BLOOD UTILIZATION AND TRANSFUSION REACTIONS UPDATE AND SOME OTHER STUFF AS TIME PERMITS (WB, SCD MGMT, AND MTP)

ERIC FILLMAN, MD

THURSDAY, FEBRUARY 17, 2022-12-1PM



PRIMARY LEARNING OBJECTIVES

- TO GIVE A BACKGROUND ON BLOOD BANK SERVICES AND BEST PRACTICES IN BLOOD UTILIZATION.
- TO PROVIDE CURRENT RECOGNIZED AABB TRANSFUSION REACTIONS AND WAYS TO RECOGNIZE THEM.
- TO PROVIDE A BASIC BACKGROUND ON THE BLOOD SUPPORT OF SICKLE CELL PATIENTS.
- IF TIME PERMITS, BRIEF REVIEW OF OUR NEW MASSIVE TRANSFUSION PROTOCOL (MTP) AND USE OF WHOLE BLOOD IN BLEEDING PATIENTS OUTSIDE OF THE HOSPITAL SETTING.



FINANCIAL DISCLOSURES

- NO RELEVANT FINANCIAL DISCLOSURES
- THE VIEWS PRESENTED ARE NOT A REFLECTION OF CPA, GRMC, OR THE US ARMY



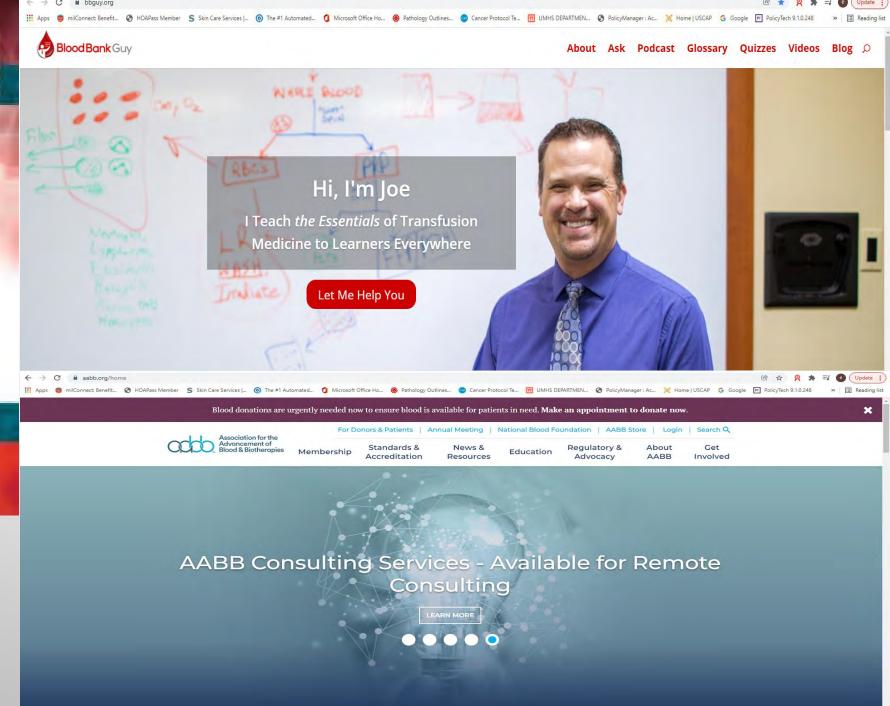


Modern Blood Banking & Transfusion Practices

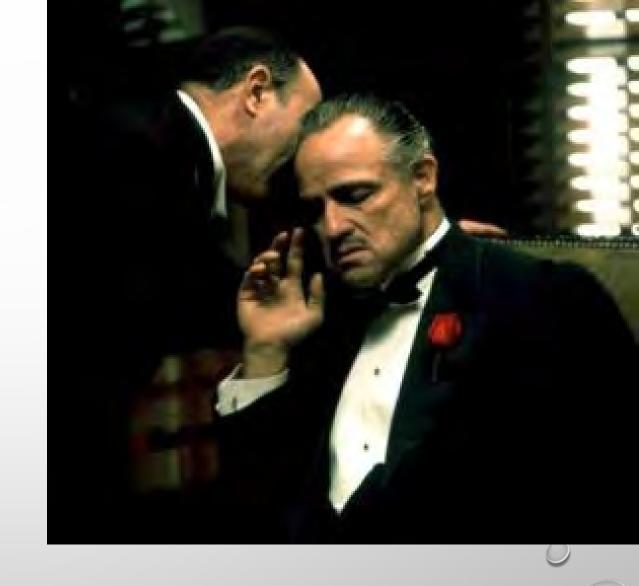
Seventh Edition



https://www.youtube.com/watch
2v=RaA6BeepiJM Dr. Steven
Frank, Med Dir at the JH Center
for Bloodless Medicine and
Surgery, Professor of
Anesthesiology and Critical
Care Medicine







My consiglieres, Ms Edna Pasadilla, MT and Ms Maria Courriveau, MT (aka Tom Hagen)





Secrets and Lies in a Silicon Valley Startup

John Carreyrou



SYLVESTER **STALLONE**



Blood Management –

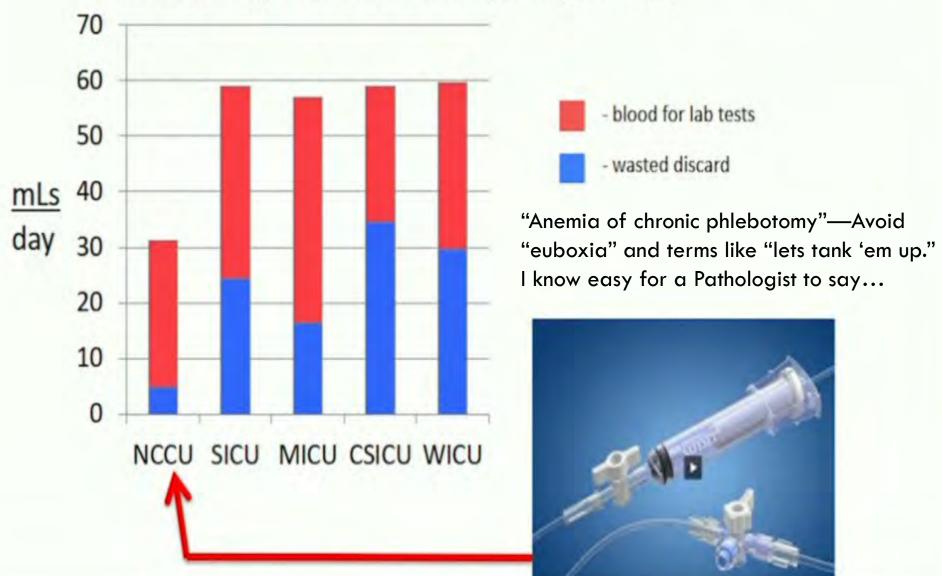
One of the few areas in medicine where all three of these can be achieved at the same time:

- Reduce Risk
- Save Cost
- Improve Outcomes

SO WHAT IS A REASONABLE HGB LEVEL TO TREAT?

- OBVIOUSLY WE DON'T TREAT A NUMBER (IE HGB); WE TREAT THE PATIENT, HOWEVER, THERE IS A REASONABLE TRANSFUSION (HGB) TRIGGER.
- RATIONALLY OPTIMIZING ANEMIA AND HEMOSTASIS.
- GOAL OF "RESTRICTING" BLOOD UTILIZATION (IE INFECTIOUS AND TRANSFUSION ASSOCIATED REACTIONS)...MORE TO FOLLOW.
- IMPROVE PATIENT OUTCOMES, EFFICIENCY, AND VALUE.
- HOWEVER, THERE ARE RELATIVELY WELL ESTABLISHED GUIDELINES FOR A "TARGET" HGB.
- REMEMBER ANEMIA IS NOT A DIAGNOSIS IT IS A SIGN OF AN UNDERLYING DISEASE/DIAGNOSIS.
- ULTIMATELY GOING BACK TO PATIENT CENTRIC MEDICINE AND TREATING THE DISEASE PROCESS.

ICU phlebotomy at Johns Hopkins
Over 1% of blood volume/day (cancels out erythropoiesis)
Cut in half using Safeset blood draw system (\$9.45 cost)



SO WHAT IS AN ESTABLISHED REASONABLE HGB TRANSFUSION TRIGGER?

A. 9 G/DL

B. 10 G/DL

C. 7 G/DL

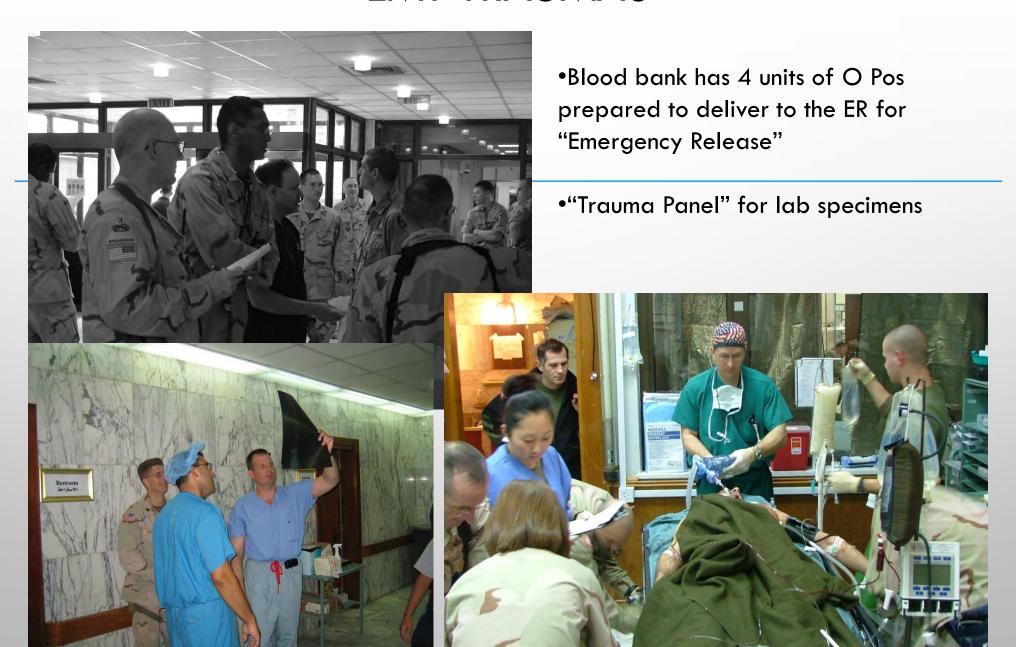
D. 8 G/DL

E. 6 G/DL



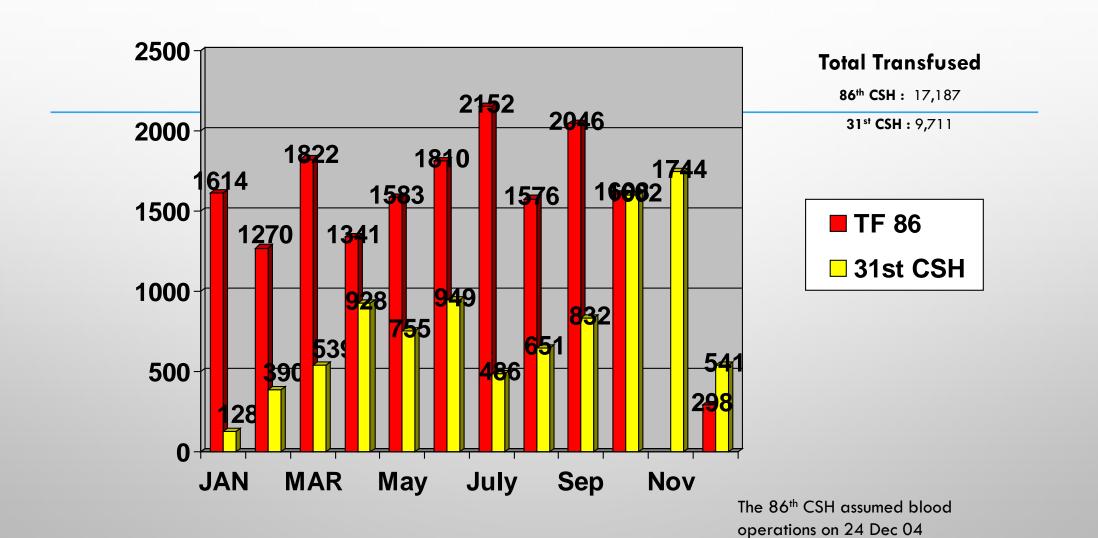


EMT TRAUMAS



Medical Task Force 86 Units Transfused

As of 11 SEP 05



CRITICAL TRAUMA PATIENTS

- 19 JUN 2005 (FATHER'S DAY)
 - RIGHT LEG TRAUMATIC AMPUTATION FOLLOWING AN IED.
 - 203 BLOOD PRODUCTS.

CRITICAL TRAUMA PATIENTS

- 7 SEP 05: IED, SHRAPNEL LEFT BUTTOCK, BACK, AND LEFT LOWER EXTREMITY
 - CRYO: 20 UNITS
 - FFP: 64 UNITS
 - PLT: 6 UNITS
 - RBC: 82 UNITS
 - WB: 35 UNITS
 - TOTAL: 207 UNITS OF BLOOD PRODUCTS
- SOLDIER IS FROM THE SAME MP UNIT WHO LOST A SOLDIER AT WRAMC AFTER RECEIVING
 203 UNITS ON 19 JUN 05



EVIDENCE BASED MEDICINE

• TRICC TRIAL ("TRANSFUSION REQUIREMENTS IN CRITICAL CARE") WAY BACK IN 1999.





The New England Journal of Medicine

Copyright, 1999, by the Massachusetts Medical Society

VOLUME 340

FERRUARY 11, 1999



A restrictive strategy of red-cell transfusion is at least as effective as and possibly superior to a liberal transfusion strategy in critically ill patients, with the possible exception of patients with acute myocardial infarction and unstable angina.



A MULTICENTER, RANDOMIZED, CONTROLLED CLINICAL TRIAL OF TRANSFUSION REQUIREMENTS IN CRITICAL CARE

PAUL C. HÉBERT, M.D., GEORGE WELLS, Ph.D., MORRIS A. BLAJCHMAN, M.D., JOHN MARSHALL, M.D., CLAUDIO MARTIN, M.D., GIUSEPPE PAGLIARELLO, M.D., MARTIN TWEEDDALE, M.D., Ph.D., IRWIN SCHWEITZER, M.SC., ELIZABETH YETISIR, M.SC., AND THE TRANSFUSION REQUIREMENTS IN CRITICAL CARE INVESTIGATORS
FOR THE CANADIAN CRITICAL CARE TRIALS GROUP*

ABSTRACT

Background To determine whether a restrictive strategy of red-cell transfusion and a liberal strategy produced equivalent results in critically ill patients, we compared the rates of death from all causes at 30 days and the severity of organ dysfunction.

Methods We enrolled 838 critically ill patients with euvolemia after initial treatment who had hemoglobin concentrations of less than 9.0 g per deciliter within 72 hours after admission to the intensive care unit and randomly assigned 418 patients to a restrictive strategy of transfusion, in which red cells were transfused if the hemoglobin concentration dropped below 7.0 g per deciliter and hemoglobin concentrations were maintained at 7.0 to 9.0 g per deciliter, and 420 patients to a liberal strategy, in which transfusions were given when the hemoglobin concentration fell below 10.0 g per deciliter and hemoglobin concentrations were maintained at 10.0 to 12.0 g per deciliter.

Results Overall, 30-day mortality was similar in the two groups (18.7 percent vs. 23.3 percent, P= 0.11). However, the rates were significantly lower with the restrictive transfusion strategy among patients who were less acutely ill - those with an Acute Physiology and Chronic Health Evaluation II score of ≤20 (8.7 percent in the restrictive-strategy group and 16.1 percent in the liberal-strategy group, P=0.03) — and among patients who were less than 55 years of age (5.7 percent and 13.0 percent, respectively; P=0.02), but not among patients with clinically significant cardiac disease (20.5 percent and 22.9 percent, respectively; P=0.69). The mortality rate during hospitalization was significantly lower in the restrictive-strategy group (22.2 percent vs. 28.1 percent, P=0.05).

Conclusions A restrictive strategy of red-cell transfusion is at least as effective as and possibly superior to a liberal transfusion strategy in critically ill patients, with the possible exception of patients with acute myocardial infarction and unstable angina. (N Engl J Med 1999;340:409-17.)

@1999, Massachusetts Medical Society.

ED-cell transfusions are a cornerstone of critical care practice,¹ but there are divergent views on the risks of anemia and the benefits of transfusion in this setting. One important concern is that anemia may not be well tolerated by critically ill patients.²³ Indeed, two recent studies suggested that anemia increases the risk of death after surgery in patients with cardiac disease² and in critically ill patients.³ Red-cell transfusions are used to augment the delivery of oxygen in the hope of avoiding the deleterious effects of oxygen debt.⁴ This view prompted the routine use of transfusion in patients with hemoglobin concentrations that were often more than 10.0 g per deciliter in studies evaluating resuscitation protocols.⁵.6

Critically ill patients may, however, be at increased risk for the immunosuppressive⁷⁸ and microcirculatory^{9,10} complications of red-cell transfusions. In addition, concern about the supply and safety of blood has also encouraged a conservative approach to transfusions. For these reasons, the optimal transfusion practice for various types of critically ill patients with anemia has not been established.

To elucidate the potential risks of anemia and possible benefits of transfusions in critically ill patients, we conducted a randomized, controlled, clinical trial to determine whether a restrictive approach to redcell transfusion that maintains hemoglobin concentrations between 7.0 and 9.0 g per deciliter is equiv-

From the Critical Care Program (P.C.H., G.P.) and the Clinical Epidemiology Unit (P.C.H., G.W., L.S., E.Y.), University of Ortuwa, Ortawa; the Department of Pathology, McMaster University, Hamilton, Ont. (M.A.B.); the Critical Care Program, University of Toronto, Toronto (J.M.); the Critical Care Program, University of Western Omario, London (C.M.); and the Critical Care Program, University of British Columbia, Vancouver (M.T.) — all in Canada. Address reprint requests to Dr. Hébert at the Department of Medicine, Ottawa General Hospital, 501 Smyth Rd., Box 205, Ottawa, ON K1H 8L6, Canada.

TRICC Study

The "Transfusion Requirements in Critical Care" study, published by Hebert et al in the New England Journal of Medicine in 1999 (NE/M 1999;340:409-17), is one of the most important studies in transfusion medicine in the last 50 years. The authors performed a randomized, prospective study dividing patients in the intensive care unit into those who received transfusions "liberally" (only with hemoglobin below 10.0 g/dL) and on a "restricted" basis (only with hemoglobin levels below 7.0 g/dL). The study showed that the restrictive transfusion strategy was at least as effective as the liberal, with significantly less in-house mortality in the restrictive group (with the exception of those with serious cardiac disease). This is one of the landmark studies in this field, and it is quoted often to help justify the use of lower thresholds for transfusion of red cells.

^{*}Study investigators are listed in the Appendix.

AABB

Annals of Internal Medicine

CLINICAL GUIDELINE

Red Blood Cell Transfusion: A Clinical Practice Guideline From the AABB*

Jeffrey L. Carson, MD; Brenda J. Grossman, MD, MPH; Steven Kleinman, MD; Alan T. Tinmouth, MD; Marisa B. Marques, MD; Mark K. Fung, MD, PhD; John B. Holcomb, MD; Orieji Illoh, MD; Lewis J. Kaplan, MD; Louis M. Katz, MD; Sunil V. Rao, MD; John D. Roback, MD, PhD; Aryeh Shander, MD; Aaron A.R. Tobian, MD, PhD; Robert Weinstein, MD; Lisa Grace Swinton McLaughlin, MD; and Benjamin Djulbegovic, MD, PhD, for the Clinical Transfusion Medicine Committee of the AABB

Description: Although approximately 85 million units of red blood cells (RBCs) are transfused annually worldwide, transfusion practices vary widely. The AABB (formerly, the American Association of Blood Banks) developed this guideline to provide clinical recommendations about hemoglobin concentration thresholds and other clinical variables that trigger RBC transfusions in hemodynamically stable adults and children.

Methods: These guidelines are based on a systematic review of randomized clinical trials evaluating transfusion thresholds. We performed a literature search from 1950 to February 2011 with no language restrictions. We examined the proportion of patients who received any RBC transfusion and the number of RBC units transfused to describe the effect of restrictive transfusion strategies on RBC use. To determine the clinical consequences of restrictive transfusion strategies, we examined overall mortality, nonfatal myocardial infarction, cardiac events, pulmonary edema, stroke, thromboembolism, renal failure, infection, hemorrhage, mental confusion, functional recovery, and length of hospital stay.

Recommendation 1: The AABB recommends adhering to a restrictive transfusion strategy (7 to 8 g/dL) in hospitalized, stable patients (Grade: strong recommendation; high-quality evidence).

Recommendation 2: The AABB suggests adhering to a restrictive strategy in hospitalized patients with preexisting cardiovascular disease and considering transfusion for patients with symptoms or a hemoglobin level of 8 g/dL or less (Grade: weak recommendation; moderate-quality evidence).

Recommendation 3: The AABB cannot recommend for or against a liberal or restrictive transfusion threshold for hospitalized, hemodynamically stable patients with the acute coronary syndrome (Grade: uncertain recommendation; very low-quality evidence).

Recommendation 4: The AABB suggests that transfusion decisions be influenced by symptoms as well as hemoglobin concentration (Grade: weak recommendation; low-quality evidence).

www.annals.org

Ann Intern Med. 2012;157:49-58.
For author affiliations, see end of text.
This article was published at www.annals.org on 27 March 2012.

Nine Landmark Randomized Clinical Trials Supporting Hb Triggers of 7-8 g/dL (Less is More)

Randomized Trials:

- all supporting Hb triggers of 7 or 8 g/dL
- Hebert PC, et al: NEJM 1999 Critically ill MICU pts.
- Lacroix J, et al: NEJM 2007 Critically ill PICU pts.
- Villanueva C, et al: NEJM 2013 Severe GI Bleeding
- Holst LB, et al: NEJM 2014 Septic Shock
- Robertson CS. et al: JAMA 2014 Traumatic Brain Injury Same/Worse
- 8 | Carson JL, et al: NEJM 2011 Elderly orthopedic Surg.
- Hajjar LA, et al: JAMA 2010 Cardiac surgery pts.
- 7.5 Murphy GJ, et al: NEJM 2015 Cardiac surgery pts.
- 7.5 Mazer CD, et al: NEJM 2017 Cardiac surgery pts.

Higher

Triggers

(9-10 g/dL)

- Same/Worse
 - Same
 - Worse
 - Same

 - Same
 - Same
 - Same
 - Same/Worse

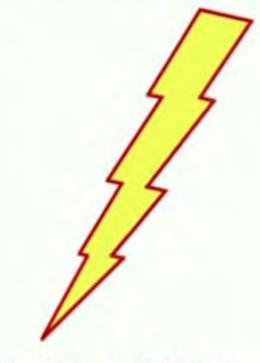
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Three Categories of Risks / Adverse Effects from Blood Transfusion

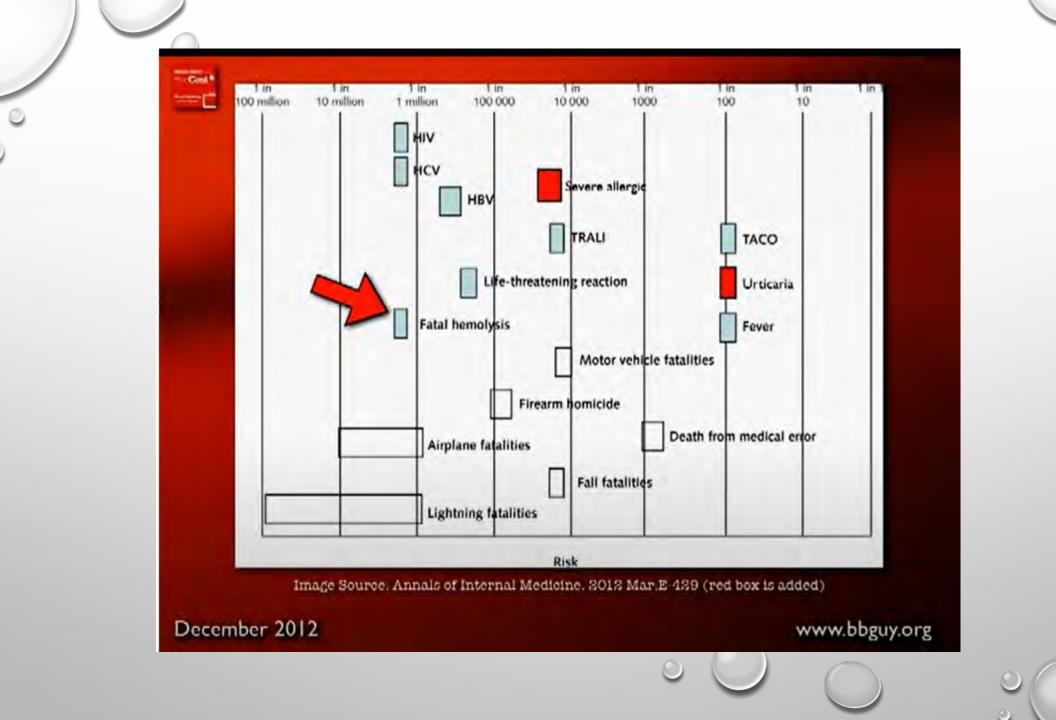
Clinical Event	Risk / Unit		
Allergic/Urticaria	1 in 100	1	
RBC Alloimmunization	1 in 100	Common	
TACO	1 in 100	Jedininen	
TRALI	1 in 5,000	1	
Hemolytic Rxn	1 in 6,000	Not so Rare	
Wrong Unit Given	1 in 15,000	J	
Hepatitis B	1 in 400,000	1.	
Hepatitis C	1 in 2,000,000	Rare	
HIV 1 and HIV 2	1 in 2,000,000	J	

Three Categories of Risks / Adverse Effects from Blood Transfusion

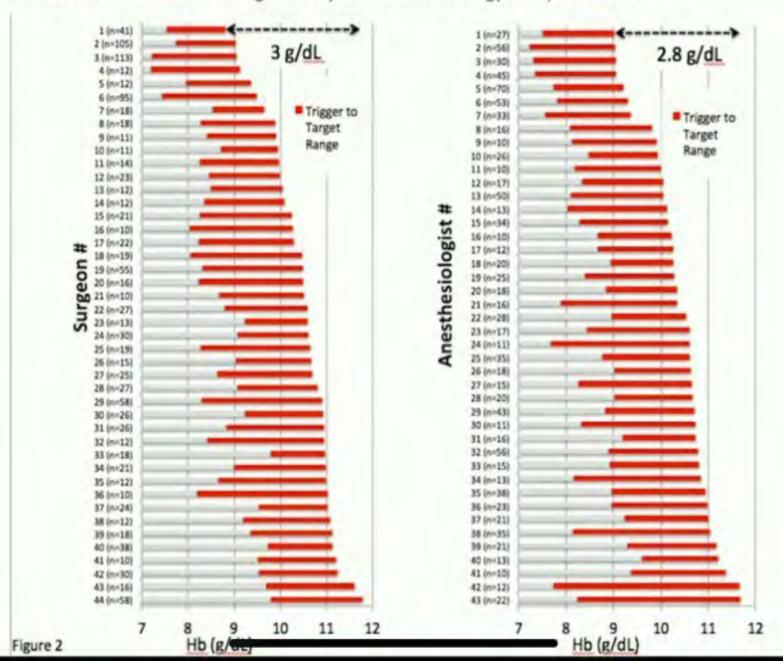
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Hepatitis C	1 in 2,000,000
HIV 1 and HIV 2	1 in 2,000,000

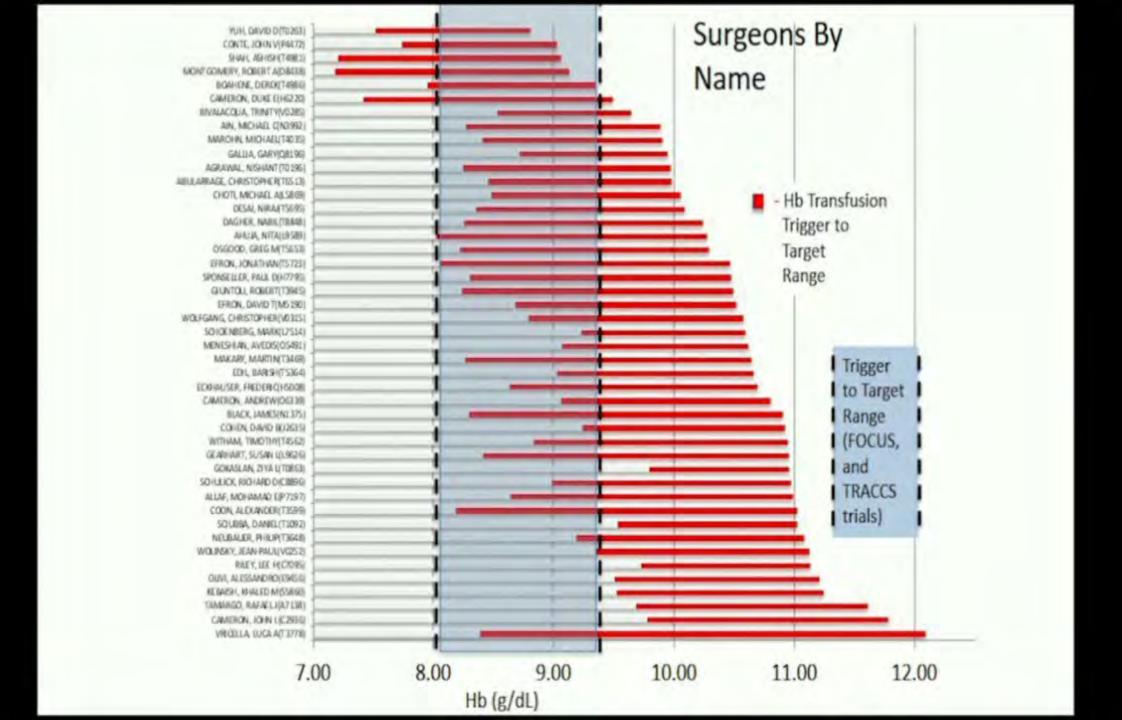


Hep C and HIV =
Death by
Lightning Strike



Frank SM, et al. Variability in blood and blood component utilization as assessed by an anesthesia information management system. Anesthesiology 2012;117:99-106



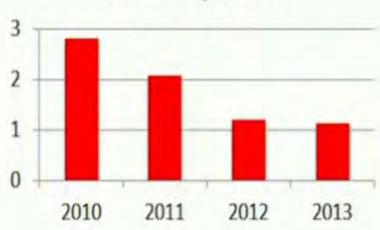




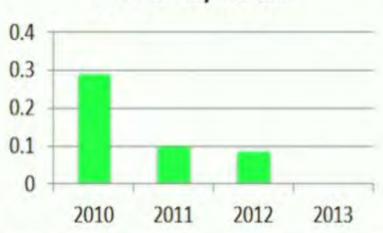
Public Display

Surgeon #44 (with permission)

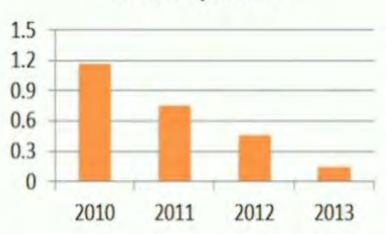
RBC Units/Patient



PLTS Units/Patient



FFP Units/Patient





6 Societies have aims to Reduce unnecessary transfusion

An initiative of the ABIM Foundation

Society of Critical Care Medicine

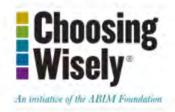
American Society of Anesthesiologists

American Society of Hospital Medicine

American Society of Hematology

American College of Obstetricians and Gynecologists

American. Assn. of Blood Banks (AABB)



Our Mission

Clinician Lists

For Patients

Getting Started

Success Stories

Q

Getting Started

Lists of Recommendations

Search Recommendations

Clinician Lists

Complete lists of recommendations by society can be found by clicking the society name or via individual recommendation pages.

Your search returned 5 results Society Recommendation Don't transfuse more units of red blood cells or other American Association of **Blood Banks** components than absolutely necessary. Don't transfuse red blood cells for iron deficiency American Association of **Blood Banks** without hemodynamic instability. American Association of Don't routinely use blood products to reverse **Blood Banks** warfarin. Don't perform serial blood counts on clinically stable American Association of **Blood Banks** patients. Don't transfuse O negative blood except to O American Association of negative patients and in emergencies for women of **Blood Banks** child bearing potential with unknown blood group.

	ion of Blood
- filter by -	
SEARCH	Clear Filters
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https://www.choosingwisely.org/







Our Mission

Clinician Lists

For Patients

Getting Started

Success Stori

Getting Started

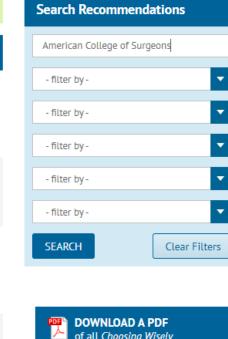
Lists of Recommendations

Search Recommendations

Clinician Lists

Complete lists of recommendations by society can be found by clicking the society name or via individual recommendation pages.

Your search returned 5 results		
Society	Recommendation	
American College of Surgeons	Don't perform axillary lymph node dissection for clinical stages I and II breast cancer with clinically negative lymph nodes without attempting sentinel node biopsy.	
American College of Surgeons	Avoid the routine use of "whole-body" diagnostic computed tomography (CT) scanning in patients with minor or single system trauma.	
American College of Surgeons	Avoid colorectal cancer screening tests on asymptomatic patients with a life expectancy of less than 10 years and no family or personal history of colorectal neoplasia.	
American College of Surgeons	Avoid admission or preoperative chest x-rays for ambulatory patients with unremarkable history and physical exam.	
American College of Surgeons	Don't do computed tomography (CT) for the evaluation of suspected appendicitis in children until after ultrasound has been considered as an option.	





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Download the Choosing Wisely



Clinician Lists

Complete lists of recommendations by society can be found by clicking the society name or via individual recommendation pages.

Search Recommendations

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SEARCH

App

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American Society of Anesthesiologists

Clear Filters

Your search returned 5 results

Society	Recommendation
American Society of Anesthesiologists	Don't obtain baseline laboratory studies in patients without significant systemic disease (ASA I or II) undergoing low-risk surgery – specifically complete blood count, basic or comprehensive metabolic panel, coagulation studies when blood loss (or fluid shifts) is/are expected to be minimal.
American Society of Anesthesiologists	Don't routinely administer colloid (dextrans, hydroxylethyl starches, albumin) for volume resuscitation without appropriate indications.
American Society of Anesthesiologists	Don't administer packed red blood cells (PRBCs) in a young healthy patient without ongoing blood loss and hemoglobin of ≥6 g/dL unless symptomatic or hemodynamically unstable.
American Society of Anesthesiologists	Don't use pulmonary artery catheters (PACs) routinely for cardiac surgery in patients with a low risk of hemodynamic complications (especially with the concomitant use of alternative diagnostic tools (e.g., TEE).
American Society of Anesthesiologists	Don't obtain baseline diagnostic cardiac testing (trans-thoracic/esophageal echocardiography – TTE/TEE) or cardiac stress testing in asymptomatic stable patients with known cardiac disease (e.g., CAD, valvular disease) undergoing low or moderate risk non-cardiac surgery.





An initiative of the ABIM Foundation

https://www.sabm.org/wp-content/uploads/2018/08/SABM-

Choosing-Wisely-List.pdf

Society for the Advancement of Blood Management



Five Things Physicians and Patients Should Question

Don't proceed with elective surgery in patients with properly diagnosed and correctable anemia until the anemia has been appropriately treated.

Anemia is common, presenting in approximately one-third of patients undergoing elective surgery. There is often the misconception that anemia is harmless, when, in fact, it is independently associated with significant morbidity and mortality that can be as high as 30-40% in certain patient populations. Treatment of anemia improves patient readiness for surgery, aids in management of comorbid conditions, decreases length of stay and readmission rates, and reduces transfusion risks. Treatment modalities may include nutritional supplementations, such as iron, B12 and folate, changes in medication, management of chronic inflammatory conditions or previously undiagnosed malignancy, or other interventions based on the etiology.

7

Don't perform laboratory blood testing unless clinically indicated or necessary for diagnosis or management in order to avoid iatrogenic anemia.

Up to 90% of patients become anemic by day 3 in the intensive care unit. Although laboratory testing can aid in diagnosis, prognosis and treatment of disease, a significant number of tests are inappropriate or unnecessary. Anemia secondary to iatrogenic blood loss causes an increased length of stay and mortality. Increased phlebotomy for laboratory testing also increases the odds for transfusion and its associated risks. Unnecessary laboratory testing adds to the cost of care through laboratory test charges and also by increasing downstream costs due to unnecessary interventions, prescriptions, etc. Thus judicious use of laboratory testing is recommended, and testing should not be performed in the absence of clinical indications.

3

Don't transfuse plasma in the absence of active bleeding or significant laboratory evidence of coagulopathy.

Recent studies demonstrate that plasma is often transfused inappropriately. In the absence of active bleeding or clear evidence of coagulopathy, current literature shows no reduction in blood loss or transfusion requirements with the use of plasma, but shows increased risk of transfusion-associated adverse events such as transfusion-related acute lung injury, transfusion-associated circulatory overload and allergic reactions. These transfusion-associated adverse events lead to poorer outcomes and increased cost of care.



5

Avoid transfusion when antifibrinolytic drugs are available to minimize surgical bleeding.

Antifibrinolytic pharmacologic therapy has been shown to reduce blood loss and transfusion requirements in orthopedic and cardiovascular surgeries. Early administration of tranexamic acid, specifically within three hours, in trauma and obstetric hemorrhage significantly reduces mortality and bleeding.

Avoid transfusion, outside of emergencies, when alternative strategies are available as part of informed consent; make discussion of alternatives part of the informed consent process.

Informed choice/consent regarding transfusion and other effective methods should be standardized and consistently delivered. Throughout the world, there is wide variation among medical practitioners and hospitals with regard to medical knowledge about the true risks of transfusion, alternatives to transfusion, and the delivery of this information to patients. Outside of the truly emergent clinical situation, transfusion should be avoided or limited when other interventions are available. Alternative strategies include, but are not limited to pharmacologic agents, cell salvage, normovolemic hemodilution and minimally-invasive surgical techniques.

https://www.sabm.org/wp-content/uploads/2018/08/SABM-Choosing-Wisely-List.pdf

"Right dose, right product, right patient, right time"

- Preop anemia treatment –
 A \$5 bottle of iron pills beats \$500 of blood
 IV iron, EPO as needed preop and postop
- Good surgery, less invasivelaparoscopic, robotic, endovascular
- Blood Salvage (Cell Saver)The "Centerpiece" of blood conservation
- 4. Topical hemostatics and newer cautery devices
- 5. Minimize phlebotomy
- 6. CPOE with clinician decision support
- 7. Antifibrinolytics (Tranexamic acid, Amicar)
- 8. Point of care testing (TEG, rapid turnaround)
- 9. Audits with feedback
- 10. Education

MORE VIDEOS



PRE-OPERATIVE ANEMIA

- PRE-OPERATIVE ANEMIA-PT. CENTRIC: A 62 YO GOING FOR A THA VERSUS A 28 YO FEMALE GOING FOR AN "X" PROCEDURE (FE DEFICIENCY ANEMIA).
- CONSIDER 28-30 D PRE-OP WORK-UP.
- IT IS WELL KNOWN THAT PTS. WITH PRE-OPERATIVE ANEMIA ARE ASSOCIATED WITH INCREASED LENGTH OF STAY, ADVERSE OUTCOMES, AND WOUND INFECTIONS.
- DON'T "TANK-UP" THE PATIENT PRIOR TO SURGERY. REMEMBER ANEMIA IS A SIGN OF AN UNDERLYING DISEASE NOT A SPECIFIC DIAGNOSIS.

American Association of Blood Banks

View all recommendations from this society

Released April 24, 2014; Updated January 28, 2022

Don't transfuse more units of red blood cells or other components than absolutely necessary.

- For red blood cells, a restrictive threshold (7.0-8.0g/dL) should be used for the vast majority of hospitalized, stable patients without evidence of inadequate tissue oxygenation (evidence supports a threshold of 8.0g/dL in patients with pre-existing cardiovascular disease). Transfusion decisions should be influenced by symptoms and hemoglobin concentration. Single unit red cell transfusions should be the standard for non-bleeding, hospitalized patients. Additional units should only be prescribed after re-assessment of the patient and their hemoglobin value.
- For plasma, do not transfuse plasma to correct coagulopathy in non-bleeding patients or patients.
- Do not transfuse platelets without laboratory guidance outside of fixed-ratio massive transfusions.

These items are provided solely for informational purposes and are not intended as a substitute for consultation with a medical professional. Patients with any specific questions about the items on this list or their individual situation should consult their physician.

How The List Was Created

Sources

Carson JL, Grossman BJ, Kleinman S, Tinmouth AT, Marques MB, Fung MK, Holcomb JB, Illoh O, Kaplan LJ, Katz LM, Rao SV, Roback JD, Shander A, Tobian AA, Weinstein R, Swinton McLaughlin LG, Djulbegovic B; Clinical Transfusion Medicine Committee of the AABB. Red blood cell transfusion: a clinical practice guideline from the AABB. Ann Intern Med. 2012 Jul 3;157(1):49–58.



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BLOOD SAVES LIVES WHEN YOU NEED IT, BUT ONLY INCREASES RISKS AND COSTS WHEN YOU DON'T!

STEVEN FRANK, MD

I'm long on record as stating that in "the old days," when people used to use a trigger of 10 g/dL of hemoglobin to transfuse people, that was silly. But now, I fear that the pendulum has swung the other way, and I hear people saying very strongly, "Any transfusion with a hemoglobin over 7 is obviously automatically inappropriate," and I think that's just as silly. So again, I know this is a sidebar, Aryeh, but I wonder if I could get your take on this? My position is, those are numbers and we're not treating numbers, we are treating patients, and I don't want to influence your answer (obviously I couldn't), but how do you feel about that argument?

"Why give 2 when 1 will do?"

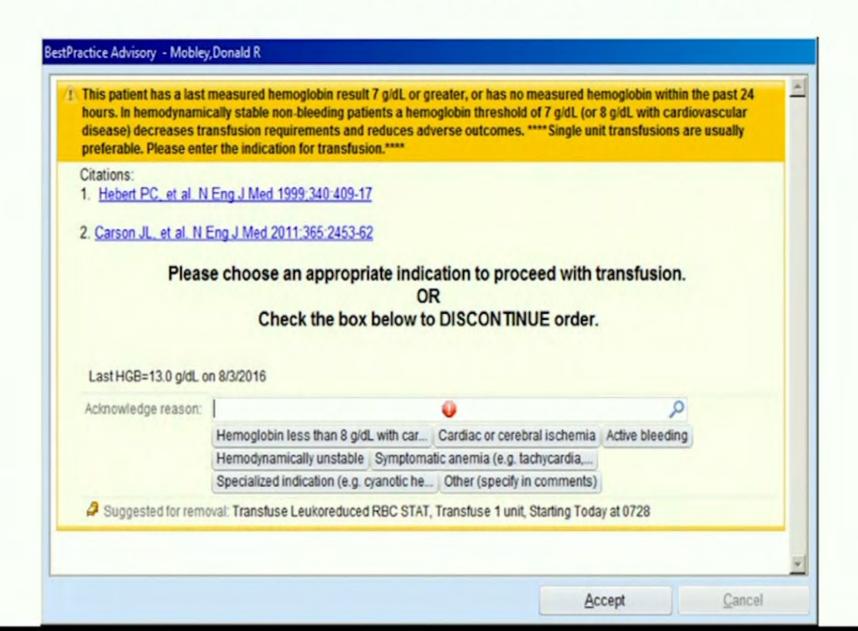
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THE REAL QUESTION IS NOT WHETHER A TRANSFUSION WAS
"APPROPRIATE."

THE REAL QUESTION IS WHETHER A TRANSFUSION WAS
"AVOIDABLE."

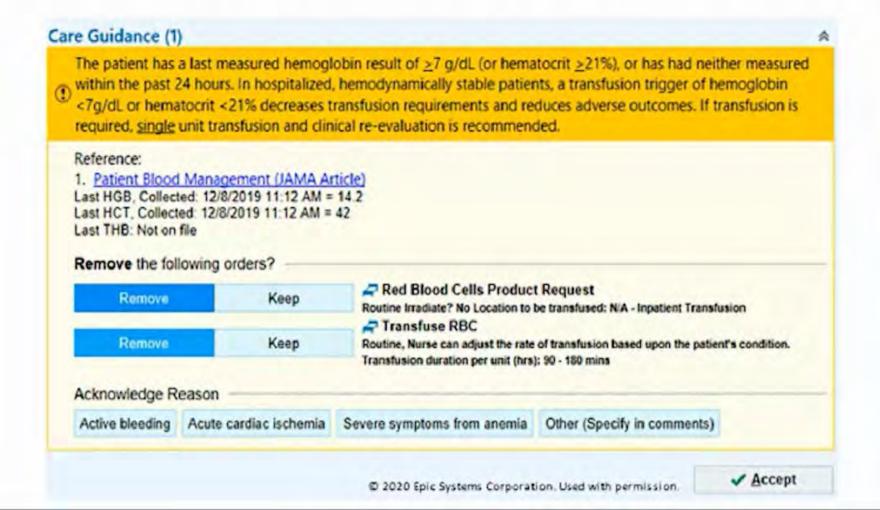
ARYEH SHANDER, MD

Best Practice Advisory triggered on Hb ≥ 7 g/dL



Hemoglobin/hematocrit BPA:

End user action to remove order 30% (very successful)

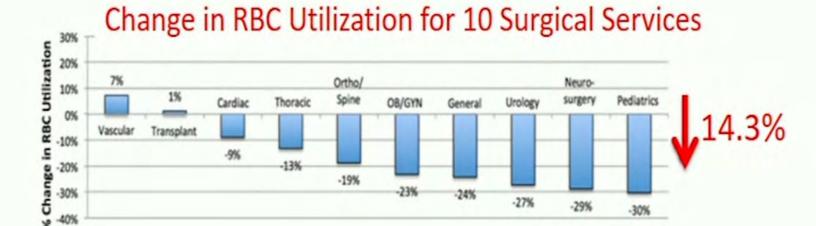


Zuckerberg GS, et al. TRANSFUSION, 2015

"Efficacy of Education Followed by Computerized Provider Order Entry with Clinician Decision Support to Reduce Red Blood Cell Utilization"

Monthly number of RBC units w/ preceding Hb > 8







Bayview Hip and Knee Replacement FY13 – FY15 Tranexamic acid - a "game-changer"

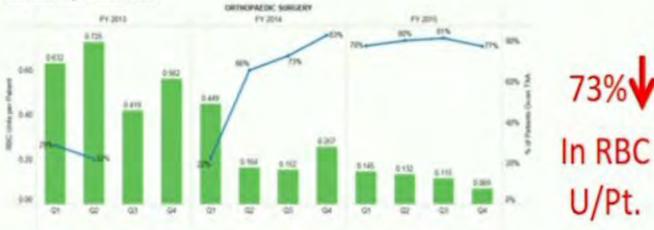
Orthopedic, Cardiac, Trauma, and PPH cases "game changer". No evidence to date that it increases the risk for development of clots. Caveat is in those pts with SAH!!

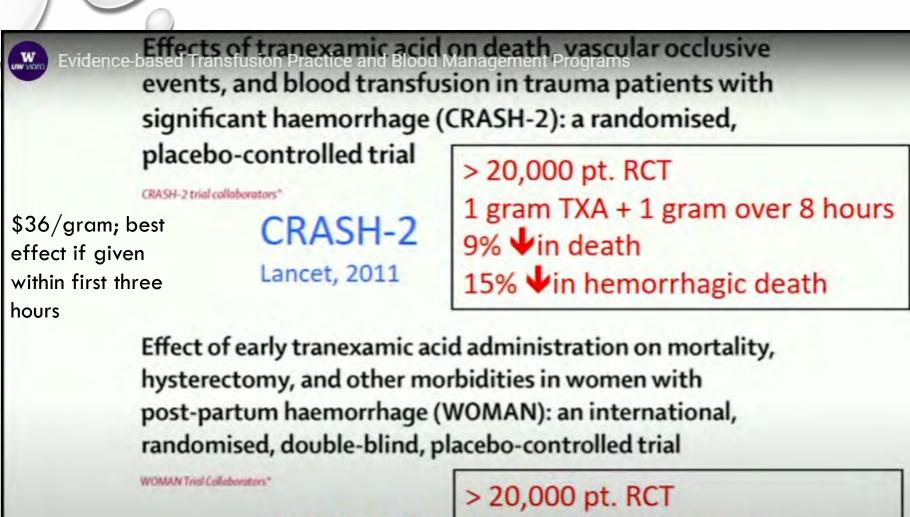
TXA began

Bayview - Total RBC Units and % of Patients Transfused by Attending Physician Orthopedics Service by Quarter (7/12-5/15 Discharge Dates) for Hip & Knee Replacement APR-DRGs



Bayview - Average RBC Units per Discharged Patient and % of Patients Given TXA Drug for Hip and Knee Replacement DRGs





MORE VIDEOS

WOMAN

Lancet, 2017

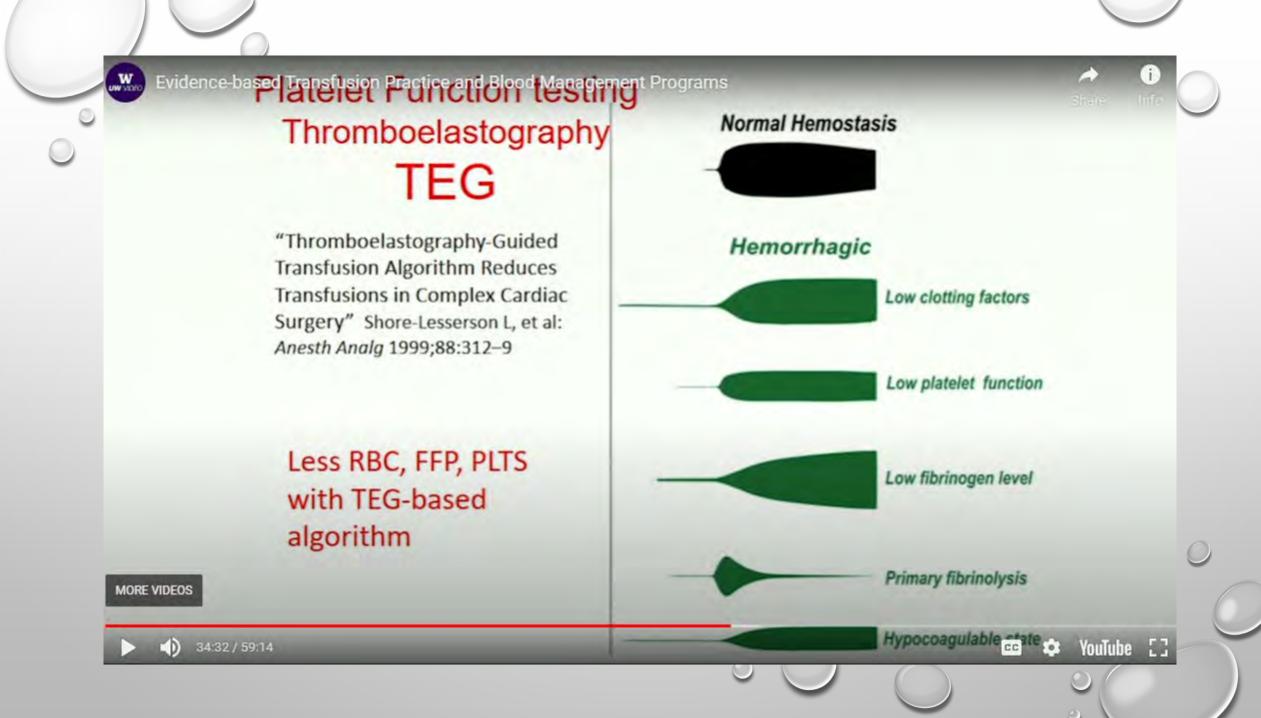
1 gram TXA ± 1 more gram 19% ♥ in death from bleeding 31% if TXA given early (<3 hrs)

YouTube []





28:14 / 59:14











Ways to Decrease Blood Utilization

Provided as guidance by South Texas Blood & Tissue Center

Red Blood Cells/Platelets:

- When possible, split platelet units into smaller doses, which is an evidence-based, practice-based method for adult oncology inpatients based on the PLADO study (Slichter et al. N Engl J Med 2010;362:600-613).
- Avoid prophylactic transfusion of thrombocytopenic patients not actively bleeding, so platelets are available for actively bleeding patients needing therapeutic platelet transfusion.
- Consider using platelet counts of 5 x 10³ as the trigger for prophylactic platelet transfusions rather than 10 x 10³.

Plasma/Cryoprecipitate:

- Evaluate a more permissive policy for Prothrombin Complex Concentrates (e.g., Kcentra).
- Consider using tranexamic acid when appropriate.
- Implement off-label use of fibrinogen concentrate (RiaSTAP or other FDA approved similar products) to reduce the use of cryoprecipitate.

"The average decrease in blood donations was found to be 38%, with some regions showing up to 67% decrease."

The Impact of COVID-19 on Blood Transfusion Services: A Systematic Review and Meta-Analysis

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Keywords

COVID-19 · Blood transfusion · Donations · RNAaemia

Abstract

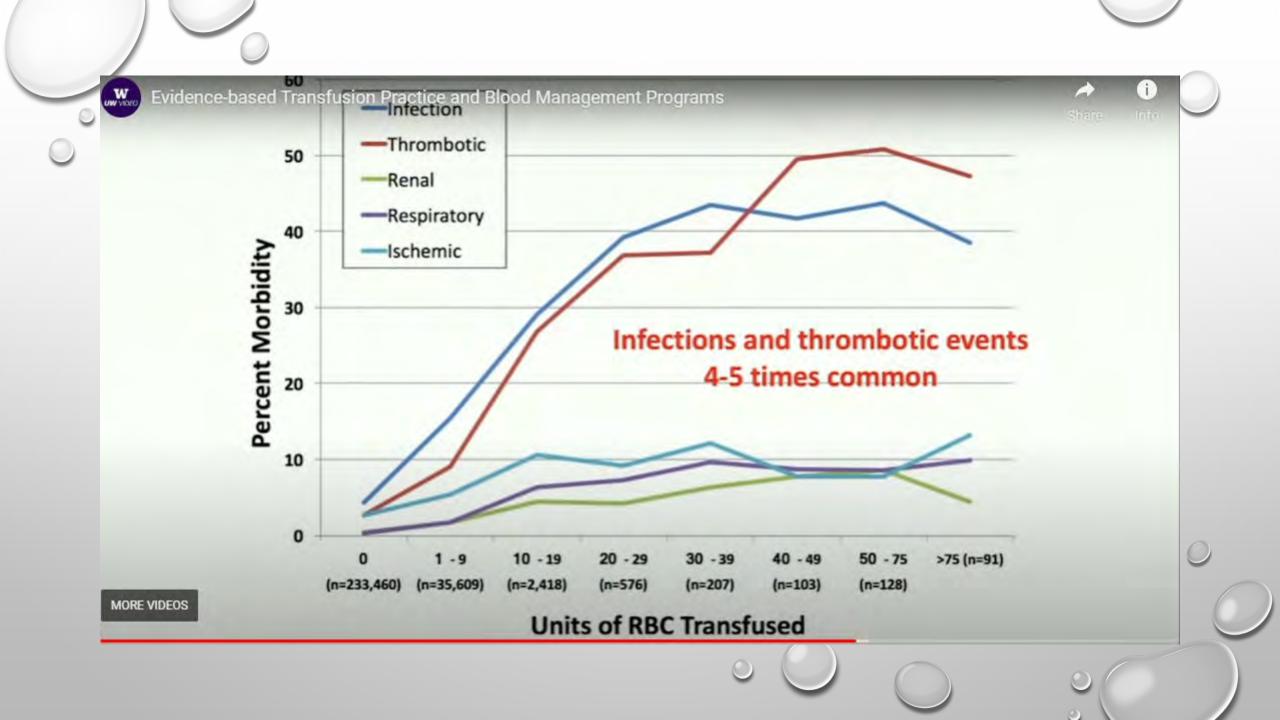
Introduction: While SARS-CoV-2's main transmission route is through respiratory droplets, research has found that viral RNA could be detected in blood samples, causing concerns over the safety of blood donations and blood products. This paper therefore aims to systematically search for studies that have addressed their country's lack of donations and analyse the risk of blood transfusion-transmission. As such, it will answer the question "should blood services focus more on donation vigilance or worry more about the risks of transmission through blood products?" Methods: 38 articles were identified through a systematic review adopting the PRISMA and STROBE guidelines. Meta-analysis was conducted using Open-Meta software. Results: The average decrease in blood donations was found to be 38%, with some regions showing up to 67% decrease. To assess the risk of actual blood transfusiontransmission, three datasets were analysed. Firstly, the viral load in COVID-19 patients was studied and found to have less than 1% detection rate (ARD = -0.831, 95% -0.963, -0.699). Secondly, the prevalence of finding viral RNA in a pool of donations was nearly -1.503 (ARD = -1.538, -1.468). Lastly, recipients who were given blood products of positive donors were found to be -0.911 (ARD 95% = -1.247, -0.575). Discussion/Conclusion: Blood centres should focus more on launching initiatives and policies that would increase their countries' blood supply as the virus has no direct threat to blood safety.

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Introduction

Blood transfusion is considered an integral part of medicine that treats thousands of patients every year, making blood products' management, safety, and storage fundamental in every country's national healthcare policy [1, 2]. Maintaining an adequate supply of blood products is no easy feat considering blood components have a short shelf life ranging from 5 to 42 days, and the fact that blood collection relies entirely on generous donations from the public [3]. However, not everyone can or is willing to donate; hence, constant encouragement from blood centres is therefore a necessary prosocial behaviour so that blood can be collected from all ethnicities and blood types. Due to its reliance on donations, the supplies can easily be diminished when disaster strikes especially in the cases of pandemics [4, 5].

Since late 2019, the world was plagued by a new disease that originated from Wuhan, China, infecting over 28 million people and causing at least 900,000 deaths across 188 countries [4]. Healthcares were suddenly hit with critical shortages of PPEs, ventilators, and hospital beds, causing governments around the world to act in an unprecedented manner to mitigate the exponential rise of infected cases [6–8]. Social distancing was implemented, as well as closing down public venues, schools, universities, and any non-essential work [7, 8]. This lockdown and the fear of virus transmission has not only affected the public's health and a decline in economy, it also caused a significant drop in the number of blood donations across the world, creating shortages at various blood banks and diminishing nationwide blood supply.



- Blood saves lives when you need it - Only increases risks and costs when you don't



TRANSFUSION REACTIONS



HTTPS://WWW.AABB.ORG/DOCS/DEFAULT-SOURCE/DEFAULT-DOCUMENT-LIBRARY/RESOURCES/AABB-QUICK-REFERENCE-GUIDE-NHSN-HEMOVIGILANCE-MODULE.PDF?SFVRSN=30F1600B 0

TRANSFUSION-ASSOCIATED GRAFT VS. HOST DISEASE (TAGVHD)

The introduction of immunocompetent lymphocytes into susceptible hosts. The allogeneic lymphocytes engraft, proliferate, and destroy host cells. If performed, marrow study shows hypoplasia, aplastic anemia, or marked hypocellularity with a lymphohistiocytic infiltrate.

Definitive: A clinical syndrome occurring from 2 days to 6 weeks after cessation of transfusion characterized by:

formation.

elevated ALT, AST,

and bilirubin)

Marrow aplasia

Alkaline phosphatase,

Diarrhea

Fever

 Characteristic rash: erythematous, maculopapular eruption centrally that . Hepatomegaly and may, in severe cases, progress

spreads to extremities . Liver dysfunction (i.e., to generalized erythroderma and hemorrhagic bullous Pancytopenia

Characteristic histological appearance of skin or liver biopsy.

Probable: Meets definitive criteria

Biopsy negative or not done. Possible: N/A

POST TRANSFUSION PURPURA

Thrombocytopenia usually arising 5-12 days following transfusion of cellular blood components with findings of antibodies in the patient directed against the Human Platelet Antigen (HPA) system.

Definitive: Alloantibodies in the patient directed against HPA or other platelet specific antigen detected at or after development of thrombocytopenia

Thrombocytopenia (i.e., decrease in platelets to less than 20% of pre-transfusion count).

Probable: Alloantibodies in the patient directed against HPA or other platelet specific antigen detected at or after development of thrombocytopenia.

Decrease in platelets to levels between 20% and 80% of pre-transfusion count.

Possible: PTP is suspected, but laboratory findings and/or information are not sufficient to meet defined criteria above. For example, the patient has a drop in platelet count to less than 80% of pre-transfusion count but HPA antibodies were not tested or were negative. Other, more specific adverse reaction definitions

do not apply. UNKNOWN

TRANSFUSION-RELATED ACUTE LUNG INJURY (TRALI)

Acute hypoxemia with PaO2/fraction of inspired oxygen [FIO2] ratio of 300 mmHg or less combined with chest x-ray showing bilateral infiltrates in the absence of left atrial hypertension (i.e., circulatory overload). Onset of TRALI is abrupt in association with transfusion.

Definitive: NO evidence of acute lung injury (ALI) prior to transfusion

ALI onset during or within 6 hours of cessation of transfusion

Hypoxemia defined by any of these methods:

- PaO2/FiO2 less than or equal to 300 mmHg
- . Oxygen saturation less than 90% on room air Other clinical evidence

AND

Radiographic evidence of bilateral infiltrates

No evidence of left atrial hypertension (i.e., circulatory overload)

Probable: N/A Possible: N/A

TRANSFUSION-ASSOCIATED CIRCULATORY OVERLOAD (TACO)

Infusion volume that cannot be effectively processed by the recipient either due to high rate and/or volume of infusion or an underlying cardiac or pulmonary pathology

Definitive: New onset or exacerbation of 3 or more of the following within 12 hours of cessation of transfusion:

(At least 1 of the following from A & B:)

A. Evidence of acute or worsening respiratory distress (dyspnea, tachypnoea, cyanosis and decreased oxygen saturation values in the absence of other specific causes) and/or B. Radiographic or clinical evidence of acute or worsening pulmonary edema (crackles on lung auscultation, orthopnea, cough, a third heart sound and pinkish frothy sputum in severe cases); or both

- Elevated brain natriuretic peptide (BNP) or NT-pro BNP relevant biomarker
- Evidence of cardiovascular system changes not explained by underlying medical condition (Elevated central venous pressure, evidence of left heart failure including development of tachycardia, hypertension, widened pulse pressure, jugular venous distension, enlarged cardiac silhouette and/or peripheral edema)
- · Evidence of fluid overload Probable: N/A Possible: N/A

TRANSFUSION-TRANSMITTED INFECTION (TTI)

A bacteria, parasite, virus, or other potential

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QUICK REFERENCE GUIDE

NHSN Hemovigilance Module: **Adverse Reaction Definitions**

ALLERGIC REACTION

The result of an interaction of an allergen with preformed antibodies. In some instances, infusion of antibodies from an atopic donor may also be involved. It may present with only muccocutaneous signs and symptoms.

Note: Minor allergic reactions (non-severe) do not have to be reported to NHSN.

Definitive: 2 or more of the following occurring during or within 4 hours of cessation of transfusion:

- Conjunctival edema
 Localized
- Edema of lips, tongue angioedema and uvula
- Maculopapular rash Erythema and edema Pruritus (itching)
- of the periorbital area Respiratory distress:
- Generalized flushing bronchospasm
- Hypotension Urticaria (hives)
- Probable: ANY 1 of the following occurring during or within 4 hours of cessation of transfusion:
- Conjunctival edema
 Localized
- Edema of lips, tongue
- angioedema Maculopapular rash and uvula
- Erythema and edema Pruritus (itching)
- of the periorbital area . Urticaria (hives)

Possible: N/A

TRANSFUSION ASSOCIATED

DYSPNEA (TAD)

Respiratory distress within 24 hours of cessation of transfusion that does not meet the criteria for TRALI, TACO, or allergic reaction, Respiratory distress should not otherwise be explained by a patient's underlying or pre-existing medical condition.

Definitive: Acute respiratory distress occurring within 24 hours of cessation of transfusion

Allergic reaction, TACO, and TRALI definitions are not applicable.

Probable: N/A Possible: N/A

UNKNOWN :----

Use this category if the patient experienced transfusion-related symptoms, but the medical event that caused those symptoms could not be classified.

A bacteria, parasite, virus, or other potential pathogen transmitted in donated blood to transfusion recipient.

Definitive: Laboratory evidence of a pathogen in the transfusion recipient.

Probable: N/A

Possible: Temporarily associated unexplained clinical illness consistent with infection, but no pathogen is detected in the recipient. Other, more specific adverse reactions are ruled out. Note: Possible cases cannot meet the definite or probable imputability criteria.

Possible: N/A

Use this option if the recipient experienced an adverse reaction that is not defined in the Hemovigilance Module Surveillance Protocol (e.g., transfusion-associated acute gut injury (TRAGI), transfusion-associated immunomodulation (TRIM), iron overload, microchimerism, hyperkalemia, thrombosis).

FEBRILE NON-HEMOLYTIC TRANSFUSION REACTION (FNHTR)

Fever and/or chills without hemolysis occurring in the patient during or within 4 hours of cessation of transfusion. If transfusion-related, the most common cause is a reaction to passively transfused cytokines or a reaction of recipient antibodies and leukocytes in the blood product. If blood culture of patient or residual component is performed, the results should be negative. Laboratory findings should show no evidence of acute hemolysis.

Note: Reactions may be classified as FNHTRs in the absence of fever if chills or rigors occur.

Definitive: Occurs during or within 4 hours of cessation of transfusion

AND EITHER

 Fever (greater than or equal to 38°C/100.4°F oral and a change of at least 1°C/1.8°F from pre-transfusion value)

Chills/rigors are present.

Probable: N/A

Possible: FNHTR is suspected, but reported symptoms and/or available information are not sufficient to meet the criteria defined above. Other, more specific adverse reaction definitions do not apply.

HYPOTENSIVE TRANSFUSION REACTION ==

A drop in blood pressure occurring during or within 1 hour of cessation of transfusion. Other symptoms, such as facial flushing, dyspnea, or abdominal cramps may occur, but usually hypotension is the sole manifestation.

Definitive: All other adverse reactions presenting with hypotension are excluded

Hypotension occurs during or within 1 hour after cessation of transfusion

 Adults (≥18 years): Drop in systolic BP of greater than or equal to 30 mmHg and systolic BP less

DELAYED HEMOLYTIC TRANSFUSION REACTION (DHTR) =:

The recipient develops antibodies to RBC antigen(s) between 24 hours and 28 days after cessation of transfusion. Clinical signs of hemolysis are usually present. If performed, post-transfusion LDH and bilirubin levels increase and subsequently fall back to baseline in the following days.

Note: Report all hemolytic reactions, including when the recipient is intentionally transfused with incompatible blood components.

Definitive: Positive direct antiglobulin test (DAT) for antibodies developed between 24 hours and 28 days after cessation of transfusion

AND EITHER

· Positive elution test with alloantibody present on the transfused red blood cells

· Newly-identified red blood cell alloantibody in recipient serum

AND EITHER

 Inadequate rise of post-transfusion hemoglobin level or rapid fall in hemoglobin back to pretransfusion levels

OR

 Otherwise unexplained appearance of spherocytes.

Probable: Newly-identified red blood cell alloantibody demonstrated between 24 hours and 28 days after cessation of transfusion

Incomplete laboratory evidence to meet definitive case definition criteria.

Note: Patient may be asymptomatic or have symptoms that are similar to but milder than AHTR; symptoms are not required to meet case definition criteria.

Possible: DHTR is suspected, but reported symptoms, test results, and/or available information are not sufficient to meet the criteria defined above. Other, more specific adverse

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ACUTE HEMOLYTIC TRANSFUSION REACTION (AHTR)

Rapid destruction of red blood cells during, immediately after, or within 24 hours of cessation of transfusion. Clinical and laboratory signs of hemolysis are present.

Note: Report hemolytic reactions resulting from immune or non-immune causes, including when the recipient is intentionally transfused with incompatible blood components.

Definitive: Occurs during, or within 24 hours of cessation of transfusion with new onset of ANY of the following signs/symptoms:

- Back/flank pain
- Chills/rigors
- Disseminated intravascular
- coagulation (DIC) Epistaxis
- Fever
- Pain and/or oozing at IV site

blood film

Hypotension

Oliquria/anuria

Hematuria (gross

visual hemolysis)

Plasma discoloration

· Renal failure

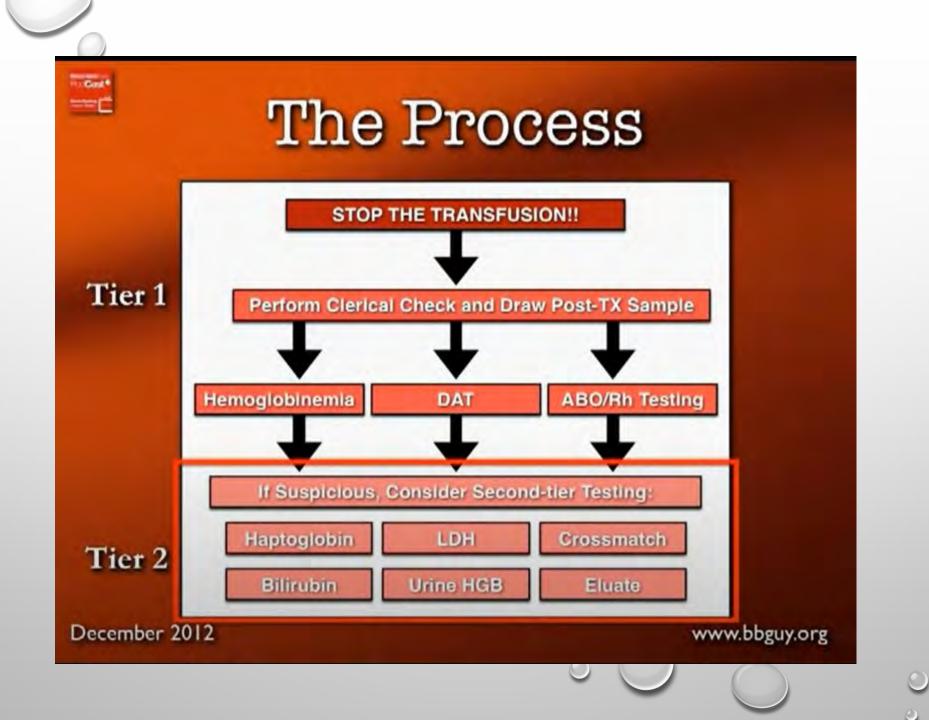
AND 2 or more of the following:

Decreased fibrinogen
 Hemoglobinuria

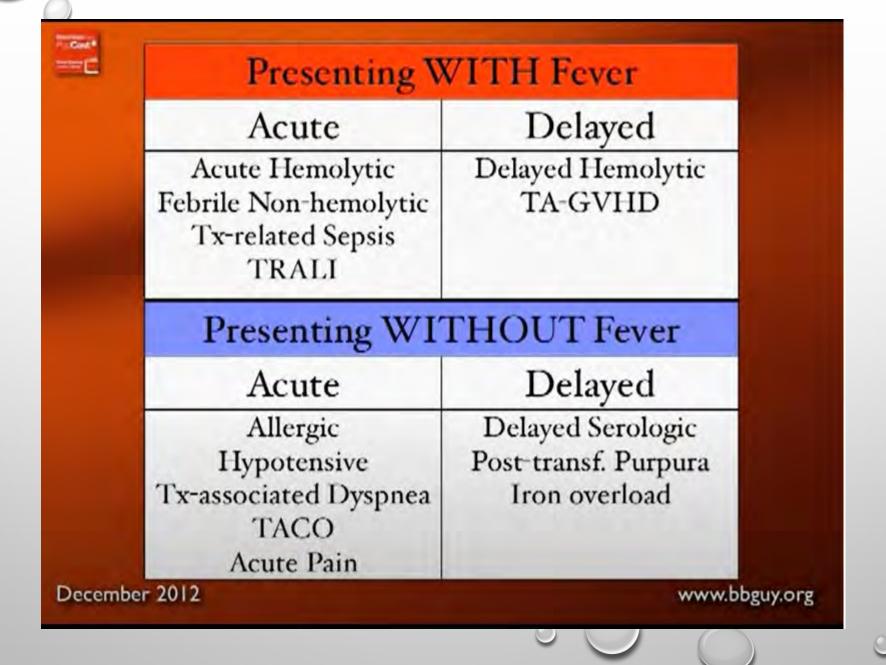
- Decreased
- haptoglobin c/w hemolysis Elevated bilirubin Spherocytes on
- Elevated LDH
- Hemoglobinemia

AND EITHER IMMUNE MEDIATED

 Positive direct antiglobulin test (DAT) for anti-lgG or anti-C3



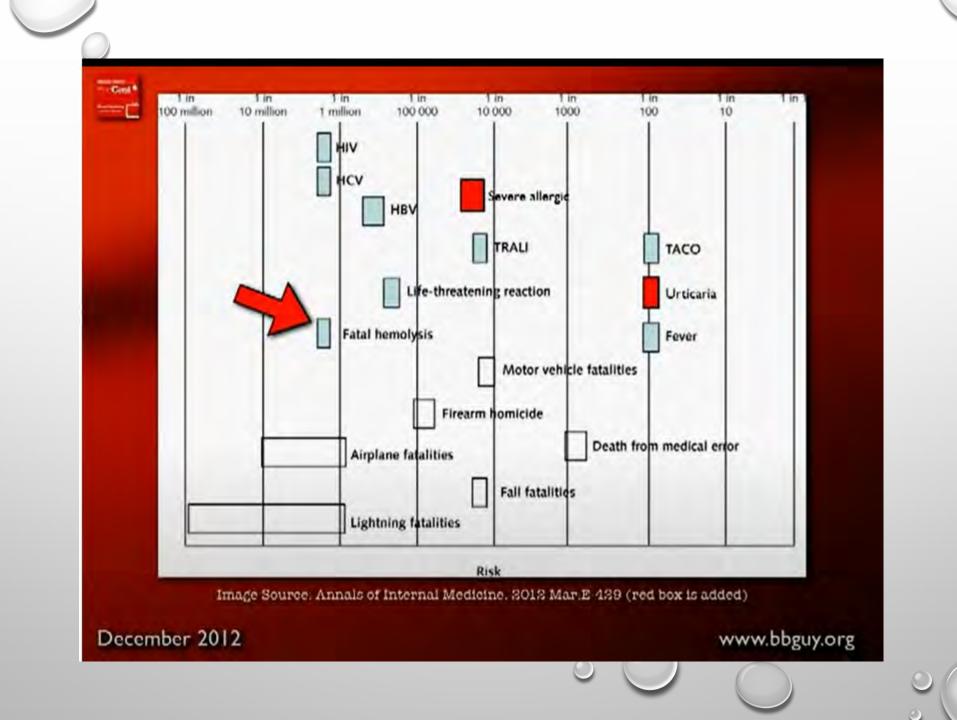
Less than 24 hours versus Greater than 24 hours is the cut point for acute and delayed.





TRANSFUSION REACTIONS

- WHEN IN DOUBT STOP THE TRANSFUSION!!!!!
- ACUTE HEMOLYTIC TRANSFUSION REACTION (AHTR).***
- DELAYED HEMOLYTIC TRANSFUSION REACTIONS (DHTR).
- FEBRILE NON-HEMOLYTIC TRANSFUSION REACTION (FNHTR). MOST COMMON.
- ALLERGIC REACTION.
- TRANSFUSION-ASSOCIATED CIRCULATORY OVERLOAD (TACO).
- TRANSFUSION-ASSOCIATED LUNG INJURY (TRALI).
- https://www.aabb.org/docs/default-source/default-document-library/resources/aabb-quick-reference-guide-nhsn-hemovigilance-module.pdf?sfvrsn=30f1600b_4



ACUTE HEMOLYTIC TRANSFUSION REACTION (AHTR).***

- DON'T MISS THIS ONE.
- MOST COMMON REASON: CLERICAL ERROR/HUMAN ERROR; WE ARE OUR OWN WORST ENEMY.
- DEFINITION IS ITS NAME: RAPID DESTRUCTION OF RED BLOOD CELLS DURING, IMMEDIATELY AFTER, OR WITHIN 24 HOURS OF CESSATION OF TRANSFUSION. CLINICAL AND LABORATORY SIGNS OF HEMOLYSIS ARE PRESENT.
- ACCOMPANIED BY: BACK/FLANK PAIN, CHILLS/RIGORS, DIC, EPISTAXIS, FEVER, HEMATURIA, HYPOTENSION, OLIGURIA/ANURIA, PAIN AND/OR OOZING AT IV SITE.
- 1:76000 (1/1.8 M FATAL).
- 80% ALONE PRESENT WITH SIMPLY FEVER AND CHILLS; EARLY, EARLY EARLY (WITHIN THE FIRST 15 MINUTES—NURSING STAFF START THE TRANSFUSION SLOWLY AND BE COGNIZANT OF TEMP.

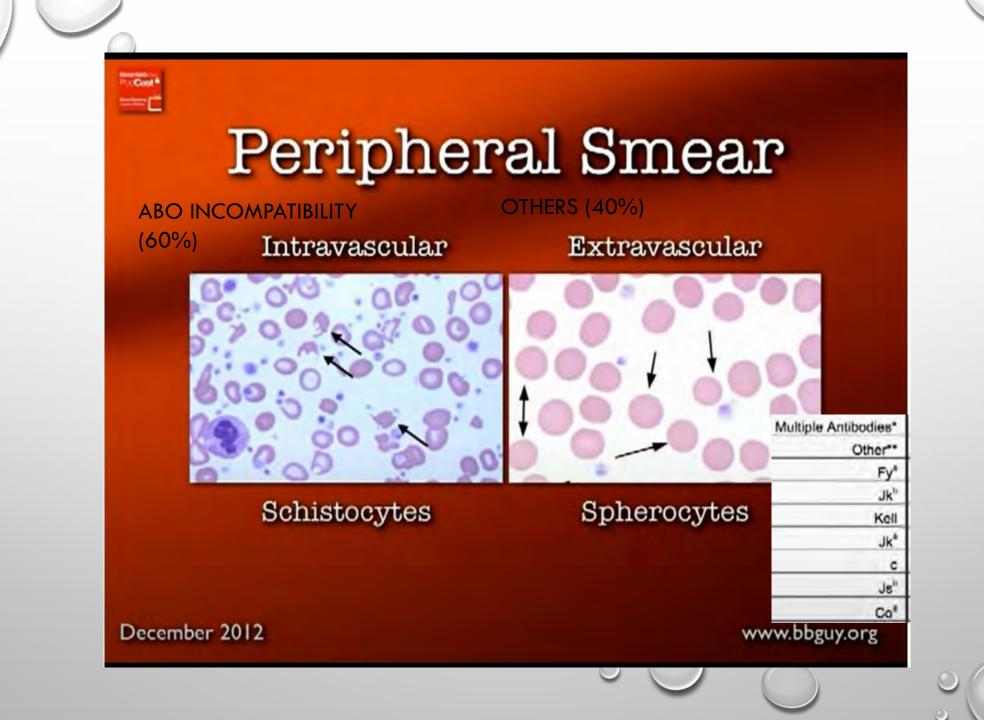


Acute Hemolytic Rxns

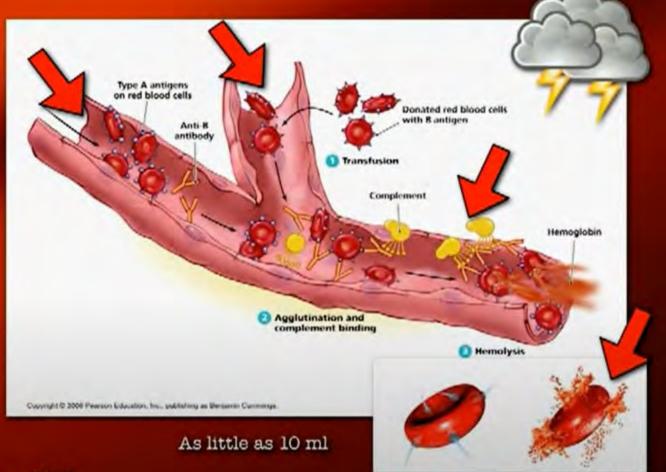
- Lab findings
 - Hemoglobinemia (gone in hrs), then -uria (gone in day)
 - + DAT (unless RBCs destroyed)
 - Hyperbilirubinemia
 - DIC labs (D-dimers, FSPs)

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CYTOKINE (PYROGENIC) STORM AND COAGULATION ACTIVATION AS WELL AS FIBRINOLYTIC CASCADES....CYTOKINES, CYTOKINES. "C'MON GUYS, IT'S ALL CYTOKINES (BALLBEARINGS) THESE DAYS!"

GEORGE CLOONEY MARK WAHLBERG AWOLFGANG PETERSENHUM PERFECT STORA



Acute Hemolytic Rxns

- Treatment
 - Hydration and diuresis
 - Maintain urine output
 - At least 1 mL/Kg/hr (furosemide)
 - Low-dose Dopamine (+/-)
 - Heparin for early DIC?
 - RBC exchange?

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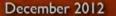
WHAT'S THE PHYSIOLOGIC ISSUE HERE? FLUID OVERLOAD ESPECIALLY IF PATIENTS BECOME ANURIC.

DO THE RIGHT THING. DON'T EVER ASK THAT A MISLABELED SPECIMEN, EVEN ON A HARD STICK EVER BE USED...REDRAW! "CHIP, I WILL COME AT YOU LIKE A SPIDER MONKEY"



Acute Hemolytic Rxns

- Prevention
 - Phlebotomy, labeling, processing, issue, and administration
 - Two separate types done by some
 - Technology (RFID, bar codes, etc)





- Standard Definition
 - NHLBI, Canadian Consensus Conf.
 - New ALI <6 hrs after transfusion
 - ALI: Hypoxemia (PaO₂/FiO₂ ≤300 or O₂ sat <90%), bilateral CXR infiltrates
 - No <u>other</u> known risk factors for pulmonary edema

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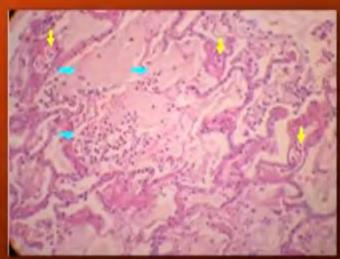
Pathophysiology is simply priming of neutrophils in the lung by usually anti-HLA and/or anti-neutrophil antibodies in the donor units. 30% of our body's neutrophils live in the lungs. Donor females pregnant and/or prior blood transfusion recipient donors.



TRALI







Images courtesy of Dr. Chris Silliman

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This is a transfusion reaction! Consider a pt with CHF or an old frail patient who comes in with chronic symptomatic anemia and you give he or she two or three units of pRBCs for an H/H of 6/18. What's gonna happen? Young and old; Renal failure.



TACO

- Acute CHF due to transfusion
 - Typical CHF physical exam
 - Dyspnea, orthopnea, rales, hypoxia
 - Systolic HTN (widened pulse pressure), JVD, headache
 - Afebrile usually

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Care Guidance (1)

The patient has a last measured hemoglobin result of ≥7 g/dL (or hematocrit ≥21%), or has had neither measured within the past 24 hours. In hospitalized, hemodynamically stable patients, a transfusion trigger of hemoglobin <7g/dL or hematocrit <21% decreases transfusion requirements and reduces adverse outcomes. If transfusion is required, single unit transfusion and clinical re-evaluation is recommended.





TACO VS. TRAIL (DIFFICULT)

TRANSFUSION ASSOCIATED CIRCULATORY OVERLOAD (TACO)

- AT RISK: CHF, VERY YOUNG/VERY OLD, RF, CHRONIC COMPENSATED ANEMIAS. "WHY DO 2 WHEN 1 WILL DO?" TRANSFUSE SLOW.
- NO FEVER; USUALLY.
- HYPERTENSION.
- ELEVATED BNP.
- DYSPNEA OFTEN OVER THE LENGTH OF THE TRANSFUSIONS.
- DIURETIC RESPONSIVE.
- TX: LIKE CHF; SLOW OR SPLIT TRANSFUSION.

TRANSFUSION-RELATED ACUTE LUNG INJURY (TRALI)

- FEVER.
- HYPOTENSION.
- ACUTE DYSPNEA.**
- DIURETIC UNRESPONSIVE.



TRALI

- Proving diagnosis usually difficult and time-consuming
 - Antibodies, ruling out other stuff
- Wet CXR, low O2 sat, no failure signs
 - BNP levels may help
- Respiratory support
- Mortality 5-25%; 80% rapid recovery

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TRALI

- Prevention
 - Strategies:
 - All/predom. male plasma/PLTs
 - Male/nullip female plasma/PLTs
 - Screening female PLT donors
 - AABB mandates
 - Defer implicated donors if Abs found

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Mild Allergic Reactions

- 1-3% incidence
- Urticaria (hives); local typical
 - Angioedema
- Type I hypersensitivity
- Benadryl prevents (+/-) and treats
- May restart transfusion
- Upper or lower resp sx = "moderate"

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Severe Allergic RXNs

- Definitions:
 - Anaphylactic: Shock, hypotension
 - Anaphylactoid:
 - Some use for milder anaphylactic
 - Respiratory symptoms with hypotension and GI complaints
 - ACEi-related reactions
 - Really loose distinction

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Severe Allergic RXNs

- Classic: IgA deficiency
 - Generally in those with severely decreased/absent IgA (<0.05 mg/dl)
- IgE-type anti-IgA is difficult to detect (anti-IgA in labs is IgG version)
- VERY few with RXN have anti-IgA





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Consider testing for IgA and/or haptoglobin deficiency and if severely low than do anti-lgA testing. VERY RARE. Treat like any other severe (anaphylactic) reaction— Epinephrine, yada yada yada. PREVENTION: IgA deficient products or washed RBCs; autologous donations.



DDX

- Acute HTR
- Septic transfusion reaction
- Acute hypotensive reaction (handout)
- Coincidental anaphylactic RXN
- Pulmonary embolus, acute MI, others
- NOTE: Lack of fever and skin findings may be VERY helpful





DELAYED HEMOLYTIC REACTIONS



Delayed Hemolytic Rxns

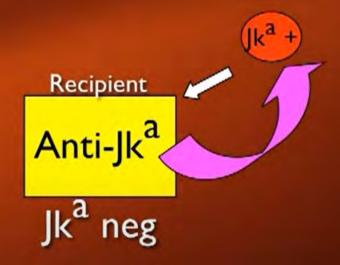
- Hemolysis occurring >24 hrs to <28 days after transfusion (per CDC)
- Anamnestic response is typical
 - Previously formed but currently undetectable antibody
 - Kidd, Duffy, Kell most common
- Primary rarely

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Anamnestic Response





Delayed Hemolytic Rxns

- Extravascular (except with Kidd)
- Signs/symptoms
 - Often none
 - Fever/anemia of unknown origin
 - Unexplained jaundice and/or scleral icterus



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Delayed Hemolytic Rxns

- Lab
 - Icteric serum
 - + DAT ("mf")
 - Anemia
 - Newly positive antibody screen
 - Spherocytes
 - LDL, bili, hapto

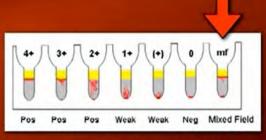


Image source: www.mirrscitech.co.kr

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TA-GVHD; NOTE THE ABSENCE OF SS DZ.



Indications



- Immunosuppression
 - T-cell defects (including drugs)
 - Stem cell/marrow transplants
 - Aplastic anemia
- Intrauterine/preemie transfusions
- Heme malignancies (esp HD)
- Granulocytes
- 1st-degree relatives or HLA-matched

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WHOLE BLOOD (O+ LOW TITER)

- I THINK WHOLE BLOOD IS GREAT IN PRE-HOSPITAL TRAUMA SITUATIONS.
- HOWEVER, THE VERDICT IS STILL NOT OUT ON ITS RISK/BENEFIT, HOWEVER, I BELIEVE THAT IT WILL BE THE FUTURE OF PRE-HOSPITAL TRANSFUSION PRACTICE.
- PHYSIOLOGICALLY, IT HAS BEEN ESTABLISHED THAT THESE WB UNITS HAVE A DEFINITE COAGULATIVE BENEFIT (STICKY PLATELETS).
- FROM A COST/BENEFIT PERSPECTIVE A LEVEL III-IV TRAUMA CENTER WOULD BURN THRU AN INORDINATE AMOUNT OF WB (COST CAN BE ANYWHERE FROM \$200-300 OR HIGHER DEPENDING ON GEOGRAPHIC REGION)
- LASTLY, WHOLE BLOOD (O+LOW TITER)...CAN ANYONE THINK OF WHY WHEN COMPONENT THEREAPY IS AVAILABLE WHY THIS WOULD BE A BAD IDEA IN A FEMALE OF CHILD BEARING AGE?

QUOTE FROM OCTOBER 2005

"THEY ONLY DO WHOLE BLOOD DRIVES WHEN ABSOLUTELY NECESSARY-THEY CAN TEST USING IMMUNOASSAY-HCV, HIV, HBV, RPR. THESE ARE QUICK SCREENING TESTS. THEY HAVE LIMITED CAPABILITY TO DETERMINE THE SPECIFICITY OF A POSITIVE DAT, BUT WE DO HAVE SOME ANTIBODY SCREENING CAPABILITY. THE LAST PATHOLOGIST GAVE THEM THE OK TO KEEP PLATELETS 7 DAYS, AS LONG AS THE CULTURES ARE NEGATIVE-THEY ARE ALL CULTURED. THERE ARE NUMEROUS THINGS/ISSUES FROM A BLOOD BANK PERSPECTIVE THAT YOU WILL SEE HERE THAT WAS ALIEN TO ME AT FIRST: WHOLE BLOOD DRIVES, USE OF THE TEG, ETC."

Whole blood transfusion versus component therapy in adult trauma patients with acute major haemorrhage

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ABSTRACT

Objective In the era of damage control resuscitation of trauma patients with acute major haemorrhage, transfusion practice has evolved to blood component (component therapy) administered in a ratio that closely approximates whole blood (WB). However, there is a paucity of evidence supporting the optimal transfusion strategy in these patients. The primary objective was therefore to establish if there is an improvement in survival at 30 days with the use of WB transfusion compared with blood component therapy in adult trauma patients with acute major haemorrhage.

Methodology A systematic literature search was performed on 15 December 2019 to identify studies comparing WB transfusion with component therapy in adult trauma patients and mortality at 30 days. Studies which did not report mortality were excluded. Methodological quality of included studies was interpreted using the Cochrane risk of bias tool, and rated using the Grading of Recommendations Assessment, Development and Evaluation approach. Results Search of the databases identified 1885 records, and six studies met the inclusion criteria involving 3255 patients. Of the three studies reporting 30-day mortality (one randomised controlled trial (moderate evidence) and two retrospective (low and very low evidence, respectively)), only one study demonstrated a statistically significant difference between WB and component therapy, and two found no statistical difference. Two retrospective studies reporting in-hospital

Key messages

What is already known on this subject

- Transfusion practice in trauma has evolved to administration of blood component therapy (red cells, plasma and platelets) in a ratio that closely approximates whole blood (WB).
- However, it has not been determined if WB versus component therapy is superior.

What this study adds

- In this systematic review, we found six studies directly addressing WB versus component therapy.
- Overall level of evidence was very low to moderate with only one randomised controlled trial.
- No studies reported worse survival with WB, however, there is insufficient evidence to support or reject the use of WB transfusion compared with component therapy for adult trauma patients with acute major haemorrhage.
- Larger prospective, randomised or adaptive trials are required to better understand if WB improves survival in adult trauma patients with acute major haemorrhage compared with component therapy.

Conclusion Recognising the limitations of this systematic review relating to the poor-quality evidence and limited number of included trials, it does not provide evidence to support or reject use of WB transfusion compared with component therapy for adult trauma patients with acute major haemorrhage.

PROSPERO registration number CRD42019131406.

erse events between groups, one study (n=354)incidence of AKI in the WB group compared therapy.30 Another study (n=369) found a of ARDS in the FWB group compared with leither study was designed to address these primary outcome, and therefore may not be onstrate an accurate difference. Both studies duced fresh WB. Both AKI and ARDS may be plex immunological mechanisms. A simplified des inflammation caused by the transfusion of d cells.30 White blood cells cause inflammation crovascular damage to the endothelium. In the ause vascular leakage into the alveolar space, onary oedema and ARDS. Of note, most WB in nd middle-income world is leucoreduced due ous diseases such as variant Creutzfeldt-Jakob

TOWAR Study Aims to Improve Mortality Rates for Trauma Patients



6/22/2021

PITTSBURGH - Researchers at the <u>University of Pittsburgh School of Medicine</u> are leading the Type O Whole blood and assessment of AGE during prehospital Resuscitation (TOWAR) Trial in an effort to better understand the intricacies of giving whole blood—blood that is not separated into its components— to trauma patients while they are in an ambulance on the way to the hospital.

"When someone donates blood, typically it gets separated it into parts—red cells, plasma, platelets—for storage and the parts can be used individually. But when someone is bleeding, they're losing all of these parts. Trauma research has shown that if you put all of these parts back together again it can be beneficial to trauma patients or patients at risk of



For Journalists

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Want to Make an Appointment or Need Patient Information? Contact UPMC at 1-800-533-8762.

Go to <u>Find a Doctor</u> to search for a UPMC doctor.

bleeding," explained Dr. Jason Sperry, a professor of surgery at Pitt and UPMC trauma surgeon.

PPOWER

Pragmatic, Prehospital group O,
Whole blood Early Resuscitation trial





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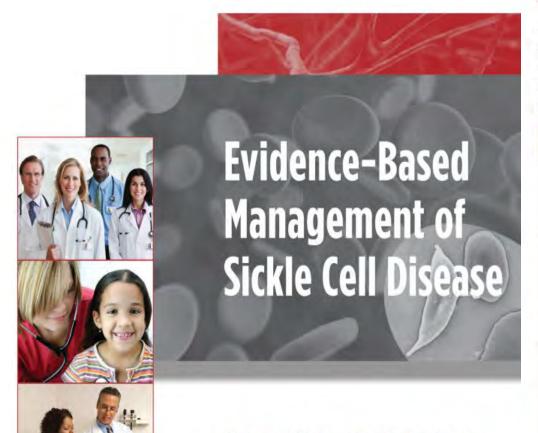


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Regional Whole Blood Program Low Titer O Positive Whole Blood (LTOWB)

STRAC is the recipient of a \$150,000 grant from the San Antonio Medical Foundation. This inter-institutional collaboration with the South Texas Blood and Tissue Center, UT Health San Antonio, University Health Systems (UHS), and the US Army Institute of Surgical Research / San Antonio Military Medical Center was formed to study and address the optimizing care of seriously injured or ill patients in STRAC region. This care need is met through the development of a cold stored whole blood product and implement transfusion of cold stored whole blood in the prehospital setting for helicopters emergency medical services. Funding from the Remote Trauma Outcomes Research Network (through the Department of Defense) allowed for an expansion to ground emergency medical services.

TRANSFUSION OF THE SICKLE CELL PATIENT



Expert Panel Report, 2014

Summary of the Evidence

An adequate systematic review of the literature with fair sensitivity and specificity for all studies indexed by SCD terms and the symptom of acute anemia was not feasible. A large and nonspecific return of studies with significant heterogeneity, high miss rate, and low-quality evidence (lack of comparative studies) was anticipated. No systematic evidence review was conducted, and the panel used a consensus process to develop a proposed strategy for triaging and promptly managing acute anemia.

Recommendations

- During all acute illnesses in people with SCD, obtain a CBC and reticulocyte count, repeat daily in all hospitalized patients, and compare the results with the patient's prior measurements.
 (Consensus-Panel Expertise)
- Assess people with SCD whose hemoglobin concentration is 2 g/dL or more below their baseline (or less than 6 g/dL when the baseline is unknown) for acute splenic sequestration, an aplastic episode, a delayed hemolytic transfusion reaction, ACS, and infection.

(Consensus-Panel Expertise)

- Use simple transfusion in people with SCD and acute anemia whose symptoms are due to anemia. (Consensus-Panel Expertise)
- Perform a CBC and reticulocyte count promptly and again 7 to 10 days later in siblings and others with SCD who are exposed to a person with an aplastic episode.

(Consensus-Panel Expertise)

 Manage aplastic events with immediate red blood cell transfusion aimed at restoring the hemoglobin to a safe (not necessarily baseline) value. Isolation of hospitalized patients (droplet precautions) is required to prevent spread of the parvovirus B19 to pregnant women and others with SCD or compromised immunity. (Consensus—Panel Expertise)



American Society of Hematology 2020 guidelines for sickle cell disease: transfusion support

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> Background: Red cell transfusions remain a mainstay of therapy for patients with sickle cell disease (SCD), but pose significant clinical challenges. Guidance for specific indications and administration of transfusion, as well as screening, prevention, and management of alloimmunization, delayed hemolytic transfusion reactions (DHTRs), and iron overload may improve outcomes.

> Objective: Our objective was to develop evidence-based guidelines to support patients, clinicians, and other healthcare professionals in their decisions about transfusion support for SCD and the management of transfusion-related complications.

> Methods: The American Society of Hematology formed a multidisciplinary panel that was balanced to minimize bias from conflicts of interest and that included a patient representative. The panel prioritized clinical questions and outcomes. The Mayo Clinic Evidence-Based Practice Research Program supported the guideline development process. The Grading of Recommendations Assessment, Development and Evaluation (GRADE) approach was used to form recommendations, which were subject to public comment.

> Results: The panel developed 10 recommendations focused on red cell antigen typing and matching, indications, and mode of administration (simple vs red cell exchange), as well as screening, prevention, and management of alloimmunization, DHTRs, and iron overload.

> Conclusions: The majority of panel recommendations were conditional due to the paucity of direct, highcertainty evidence for outcomes of interest. Research priorities were identified, including prospective studies to understand the role of serologic vs genotypic red cell matching, the mechanism of HTRs resulting from specific alloantigens to inform therapy, the role and timing of regular transfusions during pregnancy for women, and the optimal treatment of transfusional iron overload in SCD.



Sickle cell disease: when and how to transfuse

Jo Howard

Guy's and St Thomas' NHS Foundation Trust, London, United Kingdom; and King's College London, United Kingdom

Blood transfusion remains an important therapeutic intervention in patients with sickle cell disease (SCD), aiming to both increase the oxygen carrying capacity of blood and to reduce the complications of vaso-occlusion. Simple, manual exchange and automated exchange can be effective in reducing the acute and chronic complications of SCD, and the advantages and disadvantages of each methodology mean they all have a role in different situations. Evidence for the role of emergency transfusion in the management of the acute complications of SCD, including acute pain and acute chest syndrome, comes from observational data. Several important randomized controlled trials have shown the efficacy of transfusion in primary and secondary stroke prevention in patients with SCD but, outside these areas, clinical practice lacks a clear evidence base. Evidence for the role of long-term transfusion in the prevention of the non-neurologic chronic complications of SCD comes from analysis of secondary outcomes of these randomized trials and from observational data. In view of the paucity of data, the risks and benefits of transfusion should be fully discussed with patients/families before a long-term transfusion program is commenced. Evidence is only available for the role of preoperative transfusion. or for prophylactic transfusion through pregnancy in certain situations, and the role of transfusions outside these situations is discussed. Questions about when and how to transfuse in SCD remain and will need further randomized trials to provide answers.





Steps for Massive Transfusion Protocol (MTP)

ADULT

Administration of 210 FRBCS units within 24 hours Acute administration of 4-5 FRBC units within 1 hours, or Replacement of >50% of the total patient blood valume by blood products within 3 nours.

PEDIATRIC

Transfusion of one blood volume of RBC units in 24 hours

OBSTETRIC

At stage 3 of obstetrical blood loss for continued bleeding with QBL ≥1500 mL.

Any patient with abnormal vital signs/labs or oligues, or Sign/symptoms of hypovolemic shock OR



OTHE

Systolic blood pressure of \$ 90 mmHg.
Heart rate >120 beats per minute.
Evidence of penetrating torse trauma.
Patient who are expressed for require the replacement of a patient's TBV in less than 28 hours.
Assite edministration of 4-5 units within 1 hours, or Replacement of >50% of the total patient blood volume by blood products within 3 nours.

-2. Activate MTP Protocol--



After Emergency Release

Activation, MTP autivated by

the surgeon, physicians,

anesthesrologist and/or ED

obysician if MTP criteria met.





Activate MTP using RED phone that directly connects

to Blood Bank

AND

Clearly state to Blood Bank:

MTP is being initiated,

Patient Name

D.O.B. and

Patient Location

AND.

Notify RBX of MTP Activation and location







PB:

Overhead pages "Attention, Attention, Massive Transfusion Protocol Activation to (location)" three times.

Calls Vocera (7888), asks for group "Massive Transfusion", announcing the activation information,

Call House Supervisor cell phone to notify of Massive Transfusion Protocol activation and location.

If Birthing Center MTP, cad Anesthesiologist cellphone to notify of Massive Transfusion Protocolactivetion and location.

MTP Team Responds

Charge Nurse,
Illouse Supervisor,
Code Blue Team,
Attending Physician (if in
hospital)
Pastoral Care,
Anesthesiologist (if BC
location)

tab.

------3. Optimize intrinsic coagulation------3.







For OB Hemorrhage, follow OB Hemorrhage Management Plan Flow Chart (e.g., Yestotoxics (Pitocin, Methessius, Hemorrhage Catases) and Anti-Fibrinolytic (TXA)).

Avoid hypothermia, use blood warmer

------4. Arrival of cross matched blood------











Dantinge the process with 25% 45 MTP pack, set-

2rd MTP Pack (1) Cooler 5 FRBC, 6 FFP, 0 Platelet 3rd MTP Pack (1) Cooler 5 FRBC, 6 FFP, 1 Platelet 4rd MTP Pack (1) Cooler 5 FRBC, 6 FFP, 0 Platelet

For Crycoresip sate: Provider must indicate when to thow. Cryo comes in a pool of 5

Blood Bank calls RCD phone that 1" small cooler from 1" MTP Pack is ready for pick-up-

. . .

Runner brings patient sticker and picks-up (1)

Initial 1th MTP Pack Cooler (1)

Runner brings patient sticker and picks-up (1) cooler: <u>Second 1th MTP Pack Cooler (1)</u> -6 FFP & 1 Platelet

Blood Bank Calls RED phone that 2" MaxPlus cooler from 1" MTP Pack is

ready for pick-up

AND

r naskask 0 sassis

-5. Recheck & Monitor-----



-After every MTP Pack completion, re-draw labs; BNP, CBC, PT/INR, PTT, LDH. Blood Gases, and Fibrinogen
-Monitor while the transfusion is going: Temperature >35°C, pH: 7.35-7.45, Base Excess; <-8 Lactate, Hemoglobin 7-9g/gt, INR <1.5, PTT <42s, and Fibrinogen >100mg/gt,

I AM BEING CALLED ABOUT A MASSIVE TRANSFUSION, WHAT TO I DO?

- POSSIBLE MASSIVE
- MASSIVE TRANSFUSION REQUESTED.

POSSIBLE MASSIVE

- POSSIBLE MASSIVE COMES AS A RESULT OF CRITERIA GIVEN TO THE TECH TO CALL THE PATHOLOGIST.
- I.E. THE PATIENT HAS RECEIVED X AMOUNT OF RED BLOOD CELLS, PLASMA, PLATELETS CRYO.

POSSIBLE MASSIVE

- ASK FOR ALL THE BLOOD PRODUCTS THAT HAVE BEEN GIVEN.
- MAKE SURE THEY ARE "ON TRACK" WITH A RATIO OF 6 RBC, 6 FFP, 1 DOSE OF CRYO AND 1 PLATELET.
 - EX. IF THEY HAVE RECEIVED ALL RED BLOOD CELLS AND NO FFP, CRYO OR PLATELET, YOU MAY CALL TO SUGGEST THESE PRODUCTS.
- CALL TO THE ORDERING CLINICIAN, IF ORDERING LOOKS APPROPRIATE IS NOT REQUIRED.

- IF THE TECH CALLS TO NOTIFY THAT A MASSIVE TRANSFUSION HAS BEEN REQUESTED,
 INVOLVEMENT OF THE PATHOLOGIST IS REQUIRED.
- OBTAIN THE FOLLOWING INFORMATION FROM THE TECH:
 - NAME, DOB, AGE AND GENDER OF PATIENT.
 - INVENTORY OF O PRODUCTS (NEGATIVE AND POSITIVE) AND AB PLASMA.
 - WHETHER TESTING HAS BEEN PERFORMED ON NOT/BLOOD TYPE IF AVAILABLE.
 - LOCATION OF PATIENT, ORDERING PERSON IF KNOWN AND PHONE NUMBER.

- CALL THE ORDERING CLINICIAN OR MEMBER OF THE TEAM AND OFFER TO AID WITH COMMUNICATION TO THE BLOOD BANK.
- LET THEM KNOW THAT THE PRODUCTS WILL BE UNCROSSMATCHED O NEGATIVE (OR POSITIVE) RED BLOOD CELLS AND AB NEGATIVE PLASMA.
 - IF THEY CHOOSE NOT TO RECEIVE UNCROSSMATCHED PRODUCTS, LET THEM KNOW THAT THEY WILL
 NEED TO PUT IN AN ORDER FOR QUANTITY OF TYPES OF PRODUCTS DESIRED AND THAT A READ-BACK
 WILL BE REQUIRED.
 - IF THEY ASK HOW LONG IT WILL TAKE TO RECEIVE CROSSMATCHED PRODUCTS, THIS WILL DEPEND ON WHETHER A TYPE AND RE-TYPE IS COMPLETED.
 - IF COMPLETE, APPROXIMATELY 20 MINUTES.
 - IF INCOMPLETE, AT LEAST 1 HOUR FOR TESTING AND ISSUING BLOOD.

- REMIND THEM THAT THEY WILL CONTINUE TO RECEIVE BLOOD PRODUCTS IN THE FOLLOWING RATIO UNTIL THEY TELL UP TO STOP: 6 UNITS RBC, 6 UNITS FFP, 1 DOSE CRYO, 1 APHERESIS PLATELET.
- REMIND THEM TO LET THE BLOOD BANK KNOW WHEN THEY WOULD LIKE TO DISCONTINUE THE MASSIVE TRANSFUSION PROTOCOL.
- CALL THE BLOOD BANK TECH BACK REGARDLESS TO LET THEM KNOW OF THE RESULT OF THE CONVERSATION.
- ASSESS INVENTORY AND DETERMINE WHETHER ADDITIONAL PRODUCTS NEED TO BE ORDERED OR REQUESTED FROM ANOTHER HOSPITAL.

REMEMBER TO STAY CALM AND THAT YOU ARE NOT

ALONE.



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- <u>E.FILLMAN@CLINPATHASSOC.COM</u>
- GRMC BLOOD BANK: 830-401-7541
- OFFICE: 830-401-7899
- CELL: 210-823-4500
- HOME: 210-368-2011 (WE IGNORE THIS PHONE SINCE ONLY TELEMARKETERS CALL ANYMORE)

CRITICAL TRAUMA PATIENTS

- 19 JUN 2005 (FATHER'S DAY)
 - RIGHT LEG TRAUMATIC AMPUTATION FOLLOWING AN IED.
 - 203 BLOOD PRODUCTS.

https://thefallen.militarytimes.com/army-pfc-timothy-j-hines-

/980492

HONOR THE FALLEN

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Honoring those who fought and died in Operation Enduring Freedom, Operation Iraqi Freedom and Operation New Dawn



NATO KOSOVO FORCE

OPERATION ALLIES REFUGE

OPERATION ENDURING FREEDOM

OPERATION FREEDOM'S SENTINEL

OPERATION INHERENT RESOLVE

OPERATION IRAQI FREEDOM

OPERATION NEW DAWN

OPERATION OCTAVE SHIELD

OPERATION ODYSSEY LIGHTNING

OPERATION SPARTAN SHIELD

TASK FORCE SINAL

U.S. AFRICA COMMAND OPERATIONS





21, of Fairfield, Ohio; assigned to

Army Pfc. Timothy J. Hines Jr.
Died July 14, 2005 Serving During Operation Iraqi Freedom

21, of Fairfield, Ohio; assigned to the 64th Military Police Company, 720th Military Police Battalion, 89th Military Police Brigade, Fort Hood, Texas; died July 14 at the Walter Reed Army Medical Center, Washington, D.C., of wounds sustained on June 19 when an improvised explosive device detonated near his Humvee in Baghdad.

Fort Hood soldier dies of Iraq wounds

Associated Press

A Fort Hood soldier has died of injuries from a bomb explosion last month in Baghdad, the Pentagon confirmed Sunday.

Pfc. Tim Hines, 21, of Fairfield, Ohio, died Thursday night at Walter Reed Army Medical Center in Washington. His grandmother, Florence Thomas, said Friday that his wife, mother and other family members were with him.

An anonymous donor has given \$130,000 to pay private school tuition for Hines' two-yearold daughter and unborn son, The Cincinnati Enquirer reported in Monday's editions.

Family members said Hines was the gunner on a Humvee in a convoy when a bomb detonated on a Baghdad highway on Father's Day. He suffered kidney and tissue damage and internal bleeding, and his right leg was amputated in a Baghdad hospital. He was having emergency surgery Thursday when he died.

Hines enlisted in December 2003 and was stationed at Fort Hood, Texas, with the 720th Battalion, 89th Military Police Brigade, 64th Military Police Unit, before shipping out to Iraq in February.

Hundreds mourn soldier who died of Iraq injuries

SPRINGDALE, Ohio — If Pfc. Tim Hines Jr. could have held on for another few weeks, he would have been able see his second child.

The 21-year-old from Fairfield died last week at Walter Reed Army Medical Center in Washington, D.C., after battling injuries he received in a bomb explosion in Baghdad for almost a month.

More than 400 family members and friends gathered for his funeral Friday in this Cincinnati