

Updates in Transfusion²⁰²⁴



Dr. S. Raza, MD, FRCPC
Transfusion Fellow
April 5, 2024



Disclosures

None



Learning objectives

- To appreciate the use of GRADE Methodology for appraising literature in transfusion medicine
- To appreciate emerging evidence in transfusion over the last year or so

Outline

RBC
N=6

**Plasma
Deriv.**
N=5

Plt
N=1

RHIG
N=1

TXA
N=2



Outline

RBC
N=6

Plasma
Deriv.
N=5

Plt
N=1

RHIG
N=1

TXA
N=1

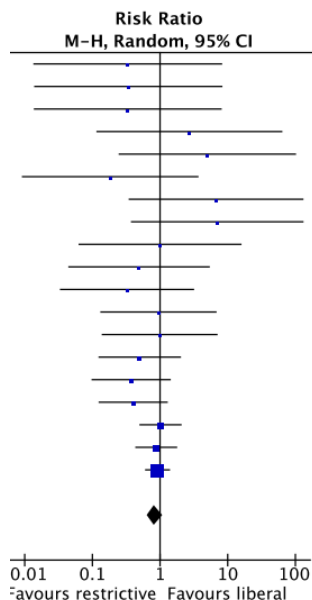
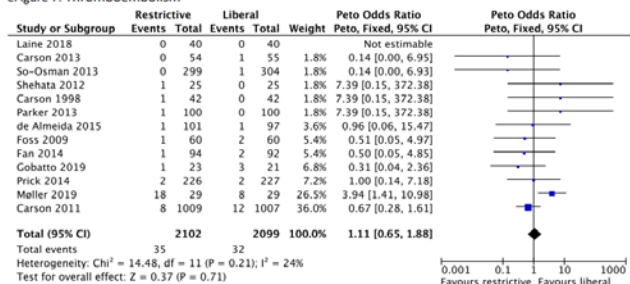


Red Blood Cell Transfusion 2023 AABB International Guidelines

Methods

1. Systematic Review

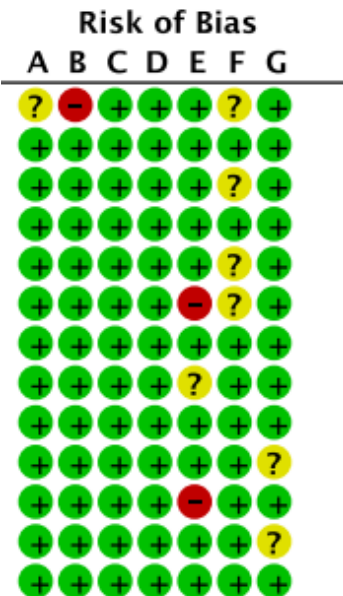
eFigure 7: Thromboembolism



2. Risk of Bias Assessment

Risk of bias legend

- (A) Random sequence generation (selection bias)
- (B) Allocation concealment (selection bias)
- (C) Blinding of participants and personnel (performance bias)
- (D) Blinding of outcome assessment (detection bias): Objective measures
- (E) Incomplete outcome data (attrition bias)
- (F) Selective reporting (reporting bias)
- (G) Other bias



3. GRADE Methodology

Grade of recommendation	I Strong recommendation to do	Ia Moderate recommendation to do	Ib Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence	Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate / Low level of evidence	Evidence from studies or systematic reviews with few important limitations	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence	Evidence from studies with serious flaws. Only expert opinion, or standards of care	Moderate recommendation based on very low level of evidence	Weak recommendation based on very low level of evidence	Recommendation based on very low level of evidence

Wording in recommendations:

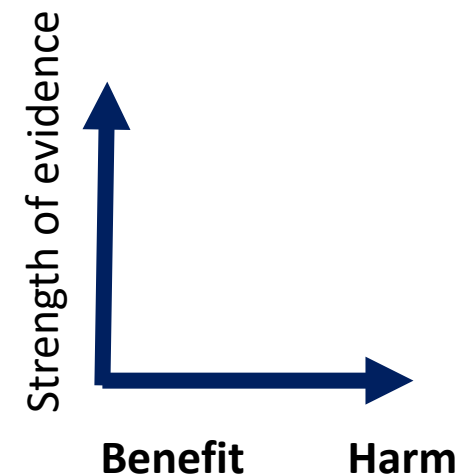
I Strong recommendation to do	Ia Moderate recommendation to do	Ib Weak recommendation to do	III Recommendation not to do
We recommend	We suggest	We might suggest	We do not recommend
We should	Is reasonable	Might be reasonable	Should not be performed
Is recommended	Is probably recommended	Might be considered	Is not useful
Is indicated	Can be useful	Usefulness is unknown	Is not beneficial
Is useful	Can be beneficial		Is not effective
Is beneficial	Can be effective		Is potentially harmful
Is effective			



A Detour for GRADE



Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)		Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care		Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	



Wording in recommendations:

We recommend	We suggest	We might suggest	We do not recommend
We should	Is reasonable	Might be reasonable	Should not be performed
Is recommended	Is probably recommended	Might be considered	Is not useful
Is indicated	Can be useful	Usefulness is unknown	Is not beneficial
Is useful	Can be beneficial		Is not effective
Is beneficial	Can be effective		Is potentially harmful
Is effective			



A Detour for GRADE

Recommendation 1	<p>For hospitalized adult patients who are hemodynamically stable, the panel recommends...less than 7 g/dL (strong recommendation, moderate certainty evidence).</p> <p>Clinicians may choose a threshold of ...7.5 g/dL for patients undergoing cardiac surgery ...8 g/dL for those undergoing orthopedic surgery or those with preexisting cardiovascular disease.</p>
Recommendation 2	<p>For hospitalized adult patients with hematologic and oncologic disorders...less than 7 g/dL (conditional recommendations, low certainty evidence).</p>
Recommendation 3	<p>For critically ill children and those at risk of critical illness who are hemodynamically stable and without a hemoglobinopathy, cyanotic cardiac condition, or severe hypoxemia...less than 7 g/dL (strong recommendation, moderate certainty evidence).</p>
Recommendation 4	<p>For hemodynamically stable children with congenital heart disease ...7 g/dL (biventricular repair) ...9 g/dL (single-ventricle palliation) ...7 to 9 g/dL (uncorrected congenital heart disease)</p> <p>(conditional recommendation, low certainty evidence).</p>

ADULT

ADULT

PEDIATRICS

PEDIATRICS



ORIGINAL ARTICLE

MINT

3506 Pts Randomized

Characteristic	All Patients (N=3504)	Restrictive Strategy (N=1749)	Liberal Strategy (N=1755)
NSTEMI	2848 (81.3)	1430 (81.8)	1418 (80.8)
Type 1	1460 (41.7)	730 (41.7)	730 (41.6)
Type 2	1955 (55.8)	967 (55.3)	988 (56.3)
White	2474 (70.6)	1229 (70.3)	1245 (70.9)
Black	440 (12.6)	217 (12.4)	223 (12.7)
Other	244 (7.0)	129 (7.4)	115 (6.6)
Missing	346 (9.9)	174 (9.9)	172 (9.8)

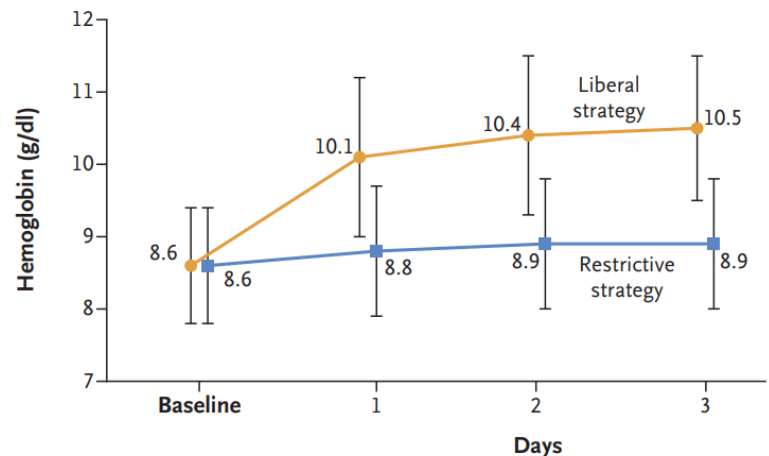
Restrictive or Liberal Transfusion Strategy in Myocardial Infarction and Anemia

Intervention

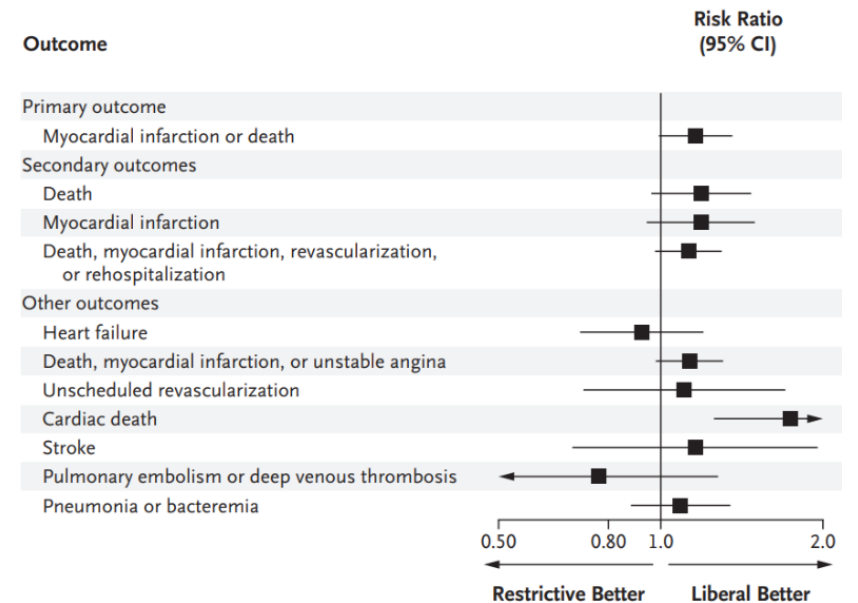
Restrictive strategy: Hb 80
 Liberal strategy: Hb 100

Open Label

Hemoglobin Level



Results



Cardiac death not adjudicated
 Not powered for harms



ORIGINAL ARTICLE

Restrictive or Liberal Transfusion Strategy in Myocardial Infarction and Anemia

► Use case-by-case, patient-centered judgement for patients with active myocardial ischemia, as some may benefit from a liberal transfusion strategy

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

Small-Volume Blood Collection Tubes to Reduce Transfusions in Intensive Care

The STRATUS Randomized Clinical Trial

Deborah M. Siegal, MD; Emilie P. Belley-Côté, MD, PhD; Shun Fu Lee, PhD; Stephen Hill, MD, PhD; Frédéric D'Aragnon, MD;

25 ICUs Randomized

Intervention

Results

No. clusters in sequence	Study period												
	1	2	3	4	5	6	7	8	9	10	11	12	13
2	n=138 99 (28)	n=148 77 (44)	n=136 88 (58)	n=151 76 (42)	n=146 73 (42)	n=162 81 (42)	n=171 85 (32)	n=167 84 (33)	n=187 94 (32)	n=56 28 (7)	n=161 81 (23)	n=179 90 (46)	n=214 107 (49)
2	n=160 80 (7)	n=134 67 (4)	n=124 62 (8)	n=145 73 (12)	n=135 68 (1)	n=151 76 (4)	n=174 87 (3)	n=170 85 (3)	n=159 80 (25)	n=50 25 (4)	n=150 75 (7)	n=173 87 (12)	n=143 72 (2)
2	n=97 49 (9)	n=116 58 (1)	n=92 46 (4)	n=84 42 (1)	n=92 46 (3)	n=101 51 (1)	n=90 45 (6)	n=90 45 (11)	n=95 48 (1)	n=28 14 (1)	n=102 51 (4)	n=111 56 (4)	n=98 49 (0)
2	n=229 115 (50)	n=228 114 (55)	n=216 108 (41)	n=188 94 (38)	n=147 74 (33)	n=193 97 (28)	n=188 94 (44)	n=140 70 (21)	n=225 113 (54)	n=69 35 (18)	n=218 109 (38)	n=181 92 (33)	n=202 101 (54)
2	n=140 70 (31)	n=143 71 (40)	n=155 78 (53)	n=129 65 (47)	n=97 49 (12)	n=112 56 (20)	n=125 78 (38)	n=140 70 (25)	n=154 77 (35)	n=32 26 (11)	n=129 65 (36)	n=151 77 (33)	n=130 65 (36)
2	n=127 64 (36)	n=137 69 (25)	n=140 70 (30)	n=126 63 (37)	n=160 80 (25)	n=151 76 (18)	n=127 64 (26)	n=162 81 (24)	n=148 74 (37)	n=50 25 (3)	n=130 65 (23)	n=135 68 (39)	n=160 80 (51)
2	n=113 57 (42)	n=114 57 (45)	n=122 61 (30)	n=106 53 (37)	n=103 52 (30)	n=98 49 (34)	n=102 51 (27)	n=84 42 (18)	n=106 53 (31)	n=31 16 (6)	n=105 53 (28)	n=133 67 (25)	n=124 62 (24)
2	n=133 67 (31)	n=140 70 (35)	n=126 63 (32)	n=122 61 (66)	n=140 70 (56)	n=138 69 (61)	n=154 77 (78)	n=148 74 (83)	n=134 67 (69)	n=43 22 (18)	n=135 68 (71)	n=143 72 (83)	n=147 74 (74)
2	n=58 29 (13)	n=116 58 (4)	n=98 49 (3)	n=102 51 (4)	n=92 46 (3)	n=117 59 (6)	n=98 49 (8)	n=104 52 (13)	n=102 51 (8)	n=16 8 (3)	n=73 37 (2)	n=79 40 (26)	n=87 44 (13)
2	n=173 87 (29)	n=176 88 (29)	n=166 83 (20)	n=157 79 (32)	n=173 87 (40)	n=173 87 (3)	n=166 83 (4)	n=162 81 (24)	n=171 86 (18)	n=64 32 (13)	n=148 74 (20)	n=185 93 (12)	n=191 96 (25)
2	n=178 89 (11)	n=159 80 (5)	n=174 87 (8)	n=156 78 (16)	n=159 80 (13)	n=180 90 (11)	n=173 87 (12)	n=151 76 (16)	n=171 86 (15)	n=56 28 (3)	n=159 80 (2)	n=130 65 (17)	n=133 67 (19)
3	n=147 49 (21)	n=170 57 (24)	n=148 49 (28)	n=144 48 (27)	n=149 50 (11)	n=127 42 (34)	n=175 58 (24)	n=158 53 (25)	n=163 54 (29)	n=51 17 (7)	n=146 49 (22)	n=156 52 (20)	n=149 50 (14)

Cluster under control condition Cluster under intervention condition

Switch to pediatric tubes

Standard-volume (4.0-6.0 mL)
Small-volume (1.8-3.5 mL)

In a prespecified secondary analysis (n = 27 411 patients), RBC units per patient per ICU saw an absolute reduction

9.84 RBC/100 patients



Red Cells

JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Small-Volume Blood Collection Tubes to Reduce Transfusions in Intensive Care The STRATUS Randomized Clinical Trial

Deborah M. Siegal, MD; Emilie P. Belley-Côté, MD, PhD; Shun Fu Lee, PhD; Stephen Hill, MD, PhD; Frédéric D'Aragnon, MD;

► Use small-volume blood collection tubes. No brainer!

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion





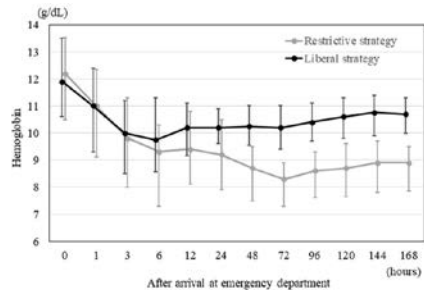
The Restrictive Red Blood Cell Transfusion Strategy for Critically Injured Patients (RESTRIC) trial: a cluster-randomized, crossover, non-inferiority multicenter trial of restrictive transfusion in trauma

Mineji Hayakawa^{1*}, Takashi Tagami^{2,3}, Daisuke Kudo⁴, Kota Ono⁵, Makoto Aoki⁶, Akira Endo⁷, Tetsuya Yumoto⁸, Yosuke Matsumura⁹, Shiho Irino¹⁰, Kazuhiko Sekine¹⁰, Noritaka Ushio¹¹, Takayuki Ogura¹², Sho Nachi¹³, Yuhei Irie¹⁴, Katsura Hayakawa¹⁵, Yusuke Ito¹⁶, Yuko Okishio¹⁷, Tomohiro Muroi¹⁸, Yoshinori Kosaki⁸, Kaori Ito¹⁹, Keita Nakatsutsumi²⁰, Yutaka Kondo²¹, Taichiro Ueda²², Hiroshi Fukuma²³, Yuichi Saisaka²⁴, Naoki Tominaga²⁵, Takeo Kurita²⁶, Fumihiko Nakayama²⁷, Tomotaka Shibata²⁸ and Shigeki Kushimoto⁴

RESTRIC Trial – N=511

Randomized to one of:

- Restrictive: 70-90 g/L
- Liberal: 10–12 g/L



Inclusion Criteria:

Adults with trauma
Patients who had:

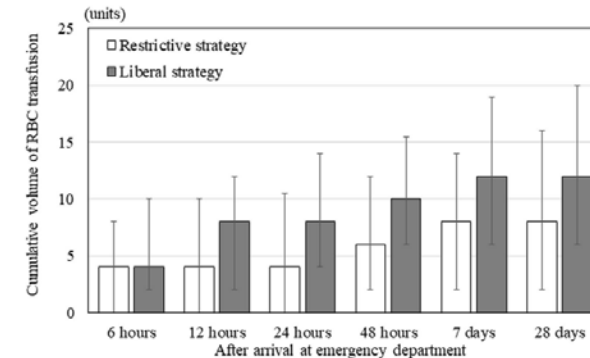
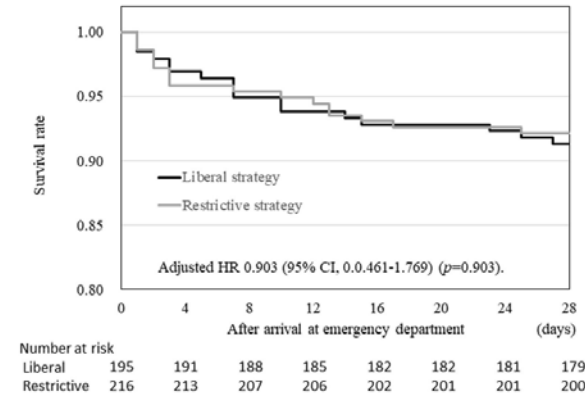
- Severe (or suspected) bleeding with shock
- Potential for severe bleeding postoperatively

Exclusion Criteria:

Cardiac arrest, palliation
Transfer
Severe burns, pregnancy
Chronic anemia

Results

- Restrictive RBC transfusion strategy did not demonstrate non-inferiority to liberal
- 28-day survival rate and survival time were similar (ICU free days, renal failure, MI...)
- Restrictive arm had smaller RBC transfusion volumes

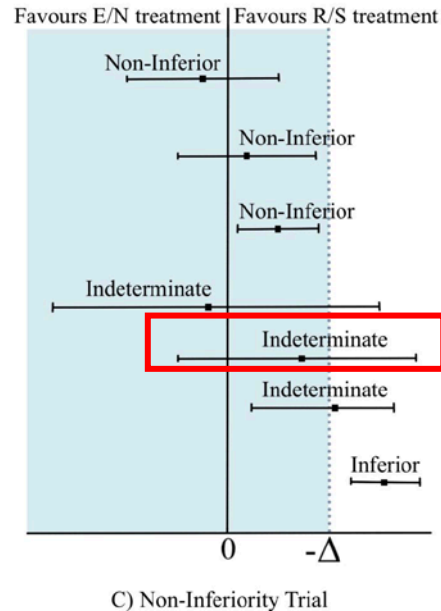


RESEARCH

Open Access



The Restrictive Red Blood Cell Transfusion Strategy for Critically Injured Patients (RESTRIC) trial: a cluster-randomized, crossover, non-inferiority multicenter trial of restrictive transfusion in trauma



► Prefer restrictive transfusion thresholds for critically injured trauma patients

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion



Orthopaedic Trauma and Anemia: Conservative versus Liberal Transfusion Strategy: A Prospective Randomized Study

Brian H. Mullis, MD,^a Leilani S. Mullis, MD,^a Laurence B. Kempton, MD,^a Walter Virkus, MD,^a James E. Slaven, MS,^a and Jennifer Bruggers, MD^b

Intervention: Restrictive threshold 55 g/L

Control: Liberal threshold 70 g/L

Outcome: Infection at 1 year follow-up

Patients (n=65): Ortho trauma past initial resuscitation phase, hemodynamically stable, aged 18-50, Hb<90 g/L



Lower transfusion rate after randomization – 46% vs. 94%

Lower infection rate – 6 vs. 25%, p=0.012

Longer length of stay – 11.5 vs. 9 days, p=0.04

No differences in any other outcome
*Small study, loss to follow-up



ORIGINAL ARTICLE

Orthopaedic Trauma and Anemia: Conservative versus Liberal Transfusion Strategy: A Prospective Randomized Study

Brian H. Mullis, MD,^a Leilani S. Mullis, MD,^a Laurence B. Kempton, MD,^a Walter Virkus, MD,^a James E. Slaven, MS,^a and Jennifer Bruggers, MD^b

► Prefer restrictive transfusion thresholds for young injured trauma patients

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

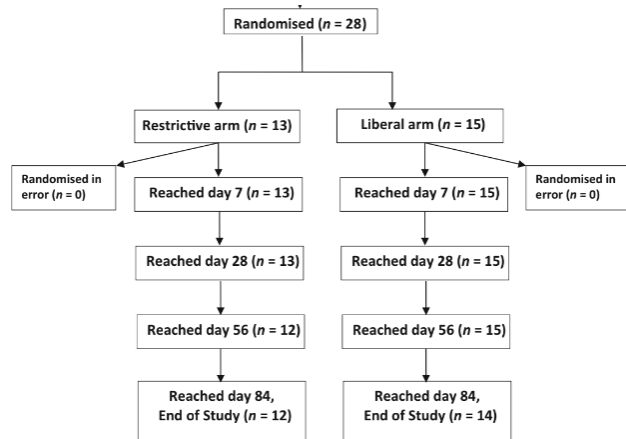


Red cell transfusion thresholds in outpatients with myelodysplastic syndromes: Results of a pilot randomized trial RBC-ENHANCE

28 Pts Randomized

Intervention

Results



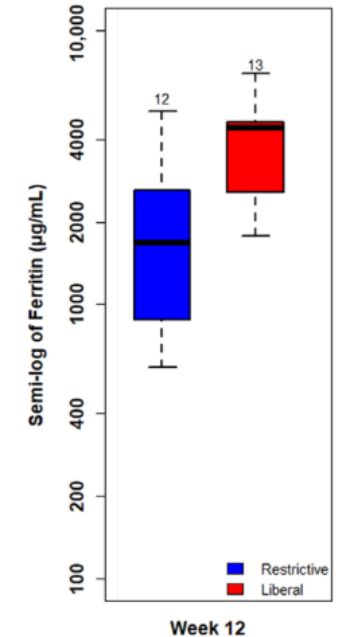
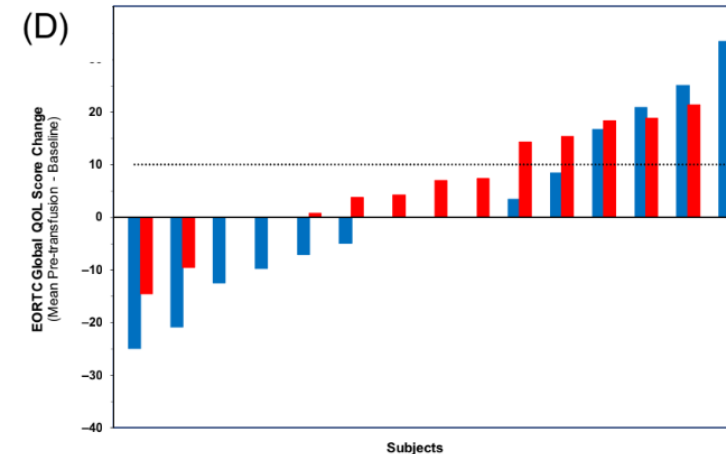
Randomized multi-center trial

Liberal (110-120 g/L) vs. Restrictive (85-105 g/L)

Outcomes

Mean Hb (worked?)
QOL scores (helped?)
Ferritin (harmed?)

Liberal Arm
Mean Hb (98 vs 86 g/L)
QOL scores (improved)
Ferritin (higher)



Red cell transfusion thresholds in outpatients with myelodysplastic syndromes: Results of a pilot randomized trial RBC-ENHANCE

Rena Buckstein¹ | Jeannie Callum² | Anca Prica³ | David Bowen⁴ | Richard A. Wells¹ | Brian Leber⁵ | Nancy Heddle⁶ | Lisa Chodirker¹ | Matthew Cheung¹ | Lee Mozessohn¹ | Karen Yee³ | Jennifer Gallagher¹ | Anne Parmentier¹ | Erin Jamula⁵ | Liying Zhang⁷ | Alex Mamedov¹ | Simon J. Stanworth^{8,9,10} | Yulia Lin¹¹

► Use a symptom-guided, case-by-case approach for transfusing patients with myelodysplastic syndrome

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion



Takeaways

New red cell transfusion guidelines are available!

Switch to small-volume tubes for blood collection

Hb thresholds <70 g/L may be safe for some young
MSK trauma patients

Higher Hb thresholds may be judiciously chosen for
symptomatic MDS patients on a case-by-case basis



Outline

RBC
N=6

Plt
N=1

Plasma
Deriv.
N=5

RHIG
N=1

TXA
N=1



Title

Red Cells

Platelets

Plasma Deriv.

RHIG

TXA

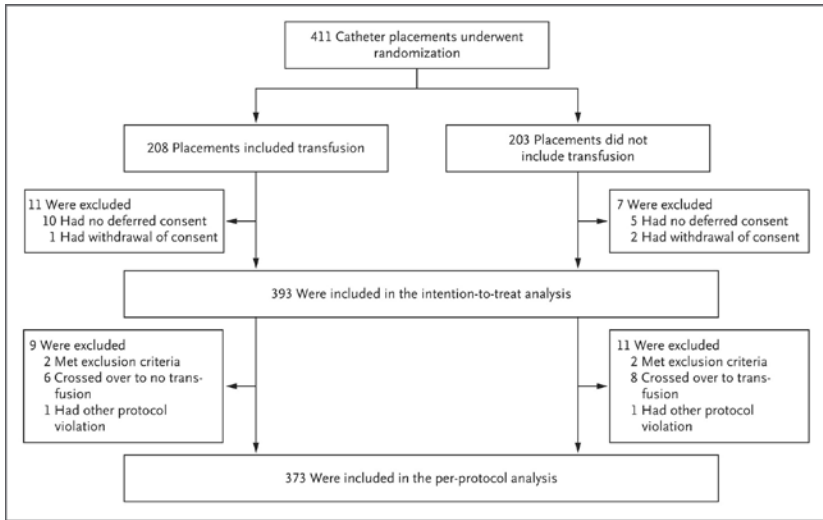
Questions

RESEARCH SUMMARY

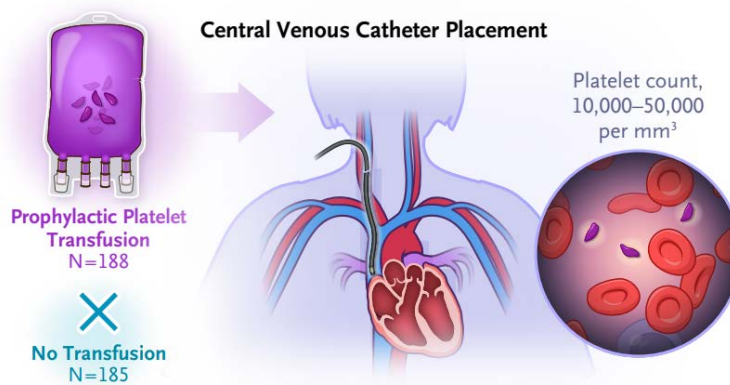
Platelet Transfusion before CVC Placement in Patients with Thrombocytopenia

van Baarle FLF et al. DOI: 10.1056/NEJMoa2214322

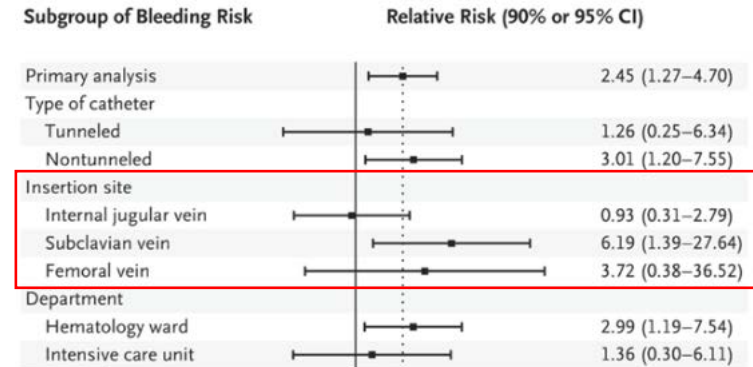
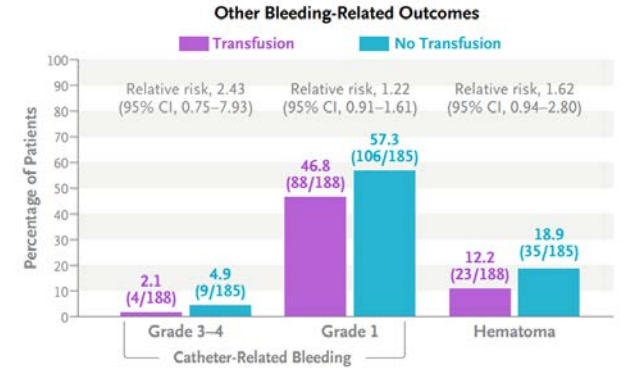
338 Patients Randomized



Intervention



Results



Platelets

RESEARCH SUMMARY

Platelet Transfusion before CVC Placement in Patients with Thrombocytopenia

van Baarle FLF et al. DOI: 10.1056/NEJMoa2214322

► Consider a platelet threshold of >50 for subclavian line placement in the ICU. For other lines, consider expert consultation to determine bleeding risk and platelet transfusion thresholds

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

Outline

RBC
N=6

Plt
N=1

Plasma
Deriv.
N=5

RHIG
N=1

TXA
N=1



Title

Red Cells

Platelets

Plasma Deriv.

RHIG

TXA

Questions

Use of Intravenous Albumin

A Guideline From the International Collaboration for Transfusion Medicine Guidelines

Jeannie Callum, MD; Nikolaos J. Skubas, MD; Aarti Bathla, MPharm, MPH; Homa Keshavarz, PhD; Edward G. Clark, MD; Bram Rochweg, MD; Dean Fergusson, PhD; Sesmu Arbous, MD; Seth R. Bauer, PharmD; Louise China, MD; Mark Fung, MD; Rachel Jug, MD; Michael Neill; Cary Paine, MD; Katerina Pavenski, MD; Prakesh S. Shah, MD; Susan Robinson, MD; Hua Shan, MD; Zbigniew M. Szczepiorkowski, MD, PhD; Thierry Thevenot, MD; Bovey Wu; Simon Stanworth, MD, PhD; and Nadine Shehata, MD; on behalf of the International Collaboration for Transfusion Medicine Guidelines Intravenous Albumin Guideline Group*

Group	Population	Indication	Stance	Evidence
Cirrhosis	Patients with cirrhosis and spontaneous bacterial peritonitis	Reduce mortality	For	Low
	Recommended dose: 1.5 g/kg on Day 1, 1.0 g/kg on Day 3			
Cirrhosis	Patients with cirrhosis and ascites undergoing large volume paracentesis (>5 liters)	Prevent paracentesis-induced circulatory dysfunction	For	Very low
	Recommended dose: 2-8 g/L of fluid removed			
Cirrhosis	Outpatients with cirrhosis and uncomplicated ascites despite diuretic	Reduce complications associated with cirrhosis	Against	Low
Cirrhosis	Hospitalized patients with decompensated cirrhosis with hypoalbuminemia (<30 g/L)	Reduce infection, kidney dysfunction or death	Against	Low
Cirrhosis	Patients with cirrhosis and extraperitoneal infections	Reduce mortality or kidney failure	Against	Low



Use of Intravenous Albumin

A Guideline From the International Collaboration for Transfusion Medicine Guidelines

*Jeannie Callum, MD; Nikolaos J. Skubas, MD; Aarti Bathla, MPharm, MPH; Homa Keshavarz, PhD; Edward G. Clark, MD; Bram Rochweg, MD; Dean Fergusson, PhD; Sesmu Arbous, MD; Seth R. Bauer, PharmD; Louise China, MD; Mark Fung, MD; Rachel Jug, MD; Michael Neill; Cary Paine, MD; Katerina Pavenski, MD; Prakesh S. Shah, MD; Susan Robinson, MD; Hua Shan, MD; Zbigniew M. Szczepiorkowski, MD, PhD; Thierry Thevenot, MD; Bovey Wu; Simon Stanworth, MD, PhD; and Nadine Shehata, MD; on behalf of the International Collaboration for Transfusion Medicine Guidelines Intravenous Albumin Guideline Group**

Group	Population	Indication	Stance	Evidence
ICU	Critically ill adults (excluding thermal injuries and ARDS)	First-line volume replacement or increase serum albumin levels	Against	Moderate
ICU	Critically ill adults	Removal of extravascular fluid with diuretics	Against	Very low
ICU	Critically ill adults with thermal injuries or ARDS	Volume replacement or increase serum albumin level	Against	Very low
Renal	Patients undergoing renal replacement therapy	Prevention or treatment of intradialytic hypotension or improving ultrafiltration	Against	Very low



Use of Intravenous Albumin

A Guideline From the International Collaboration for Transfusion Medicine Guidelines

Jeannie Callum, MD; Nikolaos J. Skubas, MD; Aarti Bathla, MPharm, MPH; Homa Keshavarz, PhD; Edward G. Clark, MD; Bram Rochweg, MD; Dean Fergusson, PhD; Sesmu Arbous, MD; Seth R. Bauer, PharmD; Louise China, MD; Mark Fung, MD; Rachel Jug, MD; Michael Neill; Cary Paine, MD; Katerina Pavenski, MD; Prakesh S. Shah, MD; Susan Robinson, MD; Hua Shan, MD; Zbigniew M. Szczepiorkowski, MD, PhD; Thierry Thevenot, MD; Bovey Wu; Simon Stanworth, MD, PhD; and Nadine Shehata, MD; on behalf of the International Collaboration for Transfusion Medicine Guidelines Intravenous Albumin Guideline Group*

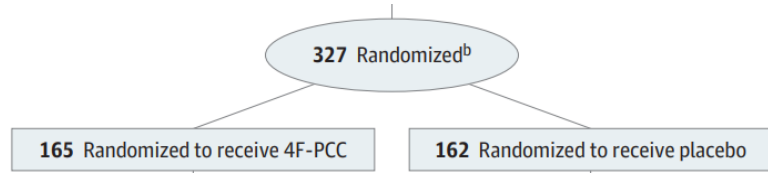
Group	Population	Indication	Stance	Evidence
Peds	Pediatric patients with infection and hypoperfusion	Reduce mortality	Against	Low
Neonate	Preterm neonates (<36 weeks) with low serum albumin levels and respiratory distress	Improve respiratory function	Against	Very low
Peds	Pediatric patients undergoing cardiovascular surgery	Priming the cardiovascular bypass circuit or volume replacement	Against	Very low
Neonate	Preterm neonates (<32 weeks or <1,500 g) with or without hypoperfusion	Volume replacement	Against	Very low



Efficacy and Safety of Early Administration of 4-Factor Prothrombin Complex Concentrate in Patients With Trauma at Risk of Massive Transfusion

The PROCOAG Randomized Clinical Trial

327 Pts Randomized



Characteristic	Median (IQR) [total No.]	
	4F-PCC (n = 164)	Placebo (n = 160)
Transfusion of ≥10 U of RBCs within the first 24 h	42 (26)	43 (28)
Fibrinogen concentrate treatment	141 (86)	129 (81)

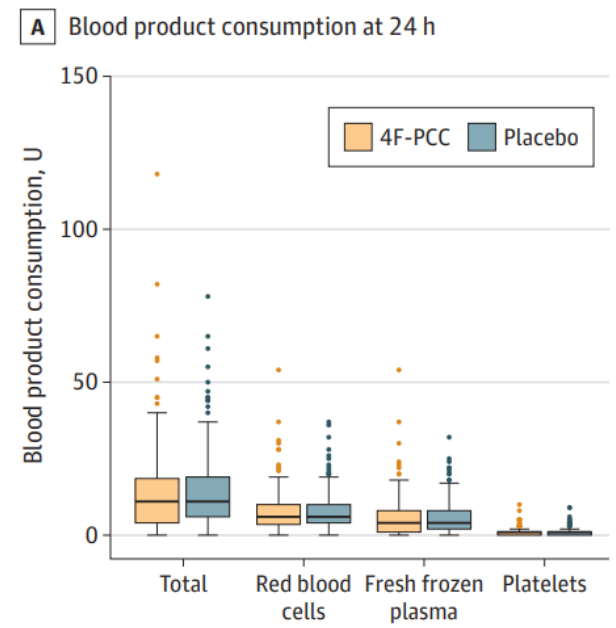
Intervention

Double-blind RCT

4F-PCC 25 IU of factor IX per kg (1 mL/kg)

Or 1 mL/kg of 0.9% saline solution

Results



Thrombosis:
 4F-PCC group, 56 patients (35%)
 Saline group, 37 patients (24%) in the placebo group [95% CI, 1%-21%]



JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Efficacy and Safety of Early Administration of 4-Factor Prothrombin Complex Concentrate in Patients With Trauma at Risk of Massive Transfusion

The PROCOAG Randomized Clinical Trial

► Do not give early empiric 4-Factor PCC for patients with trauma and massive hemorrhage unless specifically indicated

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

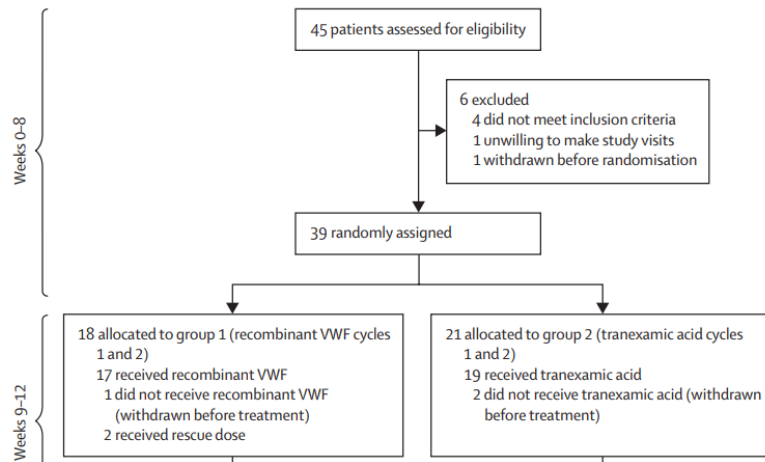


vWF Concentrate

Recombinant von Willebrand factor and tranexamic acid for heavy menstrual bleeding in patients with mild and moderate von Willebrand disease in the USA (VWDMin): a phase 3, open-label, randomised, crossover trial

Margaret V Ragni, Scott D Rothenberger, Robert Feldman, Danielle Nance, Andrew D Leavitt, Lynn Malec, Roshni Kulkarni, Robert Sidonio Jr,

45 Pts Randomized



Intervention

VWF, 40 IU/kg over 5–10 min on day 1

Oral TXA 1300 mg three times daily on days 1–5

Results

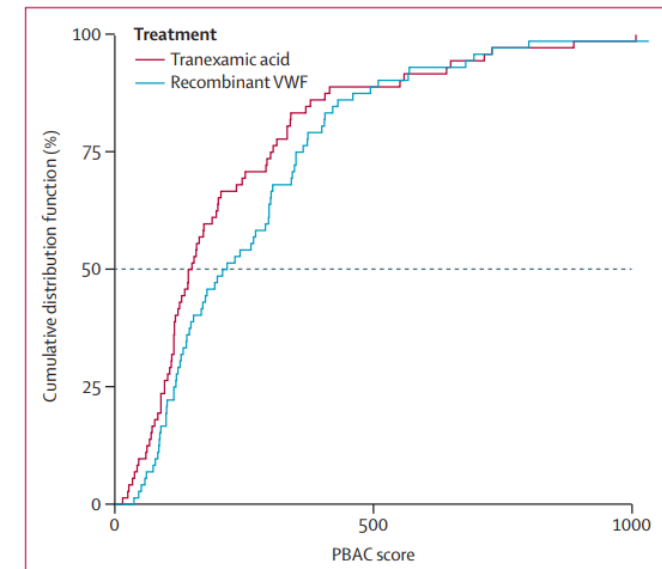


Figure 2: PBAC score for blood loss by treatment

Cumulative distribution function comparing PBAC score after tranexamic acid versus recombinant VWF. PBAC=pictorial blood assessment chart. Median PBAC score on tranexamic acid (ie, where the curve for tranexamic acid crosses 50% cumulative distribution function) is 146 (95% CI 117–199), while the median PBAC score on recombinant VWF is 213 (152–298; p=0.039).



vWF Concentrate

Recombinant von Willebrand factor and tranexamic acid for heavy menstrual bleeding in patients with mild and moderate von Willebrand disease in the USA (VWDMin): a phase 3, open-label, randomised, crossover trial

Margaret V Ragni, Scott D Rothenberger, Robert Feldman, Danielle Nance, Andrew D Leavitt, Lynn Malec, Roshni Kulkarni, Robert Sidonio Jr,

► Do not use vWF for heavy menstrual bleeding in all patients with mild-to-moderate vWD*
*trial on-going

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

Cryoprecipitate

JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Early and Empirical High-Dose Cryoprecipitate for Hemorrhage After Traumatic Injury

The CRYOSTAT-2 Randomized Clinical Trial

1604 Pts Randomized



1251 Men 330 Women

Patients 16 years or older with major trauma hemorrhage in the emergency department

Median age: 39 years

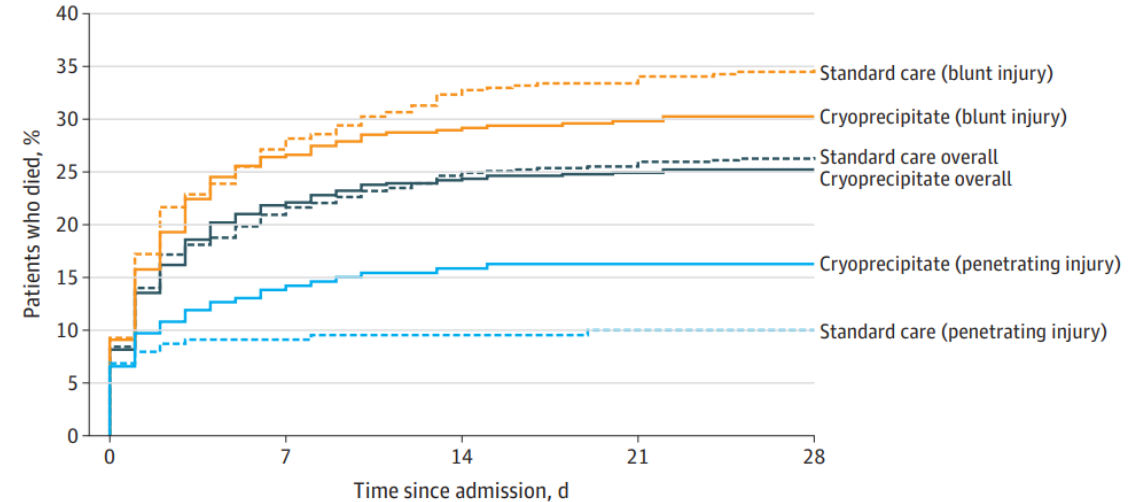
Intervention

Patients in both groups: MHP with a balanced RBC/FFP

Intervention group: 3 pools of cryoprecipitate (6-g fibrinogen equivalent) as early as possible, with the aim to start within 90 minutes of admission

vs **Standard of Care**

Results



Cryoprecipitate timing

CRYO ≤ 45m from adm	1.45 (0.91, 2.31)	0.12 ^b
CRYO 46-60m from adm	1.16 (0.78, 1.73)	0.46 ^b
CRYO 61-90m from adm	0.57 (0.38, 0.87)	0.01 ^b
CRYO > 90m from adm	1.00 (0.62, 1.60)	0.99 ^b

Cryoprecipitate

JAMA | Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Early and Empirical High-Dose Cryoprecipitate for Hemorrhage After Traumatic Injury The CRYOSTAT-2 Randomized Clinical Trial

► Do not give early empiric cryoprecipitate/fibrinogen for hemorrhage after traumatic injury (use fibrinogen levels to guide transfusion)

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

Journal Pre-proof

Use of Intravenous Albumin: A Guideline from the International Collaboration for Transfusion Medicine Guidelines.

Jeannie Callum, MD, Nikolaos J. Skubas, MD, Aarti Bathla, MPharm, MPH, Homa Keshavarz, PhD, Edward G. Clark, MD, Bram Rochweg, MD, Dean Fergusson, PhD, Sesmu Arbous, MD, Seth R. Bauer, PharmD, Louise China, MD, Mark Fung, MD, Rachel Jug, MD, Michael Neill, Cary Paine, MD, Katerina Pavenski, MD, Prakesh S. Shah, MD, Susan Robinson, MD, Hua Shan, MD, Zbigniew M. Szczepiorkowski, MD, PhD, Thierry Thevenot, MD, Bovey Wu, Simon Stanworth, MD, PhD, Nadine Shehata, MD, on behalf of the ICTMG Intravenous Albumin Guideline Group



	Population	Indication	Recommendation	Strength of Evidence
Cirrhosis	Patients with cirrhosis and spontaneous bacterial peritonitis Recommended dose: 1.5 g/kg on Day 1, 1.0 g/kg on Day 2	Reduce mortality	For	Low
Cirrhosis	Patients with cirrhosis and ascites undergoing large volume paracentesis (>5 liters) Recommended dose: 2-8 g/L of fluid removed	Prevent paracentesis-induced circulatory dysfunction	For	Very low
ICU	Critically ill adults (excluding thermal injuries and ARDS)	First-line volume replacement or increase serum albumin levels	Against	Moderate
CV Surg	Adult patients undergoing cardiovascular surgery	Priming the cardiovascular bypass circuit or volume replacement	Against	Moderate
Cirrhosis	Outpatients with cirrhosis and uncomplicated ascites despite diuretic therapy	Reduce complications associated with cirrhosis	Against	Low
Cirrhosis	Hospitalized patients with decompensated cirrhosis with hypoalbuminemia (<30 g/L)	Reduce infection, kidney dysfunction or death	Against	Low
Peds	Pediatric patients with infection and hypoperfusion	Reduce mortality	Against	Low
Cirrhosis	Patients with cirrhosis and extraperitoneal infections	Reduce mortality or kidney failure	Against	Low
Neonate	Preterm neonates (<36 weeks) with low serum albumin levels and respiratory distress	Improve respiratory function	Against	Very low
Peds	Pediatric patients undergoing cardiovascular surgery	Priming the cardiovascular bypass circuit or volume replacement	Against	Very low
Neonate	Preterm neonates (<32 weeks or <1,500 g) with or without hypoperfusion	Volume replacement	Against	Very low
ICU	Critically ill adults	Removal of extravascular fluid with diuretics	Against	Very low
ICU	Critically ill adults with thermal injuries or ARDS	Volume replacement or increase serum albumin level	Against	Very low
Renal	Patients undergoing renal replacement therapy	Prevention or treatment of intradialytic hypotension or improving ultrafiltration	Against	Very low



Journal Pre-proof

Use of Intravenous Albumin: A Guideline from the International Collaboration for Transfusion Medicine Guidelines.

Jeannie Callum, MD, Nikolaos J. Skubas, MD, Aarti Bathla, MPharm, MPH, Homa Keshavarz, PhD, Edward G. Clark, MD, Bram Rochweg, MD, Dean Fergusson, PhD, Sesmu Arbous, MD, Seth R. Bauer, PharmD, Louise China, MD, Mark Fung, MD, Rachel Jug, MD, Michael Neill, Cary Paine, MD, Katerina Pavenski, MD, Prakesh S. Shah, MD, Susan Robinson, MD, Hua Shan, MD, Zbigniew M. Szczepiorkowski, MD, PhD, Thierry Thevenot, MD, Bovey Wu, Simon Stanworth, MD, PhD, Nadine Shehata, MD, on behalf of the ICTMG Intravenous Albumin Guideline Group



Group	Population	Indication	Stance	Evidence
Cirrhosis	Patients with cirrhosis and spontaneous bacterial peritonitis	Reduce mortality	For	Low
	Recommended dose: 1.5 g/kg on Day 1, 1.0 g/kg on Day 3			
Cirrhosis	Patients with cirrhosis and ascites undergoing large volume paracentesis (>5 liters)	Prevent paracentesis-induced circulatory dysfunction	For	Very low
	Recommended dose: 2-8 g/L of fluid removed			
Cirrhosis	Outpatients with cirrhosis and uncomplicated ascites despite diuretic	Reduce complications associated with cirrhosis	Against	Low
Cirrhosis	Hospitalized patients with decompensated cirrhosis with hypoalbuminemia (<30 g/L)	Reduce infection, kidney dysfunction or death	Against	Low
Cirrhosis	Patients with cirrhosis and extraperitoneal infections	Reduce mortality or kidney failure	Against	Low



Journal Pre-proof

Use of Intravenous Albumin: A Guideline from the International Collaboration for Transfusion Medicine Guidelines.

Jeannie Callum, MD, Nikolaos J. Skubas, MD, Aarti Bathla, MPharm, MPH, Homa Keshavarz, PhD, Edward G. Clark, MD, Bram Rochweg, MD, Dean Fergusson, PhD, Sesmu Arbous, MD, Seth R. Bauer, PharmD, Louise China, MD, Mark Fung, MD, Rachel Jug, MD, Michael Neill, Cary Paine, MD, Katerina Pavenski, MD, Prakesh S. Shah, MD, Susan Robinson, MD, Hua Shan, MD, Zbigniew M. Szczepiorkowski, MD, PhD, Thierry Thevenot, MD, Bovey Wu, Simon Stanworth, MD, PhD, Nadine Shehata, MD, on behalf of the ICTMG Intravenous Albumin Guideline Group



Group	Population	Indication	Stance	Evidence
Peds	Pediatric patients with infection and hypoperfusion	Reduce mortality	Against	Low
Neonate	Preterm neonates (<36 weeks) with low serum albumin levels and respiratory distress	Improve respiratory function	Against	Very low
Peds	Pediatric patients undergoing cardiovascular surgery	Priming the cardiovascular bypass circuit or volume replacement	Against	Very low
Neonate	Preterm neonates (<32 weeks or <1,500 g) with or without hypoperfusion	Volume replacement	Against	Very low



Journal Pre-proof

Use of Intravenous Albumin: A Guideline from the International Collaboration for Transfusion Medicine Guidelines.

Jeannie Callum, MD, Nikolaos J. Skubas, MD, Aarti Bathla, MPharm, MPH, Homa Keshavarz, PhD, Edward G. Clark, MD, Bram Rochweg, MD, Dean Fergusson, PhD, Sesmu Arbous, MD, Seth R. Bauer, PharmD, Louise China, MD, Mark Fung, MD, Rachel Jug, MD, Michael Neill, Cary Paine, MD, Katerina Pavenski, MD, Prakesh S. Shah, MD, Susan Robinson, MD, Hua Shan, MD, Zbigniew M. Szczepiorkowski, MD, PhD, Thierry Thevenot, MD, Bovey Wu, Simon Stanworth, MD, PhD, Nadine Shehata, MD, on behalf of the ICTMG Intravenous Albumin Guideline Group



	Population	Indication	Recommendation	Strength of Evidence
Cirrhosis	Patients with cirrhosis and spontaneous bacterial peritonitis Recommended dose: 1.5 g/kg on Day 1, 1.0 g/kg on Day 2	Reduce mortality	For	Low
Cirrhosis	Patients with cirrhosis and ascites undergoing large volume paracentesis (>5 liters) Recommended dose: 2-8 g/L of fluid removed	Prevent paracentesis-induced circulatory dysfunction	For	Very low
ICU	Critically ill adults (excluding thermal injuries and ARDS)	First-line volume replacement or increase serum albumin levels	Against	Moderate
CV Surg	Adult patients undergoing cardiovascular surgery	Priming the cardiovascular bypass circuit or volume replacement	Against	Moderate
Cirrhosis	Outpatients with cirrhosis and uncomplicated ascites despite diuretic therapy	Reduce complications associated with cirrhosis	Against	Low
Cirrhosis	Hospitalized patients with decompensated cirrhosis with hypoalbuminemia (<30 g/L)	Reduce infection, kidney dysfunction or death	Against	Low
Peds	Pediatric patients with infection and hypoperfusion	Reduce mortality	Against	Low
Cirrhosis	Patients with cirrhosis and extraperitoneal infections	Reduce mortality or kidney failure	Against	Low
Neonate	Preterm neonates (<36 weeks) with low serum albumin levels and respiratory distress	Improve respiratory function	Against	Very low
Peds	Pediatric patients undergoing cardiovascular surgery	Priming the cardiovascular bypass circuit or volume replacement	Against	Very low
Neonate	Preterm neonates (<32 weeks or <1,500 g) with or without hypoperfusion	Volume replacement	Against	Very low
ICU	Critically ill adults	Removal of extravascular fluid with diuretics	Against	Very low
ICU	Critically ill adults with thermal injuries or ARDS	Volume replacement or increase serum albumin level	Against	Very low
Renal	Patients undergoing renal replacement therapy	Prevention or treatment of intradialytic hypotension or improving ultrafiltration	Against	Very low



Outline

RBC
N=6

Plt
N=1

Plasma
Deriv.
N=5

RHIG
N=1

TXA
N=1



Title

Red Cells

Platelets

Plasma Deriv.

RHIG

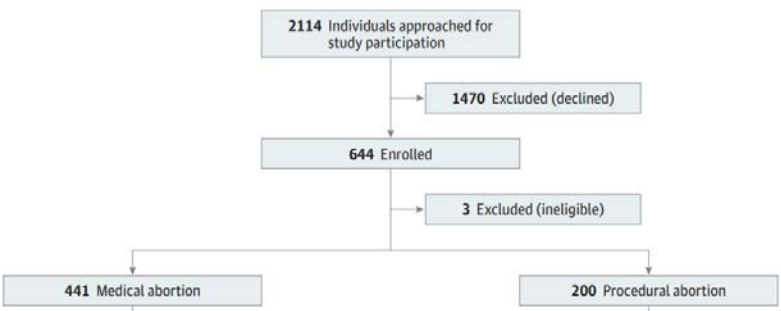
TXA

Questions

Induced Abortion and the Risk of Rh Sensitization

Is administration of Rh immunoglobulin necessary for individuals undergoing induced first-trimester abortion care?

506 Pts Randomized



All patients undergoing medication abortion at less than 12 week

Intervention

Exclusion criteria:
Sickle cell disease, β -thalassemia, hereditary persistence of fetal hemoglobin

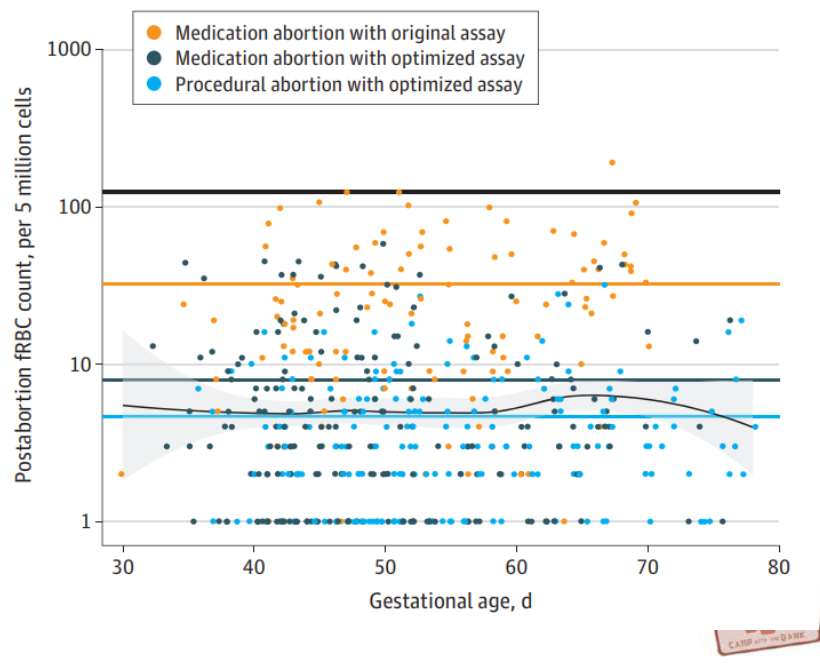
These patients had blood drawn before abortion and after abortion at 1, 2, 4, 8, 12, 24, 48, and 72 hour

The threshold for Rh sensitization used in the study was at least 0.1% fetal red blood cells* (fRBCs) in maternal circulation following induced abortion at less than 12 weeks' gestation

*measured using flow cytometry

Results

Figure 3. Fetal Red Blood Cell (fRBC) Count After Abortion by Gestational Age



JAMA | Original Investigation

Induced Abortion and the Risk of Rh Sensitization

► Have a case-by-case discussion with patients when offering RhD prophylaxis for induced abortion during the 1st trimester*
*evolving

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

Outline

RBC
N=6

Plt
N=1

Plasma
Deriv.
N=5

RHIG
N=1

TXA
N=2



Title

Red Cells

Platelets

Plasma Deriv.

RHIG

TXA

Questions

ORIGINAL ARTICLE

Prehospital Tranexamic Acid for Severe Trauma

1310 Pts Randomized Intervention

Results



Before admission: 1-g intravenous bolus dose within 3 hr after injury
After admission: 1-g infusion over 8 hr

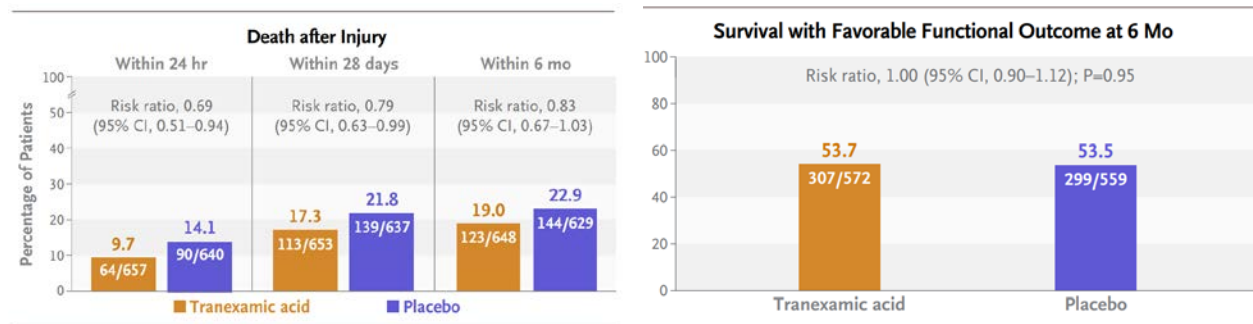
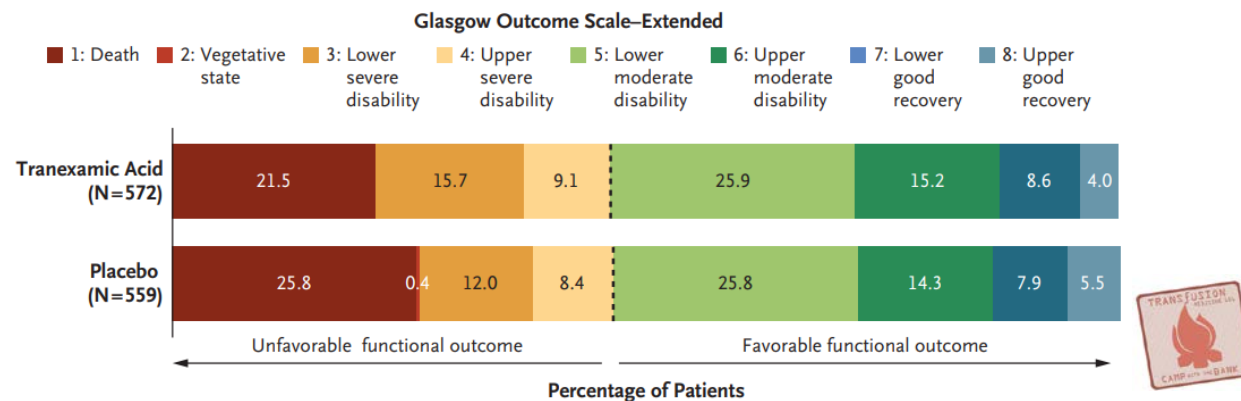


Table 1. Characteristics of the Patients at Baseline.*

Characteristic	Tranexamic Acid (N=657)	Placebo (N=643)
Age — yr	44.1±19.7	44.2±18.9
Male sex — no. (%)	459 (69.9)	459 (71.4)
Mechanism of injury — no. (%)		
Blunt	610 (92.8)	588 (91.4)
Penetrating	44 (6.7)	55 (8.6)
Burn	3 (0.5)	0



ORIGINAL ARTICLE

Prehospital Tranexamic Acid for Severe Trauma

► Give 1g prehospital TXA within 3 hours of severe trauma

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

Tranexamic Acid to Prevent Bleeding in Patients with Hematologic Malignancies and Severe Thrombocytopenia (TREATT trial). a Randomized Placebo-Controlled Trial

Lise J Estcourt, Zoe K McQuilten, Peter Bardy, Merrole Cole-Sinclair, Graham P. Collins, Philip J Crispin, Jennifer Curnow, Amber Degelia, Claire Dyer, Vanessa Fox, [Adam Friebe](#), Lajos Floro, Effie Grand, Cara Hudson, Gail Jones, Joanne Joseph, Charlotte Kallmeyer, Marina Karakantza, Paul Kerr, Sara Last, Maria Lobo-Clarke, Matthew Lumley, Mary Frances McMullin, Patrick G. Medd, Suzy M Morton, Andrew David Mumford, Maria Mushkbar, Joe Parsons, Gillian Powter, Mallika Sekhar, Richard Soutar, William S. Stevenson, Elango Subramoniapillai, Robyn Sutherland, Jeff Szer, Neil A Waters, Andrew H. Wei, David Alan Westerman, Sarah A Wexler, Erica M. Wood, Simon J Stanworth

- Patients: hematologic malignancy or stem cell transplant expected to have a platelet count $<10 \times 10^9/L$ for 5+ days at 27 centres in UK and Australia
- Intervention: TXA (1 g iv or 1.5 g oral) q8h if platelet count $< 30 \times 10^9/L$ for a maximum of 30 days
- Control: Placebo
- Outcome: death or WHO grade 2+ bleeding at 30 days
- Results: Randomized 616 patients – no difference in primary outcome, bleeding, death, or ATE/VTE events
- Results: TXA treated patients were more likely to “survive to 30 days without a red cell transfusion”* (29.4% vs. 20.5%, HR 0.82, 95% CI 0.72-0.93, $p=0.003$)



401.BLOOD TRANSFUSION | NOVEMBER 28, 2023

Tranexamic Acid to Prevent Bleeding in Patients with Hematologic Malignancies and Severe Thrombocytopenia (TREATT trial). a Randomized Placebo-Controlled Trial

Lise J Estcourt, Zoe K McQuilten, Peter Bardy, Merrole Cole-Sinclair, Graham P. Collins, Philip J Crispin, Jennifer Curnow, Amber Degelia, Claire Dyer, Vanessa Fox, Adam Friebe, Lajos Floro, Effie Grand, Cara Hudson, Gail Jones, Joanne Joseph, Charlotte Kallmeyer, Marina Karakantza, Paul Kerr, Sara Last, Maria Lobo-Clarke, Matthew Lumley, Mary Frances McMullin, Patrick G. Medd, Suzy M Morton, Andrew David Mumford, Maria Mushkbar, Joe Parsons, Gillian Powter, Mallika Sekhar, Richard Soutar, William S. Stevenson, Elango Subramoniapillai, Robyn Sutherland, Jeff Szer, Neil A Waters, Andrew H. Wei, David Alan Westerman, Sarah A Wexler, Erica M. Wood, Simon J Stanworth

► Consider TXA in patients with hematologic malignancies on a case-by-case basis, as there is some benefit (fewer transfusions) and little downside in some patients

Grade of recommendation	I Strong recommendation to do	IIa Moderate recommendation to do	IIb Weak recommendation to do	III Recommendation not to do
Conclusions of evidence	Benefits >>> risk & burdens	Benefits >> risk & burdens	Benefits >= risks & burdens	No benefit / Potentially harm
A High level of evidence Consistent evidence from well performed and high quality studies or systematic reviews (low risk of bias, direct, consistent, precise)	Strong recommendation based on high level of evidence	Moderate recommendation based on high level of evidence	Weak recommendation based on high level of evidence	Recommendation based on high level of evidence
B Moderate /Low level of evidence Evidence from studies or systematic reviews with few important limitations	Strong recommendation based on moderate/ low level of evidence	Moderate recommendation based on moderate/ low level of evidence	Weak recommendation based on moderate/ low level of evidence	Recommendation based on moderate/ low level of evidence
C Very low level of evidence Evidence from studies with serious flaws. Only expert opinion, or standards of care	Strong recommendation based on expert opinion	Moderate recommendation based on very low level of evidence Diverging expert opinions	Weak recommendation based on very low level of evidence Diverging expert opinions	Recommendation based on very low level of evidence Expert opinion

Questions

Acknowledgements

Aditi Khandelwal

Yulia Lin

Jeannie Callum

Nadine Shehata

Katerina Pavenski

Heather Vandermeulen

Casey Kapitany

Paula Nixon

