:**

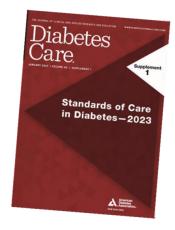
Ex: Insulin Management Inpatient





### Common challenges of DM care in the hospital

- Hypoglycemia
- Steroid induced hyperglycemia
- DKA/HHS/Euglycemic DKA
- Nutrition Tube Feed, TPN
- NPO, Perioperative Care
- Technology
- Discharge Planning

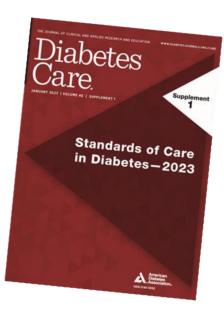


No time great detail Some cases and OSUWMC examples but more: Review summarize guidelines Can get more guidance guidelines Q&A after 1-1 more specifics

# Hypoglycemia in the hospital

# 6. Glycemic Targets: Standards of Care in Diabetes—2023

Diabetes Care 2023;46(Suppl. 1):S97-S110 | https://doi.org/10.2337/dc23-S006



# Hypoglycemia

|         | Glycemic criteria/description   |
|---------|---|
| Level 1 | Glucose <70 mg/dL (3.9 mmol/L) and ≥54 mg/dL (3.0 mmol/L)   |
| Level 2 | Glucose <54 mg/dL (3.0 mmol/L)  |
| Level 3 | A severe event characterized by altered mental and/or physical status requiring<br>assistance for treatment of hypoglycemia |

#### Inpatient risk factors:

- Medications: insulin & sulfonylureas
- Illnesses/conditions: AKI/CKD, cirrhosis, sepsis
- Nutrition: decrease in po intake, interruption in TF

Overall goal hypoglycemia prevention Outpatient greater use of continuous glucose monitor recommended (not yet FDA approved inpatient).

| Table 2: Pati | ents Who Are Alert, Able   | to Tolerate PO Intake  | , and with Intact Cognitiv  | ve Status  |  |
|---------------|--|--|---|--|--|
| BG Level*     |  | Action*  |   | Fellow-Up  |  |
| 60-79 mg/dL   | Administer 15 g oral<br>carbohydrate, choose one:<br>• 4 oz. juice (NOT OJ)**<br>or regular soda/pop<br>• 1 tbsp. jelly or sugar<br>• 3 glucose tablets<br>• 1 tube glucose 40% oral<br>gel  | If next meal within 1-2<br>hrs. also administer<br>(choose one):<br>• 3 graham crackers<br>• 6 saltine crackers<br>• 8 oz. skim milk | <ul> <li>If next meal &gt; 2 hrs. also<br/>administer (choose one):</li> <li>½ sandwich (15 g)</li> <li>3 graham crackers<br/>with one tbsp. peanut<br/>butter</li> </ul> | <ul> <li>Recheck BG q15 min<br/>following treatment<br/>and continue to treat<br/>accordingly until ≥ 80<br/>mg/dL</li> </ul>  |  |
|               | hypoglycemia (> 2 distinc  | fficer to report BG if patien<br>it events with BG < 70 mg/o<br>d glucose lowering therapy<br>if symptomatic                         | iL in past 12 hours)  | <ul> <li>Once BG ≥ 80 mg/dL,<br/>recheck BG q1h x 2,<br/>then resume POC<br/>glucose as previously</li> </ul>  |  |
| 45-59 mg/dL   | Administer 20 g oral<br>carbohydrate, choose one:<br>6 oz. juice (NOT OJ)**<br>or regular soda/pop<br>1 ½ tbsp. jelly or sugar<br>4 glucose tablets<br>1 ½ tube glucose 40%<br>oral gel  | If next meal within 1-2<br>hrs. also administer<br>(choose one):<br>• 3 graham crackers<br>• 6 saltine crackers<br>• 8 oz. skim milk | <ul> <li>If next meal &gt; 2 hrs. also<br/>administer (choose one):</li> <li>½ sandwich (15 g)</li> <li>3 graham crackers<br/>with one tbsp. peanut<br/>butter</li> </ul> | ordered<br>Patients who are<br>admitted with<br>hypoglycemia should<br>be monitored at least<br>q4h for a minimum<br>of 24h  |  |
|               | Call House Officer to report B   |  |   |  |  |
| < 45 mg/dL    | Administer 30 g oral<br>carbohydrate, choose one:       If next meal within 1-2<br>hrs. also administer<br>(choose one):         8 oz. juice (NOT OJ)**<br>or regular soda/pop       1 f next meal within 1-2<br>hrs. also administer<br>(choose one):         2 tbsp. jelly or sugar       3 graham crackers         6 glucose tablets       6 saltine crackers         2 tubes glucose 40%<br>oral gel       8 oz. skim milk |  | <ul> <li>If next meal &gt; 2 hrs. also<br/>administer (choose one):</li> <li>1 sandwich (30g)</li> <li>3 graham crackers<br/>with one tbsp. peanut<br/>butter</li> </ul>  | <ul> <li>Recheck BG q15 min<br/>following treatment<br/>and continue to treat<br/>accordingly until ≥ 80<br/>mg/dL</li> <li>Once BG ≥ 80 mg/dL,<br/>recheck BG q1h x 4,<br/>then q 4h for a</li> </ul> |  |
|               | Call House Officer to report B   | G and action taken   |   | minimum of 24h   |  |

\*Choose one item from one column based on next mealtime. If the next meal is 1-2 hours away, include complex carbohydrate as suggested by the examples. If the next meal is > 2 hours away, include protein as suggested by examples.

\*\*Orange juice not appropriate for patients with renal dysfunction or patients at risk for hypoglycemia.

#### Hypoglycemia Guideline Example - Eating

Guideline - recommended Streamline Standardize Consistency

- Treatment:
  - If patient can eat,
  - po intake is preferred



# Hypoglycemia – Guideline Example – pt not eating

| Table 1: Pati | ents Who Are Not Alert or Who Are NPO   |   |
|---------------|---|---|
| BG Level*     | Action  | Follow-Up   |
| 60-79 mg/dL   | <ul> <li>Administer 7.5 g Dextrose D50% (15 ml) IV*</li> <li>Consider calling House Officer if patient<br/>experiences recurrent BG &lt; 70 mg/dL in past 12h</li> </ul>                                      | <ul> <li>Recheck BG q 15 min following treatment and<br/>continue to treat accordingly until ≥ 80 mg/dL.</li> </ul>                                   |
| 45-59 mg/dL   | <ul> <li>Administer 12.5 g Dextrose D50% (25 ml) IV*</li> <li>Call House Officer to report BG and action taken</li> </ul>   | <ul> <li>Once BG</li></ul>  |
| < 45 mg/dL    | <ul> <li>Administer 25 g Dextrose D50% (50 ml) IV*</li> <li>Call House Officer to report BG and action taken</li> </ul>   | with hypoglycemia should be monitored at least q<br>4h for a minimum of 24h.  |
| ALL           | <ul> <li>Consider adding Dextrose 5% to maintenance IV<br/>fluids at a rate          <u>&gt;</u> 50 ml/hr OR increasing rate of<br/>existing maintenance IV if dextrose source<br/>already present</li> </ul> | <ul> <li>If &gt; 4h from initial event and BG <u>&gt;</u> 80 mg/dL for tw<br/>consecutive readings, may consider reducing IV<br/>dextrose.</li> </ul> |

\*If IV access is not available, administer 1 mg glucagon IM and contact provider to obtain IV access. Repeat BG in 30 min.

#### Treatment:

- If patient unresponsive or NPO then IV dextrose given
- If prolonged hypoglycemia, consider dextrose infusion



# Steroid induced hyperglycemia in the hospital

16. Diabetes Care in the Hospital: Standards of Care in Diabetes—2023

Diabetes Care 2023;46(Suppl. 1):S267-S278 | https://doi.org/10.2337/dc23-S016



### **Steroids**

#### Patient presents with Steroid Induced Hyperglycemia after starting chemo.

Cc: N/V, Pain, Weight loss. On IV steroids and chemo, pt is NPO, with no previous Hx of DM

BS 250-300 mg/dL

What would you do?

- A. A1C, start SQ correction insulin
- B. A1C, start IV insulin drip
- C. A1C c/s DM team
- D. A1C start basal and correction insulin





#### Inpatient Glucose Management – Steroids





### Insulin Regimen for Steroids

- The insulin regimen should reflect the duration of the steroid
- Example: a shorter acting steroid like prednisone can be covered with increased prandial insulin or an injection of NPH insulin at the same time as the prednisone is administered
- Example: a longer acting steroid like dexamethasone will require increases in basal insulin or an IV insulin infusion
- Dose adjustments for a steroid taper
  - Insulin doses should be decreased at the same percentage that the steroids are being decreased
- Patients with pre-existing diabetes require insulin dose increases from their home doses, including prandial and correctional insulin up to 40 60% more.
- Whatever insulin orders are started, adjustments to insulin dosing will need to be made based on anticipated changes in glucocorticoid dosing and POC glucose test results.

Patient presents with Steroid Induced Hyperglycemia after starting chemo.

Cc: N/V, Pain, Weight loss. <u>On IV</u> steroids and chemo, pt is NPO, with no previous Hx of DM

BS 250-300 mg/dL

#### WWYD:

- A. A1C, start SQ correction insulin
- B. A1C, start IV insulin drip
- C. A1C c/s DM team
- D. A1C start basal and correction insulin

Answer: B and C Once on oral steroids and taper could switch SQ insulin basal/ increased prandial



Inpatient Glucose Management – Steroids

### Case 2 – the plan:



# DKA – Diabetic Ketoacidosis

# HHS – Hyperosmolar Hyperglycemic State

16. Diabetes Care in the Hospital: Standards of Care in Diabetes—2023

Diabetes Care 2023;46(Suppl. 1):S267-S278 | https://doi.org/10.2337/dc23-S016

### Summary PATHOPHYSIOLOGY DKA / HHNKS

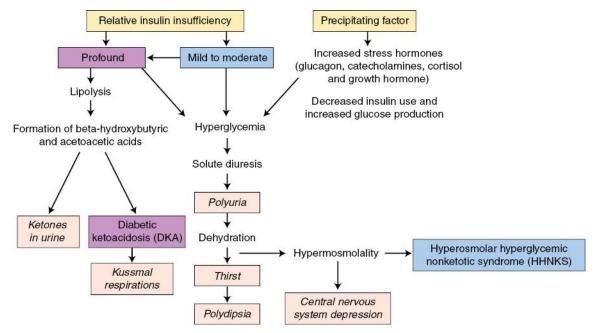


Figure 21-15 Pathophysiology of DKA and HHNKS in diabetes mellitus.

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# Comparison of DKA & HHS

|  | DKA      | HHS           |
|--|----------|---------------|
| Glucose  | >250     | >500          |
| *Osmolarity  | Variable | >350          |
| Urinary/ serum ketones<br>(BHB beta-hydroxybutyrate) | ++       | + or negative |
| рН   | < 7.3    | >7.3          |
| Bicarbonate  | <18      | >18           |
| **Anion Gap  | >12-15   | <15           |
| Precipitating illness                                | Yes      | Yes           |
| Mortality  | +        | ++            |
| Age  | Young    | Elderly       |

\*= osmolality= 2[ measured Na(mEq/L) ] +glucose/18 + BUN/2.8

\*\*Anion gap = (Na+) – ( $CI^- + HCO3^-$ )



# Diabetic Ketoacidosis (DKA)

**Profound Insulin deficiency** causes rapid mobilization of energy from stores in muscle and fat. These convert to **ketones**.

- Increased glucose causes osmotic diuresis which causes the loss of intravascular volume.-Dehydration
- Renal blood flow is then decreased and thus the kidney's ability to get rid of glucose is decreased
- Metabolic acidosis is caused
- The cycle keeps repeating until interrupted
- Prolonged acidosis can compromise CO and vascular tone and lead to CV collapse

Key features, hyperglycemia, Ketosis, acidosis, dehydration

**Present** 3Ps weak NV, fatigue, abd pain, fruity acetone breath, kussmaul respirations, tachycardia

**Precipitating** Infection, illness, not taking insulin, acute event MI CVA, Stress, Surgery, Trauma, T1D evaluate if psychosocial challenges

Plan of Care Treatment key components:

- Monitoring
- Fluid resuscitation
- Insulin and dextrose infusion (insulin drip continues till DKA resolves even if BS normal)
- Electrolyte repletion K+Potassium should be replace prior to insulin infusion in patients who present with hypokalemia (<3.5 meq/L)</li>
- Treating underlying cause

#### LIFE THREATENING

### Hyperosmolar Hyperglycemic State (HHS)

- Can sometimes occur when patients get diagnosed w/ Type 2 DM or could be if a Type 2 patient gets ill
- Occurs usually in middle-aged or elderly
- Underlying RI or CHF are common and worsen prognosis
- Precipitating event often identifiable
- Partial insulin deficiency may initiate this and then there is decreased glucose utilization by muscle, fat, and liver. Meanwhile increased liver production of glucose
- Presence of small amount insulin prevents acidosis
- Marked dehydration occurs

Key features, hyperglycemia, usually no ketosis

Present Adherence to insulin, Infection, MI, CVA, Trauma

- Plan of Care Treatment key components:
- Very similar to management of DKA
  - Fluid resuscitation is the main priority
  - Insulin drip
  - Potassium Electrolyte management
  - Follow up and Education

### LIFE THREATENING



# SGLT2 Inhibitors and Euglycemic DKA

- The safety and efficacy of SGLT2 inhibitors **have not** been established in patients with T1D Not FDA approved T1D
  - Have been cases of acidosis diabetic ketoacidosis (DKA), in patients treated with SGLT2 inhibitors
- Requires emergency room visits or hospitalization to treat the ketoacidosis.

Usually mildly elevated glucose at less than 200 mg/d – Euglycemic

**Triggers:** acute illness, UTI, reduced intake, fasting, NPO, reduced insulin dose **Risk:** Lower carb intake, lower insulin, greater lipolysis, ketosis

Buschur, E. O., Buse, J. B., Cohan, P., Diner, J. C., & Hirsch, I. B. (2015). Euglycemic diabetic ketoacidosis: a potential complication of treatment with sodium–glucose cotransporter 2 inhibition. *Diabetes care*, *38*(9), 1687-1693. Plewa, M. C., Bryant, M., & King-Thiele, R. (2020). Euglycemic diabetic ketoacidosis. StatPearls Publishing; Peters, A. L.,



# **Euglycemic DKA**

| Dia | agnosis  |
|-----|--|
| :   | Blood glucose < 250 mg/dL with: pH < 7.3, serum bicarbonate < 18 mEq/L, and anion gap > 12<br>Consider checking B-hydroxybutyrate, pH, bicarbonate, and anion gap in patients with diabetes presenting with<br>nausea/vomiting, malaise, or shortness of breath who meet one or more of the risk factors below.    |
| Ris | sk Factors   |
| :   | Sodium-glucose transport 2 (SGLTs) inhibitors (empagliflozin, canagliflozin, dapagliflozin, Ertugliflozin)<br>Pregnancy  |
| :   | Alcoholic ketoacidosis will have a very similar presentation<br>Decreased caloric intake, glycogen storage disease, stress, chronic liver disease, pancreatitis, alcohol use, and cocaine<br>intoxication  |
| Ma  | anagement  |
| •   | Initiate insulin infusion and start titration of intravenous fluids utilizing two-bag system as described above but using the titration algorithm below to address the higher risk of hypoglycemia. Refer to the <u>IV Insulin Infusion Titration Protocols -</u><br><u>Table 3</u> per the Department of Pharmacy |
| •   | Minimum insulin infusion rate (e.g., 0.3 units/hr) should be maintained, increasing dextrose rather than holding insulin infusion  |
| •   | Maintain on insulin infusion at minimum rate until resolution of DKA: pH > 7.3, bicarbonate > 18 mEq/L, anion gap < 15 mEq/L   |
|     | <ul> <li>Resolution of DKA due to SGLT2s may take days due to the duration of action of these medications</li> <li>Consider endocrinology consult</li> </ul>   |

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# Medical Nutrition Therapy in the hospital Enteral – Tube Feedings Parenteral - TPN

16. Diabetes Care in the Hospital: Standards of Care in Diabetes—2023 Diabetes Care 2023;46(Suppl. 1):5267-5278 | https://doi.org/10.2337/dc23-5016



### **Insulin Regimen for Enteral – Tube Feedings**



- Regimen should include coverage of basal, prandial, and correctional needs
- Patients with T1D should receive basal insulin even if feedings are discontinued
- Enteral bolus feedings: Give regular or fast acting subcutaneously before each feeding
- Nocturnal tube feeding: NPH insulin administered with the initiation of feeding

### **Insulin Regimen for Parenteral - TPN**

- For patients receiving parenteral nutrition, regular insulin may be added to the solution, especially if >20 units of correctional insulin required in the past 24 hours.
- Recommended to start with 1 unit of regular insulin for every 10 grams of dextrose
- Adding insulin to the TPN bag is safest way to prevent hypoglycemia if TPN is stopped.
- Continue correctional insulin subcutaneously if needed for hyperglycemia



# NPO Perioperative Care

16. Diabetes Care in the Hospital: Standards of Care in Diabetes—2023 Diabetes Care 2023;46(Suppl. 1):5267-5278 | https://doi.org/10.2337/dc23-5016

A1C target less 8% for elective surgeries Peri-op target 100 – 180 mg/dL



#### NPO/Pre-op/Pre-procedure/ Peri-op Glycemic Management

Standards of Medical Care in Diabetes 2023

#### <u>ORALS</u>

Hold the day of SGLT-2 hold 3-4 days prior Resume once patient is eating Resume meds containing <u>metformin</u> 2 days after IV contrast dye Adjust <u>insulin doses</u> based on type of DM and clinical judgement <u>NPH or mixed insulin</u> Reduce 50% the morning of Reduce 20% evening prior

<u>Long acting</u> reduce 20% <u>Insulin pump</u> reduce 20% basal rates May need additional <u>short-acting insulin</u> consider correction factor Check BG frequently during first 24 hours post-procedure

If BG control suboptimal, close follow-up with PCP recommended

Standards of Medical Care in Diabetes 2023



#### Procedure area Hyperglycemia:

What would you do?

Pt presents to procedure / peri-op area this morning with hi BS above 300 mg/dL

- Bolus with SQ insulin using peri-procedure guideline or correction factor
- Consider labs, A1C
- Consider inpt admit prn and SQ insulin or insulin drip
- Consider fluid bolus if dehydrated
- Consider canceling if chronically elevated BG and elective procedure, refer provider to get better BG control

A1C target less 8% - elective surgery BG target 100 – 180 mg/dL within 4 hour of surgery





# Case 3

Standards of Medical Care in Diabetes 2023

VEXNER MEDICAL CENTER

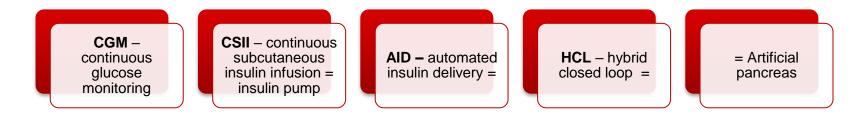
# Diabetes Technology in the hospital

7. Diabetes Technology: Standards of Care in Diabetes-2023

Diabetes Care 2023;46(Suppl. 1):S111-S127 | https://doi.org/10.2337/dc23-S007



# Compare and contrast between Types of DM technology – Terms:



*Goal:* AID, HCL, artificial pancreas all same use smart algorithms to auto adjust insulin delivery in pump

Goal more time in range (TIR)

Goal Less hyperglycemia and hypoglycemia excursions

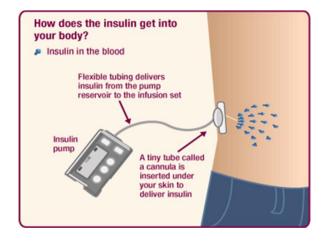
#### Continuous Subcutaneous Insulin Infusion (CSII) Technology

### Insulin Pumps Then....





### Insulin Pumps Now...



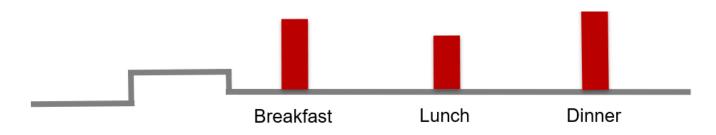






# **Insulin Pumps**

- Can be used alone or with a continuous glucose monitor
- A small mechanical device that continuously delivers insulin though a catheter that placed under the skin
  - Wear for 2-3 days
  - Delivers fast acting insulin two ways
    - Bolus (meal coverage and/or correction)
    - Basal (continuous background coverage- to act as long-acting insulin)





#### AID Insulin Pump Comparisons

|                                | <u>OmniPod ⁵</u>  | <u>Medtronic</u> 770G<br>"780G recently approved" | Tandem Control IQ                                   |  |
|--------------------------------|---|---|---|--|
| Integrated CGM                 | Dexcom G6   | Guardian 3  | Dexcom G6   |  |
| Baseline Basal<br>Patterns     | Based on insulin delivery hx Insulin delivery updates q6d |   | Programmed settings                                 |  |
| Algorithm adjust               | Based on CGM – 60 min predictive                          | 120 mg/dL   | Based on CGM – 30 min predictive                    |  |
| Algorithm and<br>Bolus targets | 110 – 150 mg/dL   | 120 – 150 mg/dL                                   | 110 – 160 mg/dL                                     |  |
| Temp Override                  | Activity 150 mg/dL  | Temp Target 150 mg/dL                             | Exercise 140 – 160 mg/dL<br>Sleep 112.5 – 120 mg/dL |  |
| Insulin Action                 | 2 – 6 hours   | 2 - 8 hours                                       | 5 hours   |  |

5-2023 FDA Approves iLet Beta Bionic Pump

### Insulin Pumps – in hospital order set recommended:



DO NOT TAKE OFF or STOP... Unless patient not alert to self manage



Need specialist to help manage to many devices for staff to know.

Need clear orders Insulin settings.

Need clear Documentation of sites and amount insulin Getting, BG Options to help with management (AID guidance still emerging):

Temporary basal rates

Suspend before low mode (Medtronic)

For automated insulin delivery:

Temporary target (Medtronic)
 Tandem: Exercise activity
 OP5: Activity



## Should have supplies at bedside

Change infusion Site every 3 dys

Caution hyperglycemia assess for kinks or any malfunctions ex: battery charger concerns.



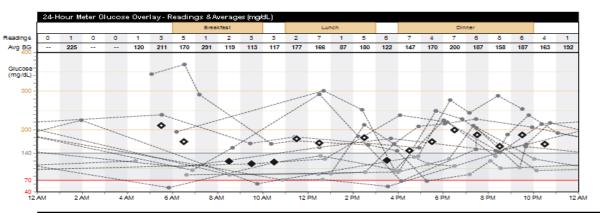
#### Disconnect for MRI, CT, if surgery more than 3 hours

Plan for alternate form of insulin if needed

# Insulin Pump: Medtronic Reports

Sensor & Meter Overview (1 of 2) 3/1/2013 - 3/13/2013

Generated: 4/17/2013 1:25:10 PM Data Sources: Paradigm Revel - 523 (443031)



| Statistics                | 3/1 - | 3/13     |
|---------------------------|-------|----------|
| Avg BG (mg/dL)            | 165 ± | :71      |
| BG Readings               | 83    | 6.8/dary |
| Readings Above Target     | 49    | 59%      |
| Readings Below Target     | 5     | 6%       |
| Sensor Avg (mg/dL)        |       |          |
| Avg AUC > 140 (mg/dL)     |       |          |
| Avg AUC < 70 (mg/dL)      |       |          |
|                           |       |          |
| Avg Daily Carbs (g)       | 184 ± | E 48     |
| Carbs/Bolus Insulin (g/U) | 9.8   |          |

Page 2 of 18

| Carbs/Bolds Fedin(g/o)      | a.o    |      |
|-----------------------------|--------|------|
| Avg Total Daily Insulin (U) | 35.1±4 | .9   |
| Avg Daily Basal (U)         | 16.3   | 47 % |
| Avg Daily Bolus (U)         | 18.8   | 53%  |

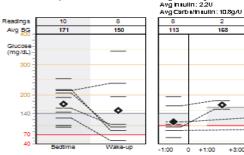
#### Meter Glucose Overlay Bedtime to Wake-Up and Meal Periods - Readings & Averages (mg/dL)

Meals Analyzed: 8

Avg Carb II: 23g







Breakfaut: 6:00 AM - 10:00 AM Lunch: 1100 AM - 300 PM Meals Analyzed: 17 Avg Carb #: 38g Avg In Julin: 390 Avg Carbi/Iniulin: 9.7g/U 12 8 2 168 106 155 145 ٥ -

-1:00

0 +1:00

+3:00 +5:00

Dinner: 400 PM - 1000 PM Meal # Analyzed : 26

9

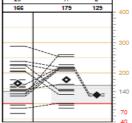
+3:00 +5:00

-1:00

144



0 +1:00



+3:00 +5:00

# Insulin Pump Omni-Pod Reports

|   | Tuesday  | Wednesday   | Thursday  | Friday  | Saturday   | Sunday  | Monday           |  |
|---|--|---|---|---|--|---|------------------|--|
| 25-   |  |   |   |   |  |   |                  | -  |
| 20-   |  |   |   | _   | _  |   |                  | 4  |
| 15-   |  |   |   |   |  |   |                  | _  |
| 10-   |  | -   |   |   |  |   |                  |  |
|   |  |   |   |   |  |   |                  |  |
| 5-  | -  |   |   |   |  |   |                  | -  |
| •   |  | 4 7   |   |   |  |   |                  |  |
|   | Tuesday  | Wednesday   | Thursday  | Friday  | Saturday   | Sunday  | Monday           | Total/Summar                                   |
| Correction  | 0.90   | 0.80  | 2.00  | 1.00  | 5.95   | 4.45  |                  | 15.10  |
| Meal Bolus  | 9.25   | 10.40   | 8.30  | 8.95  | 8.70   | 5.80  |                  | 51.40  |
| Total Bolus   | 10.15  | 11.20   | 10.30   | 9.95  | 14.65  | 10.25   |                  | 66.50  |
| Basal   | 11.75  | 12.00   | 10.60   | 10.70   | 10.65  | 11.70   |                  | 67.40  |
| Total Insulin   | 21.90  | 23.20   | 20.90   | 20.65   | 25.30  | 21.95   |                  | 133.90   |
| Avg/Day Carbs<br>(g)  | 187  | 181   | 148   | 148   | 134  | 98  |                  | 149  |
|   |  |   |   |   |  |   |                  |  |
| In  | sulin in %<br>Tuesday  | Wednesday   | Thursday  | Friday  | Saturday   | Sunday  | Monday           |  |
| in  |  | Wednesday   | Thursday  | Friday  | Saturday   | Sunday  | Monday           | ]  |
| n<br>Daily Insulin in   |  | Wednesday   | Thursday  | Friday  | Saturday   | Sunday  | Monday<br>Monday | ] Total/Summar                                 |
|   | Tuesday  |   |   | 0   |  |   |                  | Total/Summar<br>11%                            |
| Daily Insulin in<br>Corr Bol%<br>Meal Bolus%  | Tuesday<br>Tuesday<br>4%<br>42%                                | Wednesday<br>3%<br>45%                                  | Thursday  | Friday<br>5%<br>43%                               | Saturday<br>24%<br>34%                                 | Sunday<br>20%<br>26%                                |                  | 11%<br>38%                                     |
| Daily Insulin in<br>Corr Bol%   | Tuesday<br>Tuesday<br>4%                                       | Wednesday<br>3%   | Thursday<br>10%   | Friday<br>5%                                      | Saturday<br>24%  | Sunday<br>20%                                       |                  | 11%  |
| Daily Insulin in<br>Corr Bot%<br>Meal Bolus%<br>Basal%<br>Glucose<br>Statistics                                 | Tuesday<br>Tuesday<br>4%<br>42%                                | Wednesday<br>3%<br>45%                                  | Thursday<br>10%<br>40%                                  | Friday<br>5%<br>43%                               | Saturday<br>24%<br>34%                                 | Sunday<br>20%<br>26%                                |                  | 11%<br>38%                                     |
| Daily Insulin in<br>Corr Bol%<br>Meal Bolus%<br>Basal%<br>Glucose   | Tuesday<br>4%<br>42%<br>54%                                    | Wednesday<br>3%<br>45%<br>52%                           | Thursday<br>10%<br>40%<br>50%                           | Friday<br>5%<br>43%<br>52%                        | Salurday<br>24%<br>34%<br>42%                          | Sunday<br>20%<br>26%<br>54%                         | Monday           | 11%<br>38%<br>51%                              |
| Daily Insulin in<br>Corr Bol%<br>Meal Bolus%<br>Basal%<br>Glucose<br>Statistics<br>(mg/dL)                      | Tuesday<br>Tuesday<br>4%<br>42%<br>54%<br>Tuesday              | Wednesday<br>3%<br>45%<br>52%<br>Wednesday              | Thursday<br>10%<br>40%<br>50%<br>Thursday               | Friday<br>5%<br>43%<br>52%<br>Friday              | Saturday<br>24%<br>34%<br>42%<br>Saturday              | Sunday<br>20%<br>26%<br>54%<br>Sunday               | Monday           | 11%<br>38%<br>51%<br>Total/Summar              |
| Daily Insulin in<br>Corr Bol%<br>Meal Bolus%<br>Basal%<br>Glucose<br>Statistics<br>(mg/dL)<br>Highest           | Tuesday<br>Tuesday<br>4%<br>42%<br>54%<br>Tuesday<br>146       | Wednesday<br>3%<br>45%<br>52%<br>Wednesday<br>140       | Thursday<br>10%<br>40%<br>50%<br>Thursday<br>178        | Friday<br>5%<br>43%<br>52%<br>Friday<br>171       | Saturday<br>24%<br>34%<br>42%<br>Saturday<br>275       | Sunday<br>20%<br>26%<br>54%<br>Sunday<br>219        | Monday           | 11% 38% 51% Total/Summar                       |
| Daily Insulin in<br>Corr Bol%<br>Meal Bolus%<br>Basal%<br>Glucose<br>Statistics<br>(mg/dL)<br>Highest<br>Lowest | Tuesday<br>Tuesday<br>4%<br>42%<br>54%<br>Tuesday<br>146<br>57 | Wednesday<br>3%<br>45%<br>52%<br>Wednesday<br>140<br>70 | Thursday<br>10%<br>40%<br>50%<br>Thursday<br>178<br>116 | Friday<br>5%<br>43%<br>52%<br>Friday<br>171<br>81 | Saturday<br>24%<br>34%<br>42%<br>Saturday<br>275<br>85 | Sunday<br>20%<br>26%<br>54%<br>Sunday<br>219<br>104 | Monday           | 11%<br>38%<br>51%<br>Total/Summar<br>275<br>57 |

### Hybrid Closed Loop Insulin Pumps AID Automated Insulin Delivery

- Omnipod 5
- Tandem Control IQ
- Medtronic 780G





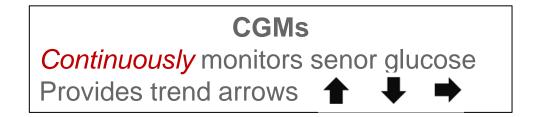




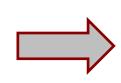
#### Continuous Glucose Monitoring (CGM) Technology

CGM should be considered in all children and adolescents with **type 1 diabetes**, whether using injections or continuous subcutaneous insulin infusion, as an additional tool to help improve glucose control.

CGMs in conjunction with insulin therapy are useful tools to lower A1C and/or reduce hypoglycemia in adults with **type 2 diabetes** who are not meeting glycemic targets.









If this was your meter reading, what would your next step be?

Тһеп.... АС &НS

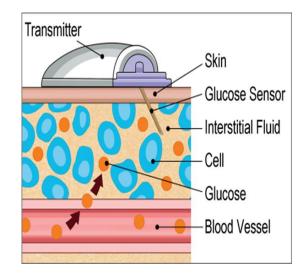
How would you respond if you saw this instead?

Now .... real time

# **Continuous Glucose Monitors**

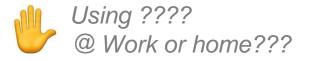
- Sensor placed under skin held in place with adhesive
- Continuously measures glucose levels in the interstitial fluid
- <u>Transmitter</u> connected to sensor sends data to <u>receiver</u> or compatible smart device displays reading every 1-5 min
- Can be used alone or with an insulin pump
- Provides downloads, reports to analyze patterns
- Optional Alert Features:
  - High or low sensor glucose alarms
  - Rising or dropping sensor glucose
  - Temporarily suspends insulin (Medtronic & T-slim)





# CGMs on the Market Now

- Freestyle Libre 14 day, 2, 3
- Dexcom G6, G7
- Eversense
- Medtronic Guardian













### THE OHIO STATE UNIVERSITY

WEXNER MEDICAL CENTER

# **Example Guideline**

Policy Name: Continuous Subcutaneous Insulin Infusion (CSII) Pumps and Continuous Glucose Monitoring (CGM)



<u>CGM not FDA approved in hospital</u> COVID – MICU – emergency use Definitions

| Term | Definition                                    |
|------|---|
| CGM  | Continuous Glucose Monitor                    |
| CSII | Continuous Subcutaneous Insulin Infusion Pump |

|                                  | Acetaminophen<br>interference | Calibrations required | Confirmatory<br>fingersticks<br>required prior to<br>treatment<br>(outpatient use)* | Remove for CT          | Remove for<br>MRI or<br>diathermy or<br>local<br>electricautery | Remove for x-<br>ray   |
|----------------------------------|-------------------------------|-----------------------|---|------------------------|---|------------------------|
| Dexcom G6®                       | N                             | N                     | N   | N                      | Y   | N                      |
| Medtronic©                       | Y                             | Y                     | Y   | N                      | Y   | N                      |
| Freestyle Libre™                 | N                             | N                     | N   | N                      | Y   | N                      |
| Freestyle Libre 2™               | N                             | N                     | N   | N                      | Y   | N                      |
| Eversense®<br>(surgical implant) | Not specified                 | Y                     | Y   | Y (remove transmitter) | Y (remove<br>sensor +<br>transmitter)                           | Y (remove transmitter) |

- CGM can have some variability esp. if glucose rapidly changing, lag.
- Interference: uric acid, galactose, xylose, ascorbic acid...
- For now: keep CGM on, still BG check, studies pending....
- · Can view reports possibly help with adjustments for discharge planning.

### For reference CGM....

#### **Resource 1. CGM Systems**



|                                  | FreeStyle<br>Libre 14<br>day®                        | FreeStyle<br>Libre 2ª                             | FreeStyle<br>Libre 3º           | Dexcom G6 <sup>10</sup>                   | Dexcom G7 <sup>11</sup>   | Medtronic<br>Guardian<br>Connect <sup>12,14</sup> | Eversense E3<br>CGM <sup>7</sup>    |
|----------------------------------|--|---|---------------------------------|---|---|---|-------------------------------------|
| Age                              | 18 and up  | 4 years and up                                    | 4 years and up                  | 2 years and up                            | 2 years and up  | 14-75 years<br>old                                | 14 years and<br>up                  |
| Wear Time                        | 14 days  | 14 days   | 14 days                         | 10 days                                   | 10 days   | 7 days  | 180 days                            |
| Calibration                      | No Need  | No Need   | No Need                         | No Need                                   | No Need   | 2x/day  | 2x/day                              |
| Insertion Site                   | Back of arm  | Back of arm                                       | Back of arm                     | Abdomen or<br>upper buttocks              | Upper arm or<br>abdomen (age<br>2+) or upper<br>buttocks (age<br>2-6) | Abdomen or<br>back of the<br>arm                  | Arm implant                         |
| Hyper/Hypoglycemic<br>Alerts     | No   | Yes   | Yes                             | Yes                                       | Yes   | Yes   | Yes                                 |
| Rapid Change in<br>Blood Glucose | Less<br>reliable                                     | Reliable  | Most reliable                   | More reliable,<br>30 minute<br>prediction | Most reliable,<br>30 minute<br>prediction                             | More reliable,<br>60 minute<br>prediction         | Reliable                            |
| Data Transmission                | Every 1<br>minute,<br>must scan<br>within 8<br>hours | Every 1<br>minute, must<br>scan within 8<br>hours | Every 1 minute<br>(no scanning) | Every 5<br>minutes via<br>Bluetooth       | Every 5<br>minutes via<br>Bluetooth                                   | Every 5<br>minutes via<br>Bluetooth               | Every 5<br>minutes via<br>Bluetooth |
| Links to App                     | +  | +   | +                               | +   | +   | +   | +                                   |

Cardi-OH-Beyond-the-A1C-Targets-for-Blood-Glucose-and-Methods-of-Measurement.

# PATTERN MANAGEMENT



- 1. Review medication taking behaviors.
- 2. Assess meal times, snacks, particularly overnight.
- 3. Assess overall glycemic status (TIR, mean glucose).9
- 4. Address hypoglycemia first if Time below Range (TBR) is above target.<sup>9</sup>
- 5. Address AM/fasting glucose.
- 6. Assess non-fasting glucose.
- 7. Evaluate patterns related to physical activity or work.

# CGM Results



- Continuous glucose monitoring (CGM) results in
  - Reduction in HbA1C<sup>1,2</sup>
  - Improved percentage of Time in Range (TIR), defined as 70-180 mg/dL<sup>2</sup>
  - Lower risk of hypoglycemia<sup>2</sup>
  - High patient satisfaction<sup>2,3</sup>
  - Lower risk of diabetes-related hospitalizations<sup>4,5</sup>
- Increasingly utilized in primary care practices as coverage and access expands.<sup>6</sup>

### CGM Advantages

- Ability to recognize early glucose fluctuations
- Detect unknown glucose excursions
- Alarms for readings out of range
- Answers the question "what is going on with my glucose?"

### CGM Disadvantages

- Cost
- Discrepancies from glucose readings
- 2-3 additional equipment pieces
- "Overwhelmed" with data
- Alarm fatigue
- Skin irritation



### **GLYCEMIC TARGETS**

## **Standardized CGM Metrics**

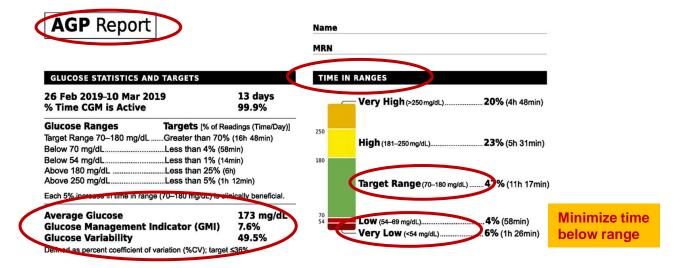
#### Table 6.2-Standardized CGM metrics for clinical care Number of days CGM device is worn (recommend 14 days) 2. Percentage of time CGM device is active (recommend 70% of data from 14 days) 3. Mean glucose 4. Glucose management indicator Glycemic variability (%CV) target ≤36%\* 6. TAR: % of readings and time >250 mg/dL (>13.9 mmol/L) Level 2 hyperglycemia 7. TAR: % of readings and time 181-250 mg/dL (10.1-13.9 mmol/L) Level 1 hyperglycemia TIR: % of readings and time 70–180 mg/dL (3.9–10.0 mmol/L) In range 9. TBR: % of readings and time 54-69 mg/dL (3.0-3.8 mmol/L) Level 1 hypoglycemia 10. TBR: % of readings and time <54 mg/dL (<3.0 mmol/L) Level 2 hypoglycemia CGM, continuous glucose monitoring; CV, coefficient of variation; TAR, time above range;

CGM, continuous glucose monitoring; CV, coefficient of variation; TAR, time above range; TBR, time below range; TIR, time in range. \*Some studies suggest that lower %CV targets (<33%) provide additional protection against hypoglycemia for those receiving insulin or sulfonylureas. Adapted from Battelino et al. (35).



#### Standardized Report Interpret BG data Ex: EKG can use different machine but reading same How download reports variable - need to standardize

**GLYCEMIC TARGETS** 



AGP ambulatory glucose profile TIR Time in Range 70-180 mg/dL GMI Glucose Management Indicator (estimate A1C from CGM)

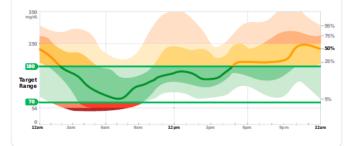
### **GLYCEMIC TARGETS**

#### AGP Report: Continuous Glucose Monitoring

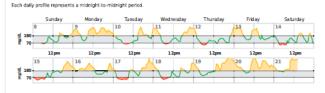


#### Ambulatory Glucose Profile (AGP)

AGP is a summary of glucose values from the report period, with median (50%) and other percentiles shown as if they occurred in a single day.



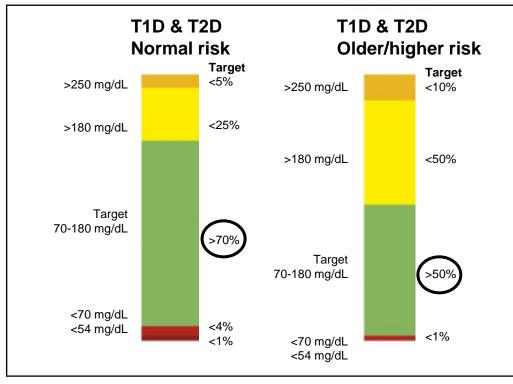
#### **Daily Glucose Profiles**



Glycemic Targets: Standards of Care in Diabetes - 2023. Diabetes Care 2023;46(Suppl. 1):S97-S110



# CGM Goals International Consensus on TIR



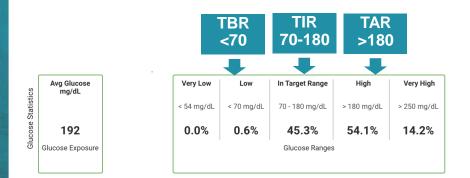


- The goal TIR for most individuals is 70% with 4% TBR and 1% of time below 54 mg/dl
- The goal TIR for older or higher risk individuals is 50% with <1% TBR</li>

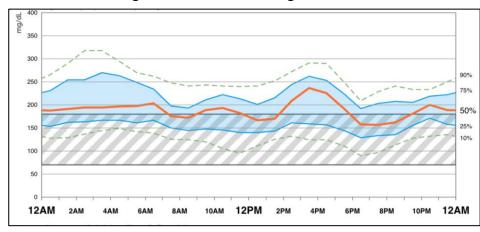
Use to guide insulin dose adjustments Reevaluate treatment plan:

T1D: Type 1 Diabetes; T2D: Type 2 Diabetes

# Ambulatory Glucose Profile



#### Check that target set to 70-180 mg/dL





% Time CGM

Active

95.2%

Data Sufficiency Cari

Goal

>70%

over 14

days

TIR (Time in Range)

**CV Goal** 

<36%

Coefficient of

Variation

30.3%

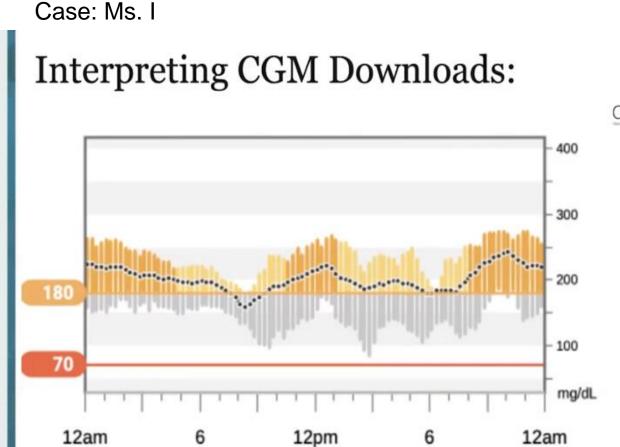
Glucose Variability

SD

mg/dL

58

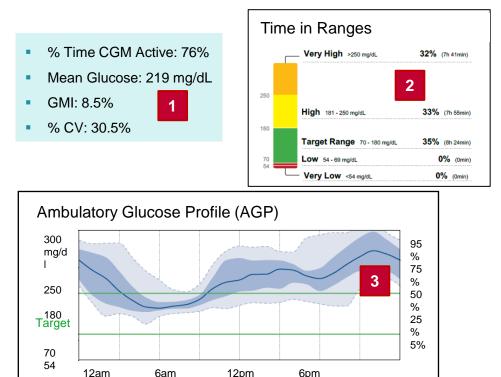
- TAR (Time above Range)
- TBR (Time below Range)





https://www.cardi-oh.org/echo/Fall-2022-Continous-Glucose-Monitoring

# Ms. O. CGM Case: Need for Prandial Insulin





#### **Patient Summary**

- 52-year-old female with T2D, no complications
- Weight: 90 kg

#### **Current Treatment**

- Metformin, Glimepiride
- Dulaglutide 1.5 mg weekly
- Glargine 60 units daily

#### CGM Interpretation (red boxes □)

- (1) Adequate amount of data
- (2) TIR is 35% (goal >70%)
- (3) Pattern is predominantly post-prandial hyperglycemia

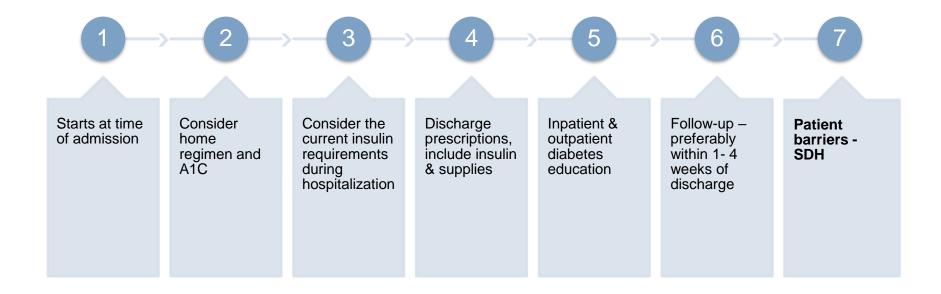
#### Plan

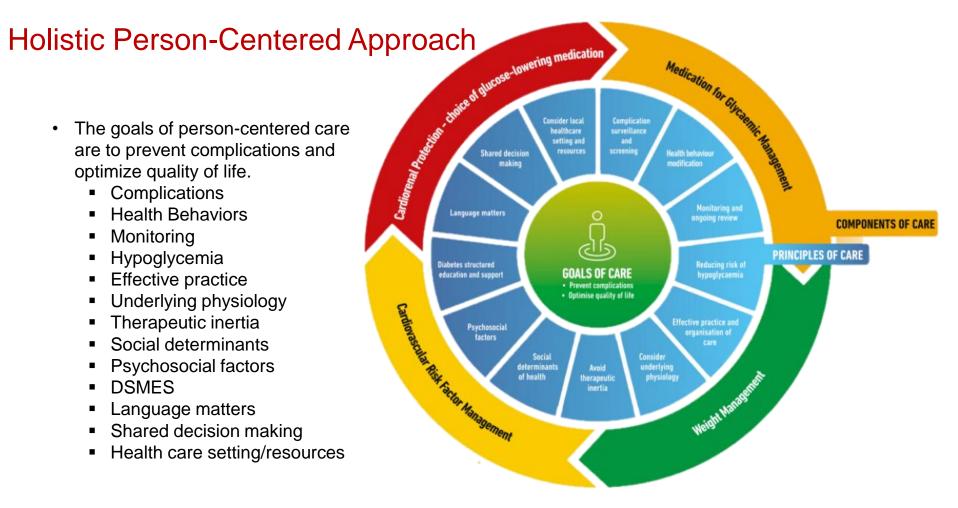
- Titrate dulaglutide to 3 to 4.5 mg
- Stop glimepiride
- Start prandial insulin at largest meal of the day
- Do not increase basal insulin because the dose is already >0.5 unit/kg and there is a high bedtime to morning differential.<sup>10</sup>

12am

## **Discharge Planning**

### **Discharge Planning**





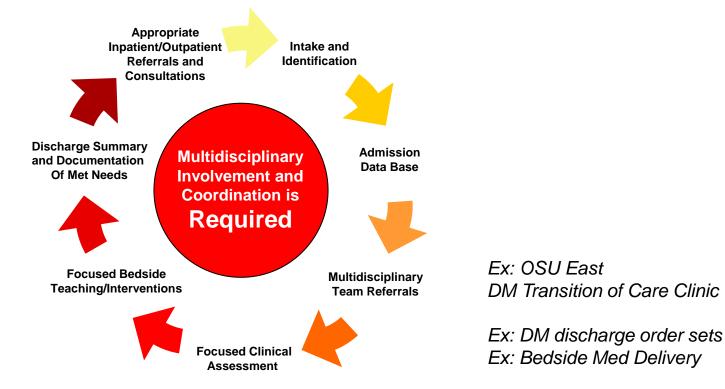
#### Be aware of resources in your area to address health disparities:

### **Social Determinants of Health**

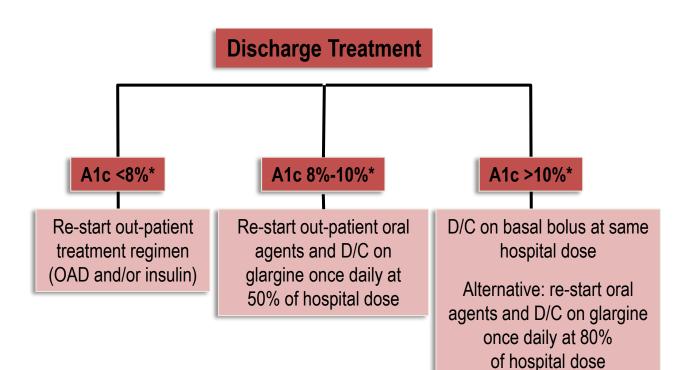
| Economic<br>Stability  | Neighborhood<br>/Environment  | Education  | Food  | Community/<br>Social Context  | Health Care<br>System  |
|--|---|--|---|---|--|
| <ul> <li>Environment</li> <li>Income</li> <li>Expenses</li> <li>Debt</li> <li>Medical bills<br/>Support</li> </ul> | <ul> <li>Housing</li> <li>Transportation</li> <li>Safety</li> <li>Parks</li> <li>Playgrounds</li> <li>Walkability</li> <li>Zip Code</li> <li>Geography</li> </ul> | <ul> <li>Literacy</li> <li>Language</li> <li>Early childhood<br/>education</li> <li>Vocational<br/>training</li> <li>Higher<br/>Education</li> </ul> | <ul> <li>Hunger</li> <li>Access to<br/>healthy<br/>options</li> </ul> | <ul> <li>Social<br/>integration</li> <li>Support<br/>systems</li> <li>Community<br/>engagement</li> <li>Discrimination</li> <li>Stress</li> </ul> | <ul> <li>Healthcare<br/>coverage</li> <li>Provider<br/>availability</li> <li>Provider<br/>linguistic and<br/>cultural<br/>competency</li> <li>Quality of<br/>care</li> </ul> |
| Health Outcomes  |   |  |   |   |  |



### Connecting Inpatient to Outpatient Support: Multidisciplinary Circle of Care



### **Revised Discharge Algorithm**



\*Use admission A1c to adjust therapy at discharge

### **Cautions and Contraindications to Oral Medications**

| Diabetes Medication Class | Drug Examples (not all<br>encompassing*)  | Cautions<br>(not all encompassing*)  | Contraindications<br>(not all encompassing*)             |
|---------------------------|---|--|--|
| Biguanides                | Metformin   | CHF, alcohol abuse   | eGFR < 46, lactic acidosis,<br>hepatic disease           |
| DPP4-Inhibitors           | Sitagliptin (Januvia)<br>Linagliptin (Tradjenta)  | CrCl < 45  | History of pancreatitis                                  |
| GLP-1 Agonists            | Exenatide (Bydureon)<br>Dulaglutide (Trulicity)<br>Liraglutide (Victoza)<br>Semaglutide (Ozempic) | Pancreatitis history, GI disease,<br>gastroparesis, renal impairment                           | Medullary thyroid carcinoma<br>history or family history |
| SGLT2 Inhibitors          | Empagliflozin (Jardiance)<br>Dapagliflozin (Farxiga)<br>Canagliflozin (Invokana)                  | eGFR 45-59, hypotension, CHF,<br>ketoacidosis risk, elderly, UTI or<br>yeast infection history | DKA, eGFR < 45   |
| Sulfonylureas             | Glimepiride   | Renal impairment, hepatic  | CrCl <50   |



What's supported in guidelines and can continue to be developed for inpatient glycemic control :

- Use of technology insulin pumps and continuous glucose monitors
- DM self management, self administer insulin and glucose monitoring
- Insulin dosing calculator to stream-line Lispro dosing process for nurses
- DM teams of specialists and educators
- Uncomplicated DKA treated with SQ insulin and fluids in ED and on monitored units with protocols in place
- Medication reconciliation with prior to arrival and hospital meds
- Receiving prescriptions prior to discharge



Before Closing .... Resources



Diabetes Med & Insulin PocketCards <u>https://diabetesed.net/pocket-cards-insulin-and-</u> <u>diabetes-medication/</u> Diabetes Education Services (DES)

| Drug Class  | Preferred   |
|---|---|
| Non-Insulin   |   |
| Metformin and combination                               | <ul> <li>Metformin in combination with         <ul> <li>Pioglitazone</li> <li>Glyburide</li> <li>Canagliflozin, empagliflozin</li> <li>Sitagliptin, linagliptin</li> <li>Repaglinide</li> </ul> </li> <li>Metformin ER (Glucophage XR)</li> </ul> |
| Sulphonylurea<br>SFU                                    | glimepiride, glipizide, glyburide   |
| Glucagon-like peptide-1<br>receptor agonist<br>GLP-1 RA | Byetta (exenatide), Trulicity (dulaglutide),<br>Victoza (liraglutide)   |
| Sodium-glucose<br>cotransporter-2 inhibitor<br>SGLT2i   | Farxiga (dapagliflozin), Invokana (canagliflozin),<br>Jardiance (empagliflozin)   |
| Dipeptidyl peptidase-4 inhibitor<br>DPP-4i              | Januvia (sitagliptin), Tradjenta (linagliptin)  |
| Thiazolidinedione<br>TZD                                | pioglitazone  |
| Alpha glucosidase inhibitor<br>AGI                      | acarbose, miglitol  |
| Glinide   | nateglinide, repaglinide  |

Table 1. 2022 Ohio Medicaid Preferred Diabetes Formulary As of July 2022

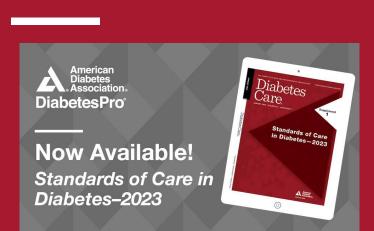
# CARDI-OH One Cardovacular and Diabetes Health Collaborative

| • | No step therapy is required for most |
|---|--------------------------------------|
|   | medications on formulary             |

 Continuous glucose monitors are now covered without the need for prior authorization

| Insulin |  |
|---------|--|
| Basal   | Lantus (glargine), Levemir (detemir), Toujeo<br>(glargine U-300), Tresiba (degludec)⁵  |
| Bolus   | Apidra (glulisine), aspart, Humalog (lispro)<br>U-100, Humulin R (regular insulin) U-500, lispro,<br>Novolog (aspart) U100   |
| Premix  | Humalog 50/50 (lispro protamine/lispro),<br>Humalog 75/25 (lispro protamine/lispro),<br>Humulin 70/30 (insulin isophane/regular<br>insulin), aspart protamine/aspart, Novolog<br>70/30 (aspart protamine/aspart) |

<sup>s</sup> Step therapy





# Consumers guide to products: <u>https://consumerguide.diabetes.org/</u>



- Full version available
- Abridged version for PCPs
- Free app, with interactive tools
- Pocket card with key figures
- Free webcast for continuing education credit

#### Professional.Diabetes.org/SOC



### **Diabetes Technology Video** and **OSUWMC Diabetes Education Resouces**



To learn more about diabetes technology, please view our *Diabetes Technology Video*. This video will direct you through the OSUWMC process in obtaining a new insulin pump, continuous glucose monitor or smart pen.







### HOW TO REALLY MAKE IT WORK



Delivery arrangements How, where and by who care is delivered Coordination of care and management of care processes Information and communication technology

Governance arrangements: Accountability for health professionals Training and certification Quality of practice Implementation strategies Health system Health care setting Health care workers



Davies MJ, Aroda VR, Collins BS, Gabbay RA, Green J, Maruthur NM, Rosas SE, Del Prato S, Mathieu C, Mingrone G, Rossing P, Tankova T, Tsapas A, Buse JB



European Association for the Study of Diabetes

Diabetes Care 2022; https://doi.org/10.2337/dci22-0034. Diabetologia 2022; https://doi.org/10.1007/s00125-022-05787-2. Copyright ADA/EASD 2022

#### **INSURANCE COVERAGE\***

**Durable Medical Equipment** 

### **Type 1 Diabetes**

### **Type 2 Diabetes**

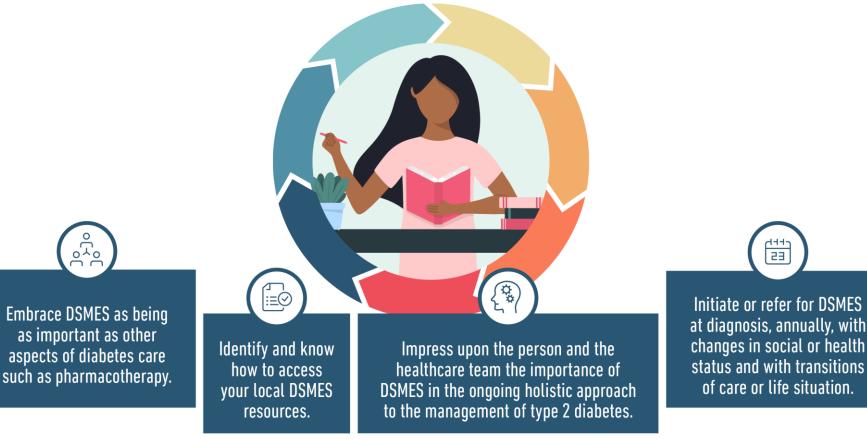
|           |  |                           | Insurance              | DME Pump              | Pharmacy                               |               |
|-----------|--|---------------------------|------------------------|-----------------------|--|---------------|
| Insurance | DME Pump<br>(Medtronic,<br>Tandem,<br>Beta Bionic) | Pharmacy<br>Pump<br>(OP5) | CGM                    | Top<br>urrent Private | (Medtronic,<br>Tandem, Beta<br>Bionic) | Pump<br>(OP5) |
| Private   | Yes  | Usually                   | Yes DME or<br>Pharmacy | Private               | Usually                                | Usually       |
| Medicare  | Yes  | Usually                   | Yes DME                | Medicare              | Yes with labs                          | Usually       |
| Medicaid  | Yes  | Yes                       | Yes -<br>Pharmacy      | Medicaid              | Yes                                    | Yes           |

| Insurance | Pump    | CGM         |
|-----------|---------|-------------|
| Private   | Usually | Usually     |
| Medicare  | Yes     | Dexcom Only |
| Medicaid  | Yes     | No          |

Bottom Yrs prior

| Insurance | Pump    | CGM    |
|-----------|---------|--------|
| Private   | Usually | Varies |
| Medicare  | No      | No     |
| Medicaid  | No      | No     |

### **DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT**





Davies MJ, Aroda VR, Collins BS, Gabbay RA, Green J, Maruthur NM, Rosas SE, Del Prato S, Mathieu C, Mingrone G, Rossing P, Tankova T, Tsapas A, Buse JB

Diabetes Care 2022; https://doi.org/10.2337/dci22-0034. Diabetologia 2022; https://doi.org/10.1007/s00125-022-05787-2.

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### Examples Lifestyle and DM Education Materials OSUWMC



https://go.osu.edu/pted3577



https://go.osu.edu/pted4524



# Person-Centered Language Recommendations

The ADA and the APA recommend language that emphasizes inclusivity and respect:

- <u>Gender</u>: Gender is a social construct and social identity; use term "gender" when referring to people as a social group. Sex refers to biological sex assignment; use term "assigned sex" when referring to the biological distinction.
- <u>Race</u>: Race is a social construct that is used broadly to categorize people based on physical characteristics, behaviors, and geographic location. Race is not a proxy for biology or genetics. Examining health access, quality, and outcome data by allows the healthcare system to assist in addressing the factors contributing to inequity.
- <u>Sexual Orientation</u>: Use the term "sexual orientation" rather than "sexual preference" or "sexual identity." People choose partners regardless of their sexual orientation; however, sexual orientation is not a choice.
- **Disability**: The nature of a disability should be indicated when it is relevant. Disability language should maintain the integrity of the individual. Language should convey the expressed preference of the person with the disability.
- <u>Socioeconomic Status</u>: When reporting SES, provide detailed information about a person's income, education, and occupation/employment. Avoid using pejorative and generalizing terms, such as "the homeless" or "poor."
- **<u>Violent Language</u>**: Avoid sayings like 'killing it,' 'pull the trigger,' 'take a stab at it,' 'off the reservation,' etc.

Flanagin A et al., 2021, JAMA; Dickinson JK et al., Diabetes Care, 2017; American Psychological Association, 2021; ODM, 2021.

CARDI•**oh** 

# Additional Cardi-OH Resources



Beyond the A1C: Targets for Blood Glucose and Methods of Measurement cardi-oh.org/best-practices/diabetes-management/beyond-the-a1c-targets-for-blood-glucose-andmethods-of-measurement

 Outpatient Diabetes Management for Primary Care Providers. Medications Intensification and Algorithm cardi-oh.org/best-practices/diabetes-management/outpatient-diabetes-management-for-primarycare-providers-medications-intensification-and-algorithm

 Optimizing the Telehealth Diabetes Visit: Glucose Monitoring Data <u>cardi-oh.org/best-practices/diabetes-management/optimizing-the-telehealth-diabetes-visit</u> (reimbursement codes, office, logistics, downloads)



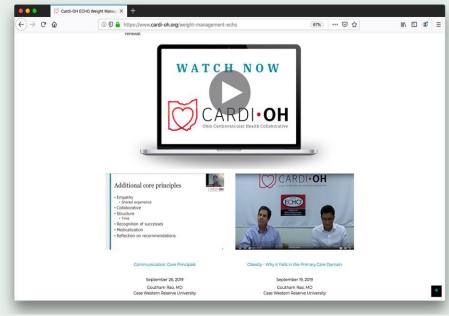
### Watch Cardi-OH TeleECHO Clinic Recordings



Register on Cardi-OH.org to watch the current clinic: <u>cardi-oh.org/user/register</u>

Current Clinic page (access recordings): <u>cardi-oh.org/echo/cardiovascular-prevention-fall-2022</u>

Access previous Cardi-OH TeleECHO Clinics: <u>cardi-oh.org/echo/archived-clinics</u>



# Future Therapies



- Once weekly basal insulin (lcodec)
- Glucose responsive insulin
- Combined peptides: GLP-1/GIP, GLP-1/glucagon receptor dual agonist, GLP-1/glucagon/GIP
- Others
  - Glucagon receptor antagonist
  - G-protein-coupled receptor ligands
  - Hormone/enzyme/receptors
  - O PPARs: insulin sensitizers
  - $\,\circ\,$  Glimins: correction of mitochondrial dysfunction

# Future Approaches



- Adult-onset DM sub-types<sup>1</sup>
- Precision medicine:<sup>2</sup>
  - Patient-level markers predict response to therapy, complications
  - $\circ$  Emphasis on clinical utility, equity
- Early combination therapy in some patients at treatment initiation to extend the time to treatment failure.<sup>3,4</sup>
- Connected devices for monitoring and treatment

<sup>1.</sup> Ahlqvist et al. Lancet Diabetes Endocrinol. 2018;6(5):361-369.

<sup>2.</sup> Nolan et al. ADA/EASD Precision Medicine in Diabetes Initiative. Diabetes Care. 2022;45(2):261-266.

<sup>3.</sup> Davies et al. ADA Standards of Care. Dia Care 2022;45(Suppl. 1):S125–S143.

<sup>4.</sup> Garber et al. AACE Consensus Statement. Endocr Pract 2019;25(1):69-100.

### Most Recent Diabetes and Technology Guidelines

| Professional Group(s)       | Year          | Title  |
|-----------------------------|---------------|--|
| AACE                        | 2023          | American Association of Clinical Endocrinology Clinical Practice Guideline: Comprehensive Type 2<br>Diabetes Management Algorithm  |
| ADA                         | 2023          | Standards of Medical Care in Diabetes  |
| ADA/EASD                    | 2022          | Management of hyperglycemia in Type 2 Diabetes. A consensus report by the American Diabetes Association and the European Association for the Study of Diabetes EASD            |
| AACE                        | 2021          | Advanced Diabetes Technology Guideline   |
| ADA/EASD                    | 2021          | The Management of Type 1 Diabetes in Adults. A Consensus<br>Report by the American Diabetes Association (ADA) and the<br>European Association for the Study of Diabetes (EASD) |
| EASD/ISPAD^                 | 2020          | Glucose management for exercise using continuous glucose monitoring (CGM) and intermittently scanned CGM (isCGM) systems in type 1 diabetes                                    |
| International<br>Consensus* | 2019          | Clinical Targets for Continuous Glucose Monitoring Data Interpretation: Recommendations From the International Consensus on Time in Range                                      |
| Endocrine Society           | 2017,<br>2018 | Endocrine Society Dosing Method Based on CGM Trend Arrows and CF   |
| EASD/ADA                    | 2017          | Improving the Clinical Value and Utility of CGM Systems: Issues and Recommendations  |

# Summary Key Take Aways

- Insulin is the preferred method of treating hyperglycemia in the hospital
- Standard protocols promote consistency in inpatient glucose management and facilitate high quality care
- Poor glycemic control in the hospital can lead to poor patient outcomes
- Treat patients with physiologic insulin including basal, bolus and correction factor and know the action of insulin
- Connect patients to outpatient follow up
- Expect continued advances in technology insulin pumps and continuous glucose monitors
- CSII and CGM can improve glucose management and there is consensus for monitoring and use of technology







# Thank you... Thank you very much....





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