

### Massive Hemorrhage in the Obstetrical Patient

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#### Faculty Disclosure

In compliance with CPD policy, Temerty Faculty of Medicine requires the following disclosures to the session audience

- This program has received no financial external support
- I have no relevant conflicts of interest to disclose

• I will use the term "woman" and "mother". I acknowledge that these terms are exclusive, and the experiences may apply to all those with the anatomy for childbirth.



#### Outline

1. The epidemiology of massive obstetrical bleeding

2. The management of the massively bleeding peripartum patient: a focus on transfusion

3. The Ontario quality indicators: how do we score on managing obstetrical bleeding?



#### Objectives

#### My job is to convince you:

- 1. Obstetrical patient ≠ trauma patient
- 2. TXA saves lives give it ASAP
- Detecting PPH early is higher yield than managing massive OB hemorrhage
- 4. Prioritize red cells + TXA + fibrinogen level
- 5. Plasma and platelets are RARELY required in OB MHP





### THE FACTS OF MASSIVE OBSTETRICAL BLEEDING



#### Every year...

14 million women experience PPH

70 000 maternal deaths

 Surviving women experience morbidity, 'lifelong reproductive disability'

2.9% USA deliveries are complicated by PPH

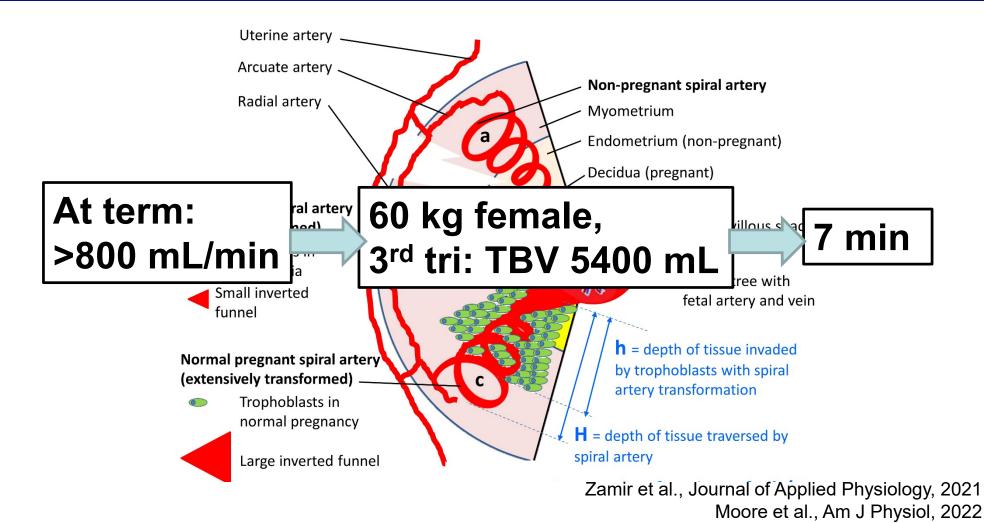


#### Etiology of PPH: 4T's

- **T**one (79%)
- **T**issue (10%)
- Thrombin (coagulopathy) (5%)
- Trauma



#### Uterine Blood Flow in Pregnancy



#### Defining 1° Postpartum Hemorrhage

#### • THEN:

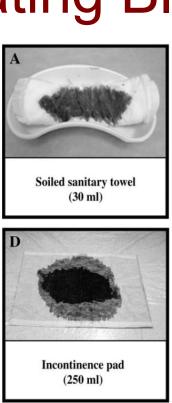
Vaginal (>500 mL) vs c-section (>1000 mL) within 24 h

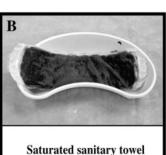
#### • NOW (2017):

 ->1000 mL within 24 h with signs/symptoms of hypovolemia (any method of delivery)

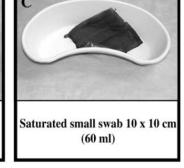


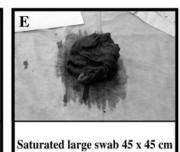
#### Estimating Blood Loss is Hard...

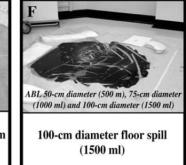


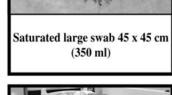


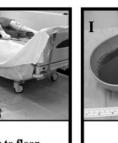
(100 ml)

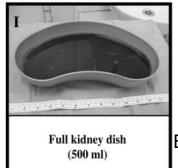


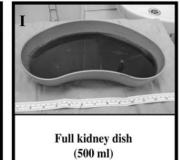












Bose et al., BJOG, 2006





PPH on bed only (1000 ml)



PPH spilling to floor (2000 ml)

Estimating Blood Loss is Hard...

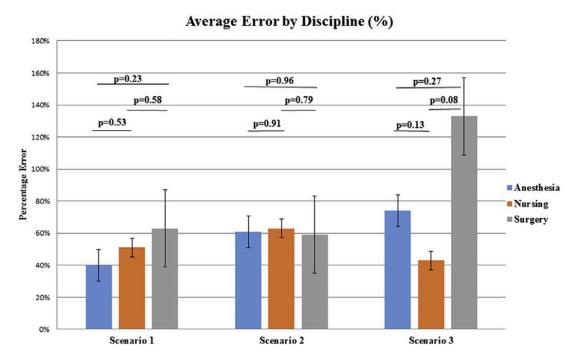


Rotherman & Lipman, Surgery, 2016



#### The Thing Is...

#### We all **EQUALLY STINK** at estimating blood loss



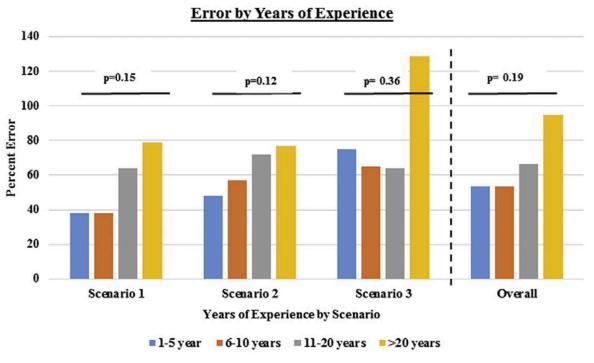
Regardless of your discipline....

Rotherman & Lipman, Surgery, 2016



#### The Thing Is...

#### We all **EQUALLY STINK** at estimating blood loss



Regardless of your experience....

Rotherman & Lipman, Surgery, 2016



#### Calibrated Drapes



### The NEW ENGLAND JOURNAL of MEDICINE

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#### Randomized Trial of Early Detection and Treatment of Postpartum Hemorrhage

Gallos, A. Devall, J. Martin, L. Middleton, L. Beeson, H. Galadanci, F. Alwy Al-beity, Z. Qureshi, G.J. Hofmeyr, N. Moran, S. Fawcus, L. Sheikh, G. Gwako, A. Osoti, A. Aswat, K.-M. Mammoliti, K.N. Sindhu, M. Podesek, I. Horne, R. Timms, I. Yunas, J. Okore, M. Singata-Madliki, E. Arends, A.A. Wakili, A. Mwampashi, S. Nausheen, S. Muhammad, P. Latthe, C. Evans, S. Akter, G. Forbes, D. Lissauer, S. Meher, A. Weeks, A. Shennan, A. Ammerdorffer, E. Williams, T. Roberts, M. Widmer, O.T. Oladapo, F. Lorencatto, M.A. Bohren, S. Miller, F. Althabe, M. Gülmezoglu, J.M. Smith, K. Hemming, and A. Coomarasamy



### Early Detection and Treatment of PPH (NEJM 2023)

#### Population:

- 80 hospitals across Africa
- 210,132 vaginal deliveries
- Intervention:
  - Calibrated blood-collection drape
  - Bundle of 1<sup>st</sup> response treatments
  - Implementation support
- Composite outcome:
  - PPH >1000 mL, laparotomy for bleeding, maternal death from bleeding

Gallos et al., NEJM 2023



#### The Bundle:

**Early Detection** Oxytocic Tranexamic Examination Massage of Uterus Acid and Trigger Criteria Drugs IV Fluids and Escalation Calibrated drape for the Massage until uterus IV fluids in addition to Ensure bladder is 10 IU IV oxytocin 1 g IV tranexamic acid the collection of has contracted or injected or diluted the infusion should injected or diluted in empty, evacuate blood, with trigger for 1 min in 200-500 ml 200 ml crystalloid be given if clinically clots, check for tears lines at 300 ml and crystalloid adminadministered over indicated for resuswith an internal 500 ml for the first hr istered over 10-min 10-min period citation and will examination and after birth period, plus a mainrequire a second placenta for com-Observations (blood tenance dose of pleteness intravenous access loss, blood flow, Escalate if bleeding 20 IU IV oxytocin uterine tone) every diluted in 1000 ml does not stop after

saline administered

over 4-hr period

800  $\mu$ g if used)

(with misoprostol

#### Implementation Strategies

15 min documented

on the blood-loss

monitoring chart

pulse monitored once in the first hr

post partum and documented on the

blood-loss moni-

toring chart

**Trigger Criteria** 

Clinical judgment

observation

Blood loss ≥500 ml Blood loss ≥300 ml

plus one abnormal

Blood pressure and

**Audit newsletters:** Sharing with all staff monthly rates of detection and bundle use, along with rates of PPH, severe PPH, blood transfusion, laparotomy, and death from PPH and giving feedback at monthly departmental meetings

**Champions:** Midwife and doctor to oversee change, troubleshoot, give feedback on audit newsletters, connect with other champions by means of chats, meetings, and websites for sharing knowledge and lessons learned

**Trolley or carry case:** Restocking of all medicines and devices used for treatment of PPH after every use and completion of a stocking checklist at the start of every shift

**Training:** Onsite, simulation-based, and peer-assisted training, lasting from 90 min to an entire workday, facilitated by the use of provider guides, flipcharts, and job aids displayed in labor wards

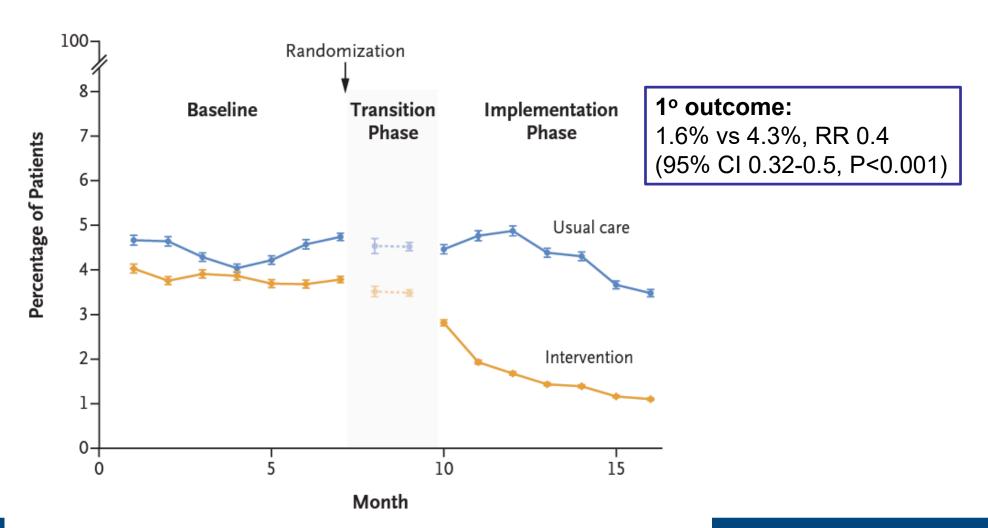


first response or

cause of bleeding

clinician is unable to identify or manage

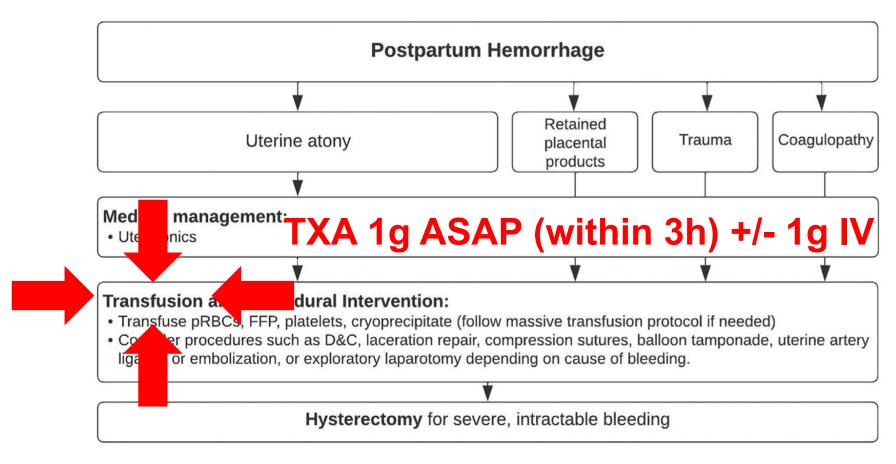
#### Patients with Primary-Outcome Event during the Baseline, Transition and Implementation Phases



#### MANAGING PPH: A FOCUS ON TRANSFUSION



#### Managing PPH



ACOG Practice Bulletin No 183, 2017 Guan et al., AJOG 2023



#### **WOMAN TRIAL:**

- An international, randomized, double blind, placebo-controlled trial
- N=20,060 women with PPH after vaginal birth or c-section
   (>500 mL or any blood loss within 24h associated with hemodynamic instability)
- TXA (1g +/- 1g for ongoing bleeding at 30 min or re-bleed within 24h) vs placebo
- Composite outcome: mortality or hysterectomy



#### **WOMAN TRIAL**

• 21 countries, mostly in Africa (12,343) and Asia (6,030)

- Primary outcome:
  - TXA (n=10,051): 5.3%
  - Placebo (n=9,985): 5.5%

RR 0.97 (95% CI 0.97-1.09), p=0.65

- Death due to hemorrhage:
  - TXA: 1.5%
  - Placebo: 1.9%

**RR 0.81** (95% CI 0.65-1.00), **p=0.045** 

Death due to hemorrhage with EARLY TREATMENT (<3h) RR 0.69 (95% CI 0.52-0.91)

WOMAN Trial Collaborators, Lancet 2017

www.transfusionontario.org



#### **Bottom line from WOMAN:**

# TXA reduces death due to bleeding when you give it early (ASAP, always <3h)

Tranexamic acid should be administered as soon as intravenous or intraosseous access is achieved but within 3 hours from time of injury or within 3 hours from MHP activation in all other patients.



WOMAN Trial Collaborators, Lancet 2017 Callum et al., CMAJ, 2019



#### The stories of women who died in WOMAN

WOMAN recruited 20,060 women with PPH

- 483 mothers died
  - 375 in Africa
  - 105 in Asia

Interviews of clinicians involved in the care of these women



### TXA is not always enough: access to blood products is critical

"There was **no more blood available** in my blood bank"

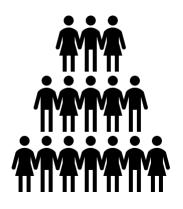
She could only get one pint of blood and **efforts to get more blood for further transfusion was on-going** when she died"

"Relatives were unable to pay or donate for further blood transfusion"

"When she became [unwell after] a blood transfusion reaction (from a transfusion which was discontinued, then found to be type-incompatible). That is the more probable cause of death."

g 🍐

#### Trauma patient ≠ OB patient







Fibrinogen in general population: 1.5-4 g/L

Fibrinogen at term: 4-6 g/L

↑ VWF, ↑ clotting factors↓ protein S

Collins, Blood, 2014



#### PTT and INR are RARELY abnormal in PPH

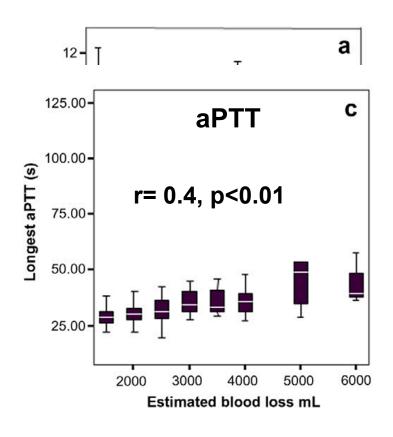
	<=2500 mL EBL (n=297)
PTT normal	296 (99.7%)
PTT abnormal	1 (0.3%)
INR normal	296 (99.7%)
INR abnormal	1 (0.3%)

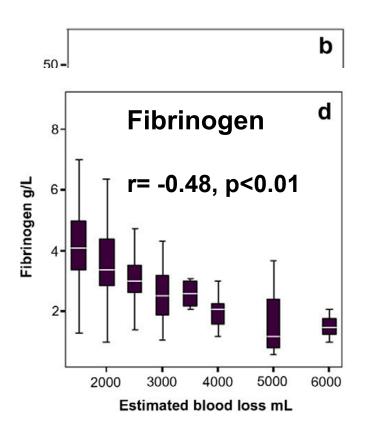
Overall: 98.8% had normal PTT and 98.2% had normal PT/INR

Collins et al., Blood, 2014



#### Fibrinogen is the best predictor of EBL in PPH





de Lloyd et al., Int Journal of Obs Anesth, 2011



#### Low Fibrinogen & Fibtem A5: Harbingers of Severe PPH

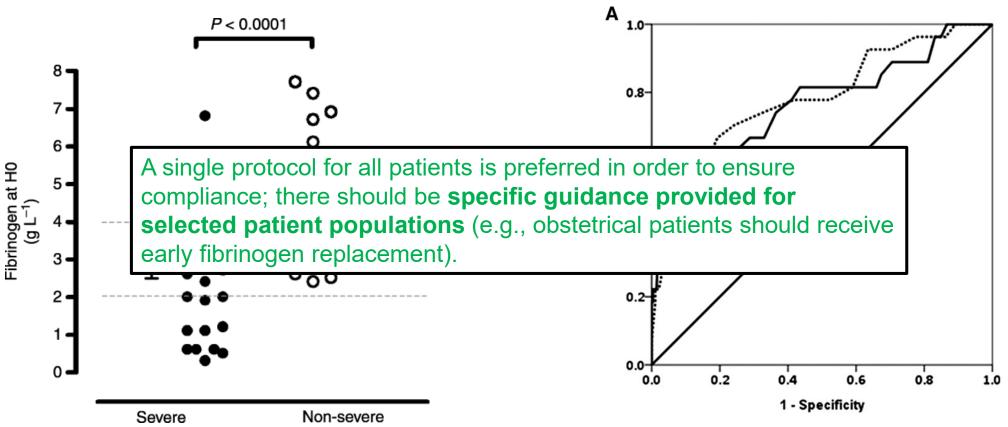


Figure 2. ROC curves for fibrinogen and Fibtem A5. (A) ROC curves for fibrinoge Progression to  $\geq$ 8 U allogeneic blood products (RBCs + FFP + platelets).

Charbit et al., JTH 2007; Collins, Blood, 2014



#### Preemptive Fibrinogen Replacement in PPH?

Randomized, placebo-controlled trial:
 249 women with early PPH

2g FC vs placebo

- RBC transfusion 20% vs 22%, p=0.88
  - Only 2.2% had fibrinogen level < 2.0 g/L</li>



#### Plasma in PPH

Ratio-based resuscitation leads to over-transfusion of plasma

- 1495 French patients with severe PPH transfused without coag.
   guidance
  - → 69% of transfused patients received FFP

Deleu et al., Int J Obstet Anesth 2019



#### Plasma in PPH

- Population: 605 women with moderate/severe PPH (>1000 mL EBL)
- Intervention: Viscoelastometric POC testing-guided FP
  - Fibtem A5 <= 15 mm + ongoing bleeding → FP</p>
  - -98% did not need plasma



#### Platelets in PPH

- Platelets rarely required for PPH
- 12/347 (3%) moderate to severe PPH required platelets

- All those transfused plt had 1 of:
  - Antenatal thrombocytopenia
  - Consumptive coagulopathy (abruption, amniotic fluid embolism)
  - >5000 mL hemorrhage



#### r7a in PPH

- ?↓ risk of invasive procedures in those non-responsive to uterotonics
- ↑ thrombotic risk
- no change in blood products, hysterectomy, EBL

#### **ISTH Guidance:**

"Consider for ongoing PPH unresponsive to standard treatment or to prevent hysterectomy"

Lavigne-Lissalde et al., JTH, 2015 Collins et al., JTH 2016



#### TEG/ROTEM

Less blood component transfusion

Timely care

• The future...?

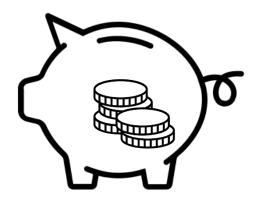
Frigo et al., Transfusion Medicine, 2020 Butwick et al., Transfusion, 2020



#### **Bottom Line**

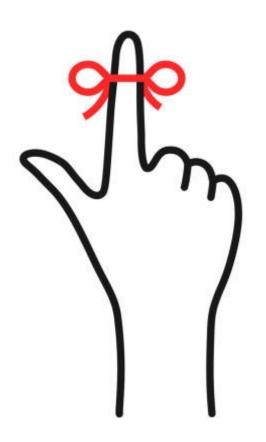
Where the money is:

- Timely recognition and care
- TXA (ideally within 1 h, always within 3h)
- Red cells with early labs to guide fibrinogen
  - Fibrinogen if <2.0 g/L</li>



#### Don't forget the G&S...

Women of child-bearing potential should be informed of the risk of red blood cell alloimmunization, which may result in hemolytic disease of the fetus and newborn, and should be counselled to undergo red blood cell antibody screening 6 weeks and/or 6 months after transfusion (many antibodies are evanescent, and there is a brief window for detection)

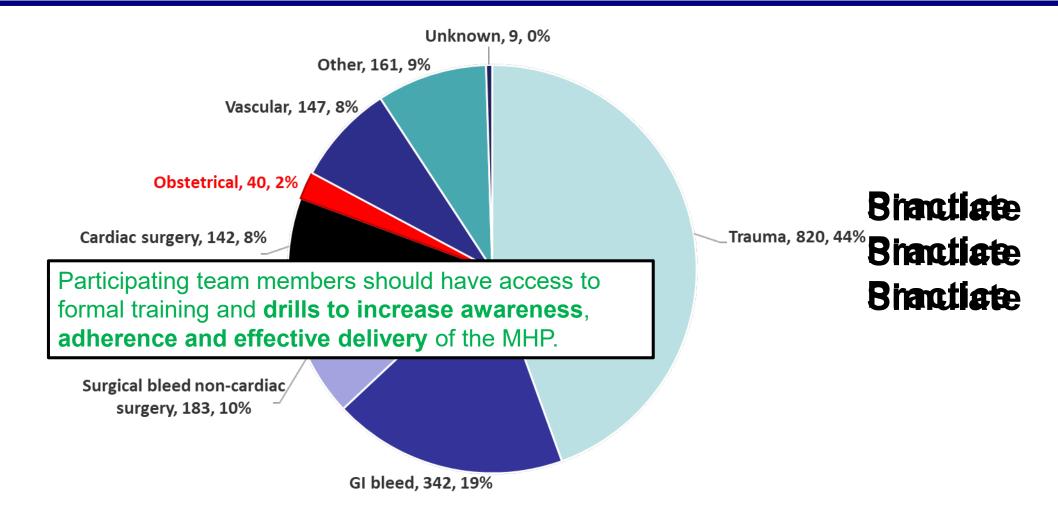


Callum et al., CMAJ, 2019



## ONTARIO DATA: QUALITY INDICATORS IN OBSTETRICAL HEMORRHAGE

#### Type of MHPs Reported in Ontario: 2019-2022



#### MHP Case Score

#### **Quality Metrics**

Did the patient receive tranexamic acid within 1 hour of activation?

Was RBC transfusion initiated within 15 minutes of activation?

Was the initiation for patient **transfer** within 1 hour of activation?

Was the patient's temperature ≥ 35°C at termination?

Was the hemoglobin maintained over 60 g/L for the first 24 hours?

Was the hemoglobin **below 110 g/L** at 24 hours?

Was the patient transitioned to group specific RBC within **90 minutes** of activation?

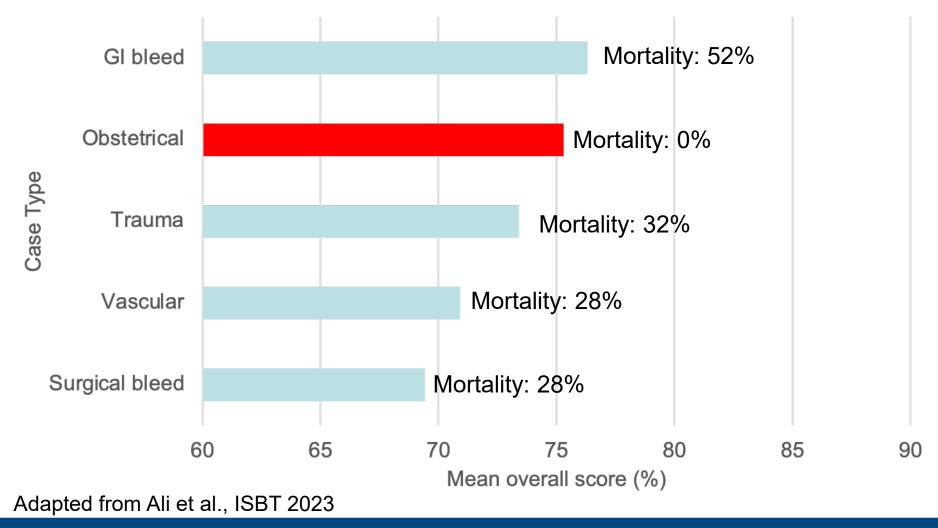
Was the MHP activation **appropriate** for this patient?

Were any blood products wasted during this activation?

SCORE: /9



#### MHP Case Score



#### ONTARIO MHP DATA IN OBSTETRICS

- This included just 18 pregnant patients
- Where do OB teams excel?
- Where do they need training?



#### **Bottom Line**

- Pregnancy hemorrhage ≠ trauma hemorrhage
  - These patients need unique mgmt. and dedicated training

- These are rare events that require simulation to maintain clinical currency
  - Simulation



#### THANK YOU FOR YOUR ATTENTION

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