

# A Silent Child's Voice at the Time of Injury: Taking on a Weighted System





Bellal Joseph, MD, FACS
Professor & Chief of Trauma and Acute Care Surgery
The University of Arizona, Tucson, AZ



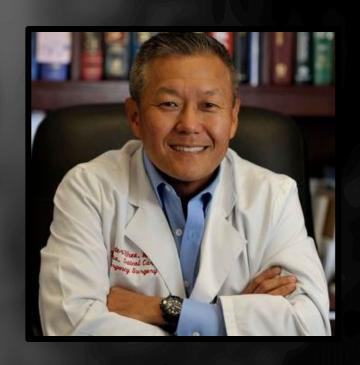
# This talk is like illusion; It alters with perspective





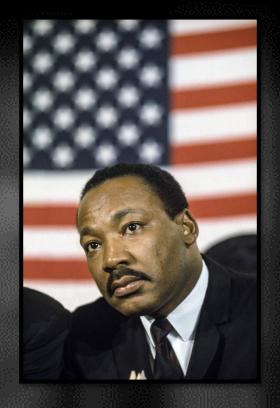


Inspiration





### Facing Our Uncomfortable Truths



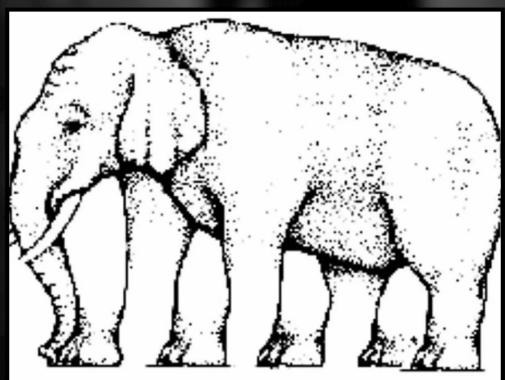
"Of all the forms of inequality,

injustice in health is the most shocking and inhumane"

- Martin Luther King Jr.

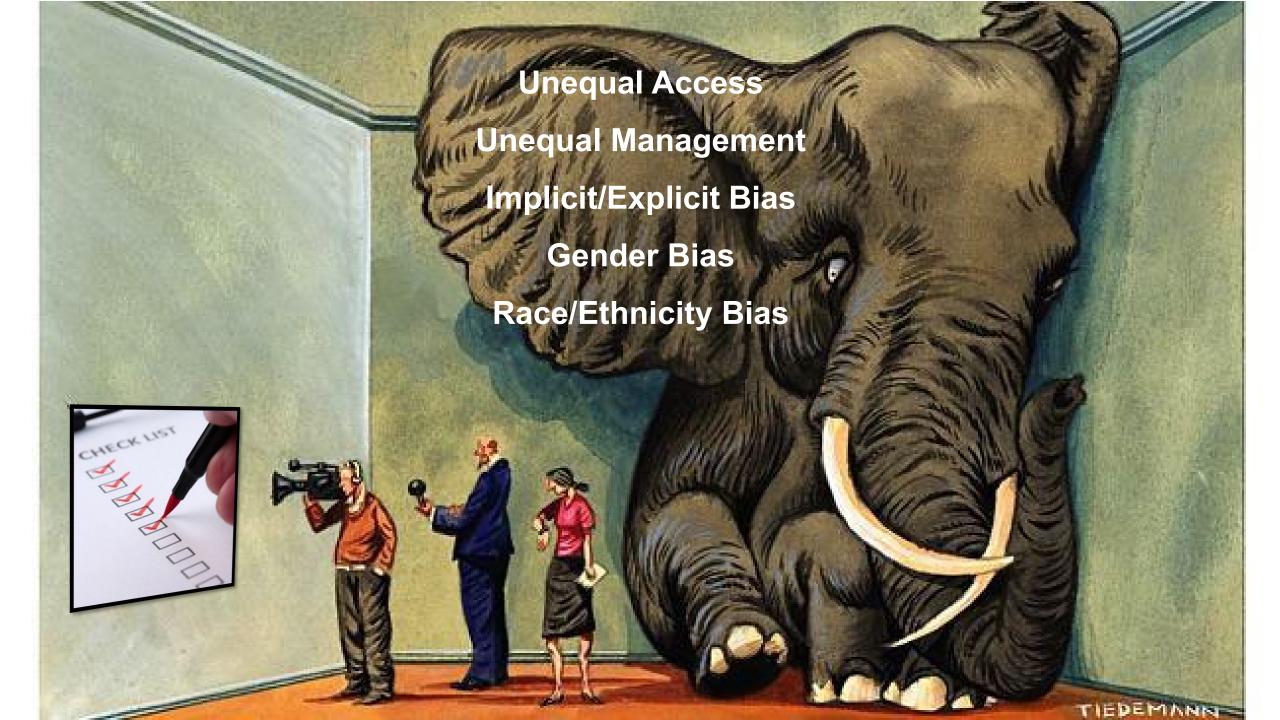
### Let's Pause for a Second













# TIME

**CODE WHITE** 





### Pediatric Surgery



- About 75% of APSA are male and white surgeons and
- Only 11% of Pediatric Surgery Fellowship Directors from 1937–2019 were Women
- First Black President of APSA → 2017 only 8 % have ever been women
- APSA DEI Committee → Established in 2018

## Exploring the gender gap: Letters of recommendation to pediatric surgery fellowship

Arika Hoffman, Rachel Ghoubrial, Melanie McCormick, Praise Matemavi, Robert Cusick



2020

- 364 LoRs for 49 female and 48 male applicants were reviewed
- Male applicant letters contain agentic terms & active possessive language
- Female applicant letters → socio-communal phrases & references to

spouse accomplishments

### Is It The Same Distance?



### **Advancements in Racial Representation**



#### **Advancements in Racial Representation**



Only 2.5% of U.S. physicians

were **Black** 



This fell to 2.2%

Advanceme

WHY?

resentation



Haruno, Lee S., et al. "Racial and Sex Disparities in Resident Attrition Among Surgical Subspecialties." JAMA surgery (2023).

# Inclusion and representation in the pediatric surgery workforce: Strategies to mitigate bias in the fellowship application process

Loren Berman a, Elizabeth Renaud b, Devon Pace a, Cynthia D. Downard c, Benedict C. Nwomeh d, Eunice Y. Huang e, Ying Z. Weatherall f, Samir K. Gadepalli g, Kevin P. Mollen h, Grace Z. Mak i, Erika Newman g, APSTPD DEI Committee



2022

Pediatric surgery workforce is 70% White

Pediatric surgeons at large do not reflect the populations they serve





## Physician-patient racial concordance and disparities in birthing mortality for newborns

Brad N. Greenwood, Rachel R. Hardeman, Laura Huang, and Aaron Sojourner

Proceedings of the National Academy of Sciences of the United States of America

National Academy of Sciences (U.S.)

Inputs both relation of the Impact both of Science of Magazine of Science of

2019

Black newborns cared for by Black physicians → 58% mortality compared to

Black newborns cared for by White physicians



### Disparities Begin as Early as From the Womb

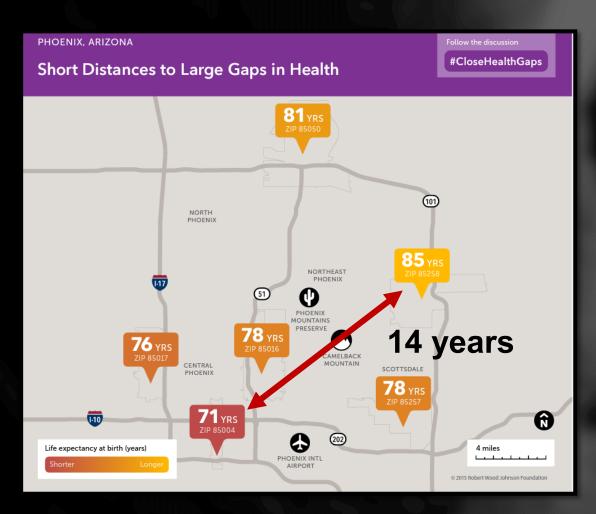




#### The Difference a Mile Can Make -











# Socioeconomic and Health Disparities Among Pediatric Trauma Patients in the United States



2021

National Inpatient Sample (2012-2015), 58,810 pediatric trauma patients

Black & Hispanics vs White











Non-Routine Discharge



#### **Care for Black Adolescents?**

- Black pts have higher mortality in hospitals with low proportion of black pts
- GSW Black adolescent pts (vs White): more likely triaged to ATCs than PTCs



- Andrade, Erin G., et al. "Racial disparities in triage of adolescent patients after bullet injury." Journal of trauma and acute care surgery 92.2 (2022)
- Kishawi, Sami K., et al. "Race and trauma mortality: The effect of hospital-level Black-White patient race distribution." Journal of Trauma and Acute Care Surgery 92.6 (2022)



## Racial/Ethnic Differences in Pediatric Emergency Department Wait Times

Jennifer R. Marin, MD, MSc; Jonathan Rodean, MPP; Matt Hall, PhD; et al



2022

- Aim: To determine racial/ethnic differences in ED wait times
- Pediatric Emergency Care Applied Research Network Registry 2016

#### **Compared to Non-Hispanic Whites**

**ED Wait Times** 



Hispanic (33%)





# Racial and Ethnic Differences in Emergency Department Diagnostic Imaging at US Children's Hospitals, 2016-2019

Jennifer R. Marin, MD, MSc; Jonathan Rodean, MPP; Matt Hall, PhD; et al



- Multicenter study of >13 million pediatric ED visits to 44 children's hospitals
- Data included from 2016-2019
- Outcome: receiving ED diagnostic imaging (X-ray, CT, US, & MRI)

Black & Hispanic children less likely to undergo ED imaging



White	Black	Hispanio
(34%)	(24%)	(26%)

# Racial and Ethnic Disparities in Pain Management of Children With Limb Fractures or Suspected Appendicitis: A Retrospective Cross-Sectional Study

frontiers in Pediatrics

2021

Romain Guedj, Maddalena Marini, Joe Kossowsky, Charles B. Berde, Amir A. Kimia and Eric W. Fleegler

- Children visiting the ED at Boston Children's Hospital (2011-2015)
- 8,347 children with limb fractures & 4,780 with suspected appendicitis
- Outcome: analgesic & opioid administration

**Compared to White non-Hispanic Children** 

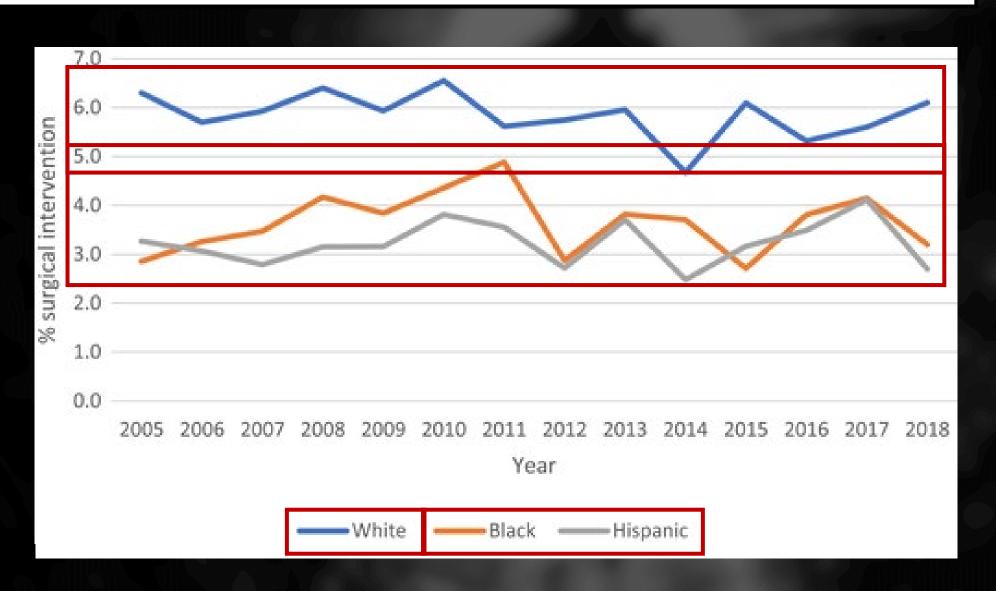




Surgical Interventions - Quantity & Quality

#### **Epidemiology of Pediatric Surgery in the United States**

Jennifer A. Rabbitts, Cornelius B. Groenewald





2021

## Patterns of Surgical Care and Health Disparities of Treating Pediatric Finger Amputation Injuries in the United States

Lee Squitieri MD, MS, Heidi Reichert MA, H. Myra Kim SCD, Justin Steggerda BA, Kevin C. Chung MD, MS



2011

#### **Digit Replantation**

**Compared to White Patients** 





Black (0.5x)



Hispanic (0.4x)



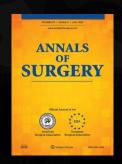


### Call For Action: Pediatric Firearm Injuries



## Pediatric Firearm Injuries and Fatalities: Do Racial Disparities Exist?

Sakran, Joseph V. MD, MPH, MPA; Nance, Michael MD; Riall, Taylor MD; Asmar, Samer MD; Chehab, Mohamad MD; Joseph, Bellal MD



2020

- Analysis of 2017 ACS-TQIP
- Pediatric (age ≤17 years) patients admitted with firearm injuries
- 3,717 pediatric firearm injuries: Blacks (67.0%) & Whites (33.0%)







Mental Health Care (x13 in Whites)

# Uncovering Gender, Racial, Ethnic, and Socioeconomic <a href="Disparities">Disparities</a> Among Adolescent Survivors of Suicide Attempts in Trauma Centers: Where Can We Do Better?





Uninsured Patients



Non-Pediatric
Trauma Centers







Mortality





## Nationwide Management of Trauma in Child Abuse

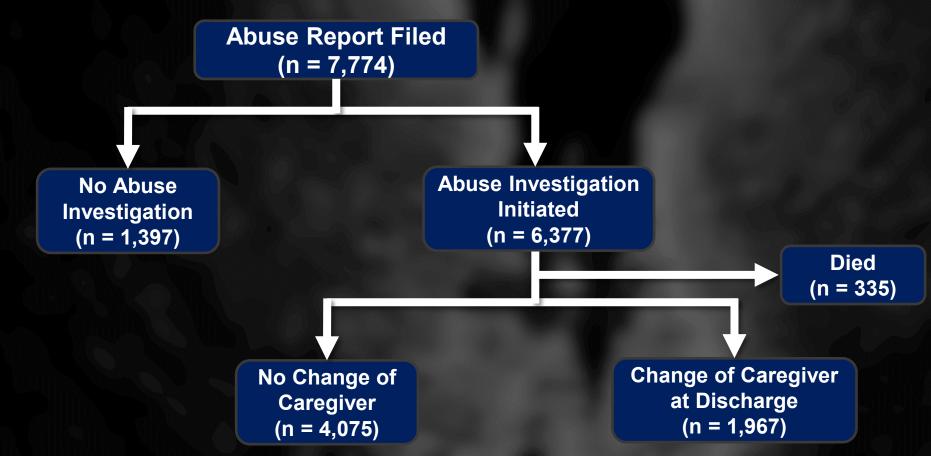
### **Exploring the Racial, Ethnic, & Socioeconomic Disparities**

Joseph, Bellal MD, FACS; Sakran, Joseph V. MD, MPH, MPA, FACS; Obaid, Omar MD; Hosseinpour, Hamidreza MD; Ditillo, Michael DO, FACS; Anand, Tanya MD, MPH; Zakrison, Tanya L. MD, MPH



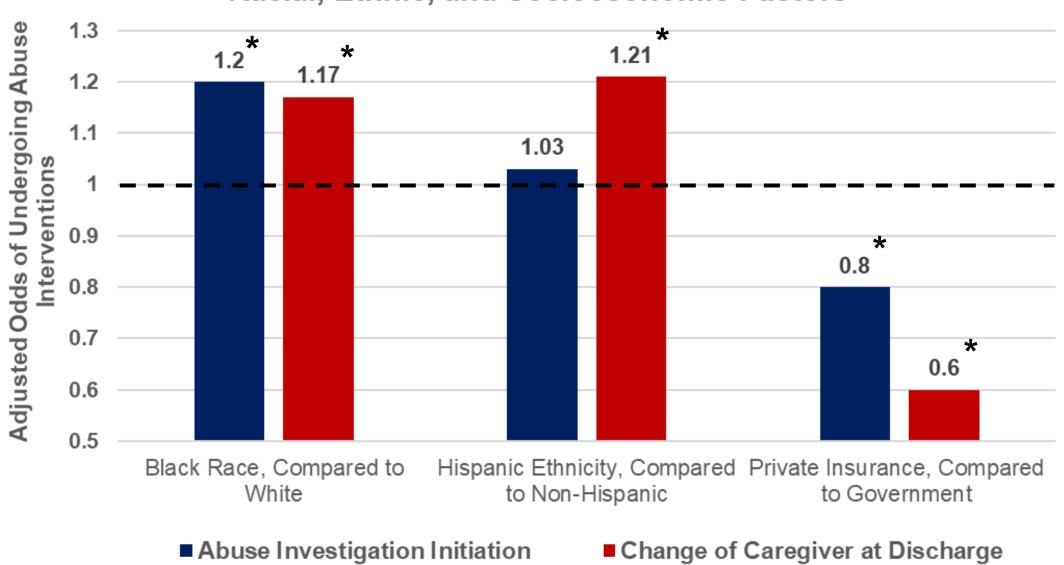
2022

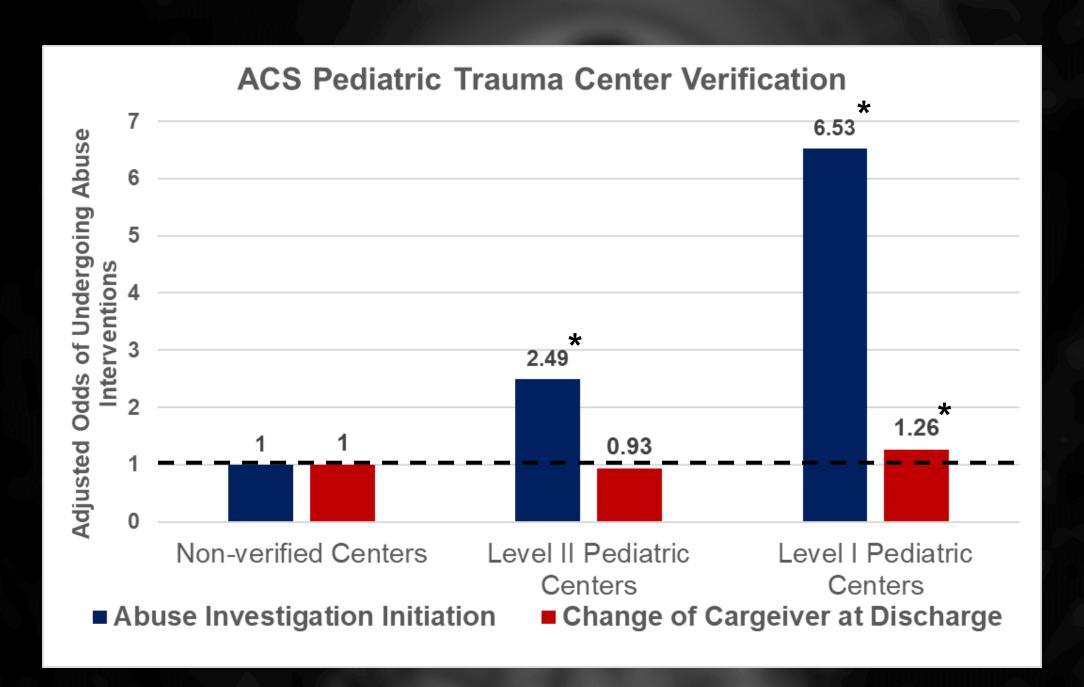
Analysis of 2017 to 2018 ACS-TQIP, child abuse victims





#### Racial, Ethnic, and Socioeconomic Factors





## Unequal access, unequal care, & unequal outcomes

**Suicide Attempts** 

**Child Abuse** 

**ED** Imaging

**Life Expectancy** 

**Wait Times** Pain Management

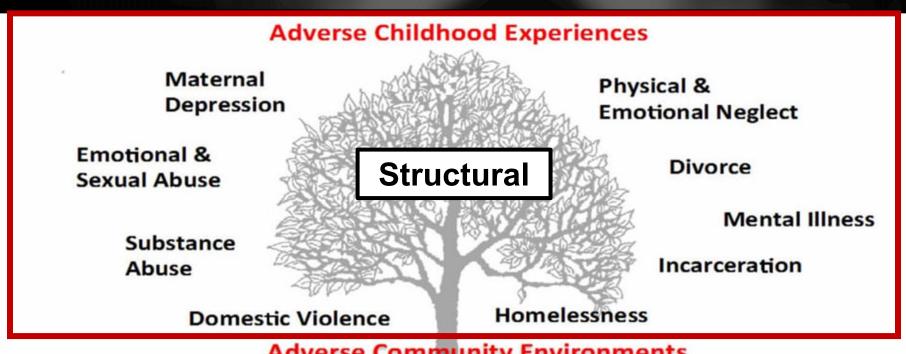
**Surgical Outcomes** 

**Gun Violence** 





## The Duo of Childhood Disparities



**Adverse Community Environments** 

**Poverty** 

Discrimination

Community Disruption

Lack of Opportunity, Economic Mobility & Social Capital

Violence

**Poor Housing** Quality & Affordability

## Adverse Childhood Experiences (ACEs)



Substance Abuse

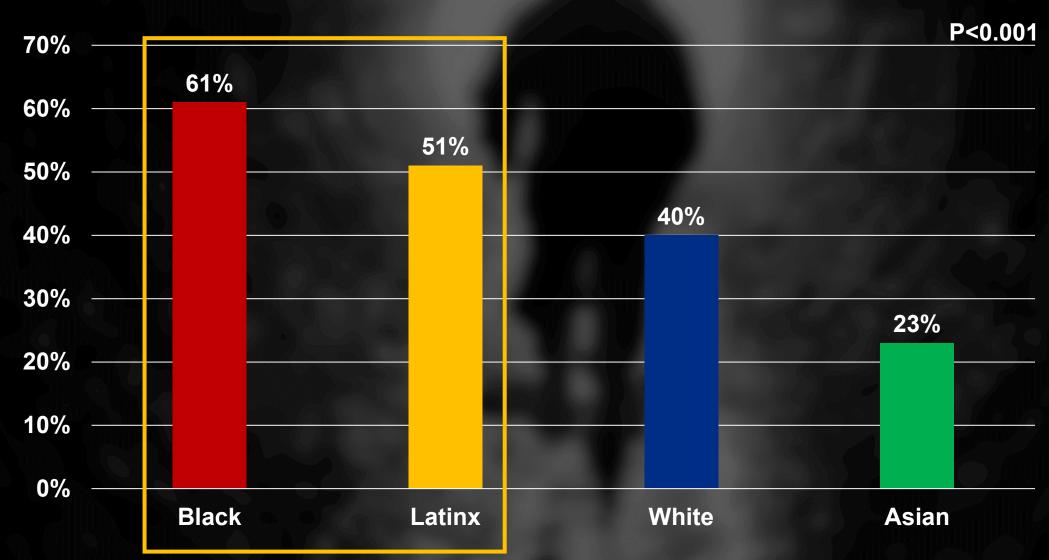
### **Traumatic Childhood Experiences**

- Experiencing or Witnessing
  - Violence, Abuse, or Neglect
- Household Challenges



## Experiencing at least one ACE – United States





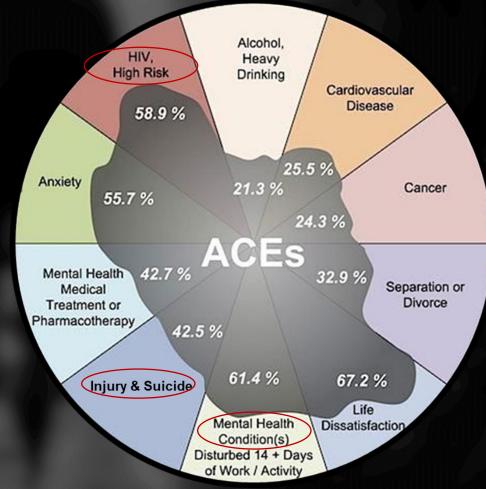
## Association of Childhood Trauma Exposure With Adult Psychiatric Disorders and Functional Outcomes



William E. Copeland, PhD; Lilly Shanahan, PhD; Jennifer Hinesley, PsyD; et al

2018

### Consequences of ACEs



## The Duo of Pediatric Disparities



Emotional & Sexual Abuse

> Substance Abuse

> > **Domestic Violence**

Physical & Emotional Neglect

Divorce

**Mental Illness** 

Incarceration

Homelessness

#### **Adverse Community Environments**

Poverty

Discrimination

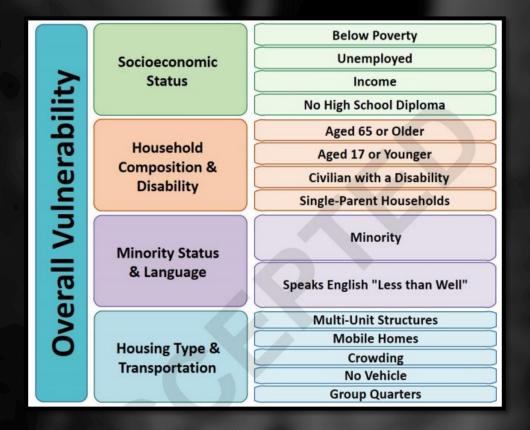
Community Disruption

Lack of Opportunity, Economic Mobility & Social Capital Violence

Poor Housing Quality & Affordability

#### Social vulnerability Index (SVI) – Four Themes

- Identifies communities vulnerable to human/economic loss after disasters
- 0 (low vulnerability) to 1 (high vulnerability) national SVI percentile rank
- Strongly Associated with worse health outcomes

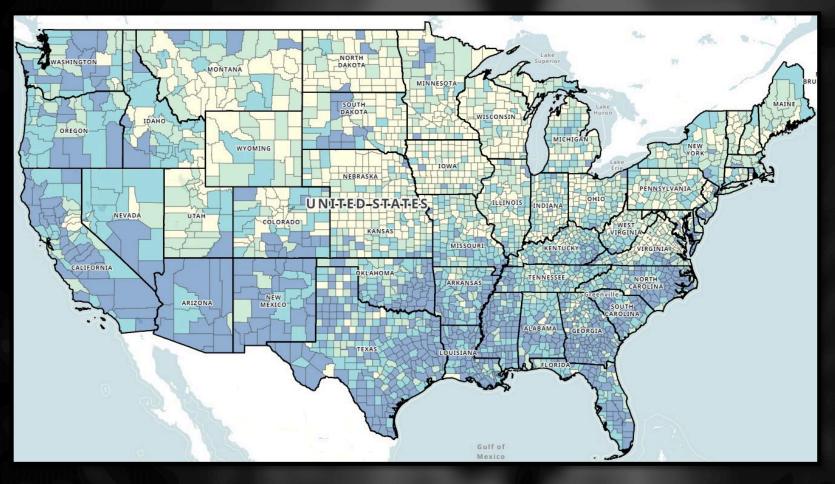






## **High-Variability** in SVI Across Counties





**Level of Vulnerability** 

Low Low-Medium Medium-High High

# Social Vulnerability Index is Strongly Associated with Urban Pediatric Firearm Violence: An Analysis of Five Major U.S. Cities

Polcari, Ann M. MD, MPH, MSGH; Hoefer, Lea E. MD; Callier, Kylie MD; Zakrison, Tanya L. MD, MPH; Rogers, Selwyn O. MD, MPH; Henry, Marion MD, MPH; Slidell, Mark B. MD, MPH; Benjamin, Andrew J. MD, MS



2023

- Included 5 Major U.S. Cities: BAL, CHI, LA, NYC, & PHL (2015-2021)
- To assess association between social vulnerability & pediatric firearm injuries



Los Angeles



**Baltimore** 



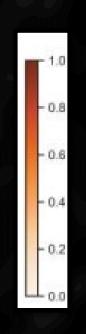
Chicago



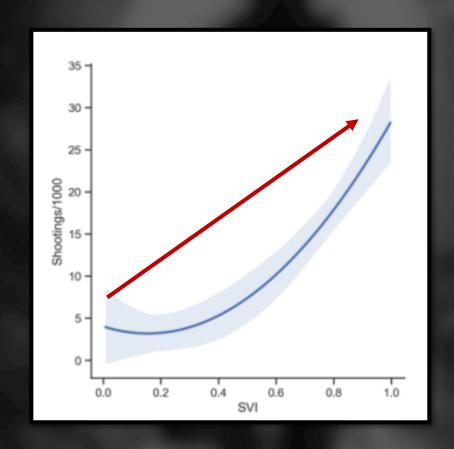
NYC



**Philadelphia** 



### **SVI & Shooting Incidents/1000**

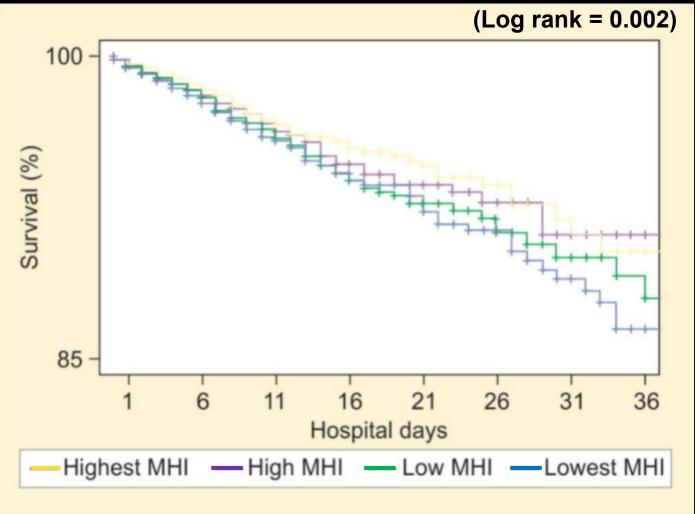


Increasing SVI was strongly associated with an increasing incidence of pediatric shooting incidents/1000

#### **Does Mon**

Bernardino C Branco, Ma

- The Arizona
- To investige
- 58,743 pts
- Lowest MH



hold

rew L Tang,



2019

mortality

ghest MHI: 20.2%



Stepwise decrease in mortality as MHI increased

## Persistency of Poverty and its Impact on Surgical Outcomes

Medicare Standard Analytical Files
Database

336,887 Medical Beneficiaries underwent lung resection, colectomy, CABG or lower extremity joint replacement



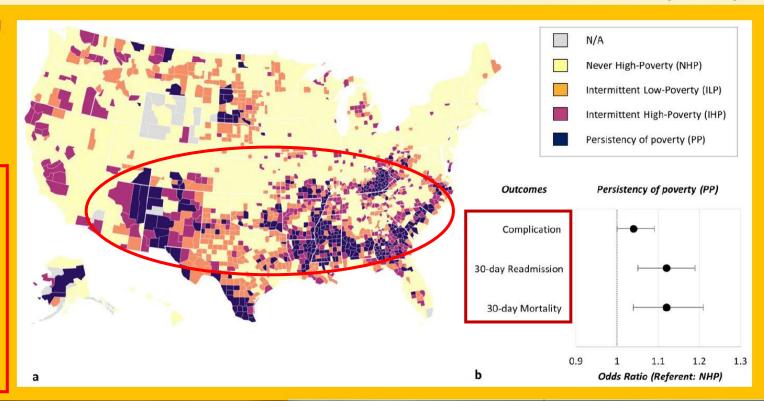
PP patients were more likely to present at younger median age, be non-White and have a higher cost of care related to surgical episode



PP patients had higher likelihood of postoperative complications, 30-day mortality, and 30-day readmission

2015-2017

County-Level Poverty Assessed with American Community Survey







## THIS IMPACTS YOU







### The Patient Protection & Affordable Care Act

- One of the most historic reforms in the US health system
- Signed into law on March 23, 2010
- Expanded access to affordable health insurance coverage to all Americans

Medicare reforms

**Expansion of Medicaid eligibility** 

Prevents insurance from denying owing to pre-existing illnesses

## Early and Periodic Screening, Diagnostic, & Treatment (EPSDT)

Medicaid's EPSDT Child Health Benefit Program

Despite federal EPSDT requirements, states vary greatly in the extent to which they cover services for children

**► What is the Impact?** 

## Counterefforts

- In 2011, the US Department of Health and Human Services (HHS) launched the largest US federal action plan for reducing healthcare disparities:
  - 1. Streamlined policy & government efforts to reduce disparities
  - 2. Assessed resource allocation
  - 3. Increased data availability to improve minority population health
  - 4. Provided incentives for quality care of minority populations





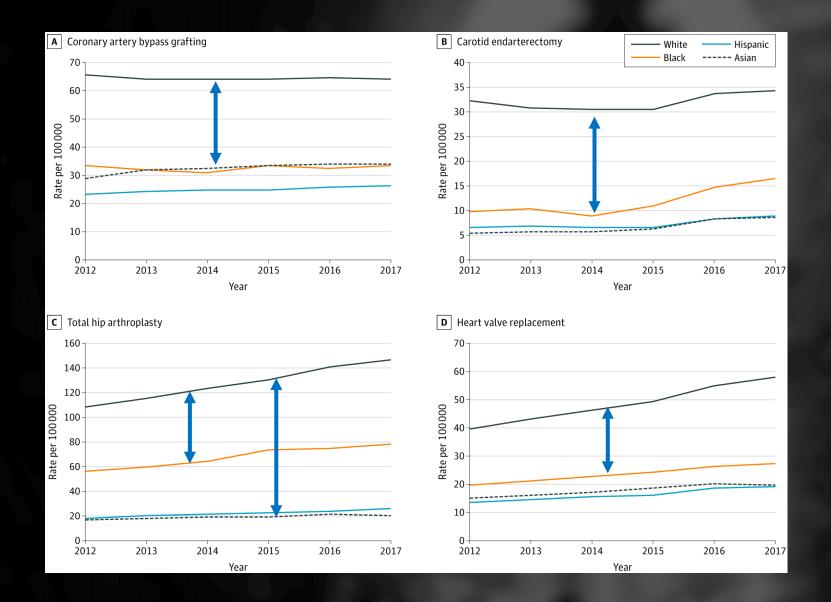
## Racial Disparities in the Use of Surgical Procedures in the US

Matthew J. Best, MD; Edward G. McFarland, MD; Savyasachi C. Thakkar, MD; Uma Srikumaran, MD, MBA, MPH



- National case-control study (2012-2017)
- To analyze whether HHS national initiatives improved:
  - Racial disparities in the use of 9 surgical procedures in the US
  - Procedures that are previously shown to have racial disparities

## By 2017, racial disparities persisted for all 9 procedures



## The impact of patient protection and Affordable Care Act on trauma care: A step in the right direction

Joseph, Bellal MD; Haider, Ansab A. MD; Azim, Asad MD; Kulvatunyou, Narong MD; Tang, Andrew MD; O'Keeffe, Terence MD; Latifi, Rifat MD; Green, Donald J. MD; Friese, Randall S. MD; Rhee, Peter MD, MPH



- Analysis of Level I TC registry (2012-2014), 9,892 patients
- To assess impact of ACA on insurance, reimbursements, & outcomes





## Does Medicaid Insurance Provide Sufficient Access to Pediatric Orthopedic Care Under the Affordable Care Act?

Global Pediatric Health



Jenny Nguyen, BS, Nidharshan S. Anandasivam, MD, Daniel Cooperman, MD, Richard Pelker, MD, PhD, and Daniel H. Wiznia, MD

Aim: Assess access to pediatric orthopedic urgent care for a child's likely

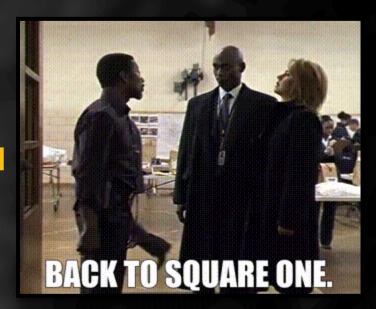
2019

- operative distal radius fracture
- Phone call to 180 pediatric orthopedic surgeons in 8 states requesting appointments for caller's fictitious 11-year-old child with distal radius fracture

**Ability to Schedule Appointments** 

Privately insured (83%)

Medicaid (38%)



## INJURY PREVENTION



## The Impact of **Disparities** in Pediatric Trauma on **Injury-Prevention** Initiatives

Fallat, Mary E. MD; Costich, Julia PhD, JD; Pollack, Susan MD



- Injury prevention initiatives have had little impact
- Interactions of race, ethnicity, language, culture, environment, socioeconomic status, and access to health care must be analyzed and understood
- Interventions should be targeted towards most vulnerable children



## Failures Everywhere

**Triage** 

**In-hospital Care** 

Interventions

**Outcomes** 

**Injury Prevention** 

**National Initiatives** 





Race Bias Religious Bias Macroaggressions **Country of Origin Bias** Comes in Many Shapes & Forms **Gender Bias** 

**Implicit & Explicit Bias** 

**Ethnicity Bias** 

## Conscious & Unconscious Bias

### **EXPLICIT BIAS**

Attitudes and beliefs that we have about a person or group on a conscious level. We are fully aware of these so they can be self reported.

**UNCONSCIOUS BIAS:** 

SIMILAR TO AN ICEBERG, ONLY
A SMALL PORTION OF THE MIND
IS OBVIOUS TO US.

Line of consciousness

### **IMPLICIT BIAS**

Unconscious attitudes that lie below the surface, but may influence our behaviors.



## THIS MAN MAKES YOU THINK



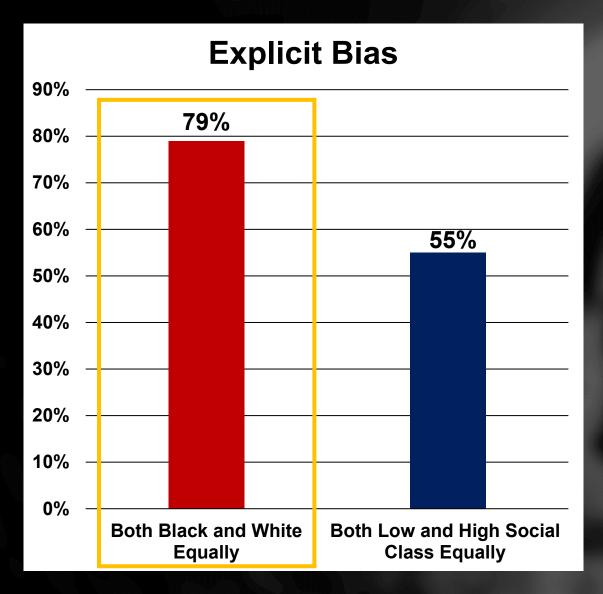
# Unconscious race and class bias Its association with decision-making by trauma and acute care surgeons

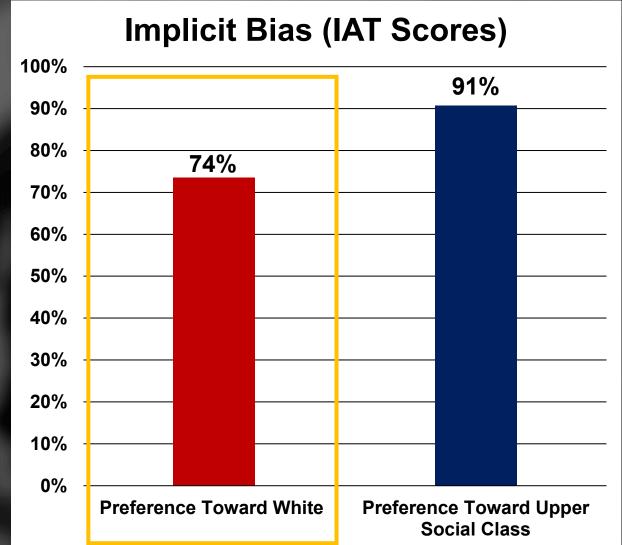
Haider, Adil H. MD, MPH; Schneider, Eric B. PhD; Sriram, N PhD; Dossick, Deborah S. MD; Scott, Valerie K. MSPH; Swoboda, Sandra M. RN; Losonczy, Lia MD, MPH; Haut, Elliott R. MD; Efron, David T. MD; Pronovost, Peter J. MD, PhD; Freischlag, Julie A. MD; Lipsett, Pamela A. MD; Cornwell, Edward E. III MD; MacKenzie, Ellen J. PhD; Cooper, Lisa A. MD, MPH



- Survey of 248 EAST members
- Nine clinical vignettes (trauma/ACS management questions)
- Race & Social Class Implicit Association Tests (IAT) completed







Clinical decisions associated with implicit bias

### **Understanding & Countering Implicit Bias**

#### **Bias We All Share**



Implicit bias is the automatic or involuntary attitudes we have about members of distinct social groups that unconsciously affect our beliefs or actions

#### **Find Your Blind Spots**



Implicit Association Testing can help identify where you might be most biased, but self-reflection to assess if your automatic reactions may be biased is also warranted

#### **Transparent Criteria**



You can't eradicate automatic thoughts, but you can develop systems around hiring, promotion, and evaluation to avoid formally codifying these implicit biases

EAST Equity, Quality and Inclusion in Trauma Surgery Practice Ad Hoc Task Force



Visual abstract by @CAHarrisMD



http://bit.ly/EAST4ALL



Eastern Association for the Surgery of Trauma Advancing Science, Fostering Relationships, and Building Careers

## **Harvard Implicit Association Test**













### The Role of Professional Societies







## We are Evolving

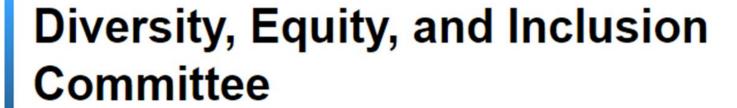
COMMITTEES

/ Committee on Diversity Issues



STATEMENTS

/ Statement on Diversity



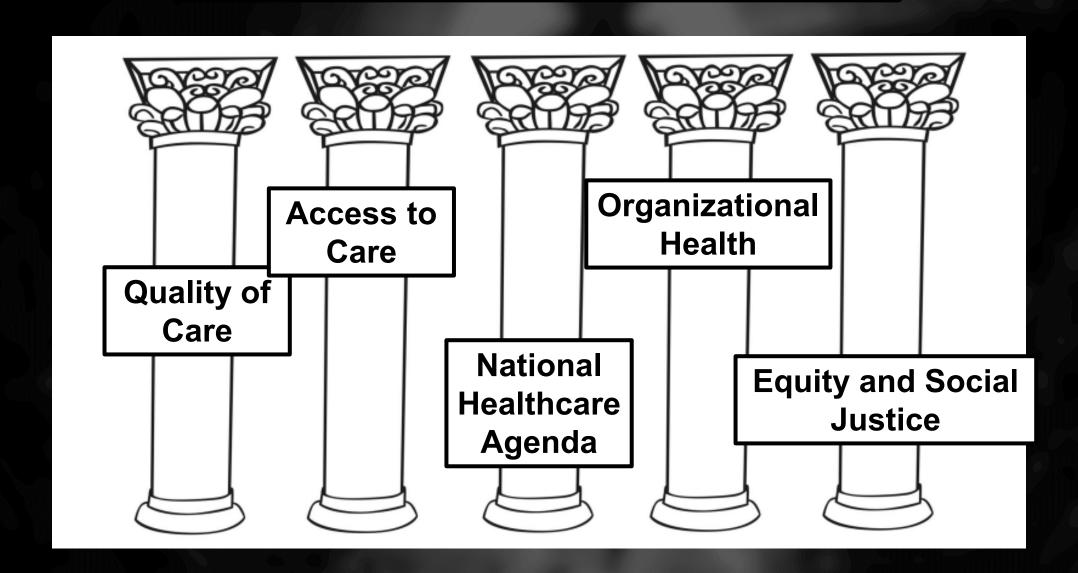




**Diversity, Equity and Inclusion Committee** 



### The Five Pillars of APSA



# **Best Practices To Mitigate Bias & Support Diverse Pediatric Surgery Applicants**



#### **DEI** subcommittee

#### **Strategies**

#### Application screening

All applications should be reviewed by a diverse team.

Remove candidate photos from ERAS (Electronic Residency Application Service) applications before screening.

Read personal statements first.

Selection criteria should center on life experiences (e.g., challenges and obstacles overcome) and attributes as well as academic performance.

Widen the lens through which you select residents to interview (identify experiences, attributes, competencies, and metrics that would add value to your program).

#### Preparing for interviews

Consider how your program presents itself to applicants. Review all online content (hospital website, ERAS, etc.) to ensure that message of inclusivity and valuing diversity is projected.

All faculty should undergo implicit bias training.

All faculty should take at least two implicit association tests.

#### **Conducting interviews**

Conduct CV-blinded interviews: faculty have access only to the applicant's personal statement.

Interviews should be structured, using a consistent list of questions for all applicants, and include attribute-based questions.

Be aware that virtual interview platforms introduce additional bias against underrepresented in medicine learners and low socioeconomic applicants.

Implement bias mitigation strategies including common identity formation, perspective taking, consider the opposite, and counter stereotypical exemplars.

Utilize an interview scoring form that encourages consideration of non-traditional metrics.

#### Creating the rank list

Maximize objective assessment and use of objective scoring systems.

When the comment is made "I'm not sure if this candidate is a good fit for our program," force the speaker to go deeper.

Avoid the practice of phone calls within our networks.

#### **Standardized Scoring Forms**

Category			SCORING			Max Possible	Relative Weight
Letters of Recommendation - 1	0 Fair Support Problems Noted	1 Average Support	2 Above Average	3 Strong Support	4 Outstanding	4	1
Letters of Recommendation - 2	0 Fair Support Problems Noted	1 Average Support	2 Above Average	3 Strong Support	4 Outstanding	4	1
Letters of Recommendation - 3	0 Fair Support Problems Noted	1 Average Support	2 Above Average	3 Strong Support	4 Outstanding	4	1
Letters of Recommendation - 4	0 Fair Support Problems Noted	1 Average Support	2 Above Average	3 Strong Support	4 Outstanding	4	1
General Surgery Residency	0 Not US or Canadian	1 Community Hospital	2 Non-University Academic	3 Academic Medical Center	4 "Top Tier"	30	7.5
Related Research, Community Work, Innovation, or Global Surgery	0 None	1 <1 year or pre-medical only	2 1 year	3 1 year - exceptional research, community work, or global surgery	4 2 years	6	1.5
Clinical or Basic Research Publications	0 None	1 1-3 papers	2 4 or More: First Authors	3 6 or More with First Authors	4 S or More with First Authors	12	3
Distance traveled (perserverance, endurance, commitment)	0 None identifiable	1 Minor setbacks	2 Adversity or hurdles	3 Unique journey or non- traditional path	4 Major setback or clear disadvantage	15	3.75
Achievement	1 Other Advance Degree	2 Grant Funding	3 Superior Academic Achievement	4 Athletic Accomplishment	5 Leadership	10	2
Hardiness, Grit & Resilience (self awareness, positive relationships, purpose and passion)	0 None identifiable, problems noted	1 Solid ability to navigate challenges	2 Demonstrated growth and dedication	3 Demonstrated endurance and high perseverance for long- term goals	4 Exceptional vision, courage, purpose or passion	15	3.75
ABSITE Scores	0 <20	1 20-40	2 >40	3 >60	4 >80	12	3
Personal Statement	0 Little Thought/Poorly Organized	1 Not insightful	2 Solid	3 Excellent	4 Exceptional	12	3
					Total	128	

# **Best Practices To Mitigate Bias & Support Diverse Pediatric Surgery Applicants**





# #EAST4ALL: An introduction to the EAST equity, quality, and inclusion task force

Bonne, Stephanie MD; Williams, Brian H. MD; Martin, Matthew MD; Kaafarani, Haytham MD; Weaver, William L. MD; Rattan, Rishi MD; Byers, Patricia M. MD; Joseph, D'Andrea K. MD; Ferrada, Paula MD; Joseph, Bellal MD; Santos, Ariel MD; Winfield, Robert D. MD; DiBrito, Sandra MD, PhD; Bernard, Andrew MD; Zakrison, Tanya L. MD



- First task force of its kind in the world of trauma & ACS
- Diverse "radically inclusive"
- To raise awareness & provide resources to combat inequities
- Chaired the committee for 4 years



Dr. Bernard

# History of Equity, Diversity, & Inclusion in Trauma Surgery for Our Patients, for Our Profession, & for Ourselves

Esther S Tseng, Brian H Williams, Heena P Santry, Matthew J Martin, Andrew C Bernard, Bellal A Joseph

# Perceptions of Equity & Inclusion in Acute Care Surgery From the #EAST4ALL Survey

Tseng, Esther S. MD; Zakrison, Tanya L. MD, MPH; Williams, Brian MD; Bernard, Andrew C. MD; Martin, Matthew J. MD; Zebib, Laura MPH; Soklaridis, Sophie PhD; Kaafarani, Haytham M. MD, MPH; Zarzaur, Ben L. MD, MPH; Crandall, Marie MD, MPH; Seamon, Mark J. MD; Winfield, Robert D. MD; Bruns, Brandon MD

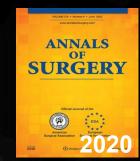
# Call to Action on the Categorization of Sex, Gender, Race, and Ethnicity in Surgical Research

Nahmias, Jeffry MD, MHPE, FACSa,\*; Zakrison, Tanya L. MD, MPH, MHSc, FACSb; Haut, Elliott R. MD, PhD, FACSd; Gurney, Onaona MD, FACSg; Joseph, Bellal MD, FACSh; Hendershot, Kimberly MD, FACSi; Ghneim, Mira MD, MS, FACSe; Stey, Anne MD, FACSc; Hoofnagle, Mark H. MD, PhD, FACSj; Bailey, Zinzi ScD, MSPHk; Rattan, Rishi MD, FACSl; Richardson, Joseph B. PhDf; Santos, Ariel P. MD, MPH, FACSm; Zarzaur, Ben MD, MPH, FACS

# EAST Statement on Structural Racism, and the Deaths of George Floyd, Ahmaud Arbery, and Breonna Taylor

Hoofnagle, Mark H. MD, PhD; Mubang, Ronnie N. MD; Joseph, D'Andrea K. MD, FACS; Joseph, Bellal A. MD, FACS; Christmas, Ashley Britton MD, FACS; Zakrison, Tanya L. MD, MPH, FACS, FRCSC















The #MGHSurgery Residency Program has, for the third consecutive year, ranked 1st in the nation by @doximity! This top ranking is recognition of our amazing residents, faculty & staff. Med students, find us on Doximity's residency navigator: bit.ly/31pMxqU #WhyMGHSurgery



MassGeneral News

6:24 AM · Aug 5, 2020 · Twitter Web App



**EQUITY AND SOCIAL JUSTICE** 

The State of Diversity in American Surgery

A Call to Action

Ellis, Danielle I. MD, MTS\*; Khi

Stanford's 2023 surgical team:

92% women (US: 50% women) 69% nonwhite (US: 38% nonwhite) 0% white men (US: 30% white men)



CYMI: Meet the newest additions to the Stanford Surgery Family! #match2023 surgery.stanford.edu/news2/Match202...



:00 AM - Mar 22, 2023 - 3.6M Views





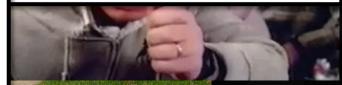


2021



**#MGHSurgerySoWhite** 

165 Bookmarks





Q 2,425

17 4,061

O 24.5K

III 8.7M



# Prevalence of unprofessional social media content among young vascular surgeons

Hardouin, Scott MDCheng, Thomas W. MSMitchell, Erica L. MDRaulli, Stephen J. MPhilJones, Douglas W. MD, MPHSiracuse, Jeffrey J. MDFarber, Alik MD, MBA



2020

#### Conclusion

- Images of residents in which they wore bikinis, held glasses of alcohol or expressed social or political views were unprofessional

Diversity, Inclusion, and Equity: Evolution of Race and Ethnicity Considerations for the Cardiology Workforce in the United States of America From 1969 to 2019



2020

Norman C. Wang

Conclusion

- "Failures of efforts to increase the number of Blacks and Hispanics have largely been attributable to the limited qualified applicant pool"
  - Called for admissions committees to abandon any diversity directives

### **Retracted Before Issue**



# JAMA Editor Placed on Leave After Deputy's Comments on Racism

After a staff member dismissed racism as a problem in medicine on a podcast, a petition signed by thousands demanded a review of editorial processes at the journal.

Diversity, Inclusion, and Equity: Evolution of Race and Ethnicity Considerations for the Cardiology Workforce in the United States of America From 1969 to 2019

### are posting photos of themselves in bathing suits

Authors apologize after uproar over study calling certain personal social media posts "potentially unprofessional."



# For Our Patients



## **Beyond Survival & Physical Healing**



#### **Trauma-informed care:**

### recognizing and resisting re-traumatization in health care

Samara Grossman, Zara Cooper, Heather Buxton, Sarah Hendrickson, Annie Lewis-O'Connor, Jane Stevens6, Lye-Yeng Wong, Stephanie Bonne



Traditional definitions of trauma as a purely physical phenomenon are prevalent





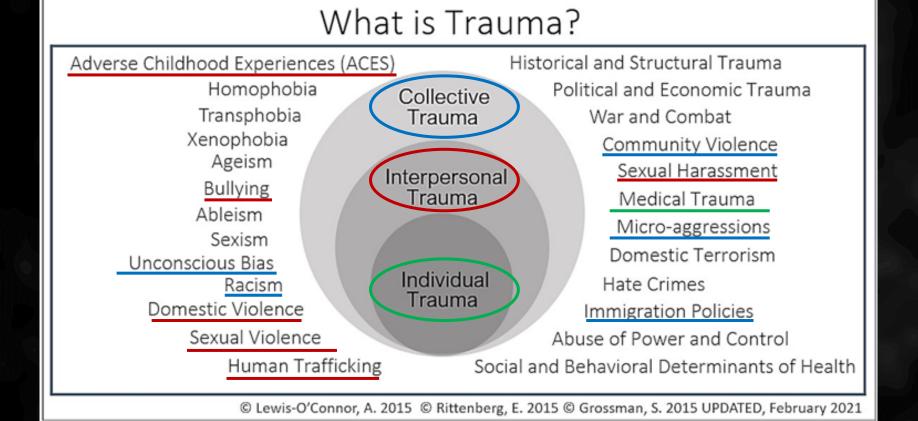
Trauma: "traumatic injury includes that from vehicular collisions, falls from heights, gunshot wounds & burns...





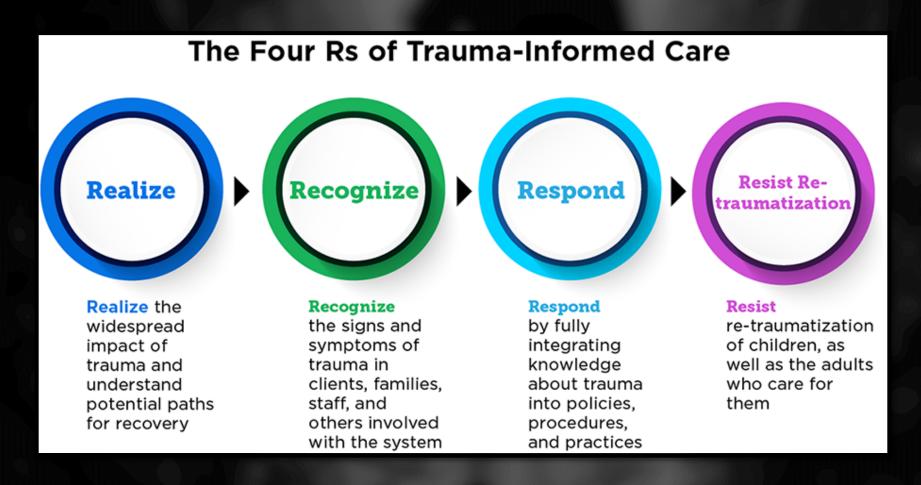
Trauma is defined as both individual and interpersonal as well as collective and structural

### What is Trauma?



### Trauma-Informed Approach

- Systematic consideration of past physical & emotional trauma
- Patient- centered, trauma- informed approach to equitable care is key



### **Building A Trauma-informed Organization -**

10 Domains of Trauma-informed Organizational Leadership







## Don't Forget the 5 S's:

Check your own biases

**Start Open Discussions** 

**Set Accountability** 

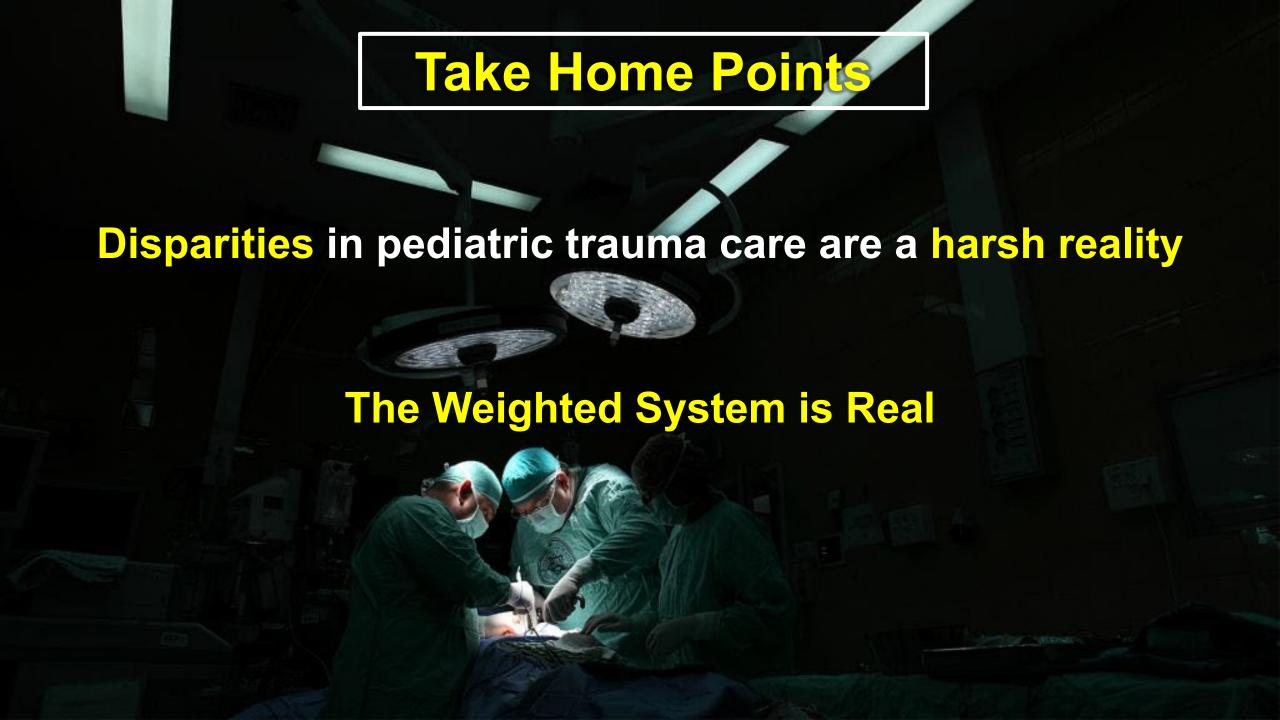
Speak Up

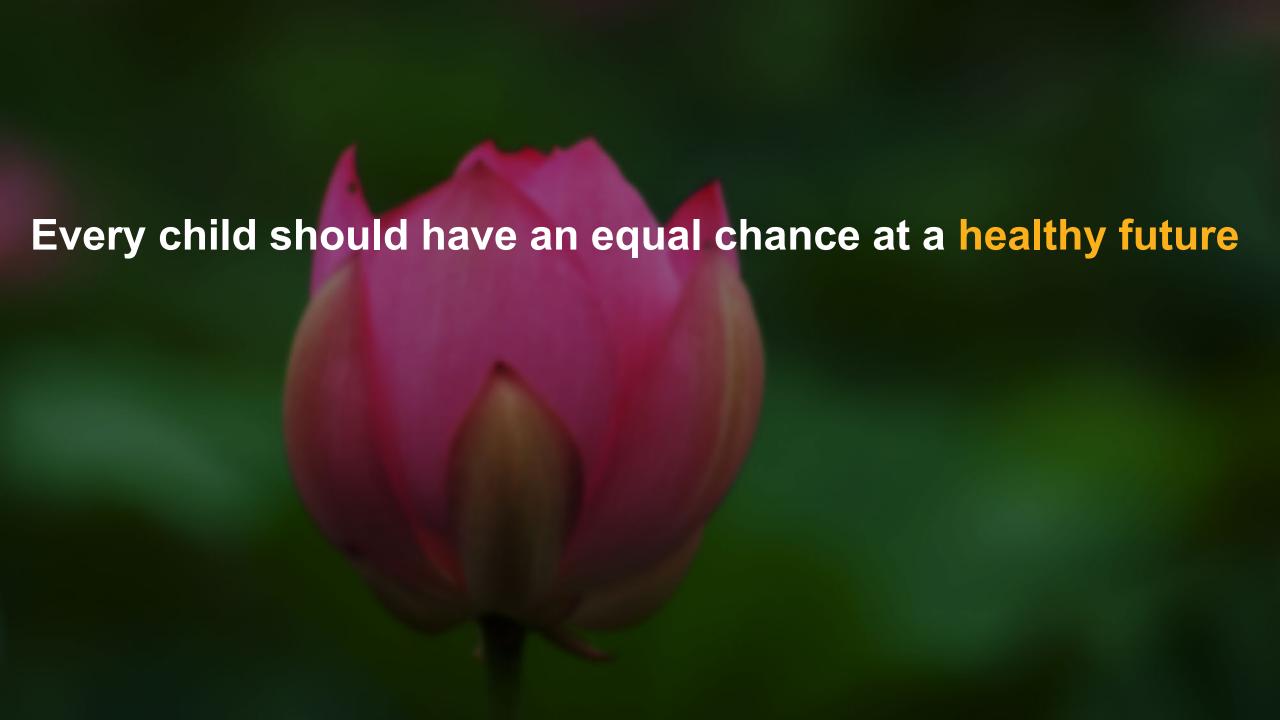
**Share The Responsibility** 

# This talk is like illusion; It alters with perspective















# Low Titer Group O Whole Blood for Pediatric Hemostatic Resuscitation

Phil Spinella, MD, FCCM
Professor, Surgery and Critical Care Medicine
University of Pittsburgh
July 2023

University of

### **Disclosures**

- Consultant
  - Hemanext, Haima, Cerus, Octapharma
- Co-Founder and Chief Medical Officer
  - Kalocyte



### **Severe Bleeding Has Poor Outcomes**

Adults 28 day Mortality<sup>2,3</sup>

- Trauma: 20-24%

Pediatric 28 day Mortality<sup>1</sup>

- Operative: 24%

Trauma: 37%

- Medical: 62%

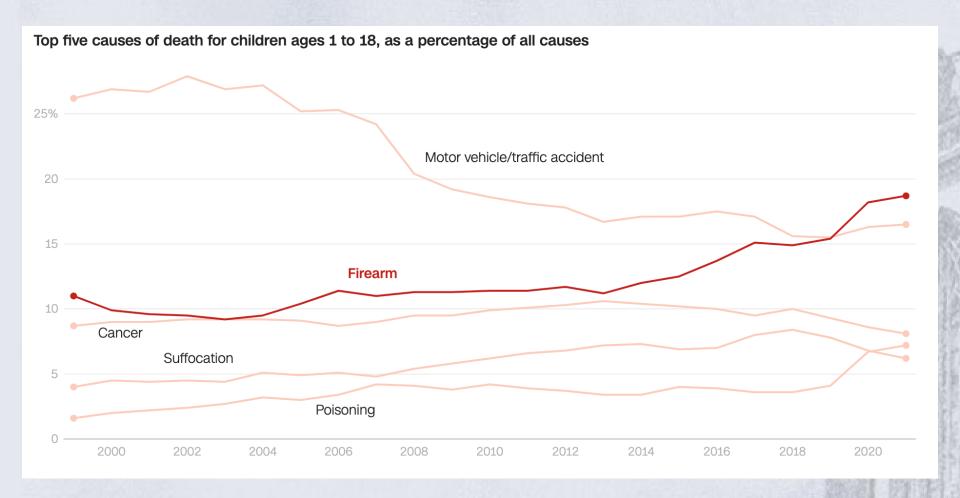
- 1. Leonard J and Spinella PC
- 2. Holcomb JB, PROMMT
- 3. Holcomb JB, PROPPR



### **Traumatic Injury Outcomes**

- Most common cause of mortality 1-46 years of age
- 30,000 preventable deaths per year in US afterinjury from traumatic bleeding
  - 2,000 preventable deaths in children per year in the US
    - Inadequate or untimely care





### Damage Control Resuscitation: Directly Addressing the Early Coagulopathy of Trauma

John B. Holcomb, MD, FACS, Don Jenkins, MD, FACS, Peter Rhee, MD, FACS, Jay Johannigman, MD, FS, FACS, Peter Mahoney, FRCA, RAMC, Sumeru Mehta, MD, E. Darrin Cox, MD, FACS, Michael J. Gehrke, MD, Greg J. Beilman, MD, FACS, Martin Schreiber, MD, FACS, Stephen F. Flaherty, MD, FACS, Kurt W. Grathwohl, MD, Phillip C. Spinella, MD, Jeremy G. Perkins, MD, Alec C. Beekley, MD, FACS, Neil R. McMullin, MD, Myung S. Park, MD, FACS, Ernest A. Gonzalez, MD, FACS, Charles E. Wade, PhD, Michael A. Dubick, PhD, C. William Schwab, MD, FACS, Fred A. Moore, MD, FACS, Howard R. Champion, FRCS, David B. Hoyt, MD, FACS, and John R. Hess, MD, MPH, FACP

J Trauma. 2007;62:307-310.

### Table 1. Damage control resuscitation principles

#### Pre-hospital

Rapid recognition of life-threatening hemorrhagic shock

Point-of-care devices: near infrared spectroscopy; INR; lactate level may be of value

Prevent hypothermia

Hemorrhage control with mechanical hemostatic adjuncts:

Tourniquet/junctional tourniquet

Pressure dressings/thrombin and fibrin-impregnated gauze

**REBOA** 

Intraabdominal foams (investigational)

Hemostatic resuscitation

Whole blood is optimal

Component therapy with plasma (dried, liquid, or thawed), RBCs, and platelets in 1:1:1 ratio

Permissive hypotension for patients without traumatic brain injury<sup>a</sup>

Avoid crystalloid resuscitation

Consider TXA administration if less than 3 h from time of injury<sup>b</sup>

Consider source of fibrinogen (fibrinogen concentrate or cryoprecipitate)

Avoid hypocalcemia

In prolonged evacuations, empiric calcium administration for every 4-6 units of RBCs or WB



## Every minute counts: Time to delivery of initial massive transfusion cooler and its impact on mortality

J Trauma Acute Care Surg Volume 83, Number 1

David E. Meyer, MD, Laura E. Vincent, RN, Erin E. Fox, PhD, Terence O'Keeffe, MBChB, Kenji Inaba, MD, Eileen Bulger, MD, John B. Holcomb, MD, and Bryan A. Cotton, MD, Houston, Texas

**TABLE 3.** Multivariate Regression Predicting 30-d Mortality

	OR	95% CI	p
Time to receipt of first cooler, min	1.05	1.01-1.09	0.016
Anatomic injury severity (ISS)	1.05	1.03-1.06	< 0.001
Disturbed arrival physiology (w-RTS)	0.61	0.53-0.69	< 0.001
Randomization group (1:1:2)	1.46	0.92 - 2.29	0.102
RI, units	1.03	0.60-1.44	0.184

Median (IQR) time from arrival to MTP activation was 9 (3-20) min Median (IQR) time from MTP activation to delivery of blood products was 8 (5-11) min



## Delay in Recognition/Treatment in Children with Life-Threatening Hemorrhage

### Life-Threatening Bleeding in Children: A Prospective Observational Study

**OBJECTIVES:** The purpose of our study was to describe children with life-threatening bleeding.

**DESIGN:** We conducted a prospective observational study of children with life-threatening bleeding events.

SETTING: Twenty-four childrens hospitals in the United States, Canada, and Italy participated.

SUBJECTS: Children 0–17 years old who received greater than 40 mL/kg total blood products over 6 hours or were transfused under massive transfusion protocol were included.

**INTERVENTIONS:** Children were compared according bleeding etiology: trauma, operative, or medical.

MEASUREMENTS AND MAIN RESULTS: Patient characteristics, therapies administered, and clinical outcomes were analyzed. Among 449 enrolled children, 55.0% were male, and the median age was 7.3 years. Bleeding etiology was 46.1% trauma, 34.1% operative, and 19.8% medical. Prior to the life-threatening bleeding event, most had age-adjusted hypotension (61.2%), and 25% were hypothermic. Children with medical bleeding had higher median Pediatric Risk of Mortality scores (18) compared with children with trauma (11) and operative bleeding (12). Median Glasgow Coma Scale scores were lower for children with trauma (3) compared with operative (14) or medical bleeding (10.5). Median time from bleeding onset to first transfusion was 8 minutes for RBCs, 34 minutes for plasma, and 42 minutes for platelets. Postevent acute respiratory distress syndrome (20.3%) and acute kidney injury (18.5%) were common. Twenty-eight-day mortality was 37.5% and higher among children with medical bleeding (65.2%) compared with trauma (36.1%) and operative (23.8%). There were 82 hemorrhage deaths; 65.8% occurred by 6 hours and 86.5% by 24 hours.

**CONCLUSIONS:** Patient characteristics and outcomes among children with life-threatening bleeding varied by cause of bleeding. Mortality was high, and death from hemorrhage in this population occurred rapidly.

**KEY WORDS:** critical care; emergency medicine; pediatric; surgery; trauma: transfusion

Julie C. Leonard, MD, MPH<sup>1</sup> Cassandra D. Josephson, MD2 James F. Luther, MA3 Stephen R. Wisniewski, PhD3 Christine Allen, MD⁴ Fabrizio Chiusolo, MD<sup>6</sup> Adrienne L. Davis, MD, MSc6 Robert A. Finkelstein, MD7 Julie C. Fitzgerald, MD. PhD. MSCE8 Barbara A. Gaines, MD<sup>9</sup> Susan M. Goobie, MD, FRCPC10 Sheila J. Hanson, MD, MS11 Hilary A. Hewes, MD12 Laurie H. Johnson, MD, MS13 Mark O. McCollum, MD14,15 Jennifer A. Muszynski, MD, MPH16 Alison B. Nair, MD17 Robert B. Rosenberg, MD, PhD18 Thomas M. Rouse, MD19 Athina Sikavitsas, DO20 Marcy N. Singleton, MSN21 Marie E. Steiner MD, MS22 Jeffrey S. Upperman, MD<sup>23,24</sup> Adam M. Vogel, MD25 Hale Wills, MD, MS26,27 Margaret K. Winkler, MS, MD<sup>28,29</sup>

Philip C. Spinella, MD30

			THE RESERVE OF THE PERSON NAMED IN COLUMN 1
	Trauma (n=210) 46%	Operative (n=174) 34%	Medical (n=97) 20%
28-day mortality	37%	24%	62%
LTH duration (hrs)	3.06 [1-6]	3.91 [2-7]	3.83 [1-7]
Time to RBC (min)	8.0 [0-42]	6.0 [0-39]	10.0 [0-45]
Time to plasma (min)	33.0 [16-75]	39.5 [12-83]	30.5 [19-65]
Time to PLTs (min)	41.0 [20-71]	42.0 [11-111]	43.0 [15-118]



### **Hemostatic Resuscitation Options**

- Empiric transfusion
  - RBCs, plasma, platelets in 1:1:1 unit ratio
  - Low Titer Group O Whole Blood

- Goal directed hemostatic resuscitation
  - TEG/ROTEM
    - Blood Products and hemostatic adjuncts
      - Antifibrinolytics and factor concentrates



## **Types of Whole Blood**

- Warm and Fresh
  - Room temp (22C)
  - Transfused within 8 hours
  - Most military data
- Cold and Stored
  - 2-6 C
  - Stored for 14-35 days
  - Civilian data



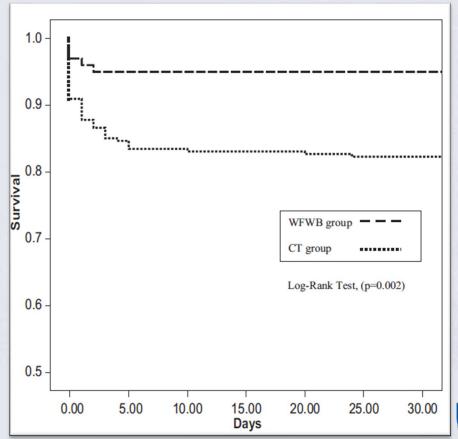
## **Types of Whole Blood**

- ABO specific
  - Military data with warm fresh whole blood
- Group O Whole Blood
  - Low titer (Anti A and B < 256)</p>
  - Mostly civilian data with cold whole blood
  - LTOWB



## Warm Fresh Whole Blood Is Independently Associated With Improved Survival for Patients With Combat-Related Traumatic Injuries J Trauma. 2009;66:S69-S76.

Philip C. Spinella, MD, Jeremy G. Perkins, MD, Kurt W. Grathwohl, MD, Alec C. Beekley, MD, and John B. Holcomb, MD





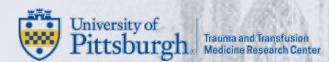
### **THOR 2018**

J Trauma Acute Care Surg Volume 84, Number 6, Supplement 1

### Raising the standards on whole blood

Mark H. Yazer, MD, Andrew P. Cap, MD, PhD, and Philip C. Spinella, MD, Pittsburgh, Pennsylvania

- 5.15 Selection of Compatible Blood and Blood Components for Transfusion
  - 5.15.1 Recipients shall receive
    - ABO group-compatible Red Blood Cell components
    - ABO group-specific Whole Blood
    - Low titer group O Whole Blood (for non group O or for recipients whose ABO group is unknown)



# Risk/Benefit Assessment LTOWB compared to blood components

Advantages of LTOWB	Risks of LTOWB
More potent product Higher Hb, plasma, platelets per volume	Incompatible plasma/immune complexes? Theoretical risk.
Cold platelets – improved hemostasis (RCT data)	Waste? Reduced/eliminated if used in non-trauma massive bleeding
Increased storage duration of platelet product	Ease of over-resuscitation
Less risk of ABO incompatible transfusion reactions than ABO compatible components	
Less bacterial contamination risk	
Logistical advantages  Quicker transfusion of balanced product  One product vs four products	
Independent association with improved survival	

### Volume and Concentrations Between Component Therapy vs. Warm Whole Blood



VS



Component Therapy: 680 mL RBC unit + PLT unit + FFP unit + Cryo unit

- Red blood cell concentration: 29%
- Platelets: 80,000
- Coagulation factors: 65%

Whole Blood: 500 mL A single WB unit

- Red blood cell concentration: 38-50%
- Platelets: 150,000-400,000
- Coagulation factor concentration: 90%



# Standard Amounts of Anti-coagulants and Additives in Reconstituted Whole Blood vs Whole Blood



### **Component Therapy per Unit:**

 $6 \times RBC (AS-5)$   $6 \times 120 \text{ ml} = 720 \text{ml}$ 

 $6 \times FFP$   $6 \times 50 \text{ ml} = 300 \text{ml}$ 

 $1 \times aPLT$   $1 \times 35 \text{ ml} = 35 \text{ml}$ 

<u>Total =1055ml</u>

WOOLS BLOOD

WASHINGTON COLORS

**Whole Blood per Unit:** 

 $6 \times 63 \text{ml} = 378 \text{ml}$ 

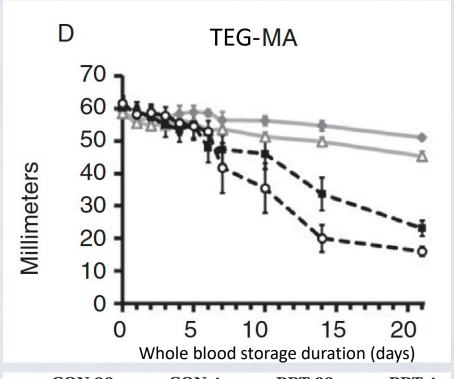
Total= 378ml

There is 3 times the volume of anticoagulant and additives with reconstituted whole blood from components compared to whole blood

## Primary hemostatic capacity of whole blood: a comprehensive analysis of pathogen reduction and refrigeration effects over time

**TRANSFUSION** 2013;53:137S-149S.

Heather F. Pidcoke, Steve J. McFaul, Anand K. Ramasubramanian, Bijaya K. Parida, Alex G. Mora, Chriselda G. Fedyk, Krystal K. Valdez-Delgado, Robbie K. Montgomery, Kristin M. Reddoch, Armando C. Rodriguez, James K. Aden, John A. Jones, Ron S. Bryant, Michael R. Scherer, Heather L. Reddy, Raymond P. Goodrich, and Andrew P. Cap



.--- = CON 22; --- = CON 4; --- = PRT 22; --- = PRT 4.



# Group O WB Less Risk of Severe Hemolytic Reactions vs Component Therapy

SHOCK, Vol. 41, Supplement 1, pp. 70-75, 2014

#### LOW TITER GROUP O WHOLE BLOOD IN EMERGENCY SITUATIONS

Geir Strandenes,\*† Olle Berséus,‡ Andrew P. Cap,§ Tor Hervig,\*<sup>||</sup> Michael Reade,¶ Nicolas Prat,§\*\*\* Anne Sailliol,†† Richard Gonzales,‡† Clayton D. Simon,§§ Paul Ness,<sup>|||</sup> Heidi A. Doughty,¶ Philip C. Spinella,§\*\*\* and Einar K. Kristoffersen\*||
\*Department of Immunology and Transfusion Medicine, Haukeland University Hospital; and †Norwegian Naval Special Operation Commando, Bergen, Norway; †Department of Transfusion Medicine, Örebro University Hospital, Örebro, Sweden; §US Army Institute of Surgical Research, FT Sam Houston, Texas; ||Institute of Clinical Science, The University of Bergen, Norway; ¶ Australian Defense Force Joint Health Command, Canberra, Australian Capital Territory; \*\*French Military Medical Service, Clamart, France; †† Commander French Military Blood Transfusion Center, Clamart, France; †‡ Director, US Army Blood Program and §§US Army Transfusion Medicine Consultant to the Surgeon General San Antonio Military Medical Center, JBSA–Fort Sam Houston, Texas; ||¶ Transfusion Medicine Division, Johns Hopkins Medical Institutions, Baltimore, Maryland; ¶ NHS Blood and Transplant, Birmingham, England, United Kingdom; and\*\*\*Division of Pediatric Critical Care, Department of Pediatrics, Washington University in St Louis, St Louis, Missouri

- Risk from incompatible plasma (LTOWB)
  - 1: 120,000 risk of mild to moderate reaction
- Risk of ABO incompatibility (RBCs)
  - 1:80,000 risk of severe (fatal) hemolytic reaction
  - Human error





### LTOWB vs CT: Mortality Data-Trauma (2020-2022)

- Williams/Cotton <sup>1</sup>
  - 350 patients, retrospective
  - 2-fold increase in 28-day survival
    - Adj OR 2.19, 1.01-4.76, p=0.047).
- Mihalko/Spinella<sup>2</sup>
  - 384 pts, Prospective/historical control
  - 2 fold reduction in 24 hour mortality
    - Adj HR, 0.51, p=0.033
- Brill/Cotton <sup>3</sup>

7 Transfusion. 2021;61:S15-S21.

- 1,377 patients, Retrospective
- 1.5 fold increased 30 day survival
  - Adj OR, 1.59, p < 0.001</li>

- Hanna/Joseph <sup>4</sup>
  - 8,494 patients, Retrospective
  - 1.2 fold increase in 24 hour survival
    - Adj OR, 0.78 (0.59-0.89), (p=0,006)
- Hazelton/Porter <sup>5</sup>
  - 1,623 patients, prospective observational
  - 2 fold reduction for in hospital mortality
    - Adj OR 0.52, p< 0.0001</li>
- Gaines/Leeper <sup>6</sup>
  - 80 children, retrospective/propensity matched
  - 2 fold reduction in 28 day mortality
    - Adj OR 0.41, P . 0.02
- Braverman/Jenkins <sup>7</sup>
  - 214 adults, retrospective/propensity matched
  - Prehospital study with reduced ED mortality (11 vs 0%), (p=0.04)
  - 6hr mortality: 22 vs 3% (p=0.08)

1 J Trauma Acute Care Surg. 2020 Jan;88(1):87-93 2 Oral Presentation, ISTH, London 2022 3 J Am Coll Surg. 2022 Apr 1;234(4):408-418 4 J Trauma Acute Care Surg. 2020 Aug;89(2):329-335 5 Ann Surg. 2022 Oct 1;276(4):579-588 6 Ann Surg. 2021 Oct 8. on line



## Outcomes with LTOWB use in Trauma Patients

- Metanalysis submitted for publication
- 15 civilian trauma studies including 14,424 patients
  - 3,446 received LTOWB
- LTOWB was defined as cold-stored RhD-positive or RhD-negative group O whole blood with low titers of anti-A and anti-B antibodies (ranging from 50 to 256).
- Studies were excluded if
  - Modified LTOWB (leukoreduced without platelet-sparing filter plus room temp plts transfused
  - Warm fresh whole blood was used.

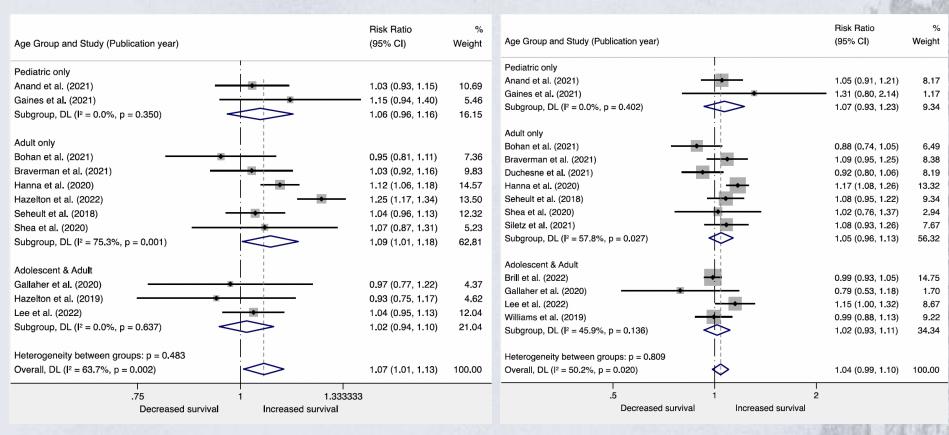


# Meta-Analysis of LTOWB studies in Trauma Patients

- LTOWB was associated with improved 24-hour survival compared to component therapy
  - RR (95% CI) =1.07 (1.01-1.13)
- No significant differences were found for later survival timepoints (28-day, 30-day, in-hospital)
  - RR(95% CI)= 1.04(0.99-1.1)
- No evidence of small study bias and all studies were graded as a moderate level of bias.



# Meta-Analysis of LTOWB studies in Trauma Patients



24 Hour Survival

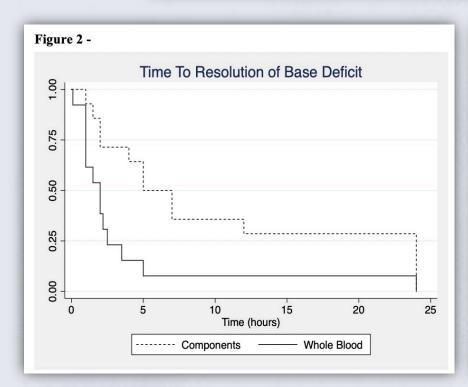
28 Day Survival

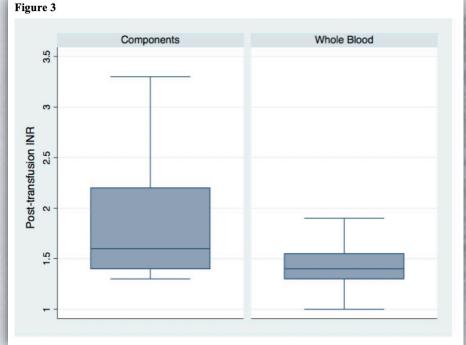


## Whole Blood is Superior to Component Transfusion for Injured Children

### A Propensity Matched Analysis

Christine M. Leeper, MD, MS,⊠ Mark H. Yazer, MD, Darrell J. Triulzi, MD, Matthew D. Neal, MD, and Barbara A. Gaines, MD





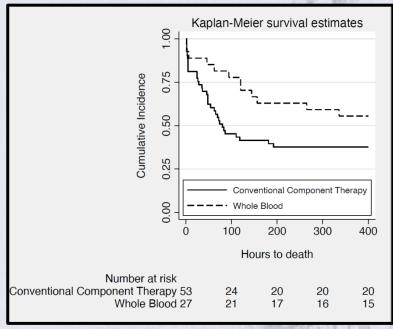


**ORIGINAL ARTICLE: PDF ONLY** 

## Low Titer Group O Whole Blood In Injured Children Requiring Massive Transfusion

Gaines, Barbara A. MD\*; Yazer, Mark H. MD†; Triulzi, Darrell J. MD†; Sperry, Jason L. MD MPH†; Neal, Matthew D. MD†; Billiar, Timothy R. MD†; Leeper, Christine M. MD MS\*

	72 hour mortality				28 day mortali	ty
	Odds Ratio	95% Confidence Interval	p value	Odds Ratio	95% Confidence Interval	p value
LTOWB	0.23	0.08-0.70	0.009	0.41	0.23-0.98	0.02
Age (years)	1.07	1.00-1.16	0.06	1.02	0.98-1.09	0.13
Total transfusion volume (mL/kg)	1.01	1.01-1.02	<0.001	1.01	1.01-1.02	<0.001
Admission Base Deficit	1.11	1.06-1.17	<0.001	1.12	1.07-1.17	<0.001
Admission INR	1.30	1.06-1.58	<0.001	1.29	1.05-1.58	0.003
Injury Severity Score	1.02	1.00-1.05	0.10	1.02	0.99-1.06	0.06
LTOWB = low titer group O whole blood						



### An assessment of the safety, hemostatic efficacy, and clinical impact of lowtiter group O whole blood in children and adolescents

Justin Gerard, Krislynn Mueck, David Lubkin, Gabrielle Hatton, Charles Wade, Bryan Cotton

### **Inverse Probability Weighted Analysis for 30 day Survival**

	Odds Ratio (95% CI)	P value
Whole Blood Group	2.48 (1.16-5.47)	0.02
Age (years)	1.004 (1.0001-1.0006)	0.003
ISS	0.93 (0.89-0.96)	<0.001
Lactate	0.76 (0.67-0.84)	< 0.001



## Whole blood hemostatic resuscitation in pediatric trauma: A nationwide propensity-matched analysis

Tanya Anand, MD, Omar Obaid, MD, Adam Nelson, MD, Mohamad Chehab, MD, Michael Ditillo, DO, Ahmad Hammad, MD, Molly Douglas, MD, Letitia Bible, MD, and Bellal Joseph, MD, FACS, Tucson, Arizona

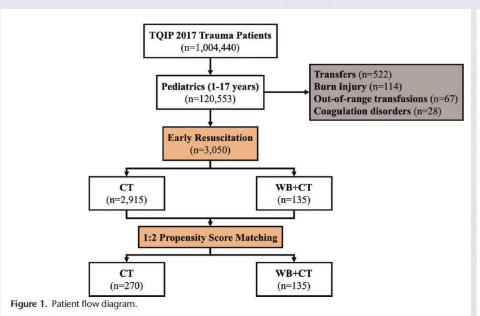


TABLE 1. Postmatch Patient and F	•		
	CT (n = 270)	WB-CT $(n = 135)$	p
Demographics			
Age, mean (SD), y	11.8 (5.3)	12.2 (4.9)	0.401
Male, n (%)	179 (66.3)	87 (64.4)	0.711
Weight, mean (SD), kg	42.1 (18.9)	44.7 (18.4)	0.247
Mechanism of injury, n (%)			0.666
Motor vehicle collision	118 (43.7)	60 (44.4)	
Pedestrian struck	37 (13.7)	19 (14.1)	
Fall	8 (3.0)	3 (2.2)	
Firearm	75 (27.8)	38 (28.2)	
Cut/pierce	17 (6.3)	8 (5.9)	
Other	15 (5.5)	7 (5.2)	
ED vital signs			
Age-adjusted shock, n (%)	136 (50.4)	71 (52.6)	0.673
Temperature, mean (SD), °C	36.1 (2.2)	35.9 (1.5)	0.516
GCS, median (IQR)	5 (3–15)	3 (3–15)	0.080
Injury characteristics, median (IQR)			
Head-AIS	2 (0–3)	2 (1–3)	0.182
Chest-AIS	3 (1–4)	3 (1–4)	0.490
Abdomen-AIS	4 (2–4)	4 (2–5)	0.209
Extremity-AIS	2 (0–3)	2 (0–3)	0.671
ISS	31 (19–41)	34 (22–45)	0.124



### LTOWB - A Blood Conservation Strategy?

	CT (n = 270)	WB-CT (n = 135)	p
h Transfusions, median (IQR), mL/kg			
PRBC	31 (22–57)	19 (11–31)	0.008
Plasma	12 (9–31)	9 (0–21)	< 0.001
Platelets	4 (4–10)	0 (0-6]	< 0.001
WB	_	13 (9–20)	_
Total blood products	48 (33–95)	35 (22–73)	0.013
4-h Transfusions, median (IQR), mL/kg			
PRBC	36 (25–71)	22 (15–53)	< 0.00
Plasma	17 (11–46)	11 (0–25)	< 0.001
Platelets	6 (4–13)	0 (0–9)	< 0.001
WB	<u> </u>	14 (10–23)	_
Total blood products	53 (36–119)	39 (24–97)	< 0.001

## Whole Blood Resuscitation is Safe in Pediatric Trauma Patients: A Multicenter Study

The American Surgeon
2023, Vol. 0(0) 1–6
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DOI: 10.1177/00031348231157864
journals.sagepub.com/home/asu

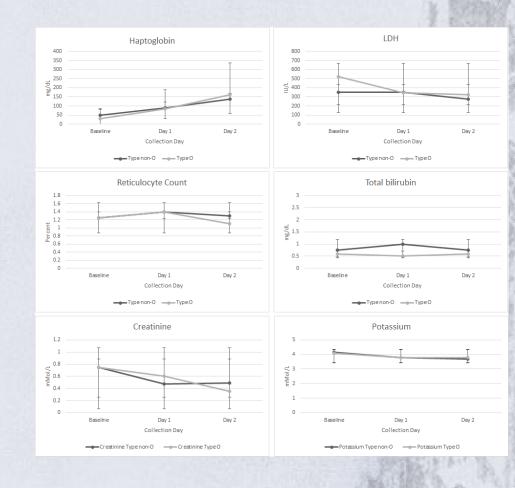
Lindsey L. Perea, DO<sup>1</sup>, Kate Moore, BA<sup>1</sup>, Courtney Docherty, DO<sup>2</sup>, Uyen Nguyen, BS<sup>3</sup>, Mark J. Seamon, MD<sup>4</sup>, James P. Byrne, MD<sup>4</sup>, Donald H. Jenkins, MD<sup>5</sup>, Maxwell A. Braverman, DO<sup>5</sup>, John M. Porter, MD<sup>6</sup>, Isabella G. Armento, BS<sup>6</sup>, Caleb Mentzer, DO<sup>7</sup>, Guy C. Leonard, BS<sup>7</sup>, Alejandro J. Luis, MD<sup>8</sup>, Matthew R. Noorbakhsh, MD<sup>9</sup>, James E. Babowice, DO<sup>9</sup>, Haytham M. A. Kaafarani, MD<sup>10</sup>, Ava Mokhtari, MD<sup>10</sup>, Matthew J. Martin, MD<sup>11</sup>, Jayraan Badiee, MPH<sup>11</sup>, Charles Mains, MD<sup>12</sup>, Robert M. Madayag, MD<sup>12</sup>, Sarah A. Moore, MD<sup>13</sup>, Kathleen Madden, MD<sup>13</sup>, and Joshua P. Hazelton, DO<sup>14</sup>

When controlling for <u>age</u>, <u>sex</u>, <u>mechanism of injury</u>, <u>and shock index</u>, there were no differences in mortality or complications between the WB and CT groups



# No Increased Risks with LTOWB in Adults or Children

- No increase in transfusion reactions
- No increase in organ failure
- No increase in hospital complications
  - AKI
  - Sepsis
  - ARDS
  - Thromboembolism
- No increase in hemolysis
  - Non Group O patients vs Group O patients who receive LTOWB



Transfusion. 2020 Jun;60 Suppl 3:S24-S30 Transfusion. 2018 Oct;58(10):2280-2288 Transfusion. 2021 Jul;61 Suppl 1:S8-S14



DOI: 10.1111/trf.16456

SUPPLEMENT ARTICLE

**TRANSFUSION** 

### Safety profile of low-titer group O whole blood in pediatric patients with massive hemorrhage

Katrina M. Morgan<sup>1</sup> | Mark H. Yazer<sup>2</sup> | Darrell J. Triulzi<sup>2</sup> | Stephen Strotmeyer<sup>3</sup> | Barbara A. Gaines<sup>3</sup> | Christine M. Leeper<sup>1</sup>

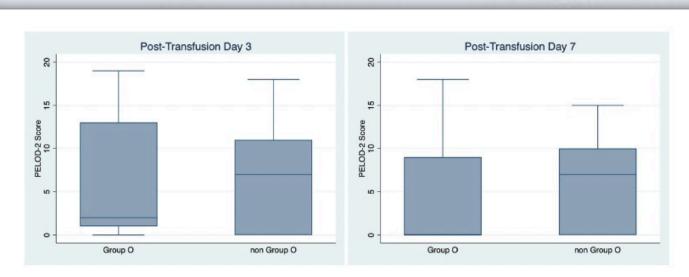


FIGURE 2 PELOD-2 scores did not significantly differ between group O and non-group O LTOWB recipients on post-transfusion days 3 and 7. On day 3, group O PELOD-2 score was 2 (1–13) and non-group O was 7 (0–11) (p = .82). On day 7, group O PELOD-2 score was 0 and non-group O was 7 (p = .48)





Comparison of the hemostatic effects of fresh whole blood, stored whole blood, and components after open heart surgery in children

CS Manno, KW Hedberg, HC Kim, GR Bunin, S Nicolson, D Jobes, E Schwartz and WI Norwood

Table 3.	Blood Loss (mL/k	g) (mean ± SE) b	v Age, Surgica	I Difficulty, and Both
		<b>27,</b> 1,	,	

Cold Whole Blood (ABO Matched)	Components in 1:1:1 unit ratio	P Value*
44.8 ± 6.0 (n = 57)	74.2 ± 8.9 (n = 52)	.03†
51.7 ± 7.4 (30)	96.2 ± 10.7 (36)	.001‡
$37.2 \pm 9.7$ (27)	$24.6 \pm 6.0$ (16)	NS

24 hr total blood used was 75.5ml/kg and 97.4ml/kg in the WB and CT groups, respectively. Absolute reduction of 22ml/kg in 16kg child (4yr old) = 352ml of blood

22% relative reduction of total blood use in 24 hrs

40% less blood loss in WB group





#### **ORIGINAL ARTICLE**



Whole blood transfusion reduces overall component transfusion in cases of placenta accreta spectrum: a pilot program

Jessian L. Munoz<sup>a,b</sup>, Alison M. Kimura<sup>a,b</sup>, Elly Xenakis<sup>a,b</sup>, Donald H. Jenkins<sup>c</sup>, Maxwell A. Braverman<sup>c</sup>, Patrick S. Ramsey<sup>a,b</sup> and Kayla E. Ireland<sup>a,b</sup>

Prospective observational study of 34 patients 16 received LTOWB and 18 received CT

Factor	Whole blood $(n=16)$	Component (n = 18)	p Value
Age	32.4 ± 5.9	31.3 ± 5.28	.57ª
BMI	$32.1 \pm 7.0$	$34.5 \pm 3.8$	.23a
Gravity	4 (3.5, 5.8)	5 (3.8, 7)	.54 <sup>b</sup> .27 <sup>b</sup>
Parity	3 (2, 3)	3.5 (2, 4.3)	.27 <sup>b</sup>
History of CD	16 (100)	16 (89)	.49 <sup>c</sup>
Number of prior CD	2 (2, 3)	3 (1.8, 4)	.62 <sup>b</sup>
Tertiary referral	14 (88)	13 (72)	.41°
Gestational age at delivery	34 (31, 34)	34 (25, 34.8)	.70 <sup>b</sup>
PAS by ultrasound			
Previa	2 (13)	5 (28)	.41°
Accreta	4 (25)	12 (67)	.02°
Increta	1 (6)	0	.47°
Percreta	9 (56)	1 (6)	.002°
Diabetes	1 (6)	1 (6)	1.0 <sup>c</sup>
Hypertension	2 (13)	2 (11)	1.0 <sup>c</sup>
Anemia	5 (31)	8 (45)	.18 <sup>d</sup>
Emergent delivery	7 (44)	5 (28)	.33°
Public insurance	9 (56)	16 (89)	.05°

BMI: body mass index; CD: cesarean delivery; PAS: placenta accreta spectrum. Values presented as mean  $\pm$  SD, median [P25, P75] or N (column %). p Values:  ${}^{a}t$ -test,  ${}^{b}$ Mann-Whitney's test,  ${}^{c}$ Fisher's exact test, and  ${}^{d}$ chi-squared. Bold values suggest p < .05.

- Total transfusion was less in the LTOWB vs CT group, 2607 ml vs. 4683 ml, (p=0.03)
- 2076 ml less blood transfused
- 44% relative reduction in total blood transfusion



Study	Outcome	Mean % Less Blood Transfused	Mean Reduced Vol Blood Transfused	Mean Reduced Vol/kg Transfused
Adult Trauma Studies				
Williams, (n=350) Retrospective	2-fold increase 28-day survival	53%, (post ED) (P=0.033)		
Brill, (N=1,377) Retrospective	1.5 fold increase 30 day survival	7%, (post ED) (P<0.001)		
Hazelton, (N=1,623) Prospective	2-fold reduction In-hosp mortality	No difference Only Recorded Units		
Hanna (N=8,494) Retrospective	1.2 fold reduction 24 hr mortality	No Difference Only Recorded Units		
Braverman (N=214) Retrospective/Propensity	Reduced death in ED (0 vs 11%)		400ml less in ED	
Mahilko, (N=384) Prospective-Unpublished	2-fold reduction 24-hour mortality	40%, (72hrs) (P<0.001)	2.5 Liters	32ml/kg
Obstetric Studies				
Munoz, (N=36) Obstetric, Prospective	44% less blood transfused	44% (p=0.03)	2.1 Liter	30ml/kg (70kg)
Pediatric Studies				
Gaines, (N=80) Peds, Retrospective	2-fold reduction 28-day mortality	18% (p=0.06)	333 ml	11 ml/kg
Annand, (N=405) Peds, Retrospective	Mortality diff not significant	26% (P<0.001)	630 ml	14ml/kg
Manno, (N=109) Ped CT surgery, RCT	40% less blood loss	40% (No statistics)	352 ml	22ml/kg

### LTOWB – Less Costs

## Transfusion-related cost comparison of trauma patients receiving whole blood versus component therapy

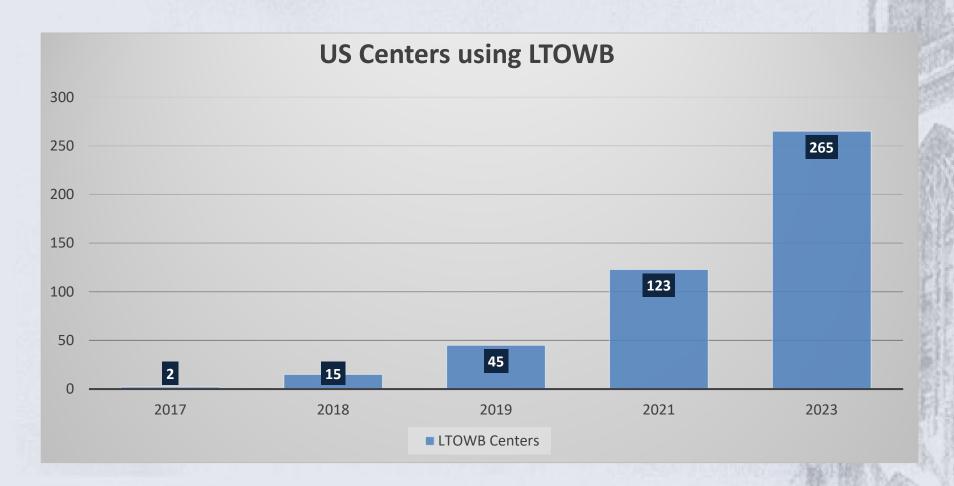
Ciaraglia, Angelo MD; Myers, John C. MD; Braverman, Maxwell DO; Barry, John BS; Eastridge, Brian MD; Stewart, Ronald MD; Nicholson, Susannah MD, MS; Jenkins, Donald MD

Author Information ⊗

Journal of Trauma and Acute Care Surgery 95(1):p 62-68, July 2023. | DOI: 10.1097/TA.000000000003933

- A retrospective review of adult and pediatric trauma patients who received either LTO+WB or CT from time of injury to within 4 hours of arrival was performed.
- Annual mean cost per unit of blood product used
- Pediatric and adult patients were analyzed separately and were compared on a cost per patient (cost/patient) and cost per patient per milliliter (cost/patient/mL) basis.
- After the initiation of the WB transfusion, the mean annual cost decreased 17.3% for all blood products
- Average net reduction in cost related to blood products was more than \$927,000.

### LTOWB in the US





### RhD Status of LTOWB

- RhD- preferred for women to prevent hemolytic disease of fetus/newborn (HDFN)
  - RhD- LTOWB very short supply
- If RhD+ LTOWB is only available, do females get it?
  - Estimates of improved survival at very least 5-10% range
- HDFN has 1-6% overall incidence
  - 0.3% risk of fetal demise
- Women and parents of girls strongly prefer to receive RhD+ LTOWB and accept risk of HDFN



### RhD+ and HDFN Risk

HEMATOLOGY 2023, VOL. 28, NO. 1, 2161215 https://doi.org/10.1080/16078454.2022.2161215



OPEN ACCESS Check for updates

Not as "D"eadly as once thought – the risk of D-alloimmunization and hemolytic disease of the fetus and newborn following RhD-positive transfusion in trauma

Mark H. Yazer<sup>a</sup>\*, Gleb Panko<sup>b</sup>\*, John B. Holcomb<sup>c</sup>, Alesia Kaplan<sup>a</sup>, Christine Leeper<sup>d</sup>, Jansen N. Seheult<sup>e</sup>, Darrell J. Triulzi<sup>a</sup> and Philip C. Spinella<sup>f</sup>

RhD PERCEPTIONS

**TRANSFUSION** 

Attitudes of American adult women toward accepting RhD-mismatched transfusions in bleeding emergencies

Gabriel Yu<sup>1</sup> | Jeffrey Siegler<sup>1</sup> | Jane Hayes<sup>2</sup> | Mark H. Yazer<sup>3</sup> | Philip C. Spinella<sup>4</sup>

SUPPLEMENT ARTICLE

**TRANSFUSION** 

SUPPL

**TRANSFUSION** 

Weighing the risk of hemolytic disease of the newborn versus the benefits of using of RhD-positive blood products in trauma

```
Jennifer Andrews<sup>1,2</sup> | Cassandra D. Josephson<sup>3,4</sup> | Pampee Young<sup>1,5</sup> | Philip C. Spinella<sup>6,7</sup> | Mark H. Yazer<sup>8</sup>
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SUPPLEMENT ARTICLE

Parent perceptions of emergent blood transfusion in children

### **ONLINE CLINICAL INVESTIGATION**

### Use of Antifibrinolytics in Pediatric Life-Threatening Hemorrhage: A Prospective Observational Multicenter Study

**OBJECTIVES:** To assess the impact of antifibrinolytics in children with life-threatening hemorrhage.

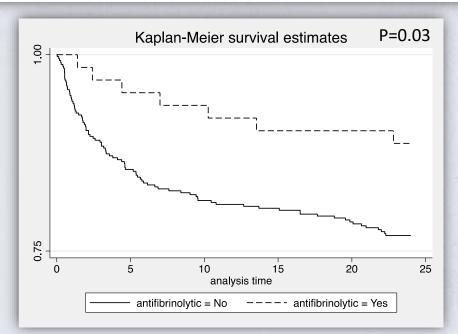
**DESIGN:** Secondary analysis of the MAssive Transfusion epidemiology and outcomes In Children study dataset, a prospective observational study of children with life-threatening bleeding events.

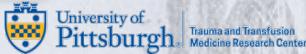
SETTING: Twenty-four children's hospitals in the United States, Canada, and Italy.

**PATIENTS:** Children 0–17 years old who received greater than 40 mL/kg of total blood products over 6 hours or were transfused under activation of massive transfusion protocol.

INTERVENTION/EXPOSURE: Children were compared according to receipt of antifibrinolytic medication (tranexamic acid or aminocaproic acid) during the bleeding event.

Philip C. Spinella, MD¹
Julie C. Leonard, MD, MPH²
Barbara A. Gaines, MD³
James F. Luther, MA⁴
Stephen R. Wisniewski, PhD⁴
Cassandra D. Josephson, MD⁵
Christine M. Leeper, MD, MS⁵
for the MAssive Transfusion
epidemiology and outcomes in
Children (MATIC) Investigators
and BloodNet





### Antifibrinolytics Independently Associated with Improved Survival

TABLE 3.

Cox Proportional Hazard Models for 6- and 24-Hour Mortalities

Measures	6-hr Mortality		24-hr Mortality	
	OR (95% CI)	р	OR (95% CI)	р
Unadjusted analysis				
Antifibrinolytic	0.29 (0.09-0.96)	0.04	0.49 (0.21-1.12)	0.08
Adjusted analysis				
Antifibrinolytic	0.29 (0.09-0.93)	0.04	0.45 (0.21-0.98)	0.04
Pediatric Risk of Mortality score	1.04 (1.02-1.06)	< 0.001	1.05 (1.04-1.07)	< 0.001
Indication operative (ref)				
Trauma	4.18 (1.90-9.17)	0.001	3.08 (1.71-5.56)	< 0.001
Medical	5.45 (2.42-12.26)	< 0.001	3.28 (1.91-6.60)	< 0.001
Age (yr)	0.94 (0.90-0.98)	0.002	0.95 (0.92-0.98)	0.002
Plasma deficit	1.01 (0.99-1.01)	0.17	1.01 (0.99-1.01)	0.12



- A pragmatic, Bayesian, group sequential, combined noninferiority/superiority, randomized, controlled, multicenter, phase III, platform trial of
  - Low titer group O whole blood vs component therapy
  - Tranexamic acid vs placebo
- Mechanisms of TIC and resuscitation effects will be investigated
  - Multi-OMICS
  - Viscoelastic assays
- Exception from Informed Consent





### **MATIC-2: General Methods**

- 20 US high volume pediatric trauma centers
- 1000 subjects < 18 years of age</li>
- Exception for Informed Consent
- A cross over cluster design
  - Sites randomly assigned to LTOWB or CT and TXA or placebo
    - Cluster stratified by #MTP/year at the 20 sites
  - Cross over for every 250 patients
- Rationale for design
  - Minimizes waste of LTOWB
  - Does not require 24/7 research coordinator coverage





### Primary Outcome – 24 Hour Mortality

### Evidence-Based and Clinically Relevant Outcomes for Hemorrhage Control Trauma Trials

John B. Holcomb, MD,\* Ernest E. Moore, MD,† Jason L. Sperry, MD, MPH,‡ Jan O. Jansen, MBBS, PhD,§ Martin A. Schreiber, MD,¶ Deborah J. del Junco, PhD,|| Philip C. Spinella, MD,\*\* Angela Sauaia, MD, PhD,†† Karim Brohi, MD,‡‡ Eileen M. Bulger, MD,§§ Andrew P. Cap, MD, PhD,¶¶ John R. Hess, MD, MPH, FACP, FAAAS,|||| Donald Jenkins, MD,\*\*\* Roger J. Lewis, MD, PhD,††† Matthew D. Neal, MD,‡‡‡ Craig Newgard, MD, MPH,§§§ Shibani Pati, MD, PhD,¶¶¶ Anthony E. Pusateri, PhD,|||||| Sandro Rizoli, MD, PhD,\*\*\*\* Robert T. Russell, MD, MPH,†††† Stacy A. Shackelford, MD,‡‡‡‡ Deborah M. Stein, MD, MPH,§§§§ Marie E. Steiner, MD, MS,¶¶¶¶ Henry Wang, MD, MS,|||||||| Kevin R. Ward, MD,\*\*\*\*\* and Pampee Young, MD, PhD†††††

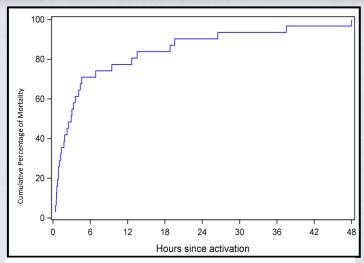


Figure 12: Mortality due to hemorrhage in children with traumatic injury and life-threatening hemorrhage





## **Eligibility Criteria**

### Inclusion criteria

- 1. Children, defined as less than 18 years of age with traumatic injury
- MTP activation for confirmed or suspected active life-threatening traumatic bleeding AND

Confirmed or suspected active life-threatening traumatic bleeding with at least 2 of 3 of the following criteria

Hypotension for age (< 5% tile)

Tachycardia for age (>95th % tile)

Traumatic injury with exam findings consistent with severe bleeding (e.g., penetrating injury, hemothorax, distended abdomen with bruising, amputation of limb).





## **Eligibility Criteria**

#### **Exclusion Criteria**

Unknown time of injury

Greater than 3 hours since time of injury

History of seizure after the injury event

Known allergy or hypersensitivity reaction to TXA

Comatose (Glasgow Coma Score of 3) with fixed and dilated pupils suggesting nonsurvivable brain injury

MTP activated but no blood products given

Patients who required an ED thoracotomy or received more than 5 consecutive minutes of cardiopulmonary resuscitation (prior to receiving randomized blood products)

Patients who are obviously pregnant on clinical examination

Known prisoners as defined in protocol

Known ward of the state

Isolated hanging, drowning or burns

Previous enrollment in MATIC-2

Prior study opt-out with bracelet





## **Study Intervention**

### LTOWB

- Group O
- Low titer is anti-A and anti-B < 200 (or supplier standard)
- Leukocyte reduced/platelet sparing filter
- Stored in CPD for up to 21 days at 1-6 Celsius
- Maximum of 8 units
- CT
  - RBC, plasma, platelets in 1:1:1 unit ratio
  - All types of manufacturing methods permitted
- Once MTP is deactivated, standard of care (CT) will be used.





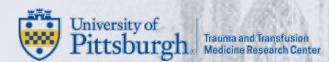
## **Study Intervention - TXA**

- Dose of 25mg/kg (2 max IV dose)
  - No maintenance infusion
- Based on pharmacokinetic data from adult RCTs
  - Modeled for children of different weights



### Conclusions

- LTOWB vs Component Therapy
  - Appears safe
  - Less expensive
  - Independently associated with reduced use of blood products and mortality
- RCTs are being performed in adult and pediatric populations



## Questions?





The risk to future pregnancies of transfusing Rh(D)-negative females of childbearing potential with Rh(D)-positive red blood cells during trauma resuscitation is dependent on their age at transfusion

Jansen N. Seheult, <sup>1,2</sup> D Michelle N. Stram, <sup>3</sup> Thomas Pearce, <sup>1</sup> Carolina B. Bub, <sup>4</sup> D Stephen P. Emery, <sup>5</sup> Jose Kutner, <sup>4</sup> D Naoko Watanabe-Okochi, <sup>6</sup> D Jason L. Sperry, <sup>7</sup> Minoko Takanashi, <sup>8</sup> Darrell J. Triulzi <sup>1,2</sup> & Mark H. Yazer <sup>1,2</sup> D

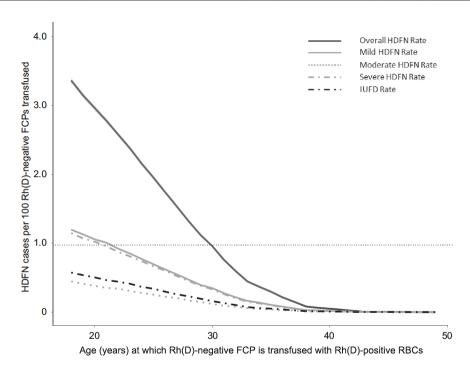


Fig. 3 Estimated rate of haemolytic disease of the fetus and newborn (HDFN), stratified by severity [mild, moderate, severe or intrauterine fetal demise (IUFD)], versus the age at which the Rh(D)-negative female of childbearing potential (FCP) is transfused with an Rh(D)-positive RBC-containing product during trauma resuscitation using assumptions for the United States. The dotted horizontal line represents the average overall HDFN rate of 0.97 HDFN cases per 100 Rh(D)-negative FCPs transfused across all ages from 18 to 49 years, assuming a uniform age distribution.

Vox Sanguinis. 2021;1-2.

# Risk of future haemolytic disease of the fetus and newborn following the transfusion of Rh(D)-positive blood products to Rh(D)-negative children

