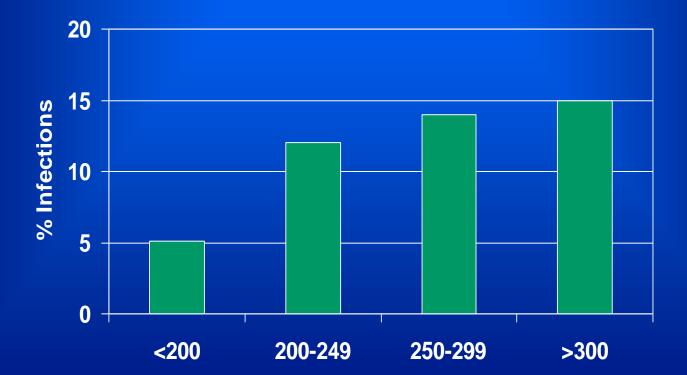
Hyperglycemia is Dangerous to <u>Non</u>Diabetics (and Diabetics)

> E. Patchen Dellinger, MD University of Washington

Glucose Control and SSIs After Median Sternotomy



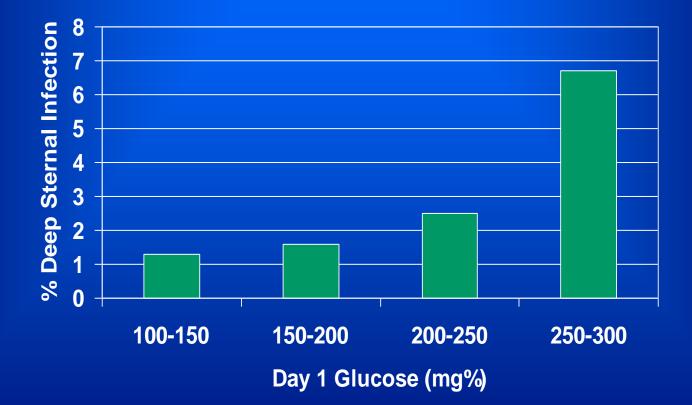
Latham. ICHE 2001; 22: 607-12

Hyperglycemia and Risk of SSI after Cardiac Operations

- Hyperglycemia doubled risk of SSI
- Hyperglycemic: 48% of diabetics 12% of nondiabetics 30% of all patients
- 47% of hyperglycemic episodes were in nondiabetics

Latham. Inf Contr Hosp Epidemiol. 2001;22:607 Dellinger. Inf Contr Hosp Epidemiol. 2001;22:604

Deep Sternal SSI and Glucose



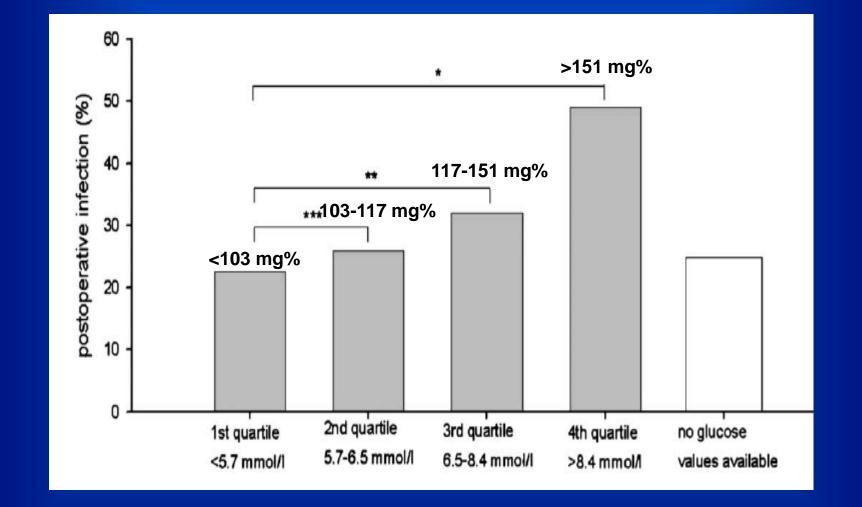
Zerr. Ann Thorac Surg 1997;63:356

SCIP only mandates glucose control for cardiac surgery

Hyperglycemia and Infection

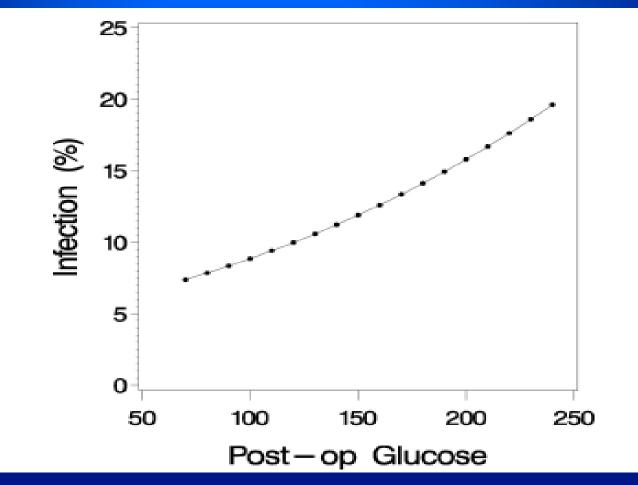
- Does it apply only to cardiac surgery?
- Do WBC struggling to work in syrup know whether they are in a median sternotomy or another incision?

Early (48h) Postoperative Glucose Levels and SSI after <u>Vascular</u> Surgery



Vriesendorp. Eur J Vasc Endovasc Surg 2004; 28:520-5

Perioperative Hyperglycemia in Noncardiac Surgical Patients



Ramos. Ann Surg 2008;248: 585-591

Mastectomy, Hyperglycemia, and SSI

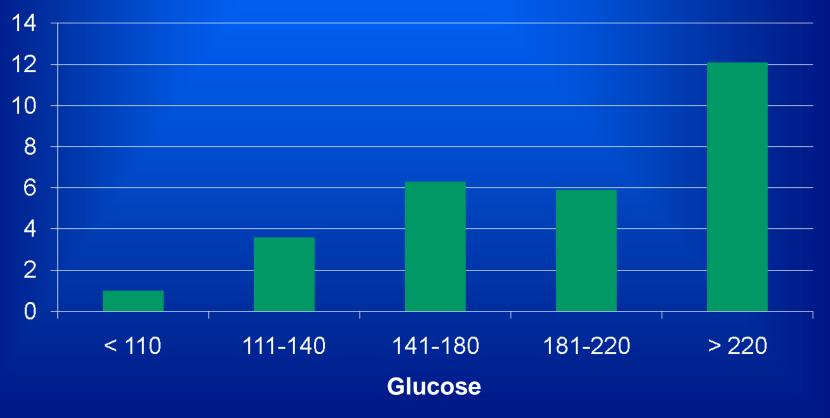
260 patients, 5 glucose determinations (pre-op, at anesthesia induction, intra-op, in PACU, at 24 hrs)

	Odds	
Risk Factor	<u>Ratio</u>	<u>C.I.</u>
Age > 50	3.7	(1.5-9.2)
Pre-Op ChemoRads	2.8	(1.4-5.8)
Any gluc <u>></u> 150 mg%	2.9	(1.2-6.2)

Villar-Compte. AJIC 2008; 36:192-8

Postop Glucose (within 48h) and SSI – <u>General Surgery</u>

Relative Risk



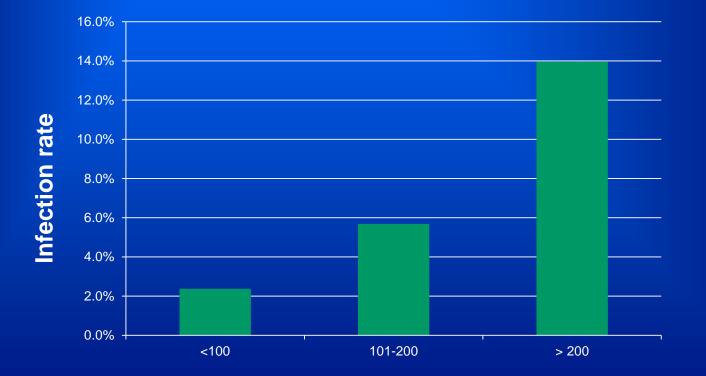
Ata. Arch Surg 2010: 145: 858-864

Postoperative Glucose and Mortality in <u>Noncardiac</u> Surgery

> Hyperglycemia in <u>non</u>diabetic patients was more dangerous than hyperglycemia in diabetics!

> > Frisch. Diabetes Care. 2010; 33: 1883-8

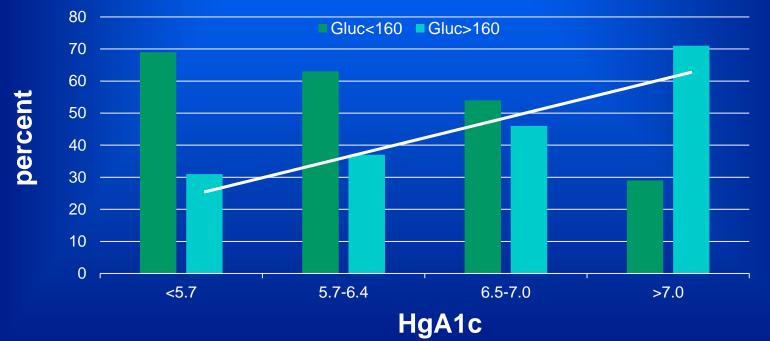
Perioperative Hyperglycemia and Total <u>Knee or Hip Arthroplasty</u> Fasting Blood Glucose POD #1



Mraovic. J Diab Science & Technol 2011; 5: 412-8

HgA1c and Postoperative Glucose – <u>Abdominal Surgery</u>

Relation of Preoperative A1c to Postoperative Glucose



Goodenough. J Amer Coll Surg 2015; 221: 854-61

Glucose & Infection Risk Non-Cardiac Surgery & Diabetes

VASQIP – 55,408 patients <u>Multivariate</u> analysis:

- All the usual risk factors significant.
- HgbA1c <u>NOT</u> significant.
- PreOp glucose <u>NOT</u> significant.
- PostOp glucose > 150 very significant.

King. Ann Surg 2011; 253:158-65

Hgb A1c vs. Glucose as Risk Factor for SSI – <u>Gastric Bypass</u> Multivariate Analysis

Odds ratio = 1.27 for every 20 mg% increase in <u>mean</u> glucose level during hospitalization (p=0.008). <u>Mean glucose</u> more significant than any single level above 200 Mg% or not. Hgb A1c <u>not</u> significant.

Perna. Surg Obes Rel Dis 2012; 8: 685-90

SCOAP Data on Perioperative Glucose Levels and Insulin Use

 Diabetic pts
 4098 (35%)

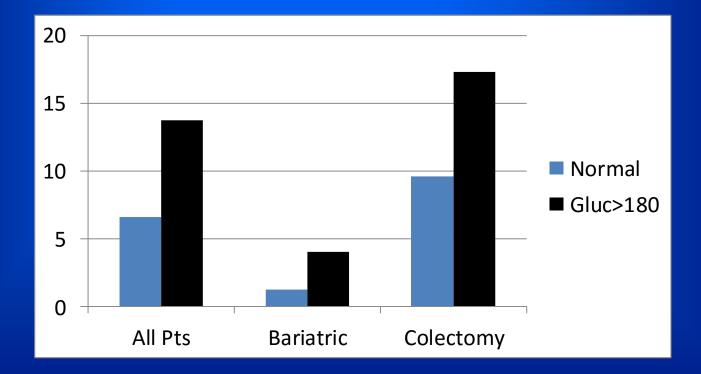
 Hyperglycemic
 2369 (58%)

 Nondiabetic pts
 7532 (65%)

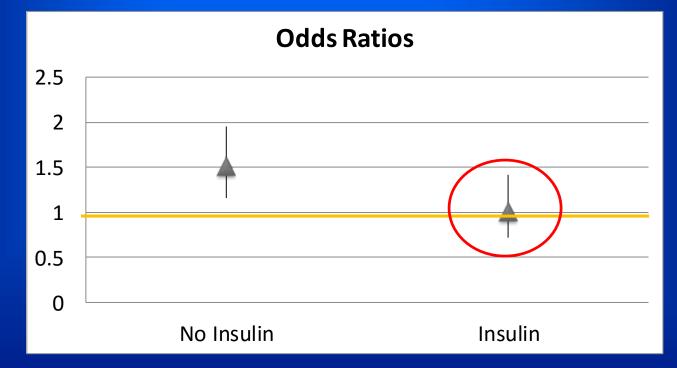
 Hyperglycemic
 1014 (13%)

<u>30% of all hyperglycemic</u> patients were not diabetic!

Composite Infection Hyperglycemia vs No Hyperglycemia Nondiabetic Patients



Composite Infection in Hyperglycemic Patients With and Without Use of Insulin



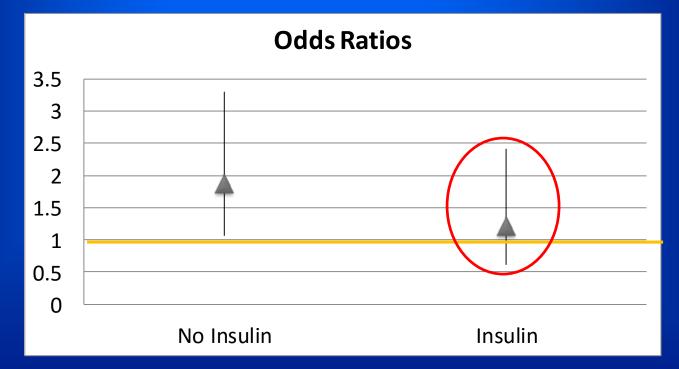
Insulin reduces risk even when glucose control is not as good as desired

Operative Reintervention in Hyperglycemic Patients With and Without Use of Insulin



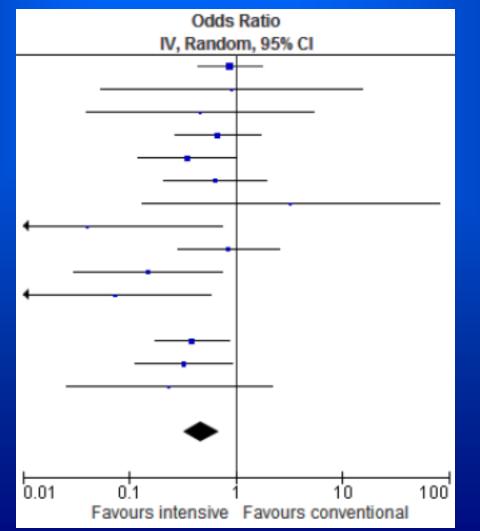
Insulin reduces risk even when glucose control is not as good as desired

<u>Mortality in Hyperglycemic</u> Patients With and Without Use of Insulin



Insulin reduces risk even when glucose control is not as good as desired

SSI with Strict vs Permissive Perioperative Glucose Control



de Vries, et al, mss in prep **Glucose Control Proven important for SSI risk: Cardiac surgery General surgery Colorectal surgery** Vascular surgery **Breast surgery Gynecologic Oncology surgery** Hepato-pancreatico-biliary surgery **Orthopedic surgery Trauma surgery**

•Regardless of the Diagnosis of Diabetes (or not) Hyperglycemia Increases

- Morbidity
- Mortality
- Length of Stay

Which Patients Are at Risk

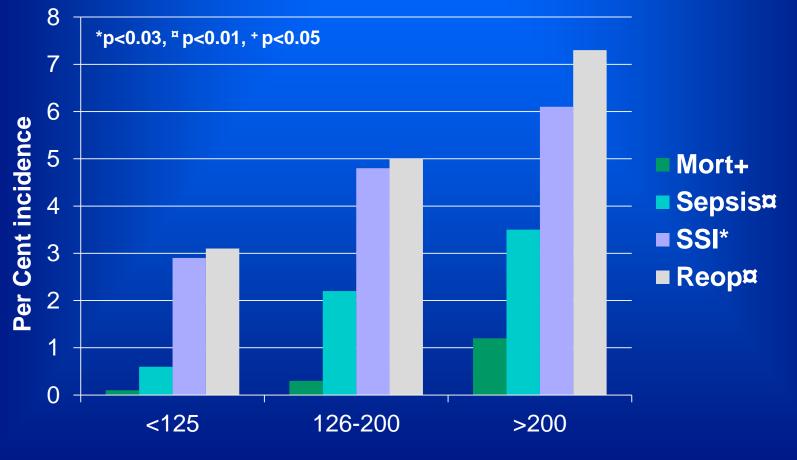
for Hyperglycemia?

Glucose in <u>Non</u>Diabetics having Colectomy at Cleveland Clinic

<u>Highest Gluc</u> ≤ 125 mg% 126-200 mg% 200 mg% All patients <u>N (%)</u> 816 (33%) 1289 (53%) 342 (14%) 2447 (100%)

Kiran, et al. Ann Surg 2013; 258:599-605

Glucose in <u>Non</u>Diabetics having Colectomy at Cleveland Clinic



Kiran, et al. Ann Surg 2013; 258:599-605

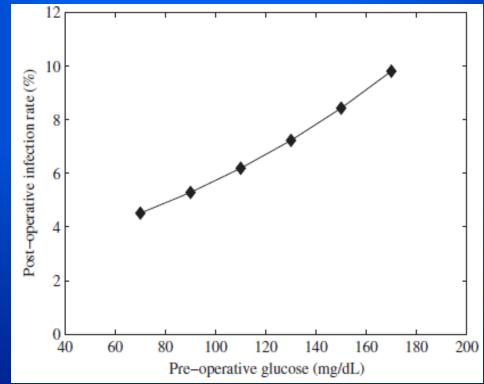
Preoperative Glucose as a Screening Tool for Patients Without Diabetes

- Random glucose within 30 days of operation
- Average 8 days before operation
- 16% within one day and 29% within 3 days
- 6683 patients

•	<70	384	pts	
•	70-99	4251	pts	
•	100-139	1801	pts	
•	140-179	187	pts	- 31%
•	>180	60	pts	

Wang. J Surg Res. 2014; 186: 371-8

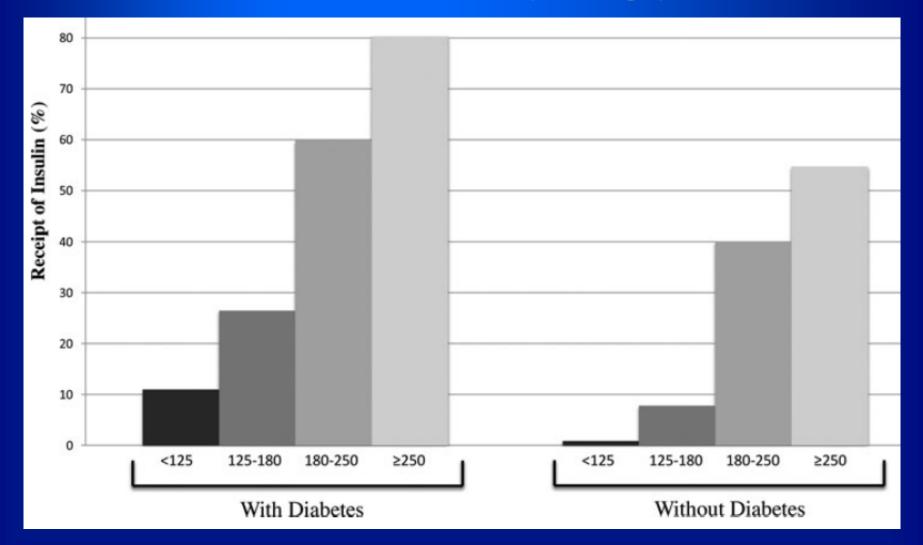
Preoperative Glucose as a Screening Tool for Patients Without Diabetes



Pre-Op Glucose vs. Post-Op Infection, adjusted for age, gender, BMI, ASA, & type of operation.

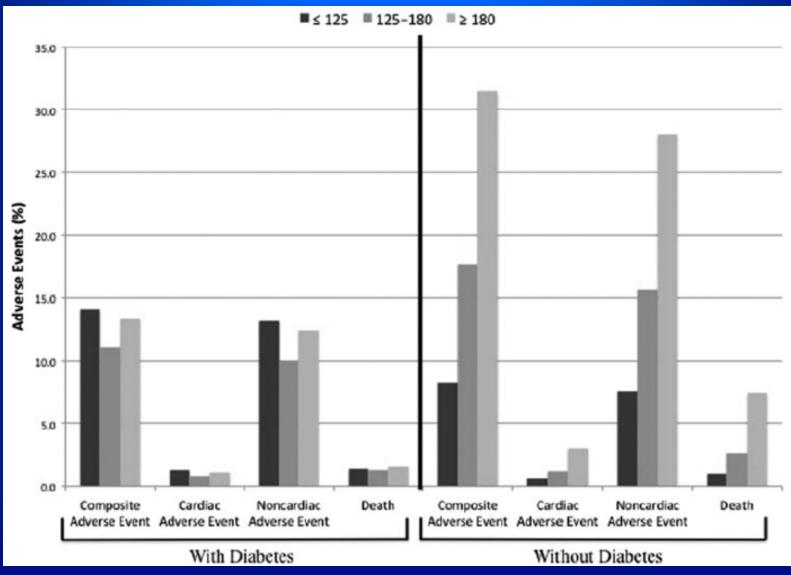
Wang. J Surg Res. 2014; 186: 371-8

SCOAP – NonDiabetics Less Likely to Receive Insulin for Hyperglycemia



Kotagal. Ann Surg 2015; 261:97-103

SCOAP Adverse Events with Hyperglycemia – Diabetics v. NonDiabetics



Kotagal. Ann Surg 2015; 261:97-103

Glucose Levels & SSI

- The exact "<u>best</u>" level of glucose control in the perioperative period is not known.
- High glucose levels unequivocally increase the risk of SSI and other perioperative infections.
- Tight glucose control in the perioperative period is tricky.
- Hypoglycemia increases the risk of morbidity and mortality.
- When algorithms are followed at UWMC hypoglycemia is very rare.

Perioperative Blood Glucose at UWMC – Amalga data

Inpatient ops, 7/1/13 – 7/30/15	11,079
Nondiabetics among those	8,974 (81%)
Nondiabetics with gluc > 140 mg%, day 0-2	4,834 (54%)
Nondiabetics with gluc > 140 mg% day of op	2,929 (33%)

Data courtesy of Ray Bunnage, Center for Clinical Excellence

Perioperative Blood Glucose at UWMC – Amalga + NSQIP

Cases both in Amalga and NSQIP5,109Cases with glucose data (pre- or intra-op)3,044Cases with gluc > 160 mg%664 (22%)Hyperglycemic cases that got insulin560 (84%)

Data courtesy of Ray Bunnage, Center for Clinical Excellence & Lucas Thornblade

I am happy to share my slides with anyone who would find them useful

patch@uw.edu

Glucose Control in the OR: Our Current State

Gail A. Van Norman MD Professor, Anesthesiology and Pain Medicine, University of Washington





For Marjorie

- OR Compliance w/established protocols
- Common mistakes/misconceptions in administering insulin and glucose
- Effectiveness of current protocols in the OR
- The Future

Kelly study 2007

Retrospective chart review of 250 pts at UWMC* with diabetes undergoing surgery to determine

- blood glucose levels
- compliance of anesthesia providers in treating intraoperative hyperglycemia

*Both SQ and IV insulin regimens were in use at the time.

< 25% of anesthesia providers used insulin in the OR

- Few patients hit target range of < 200 mg/dl
- Only 2 patients experienced hypoglycemia, defined as BS ≤ 60 mg/dl.... Both patients had been treated with SQ insulin.

Challenges to Good OR Glucose Management

- Provider Lack of knowledge about diabetes; i.e. type I diabetics need exogenous insulin despite blood glucose levels.
- Lack of task prioritization in the OR
- POC glucose testing issues—cumbersome, takes time, not automatic like BP measurements
- Lag time between glucose measurement and treatment modification—by the time pt is treated, glucose levels are rising even farther and insulin dose may already be too low
- Lack of basic treatment technique
 - Do you flush your insulin tubing or just start insulin running?
 - Do you bolus with regular insulin prior to beginning a drip?

(cont)

- Fear of overtreatment—e.g. hypoglycemia
- Patients are much more insulin resistant in the periop period:
 - Stress hormone release (epi, norepi, cortisol, etc)
 - Delivery of exogenous catecholamines is common (ephedrine, epi, dopamine, etc)
 - Underlying conditions (e.g. infection) leading to need for surgery predispose to insulin resistance
 - Anesthetic agents may induce insulin resistance (e.g. SEV, ISF both induce insulin resistance in pigs)—and yet may have an overall positive effect by lowering stress hormone levels.

Compliance with Glucose Management Protocols (Nair et al, 2015*)

- Prospective study
- Customize alert system in AIMS to nudge anesthesia providers to check a blood glucose level within ½ hour of start of surgery, and hourly thereafter
- Compliance with glucose testing and with a standard IV insulin protocol were measured before and after institution of the alert system

- Compliance with glucose testing improved from 52.6% to 72.1% with the SAM alert system
- Compliance with glucose testing was much higher in cases where SAM was enabled (81%) than when the provider disabled the SAM alert (57.4%)
- However correct insulin dosing only improved from 13.5% to 24.4%--less than ¼ of patients received treatment that matched protocol
- Target glucose levels did not improve significantly

Sathishkumar et al Anesthesiology July 2015

- 2341 patients, retrospective data study
- Excluded surgeries < 1 hr, ASA 5 and 6 pts
- 791 cases used Alert Watch: audiovisual alert system notifies of out of range glucose (≥ 200 mg/dL) and then disappears when insulin started, reminders q 90 min.

- Hypoglycemia (≤ 70 mg/dL) was no different between alert watch (AW) and controls. (approx 3-4%)
- Insulin treatment for BS ≥ 200 mg/dL improved from 52.7% to 62.7% with AW
- The odds of getting a BS rechecked after initiation of insulin therapy improved from 42.3% to 51.5% with AW

Nair, et al.*

- Retrospective review of electronic data on 2440 pts undergoing surgery at UWMC from 2011-2013.
 - At least one $BS \ge 140$ during the case
 - At least one BS available postop
 - Case duration > 1 hour

*Anesth Analg Nov 23, 2015 epub ahead of print

- Intraop glucose (all patients ≥ 140 mg/dL) 35% exceeded 180 mg/dL 45% received intraop glucose management
- Postoperative glucose
 12 hr postop glucose ≥ 180 mg/dL = 50%
 24 hr postop glucose ≥ 180 mg/dL = 53%

- For every 10 mg/dL increment of glucose over 140 mg/dL, there was a mean increase in 4.7 mg/dL in the first postoperative (≤ 1 hour) glucose, 2.6 mg/dL in the 12 hour postop glucose, and 2.4 mg/dL in the 24 hour postop glucose levels (p < 0.001 for all)
- Effect was stronger for larger BMI, pts diagnosed with diabetes mellitus, and intraop steroid use

- Postoperative hyperglycemia (≥ 180 mg/dL) was less frequent at 12 and 24 hours if intraop glucose management began at 140 mg/dL rather than 180 mg/dL (50% vs. 62% and 59% vs. 70%, respectively)
- Postop 12 and 24 hour mean glucose levels were lower by 9 and 7 mg/dL respectively if intraop insulin was initiated at 140 mg/dL compared to 180 mg/dL

Hypoglycemia (≤ 60 mg/dL)

- No difference in postop hypoglycemia whether intraop target was 140 mg/dL or 180 mg/dL.
- 15 cases of hypoglycemia intraop and within 1 hr of surgery (0.6% of all cases)
- No difference in hypoglycemia observed postop between pts with and without dx of diabetes
- Postop hypoglycemia occurred in 23/29 observed hypoglycemic cases within 12 hours (1% of all cases)
- Severe hypoglycemia (< 40 mg/dL) was seen in 2 patients intraop, and 6 patients postop (8/2440 = 0.3% of all cases)

Hypoglycemia

 In all but one of the intra and immediate postop cases, hypoglycemia occurred when intraop providers *deviated from the standard insulin protocol,* including the overdosing of insulin in at least 7 cases.

(other deviations can include failure to test hourly glucose, failure to adjust insulin infusion according to protocol)

Basic Issues

- Do all providers fill the piggyback IV tubing and then flush 22-25 cc of insulin solution through the tubing before hooking it up.
- Do all providers routinely hook insulin drip in the port closest to the skin?
- Should all providers routinely bolus with regular insulin before initiation the drip?
- Do all providers understand that type I diabetes MUST be treated with insulin even if blood glucose is NORMAL (i.e 60-100 mg/dL)? Why is this true?

The Future

- How can we further improve compliance to glucose management protocols in the intraoperative period?
 - Improved awareness of detrimental effects of perioperative hyperglycemia
 - Continued improvement of AIMs system: better compliance with protocol will allow us to refine the protocol further
 - Improved understanding/management of diabetes by perioperative providers
 - Continuous glucose monitoring tools are likely to help.

Changes to Perioperative Glucose Management

G. Alec Rooke, M.D., Ph.D. rooke@uw.edu

All Patients Get Glucose Measured

POC glucose in holding

If glucose above target range ✓ 140 for UWMC, 180 for HMC If patient will remain in-house after surgery, initiate insulin/glucose infusions If patient expected to go home, glucose management at discretion of surgical and anesthesia teams

Glucose Measurement in OR

- Regardless of glucose value in holding, repeat glucose level 30-60 minutes after incision
- Initiate glucose management if necessary
- Glucose measurements every hour if insulin initiated

Changes to Insulin Protocol in the Operating Room

 Non-diabetics and known diabetics treated the same

Insulin algorithms same as for hospital
 Includes algorithms 0.5 and 5-8

 D5 infusion at 100 ml/hr not always expected on initiation of insulin infusion

Glucose Infusion Guidelines

- Patients on chronic insulin: initiate once glucose is below 200
- Patients not on chronic insulin: initiate once glucose is below 150
- Intracranial and CPB cases will be excluded

Smart Anesthesia Monitor (SAM)

 Accesses and analyzes information: Presence of diabetes from the anesthetic record or preop evaluation Glucose values (automatically load into the anesthetic record) Detects insulin and dextrose infusion rates if entered into the anesthetic record

Provides alerts to the anesthesia team

SAM Messages

- Glucose check reminder 45 minutes after incision if not already done
- Recommends an insulin infusion once glucose levels exceed target threshold
- Hourly reminders to check glucose
 ✓ if insulin is running
 ✓ Intraop checks show increasing values
- If glucose < 100, stop insulin and check glucose every 20 minutes

Alerts For Glucose Infusion

- Patients on chronic insulin whose glucose levels fall below 200
- Patients not on chronic insulin whose glucose levels fall below 150
- Intracranial and CPB cases will be excluded

Patients Who Should Get Insulin and Dextrose Infusions

- True Type 1 diabetics
- Patients with a history of DKA
- Patients taking gliflozin drugs (Farxiga, Invokana, Jardiance, Glyxambi)
 Encourage urinary excretion of glucose
 Euglycemic DKA can occur (presenting glucose as low as 150) *Diabetes Care 2015:38;1687-93*

SAM Insulin/Dextrose Infusion Alerts

- Evidence of "Type I DM" or "DKA" in preop note
- Gliflozin in the med list, then alert for insulin and glucose

SAM Insulin Infusion Alerts

Won't recommend specific insulin rates

Will follow trends in glucose levels
 ✓ Decrease of ≥ 75/hour will suggest moving to a lower algorithm
 ✓ Failure to decrease (if still above target) will suggest moving to a higher algorithm

 Caregivers are free to change algorithms as clinically indicated

Postoperative Glucose Management

- Return to standard hospital protocol in the PACU
 ✓ Glucose infusion may then be started
- Continue/discontinue protocol on the ward/ICU as per current practice
- All patients get a fasting glucose measurement on POD #1 and #2

Thank You to the Many Multidisciplinary Team Members Who Have Worked on this Project for the Past 8 Months

Kate Curtis **Janet Kelly Christopher Kim Heather Lien Diane Matsuwaka** Alec Rooke Solina Tith **Ray Bunnage** Lucas Thornblade **Benjamin Anderson Cindy Sayre**

Diana Villaflor-Camagong David Flum Gail Van Norman Irl Hirsch Jing Chao Joan DiGiacomo John Lang Marcelo Hinojosa **Rachel Thompson** Lisa Goben **Patchen Dellinger**