#### Glytec



# Spotlight on Perioperative Glycemic Management

A Technology-Driven Approach

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#### Introduction



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## **Objectives**

- Identify the challenges of perioperative glycemic control
- Define goals for perioperative glycemic management
- Review perioperative diabetes medication management plans
  - Identify when glycemic management is best served using insulin infusion
  - Identify when glycemic management is best served utilizing subcutaneous injections
- Recognize which patient populations are at high risk for hypoglycemia



# CARE CHALLENGES

## Inpatient Glycemic Challenges

#### Hyperglycemia is common

- Occurs in > 30% of hospitalized patients & up to 40% in surgical patients
  - Severe illness and surgical stress predisposes patients to insulin resistance & hyperglycemia
- Effects of hyperglycemia:
  - Immune dysfunction
  - Impaired wound healing
  - Oxidative stress

#### Limited Support Structures

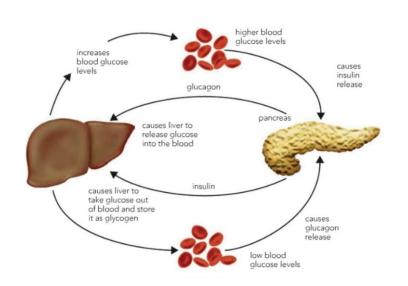
- Lack of inpatient experts: Limited endocrine support, lack CDCES support
- Inpatient prescribers varied knowledge and comfort in prescribing insulin
  - Insulin is a high-alert, high-risk medication

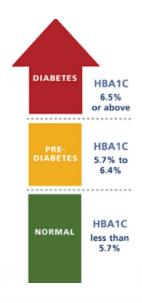
#### Challenges

- Glycemic control has focused in the critical care areas, with less focus in the general med-surg units
- Staff fear of hypoglycemia
- Patients fear insulin
- Dosing and care challenges:
  - Coordination of glucose testing, meal delivery, & insulin administration
    - Meals on demand
    - Food from home
    - Variable intakes

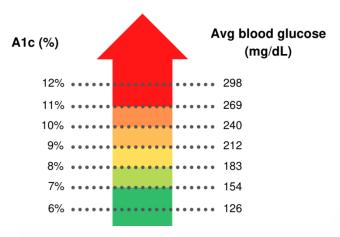


#### Perioperative Hyperglycemia = Adverse Outcomes





#### **Glycated Hemoglobin**



#### Glucose Control

- Glucose control is a finely regulated balance of hepatic glucose production and glucose utilization in peripheral tissues
  - Surgery and anesthesia alter this balance
- Studies show correction of hyperglycemia decreases mortality and reduces complications
  - Tight control may lead to hypoglycemia = increased mortality

#### Considerations

- Screen for Diabetes
- Long-term glycemic control
- Severity of hyperglycemia on admission
- Monitoring for post-operative hyperglycemia

#### Occurrence

- Peri-operative Hyperglycemia:
  - General surgery = 20-40%
  - Cardiac surgery = 80%
  - 12-30% do not have h/o diabetes = Stress Hyperglycemia





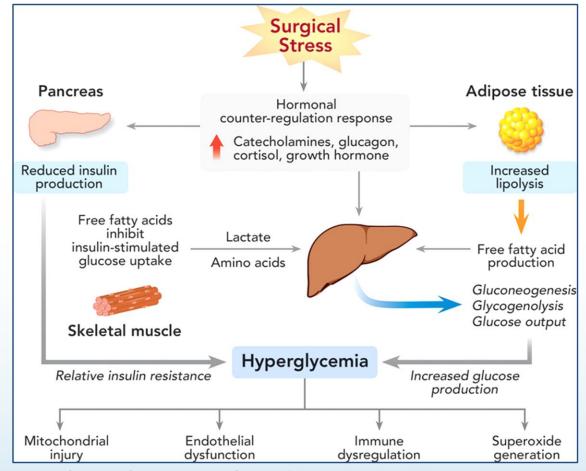
From: Perioperative Hyperglycemia Management An Update Anesthesiology. 2017;126(3):547-560. doi:10.1097/ALN.000000000001515

The magnitude of stress response relates to the severity of surgery & type of anesthesia

Thorax & abdomen General anesthesia

Laproscopic

Figure Legend: The surgical stress response



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# Enhanced Recovery After Surgery (ERAS)

- Consumption of carbohydrate-rich beverage up to 2 hours before surgery
- Preoperative carb loading
  - May counteract the state of insulin resistance (due to stress and starvation)
  - Avoids the catabolic state associated with starvation
  - Increase insulin sensitivity
  - Decreasing risk for post-op hyperglycemia
  - Associated with reduced LOS

# However, Providing pre-operative carbohydrate-containing beverages to patients with known diabetes is **not** recommended.

- https://www.endocrine.org/clinical-practice-guidelines/inpatienthyperglycemia-guideline-resources
  - Online June 2022





## Clinical Study Summary

- Preoperative glucose & mortality:
  - Retrospective analysis of 61,000 patients undergoing elective noncardiac surgery and mortality
    - Preoperative BG of 60-100 mg/dL: 3 to 5% 1-year mortality<sup>2</sup>
    - Preoperative BG >216 mg/dL: 12% mortality 1-year mortality<sup>2</sup>

- Preoperative glycemic control:
  - Preoperative A1c > 8% = wound complications (Type 2, total knee)<sup>2</sup>
  - Preoperative A1c >7% increase in infectious complications in major noncardiac surgery <sup>2</sup>
  - Preoperative A1c >7%
    higher unadjusted 5-yr
    mortality in elective CABG<sup>2</sup>

- Intraoperative glycemic control:
  - Most studies are cardiac surgery
    - For every 20 mg/dL above 100 mg/dL = 30% increase in adverse outcomes in cardiac surgery <sup>2</sup>
      - BG < 200 mg/dL = 13%,</li>>200 mg/dL = 36%,>250 mg/dL = 63%
    - Hypoglycemia = mortality risk



# Post-Operative Glycemic Management

- Inpatient hyperglycemia (> 180 mg/dL) = adverse clinical outcomes
  - Surgical site infections, delayed wound healing and increased length of stay
  - Those without prior history of diabetes (stress hyperglycemia) have been shown to have worse outcomes with elevated BG's
- Tighter control may be desired for some surgical patients but tighter control brings higher risk for hypoglycemia thus increased mortality
  - IV Glucommander very low risk for hypoglycemia



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  - IV Glucommander very low risk for hypoglycemia
  - 62.6% reduction in preventable hypoglycemia ADEs

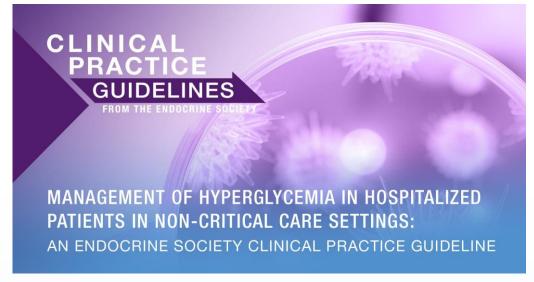


# RECOMMENDATIONS



#### A1c & Pre-Op BG

Patients with diabetes scheduled for elective surgery may have improved postoperative outcomes when preoperative hemoglobin A1c is ≤8% and when blood glucose values in the immediate pre-operative period are <180 mg/dL.

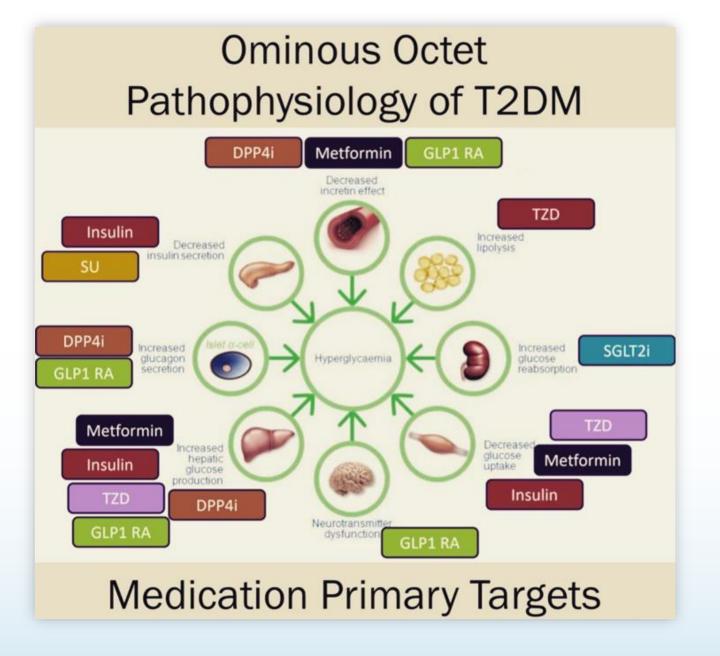


Society Guideline Recommendations for Treatment of Perioperative Hyperglycemia and Diabetes



|  | Ambulatory Surgery   | ICU  | Non-ICU  |
|--|--|--|--|
| SAMBA<br>(Society for Ambulatory<br>Anesthesia)                                  | SC rapid-acting insulin analogs are preferred over IV or SC regular insulin Treatment goal: intraoperative blood glucose levels < 180 mg/dl (10mM) |  |  |
| ADA/AACE (American Diabetes Association/American Association of Endocrinologist) |  | Initiate insulin therapy for glucose >180 mg/dl (10mM)  Treatment goal: For most patients, target a glucose level between 140 and 180 mg/dl (7.7-10mM).  Glucose target between 110 and 140 mg/dl (6.1-7.7mM) may be appropriate for select patients if achievable without significant risk for hypoglycemia | Treatment goal: If treated with insulin, premeal glucose targets should generally be < 140 mg/dl (<7.7mM), with random glucose levels < 180 mg/dl (10mM)   |
| ACP<br>(American College of<br>Physicians)                                       |  | Recommends against intensive insulin therapy in patients with or without diabetes in surgical/medical ICUs Treatment goal: Target glucose is between 140 and 200 mg/dl (7.7-11.1mM) in patients with or without diabetes   |  |
| Critical Care Society  |  | BG > 150 mg/dl (8.3 mM) should trigger insulin therapy<br>Treatment goal: Maintain glucose <150 mg/dl (8.3mM) for most patients in<br>ICU  |  |
| Endocrine Society  |  |  | Treatment goal: Target premeal blood glucose < 140 mg/dl (7.7 mM) and random glucose <180mg/dl (10mM)  Higher target glucose <200 mg/dl (11.1mM) is acceptable in patients with terminal illness and/ or with limited life expectancy or at high risk for hypoglycemia |
| Society of Thoracic Surgeons   |  | Continuous insulin infusion preferred over SC or intermittent IV boluses Treatment goal: Recommend glucose < 180 mg/dl (10mM) during surgery, <110 mg/dl (6.1mM) in fasting and premeal states   |  |
| Joint British Diabetes Societies   |  |  | Initiate insulin therapy for glucose> 10 mM (180 mg/dl)  Target blood qlucose levels in most patients are between 6 and 10mM (108-180mg/dl) with an acceptable range of between 4 and 12mM (72-216 mg/dl)  |
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- Type 1 is an insulin deficiency
- Type 2 is more complicated





## Pre-Op Glycemic Management

#### Considerations

- Type of Diabetes
- Nature and extent of the surgical procedure
- Length of pre-and post-op fasting
- State of control preceding surgery

#### Type 2 Diabetes

- Oral medications, most can be taken until the day before surgery
- Reduction in basal insulin

#### Type 1 Diabetes

- Do not hold basal insulin, may reduce dose. Hold prandial insulin morning of if NPO
- Stress of surgery may cause severe hyperglycemia or ketoacidosis



## Oral Medication Recommendations Prior to Surgery

| Oral Medication for<br>Elective Surgery | Day Before<br>Surgery | Day of Surgery if Normal Oral Intake<br>Anticipated Same Day and Minimally<br>Invasive Surgery | Day of Surgery if Reduced Postoperative<br>Oral Intake or Extensive Surgery, Anticipated<br>HD Changes and/or Fluid Shifts |
|---|-----------------------|--|--|
| Secretagogues<br>SGLT-2 Inhibitors      | Take<br>Hold          | Hold<br>Hold   | Hold<br>Hold   |
| Thiazolidinediones                      | Take                  | Take   | Hold   |
| Metformin                               | Take*                 | Take*  | Hold   |
| DPP-4 Inhibitors                        | Take                  | Take   | Take   |

\*Hold if patient having a procedure with intravenous contrast dye administration, particularly in those with glomerular filtration rate < 45 ml/min. 56
DPP = dipeptidyl peptidase-4; HD = hemodynamic; SGLT = sodium glucose cotransporter-2.



# Insulin Recommendations Prior to Surgery

| Day Dafara Curgani   | Glargine<br>or Detemir |                      | NPH or<br>70/30 Insulin |                      | Lispro, Aspart,<br>Glulisine, Regular |               | Noninsulin<br>Injectables                                   |   |
|--|------------------------|----------------------|-------------------------|----------------------|---------------------------------------|---------------|---|---|
| Day Before Surgery<br>Insulin Regimens   | ам Dose                | рм Dose              | ам Dose                 | рм Dose              | ам Dose                               | рм Dose       | ам Dose   | <sub>РМ</sub> Dose  |
| Normal diet until<br>midnight (includes<br>those permitted clear<br>liquids until 2 h before<br>surgery) | Usual<br>dose          | 80% of usual<br>dose | 80% of usual<br>dose    | 80% of usual<br>dose | Usual<br>dose                         | Usual<br>dose | Usual<br>dose   | Usual<br>dose   |
| Bowel prep (and/or clear<br>liquids only 12-24h<br>before surgery)                                       | Usual<br>dose          | 80% of usual<br>dose | 80% of usual<br>dose    | 80% of usual<br>dose | Usual<br>dose                         | Usual<br>dose | Hold when<br>starting<br>clear liquid<br>diet/bowel<br>prep | Hold when<br>starting<br>clear liquid<br>diet/bowel<br>prep |

# Insulin Recommendations Day of Surgery

| Day of Surgery Insulin<br>Regimens  | Glargine<br>or Detemir  | NPH or<br>70/30 Insulin                                       | Lispro, Aspart,<br>Glulisine, and Regular | Noninsulin<br>Injectables |
|-------------------------------------|---|---|---|---------------------------|
|                                     | 80% of usual dose if patient uses morning only or twice daily basal therapy | 50% of usual dose if BG<br>120 mg/dl* Hold for BG < 120 mg/dl | Hold                                      | Hold                      |
| *6.6 mM.<br>BG = blood glucose; NPH | = neutral protamine Hagedorn.   |   |   |                           |

#### Best Practice: Check BG prior to surgery

#### Pre-Op:

- SubQ insulin to correct hyperglycemia may be considered for:<sup>2</sup>
  - Patients undergoing ambulatory surgery
  - Procedures of short duration
    - (< 4 hours operating room time)</li>
  - Patients undergoing inpatient procedures that are minimally invasive, with expected hemodynamic stability, and allow early resumption of oral intake.

#### Pre-Op and Intra-Op:

- IV insulin infusion is recommended in patients undergoing procedures that may result in the following variables that alter subcutaneous insulin absorption or distribution:2
  - Procedures with anticipated hemodynamic changes
  - Procedures with significant fluid shifts
  - Procedures with expected changes in temperature
    - (passive hypothermia or active cooling, hyperthermic intraperitoneal chemotherapy)
  - Procedures with use of inotropes
  - Lengthy operative times
    - (>4 hours)





#### **Pearls**

- Rapid-acting insulin should not be given more frequently than every two hours to minimize the risk of insulin stacking
- IV insulin infusion is recommended in patients with anticipated hemodynamic changes, significant fluid shifts, expected changes in temperature, use of inotropes, or lengthy operative times, greater than four hours
  - These variables alter subcutaneous insulin absorption and distribution
  - Can lead to persistent hyperglycemia or sudden hypoglycemia
  - IV insulin = very short half-life and allows rapid adjustment with limited lasting effects
  - IV insulin is the best modality for patients needing ICU admission
- Same day surgery patient may resume home regimen after discharge



#### Insulin Pumps Intra-Op

#### Requires preoperative planning

- Consider insulin pump insertion site in relation to the surgical field
  - May need to inform the patient of which areas to avoid
- Closed-Loop System?
- During surgery basal rate is continued and frequent monitoring of BGs is required
  - Hypoglycemia may go unrecognized under anesthesia
- Insulin infusion may be substituted for the insulin pump
- Insulin pumps are primarily utilized with type 1 diabetes, however growing use in type 2



#### IV Insulin

- IV Insulin infusion protocols provide guidance on insulin titration and frequency of blood glucose checks
  - Target blood glucose
  - Minimize hypoglycemia
- Nursing plays a critical role in the administration of IV insulin in ICU settings
- Anesthesia in the OR

- Initiating Glucommander<sup>™</sup> requires an order for target range and multiplier
- The multiplier is an insulin sensitivity factor: 0.01-0.05
- Targets: 120-160 mg/dL or 140-180 mg/dL
  - 100-140 mg/dL CV surgery

| Blood Glucose | Multiplier | Insulin Drop Rate |
|---------------|------------|-------------------|
| 300           | 0.01       | 2.4 units/hr      |
| 300           | 0.02       | 4.8 units/hr      |
| 300           | 0.05       | 12 units/hr       |





#### Patient Management | IV Glucommander



Key takeaway:

IV insulin is responsive and can achieve and maintain BGs in target throughout the procedure

#### Technology-Driven Insulin Management: Glucommander

Benefits of an eGlycemic Management System® (eGMS):

- Quicker time to target range
- Maintenance of glucose in tighter target ranges
- Lower glycemic variability
- Lower risk for hypoglycemia
- Fewer calculation errors
- Built-in alert systems
- FDA cleared



#### Transitioning to SubQ

- First subcutaneous injection of insulin given before stopping the insulin infusion to allow adequate time for the subcutaneous insulin to take effect
- Imperative type 1 patients receive basal insulin before discontinuing IV insulin
- Glucommander makes the transition process efficient by recommending the optimal dose for a patient based on their BG and A1C values

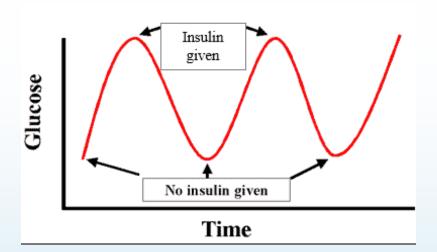




#### Subcutaneous Insulin

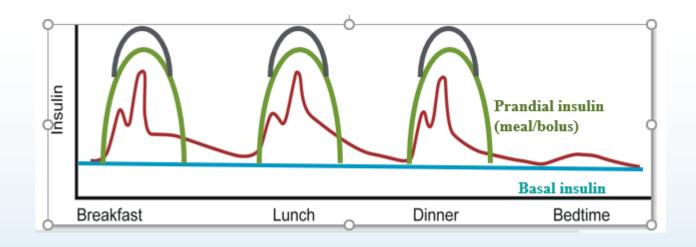
#### Sliding scale alone is discouraged

 REACTIVE—Insulin administered after hyperglycemia occurs



# Basal, prandial and correction insulin for those eating

 PROACTIVE—Insulin administered in anticipation of blood glucose spike





#### Patients on Insulin Prior to Admission

- Reduce home dose by 20 to 25%, at times more, especially for those on u500 insulin
  - High A1C may be an indicator of poor compliance with the insulin regimen
  - Diet will be controlled in house
  - Variable intakes postoperatively
- If the patient is on a high basal dosing and little to no prandial dosing, redistribute the dose between basal and meal bolus with 20 to 25% reduction

# Subcutaneous Insulin: Determining the Dosing

- 1. Calculate a total daily insulin dose. Use home insulin doses or calculate.
  - 0.3 Insulin Naive, older, renal, higher risk for hypo
  - 0.5 Most Type 2
  - 0.7 Insulin resistance, steroids
- 2. Split between basal and nutritional bolus
  - Basal once or twice daily
  - Meal bolus 3 times daily or with tube feeding bolus
    - Continuous feeding may give 4- or 6-hour doses
- 3. Correctional Insulin based on patient's insulin sensitivity
- 4. Adjust doses as needed based on response



## **Prescribing Practice**

- Order sets help drive prescribing practice
  - Include Basal, Meal Bolus, and Correction orders
  - Include Hypoglycemia Protocol
  - Include nursing notifications
  - Include BG checks

#### ▼ Basal Insulin

Insulin Glargine (LANtus) is the recommended insulin Insulin NPH may be used during pregnancy.

If the Insulin Glargine (LANtus) dose is > 100 units, consider splitting into twice a day (BID) doses.

- insulin glargine (LANtus) 100 units/mL injection
- Subcutaneous, Starting today at 1304
   Basal insulin is not held for NPO status. Contact provider with any questions or concerns.
- Transition from IV insulin using Glucommander Recommendations OR Provider Calculated Dosing
- NPH insulin

#### ▼ Meal Bolus Insulin

- insulin lispro (HumaLOG) 100 units/mL injection
  - Subcutaneous, 3 times daily with meals, First dose today at 1330
- MEAL BOLUS. HOLD IF NPO
- O CARB BASED: insulin lispro (HumaLOG) 100 units/mL injection

Subcutaneous, 3 times daily with meals, MEAL BOLUS. HOLD IF NPO or if eats < 15 grams of carbs. Give only \*\*\* units if eats 15-29 grams of carbs Give only \*\*\* units if eats 30-44 grams of carbs. Give only \*\*\* units if eats 45-59 grams of carbs.

#### ▼ Correction Scale Insulin - With Meals

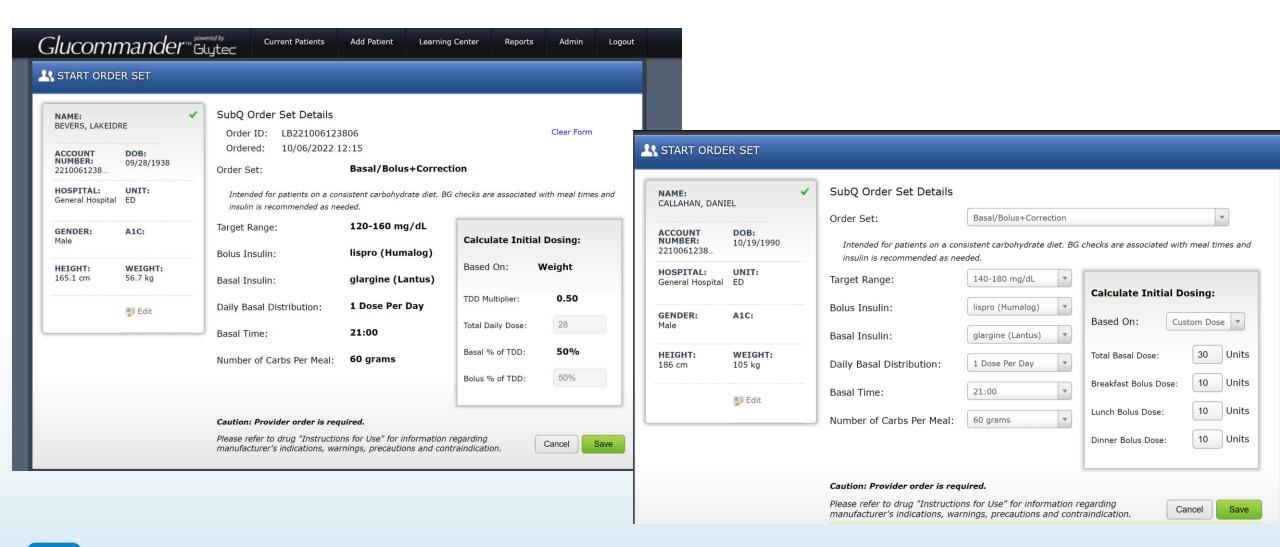
TDD = Total Daily Dose of scheduled insulin (basal + meal bolus)

- SENSITIVE (TDD less than 30) insulin lispro (HumaLOG) 100 units/mL injection CORRECTION SCALE
  - 1-5 Units, Subcutaneous, 3 times daily with meals, DO NOT HOLD IF NPO Notify provider if Blood Glucose LESS than 70 For BG 141-190 administer 1 unit For BG 191-240 administer 2 units For BG 291-340 administer 4 units For BG MORE than 340, administer 5 units AND notify provider
- O STANDARD (TDD 30-49) insulin lispro (HumaLOG) 100 units/mL injection CORRECTION SCALE
- 1-6 Units, Subcutaneous, 3 times daily with meals, DO NOT HOLD IF NPO Notify provider if Blood Glucose LESS than 70 For BG 141-180 administer 1 unit For BG 181-220 administer 2 units For BG 221-260 administer 3 units For BG 261-300 administer 4 units For BG 301-340 administer 5 units For BG MORE than 340, administer 6 units AND notify provider
- O MODERATE INSULIN RESISTANCE (TDD 50-70) insulin lispro (HumaLOG) 100 units/mL CORRECTION SCALE
- 1-8 Units, Subcutaneous, 3 times daily with meals, DO NOT HOLD IF NPO Notify provider if Blood Glucose LESS than 70 For BG 141-170 administer 1 unit For BG 171-200 administer 2 units For BG 201-230 administer 3 units For BG 231-260 administer 4 units For BG 261-290 administer 5 units For BG 291-320 administer 6 units For BG 321-350 administer 7 units For BG MORE than 350, administer 8 units AND notify provider
- SEVERE INSULIN RESISTANCE (TDD more than 70) insulin lispro (HumaLOG) 100 units/mL injection CORRECTION SCALE
- 1-11 Units, Subcutaneous, 3 times daily with meals, DO NOT HOLD IF NPO Notify provider if Blood Glucose LESS than 70 For BG 141-160 administer 1 unit For BG 161-180 administer 2 units For BG 181-200 administer 3 units For BG 201-220 administer 4 units For BG 221-240 administer 5 units For BG 241-260 administer 6 units For BG 261-280 administer 7 units For BG 281-300 administer 8 units For BG 301-320 administer 9 units For BG 321-340 administer 10 units For BG MORE than 340,



#### Glucommander Initiation

Can be started as a weight-based multiplier or as a custom start







# SPECIAL CONSIDERATIONS

#### **Steroids**

- Dexamethasone is often given intraoperatively to decrease risk for postoperative nausea and vomiting, decrease inflammation, and decrease postoperative pain
- Steroids are counter regulatory hormones
  - Impair insulin action
  - Increase insulin resistance
  - Diminish insulin secretion
- Many patients receiving > 40 mg of prednisone per day develop hyperglycemia
  - Many are not monitored for hyperglycemia
  - Once daily short-acting: effect seen 4-6 hours later, hyperglycemia seen throughout the day, BG decreases overnight = need more insulin with meals
- Long-acting: effect seen throughout the day and night = need increased basal and meal bolus
- Insulin needs decrease when steroid is tapered or stopped





## Renal Insufficiency

- Loss of renal clearance of insulin = Higher risk for hypoglycemia
  - Endogenous insulin: liver is the main site of clearance
  - Exogenous insulin: kidney is the main site of clearance
    - Exogenous bypasses the liver and directly enters systemic circulation
- Reduced hepatic and renal gluconeogenesis
- A1c may be falsely low: Anemia, Erythropoietin-stimulating agents, Transfusions

#### Recommendations:

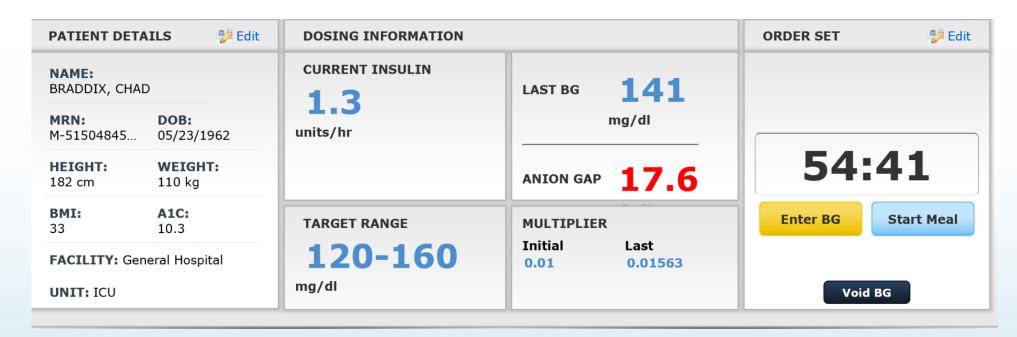
- 0.3 units/kg dosing and higher target range
- Consider reduction in insulin doses on dialysis days
  - 25% reduction in basal insulin up to 24 hours after hemodialysis
  - Greatest risk for hypoglycemia is overnight





#### **Artificial Nutrition**

- Work with the inpatient dietitians
- Tube Feeding; provide formulas low in carbohydrates when possible
  - May start with IV insulin drip to determine insulin needs
  - May use the last 4-6 drip rates to determine daily needs
    - Drip rates should be relatively stable
    - Do not use the full 24 hours rates as resistance changes and may overestimate.



#### Summary

- Hyperglycemia is associated with adverse clinical outcomes in surgical patients
- Screen patients for diabetes and pre/post-surgical hyperglycemia
- Insulin is preferred for management of hyperglycemia
  - IV insulin is very responsive and easily titrated to meet needs
  - Transition to SubQ insulin when medically stable
- Prevent Hypoglycemia
  - Reassess and modify treatment when blood glucose is <100 mg/dL</li>
- Special Considerations in populations
  - On enteral or parenteral nutrition
  - Receiving steroids
  - Renal insufficiency
  - Insulin pumps
- Glycemic targets should be modified according to clinical status



# Thank You!

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#### Resources

#### References:

- 1. American Diabetes Association (2022). Diabetes Care in the Hospital: Standards of Medical Care in Diabetes. Diabetes Care 45(Suppl. 1): S244–S253 | https://doi.org/10.2337/dc22-S016
- 2. Duggan, E., Carlson, K., Umpierrez, G. (2017). Perioperative Hyperglycemia Management: An Update. Anesthesiology. 126(3): 547–560. doi:10.1097/ALN.000000000001515.
- Best Practice Using Glucommander IV in the Periop Setting
- Internal Talking Points: Non-Glucommander Order Sets