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June

RBC Transfusion: Applying evidence-based transfusion care Jeannie Callum, MD, FRCPC (Presented by Yulia Lin May 2024)

Disclosure

J. Callum:

Research funding from Canadian Blood Services, Octapharma Canada¹, Defense Research and Development¹ and Novo Nordisk²

Y. Lin

- Research funding from Canadian Blood Services and Octapharma
- Consultant for Choosing Wisely Canada
- This is a talk about using red cell transfusions for adult patients
- Management of major hemorrhage is covered in a separate talk | Pendergrast

I. Callum

Outline

2 Cases

- Red blood cell basics
- Key risks of red blood cell transfusion
- When should you give red cell transfusions?
 - Multiple randomized trials and meta-analyses to guide your decisions

Case 1

- 27 year old female, 2 days post C-section complicated by moderate post-partum hemorrhage
- Pre-delivery hemoglobin was 10.5 g/dL
- Hemoglobin this morning is 5.7 g/dL
- Heart rate 87, blood pressure 102/56.
- She is pale and tired but no pre-syncope or lightheadedness
- You have ordered a dose of intravenous iron
- Plan is for discharge today
- Should you transfuse I unit of RBCs before sending her home?

Case 2

- 76 year old woman 4 days post-op from hip fracture surgery (after fracture from a fall)
- Plan to discharge tomorrow
- Hemoglobin 7.1 g/dL
- Asymptomatic, vitals stable
- No cardiac history
- Should you transfuse red blood cells?



Red blood cell basics

- Volume ~300 mL, hematocrit 50-65%, anticoagulant SAG-M
 - Some units may have lower volumes so ensure you include the volume required on your order (~5 mL/kg for an adult dose)
- Each unit increases hemoglobin by 1.0 g/dL
- Small amount of residual plasma
- Acceptable for transfusion for 42 days from donation
- Red cells in Rwanda are not routinely leukoreduced, but can be requested on an as-needed basis
 - Leukoreduction reduces the risk of transfusion reactions (marginally), reduces the risk of HLA alloimmunization (important only for patients undergoing aggressive chemotherapy), and prevents some infectious transmissions (HTLV, CMV)

Irradiated red blood cells

If not available, transfuse red blood cells over 14 days from collection [Kopolovic, et al. Blood 2015:16;126:406-14]

PATIENTS REQUIRING IRRADIATED BLOOD 146

- First and second degree family members or HLA-selected donors.
- Intra-uterine or neonatal exchange transfusion.
- Congenital T-cell immunodeficiency.
- Autologous stem cell transplant recipients from 7 days prior to stem cell collection to 3 months post-transplant (6 months if total body irradiation is part of the conditioning regimen).
- Allogeneic stem cell transplant from initiation of conditioning regimen and continued until over 6 months posttransplant and lymphocyte count >1x10⁹/L and patient free of chronic GvHD and off all immunosuppressive agents (otherwise continue indefinitely).
- CAR-T cell infusion from 7 days prior to collection and for 3 months after infusion.
- All patients with Hodgkin's Disease.
- Certain therapeutics in select patient populations (see box to right)

Alemtuzumab (anti-CD52) Anti-thymocyte globulin (ATG) Bendamustine Cladribine (2-CDA) Clofarabine Deoxycoformicin Fludarabine Nelarabine

http://www.bcshguidelines.com/do cuments/irrad_bcsh_072010.pdf



Risks of RBCs

- Transfusion associated circulatory overload (TACO) 1 in 50 to 1 in 100
 - 300 mL of RBCs is not the same as 300 mL of saline
- Transfusion-related acute lung injury (TRALI) 1 in 10,000
- Acute and delayed hemolytic transfusion reactions
 - ABO-immune hemolysis (by mistake) 1 in 354,000
 - RBC alloantibodies I in I3 (hemolytic disease of the newborn risk for girls and young women)
 - Delayed hemolytic transfusion reactions 1 in 2500
- Anaphylaxis 1 in 40,000
- More bleeding (from GI bleeding trials)
- HLA alloimmunization (leading to long waits for organ transplants)
- Increased risk of thromboembolic complications
- Hyperhemolysis in patients with sickle cell disease
- Association of ICH in recipients of donors who decades later had multiple ICH (cerebral amyloid angiopathy)

What about the risk of HIV, HBV or HCV?

		Sub-Saharan Africa
<1 in 1,000,000	Transmission of West Nile Virus	
1 in 2,000,000	Residual risk of hepatitis B per unit ⁸⁷	
1 in 4,000,000	Transmission of Chagas disease per unit	HBV = 1:233
1 in 12,900,000	Residual risk of human immunodeficiency virus (HIV) per unit ⁸⁷	
1 in 27,100,000	Residual risk of hepatitis C per unit ⁸⁷	HCV = 1;400
<1 in 1,000,000,000	Transmission of HTLV per unit ⁸⁸	HIV = 1:1000

Bloody easy 5.1, Ontario Transfusion Handbook, 2022.

Jayaraman S, Transfusion. 2010;50:433

Rwanda

<u>Trends and Gaps in National Blood Transfusion Services — 14 Sub-Saharan African</u> <u>Countries, 2014–2016 | MMWR (cdc.gov)</u>

				Prevalence (%) of TTIs in collected blood units								
							Other	TTIs		All TTIs		
	HIV popu	lation prev	alence (%)	HIV			HBV, H	ICV, and	syphilis	HIV, HB	V, HCV, an	d syphilis
Country	2014	2015	2016	2014	2015	2016	2014	2015	2016	2014	2015	2016
Côte d'Ivoire	3.0	2.8	2.7	0.3	0.04	0.2	8.6	9.0	8.9	9.0	9.0	9.1
Ethiopia	1.1	1.1	0.9	2.1	1.2	1.1	4.4	4.6	4.2	5.2	5.1	4.5
Ghana	1.7	1.6	1.6	0.7	0.5	0.3	9.7	7.1	11.6	11.8	8.3	12.7
Kenya	5.7	5.6	5.4	0.6	0.8	0.6	2.8	4.3	2.5	3.5	5.2	3.2
Lesotho	24.7	24.9	25	2.6	2.4	2.5	3.6	3.8	5.0	6.2	6.2	7.6
Mozambique	13.0	12.7	12.3	5.2	4.8	4.0	8.2	8.8	6.9	13.4	13.6	11.0
Nigeria	3.1	3.0	2.9	1.4	1.4	1.5	11.3	11.7	13.1	12.9	13.2	14.6
Rwanda	3.2	3.2	3.1	0.1	0.1	0.2	2.6	2.7	3.4	2.8	2.9	3.6
South Africa	18.8	18.9	18.9	0.2	0.2	0.1	0.3	0.3	0.5	0.5	0.5	0.7

Risk of donor testing positive



Figure 2. The average proportion of Voluntary non-remunerated blood donations in 46 countries of the WHO Afro Region in 2013

TACO

- Rate of TACO increases with increasing age:
 - aged 49 yr or less: 2.0%
 - aged 50-59 yr: 3.3%
 - aged 60-69 yr: 4.2%
 - aged 70-79 yr: 5.2%
 - aged 80 yr+: 7.4%
- Increased rate of TACO with increasing amount of volume transfused (P<0.001) and increasing total fluid balance (P<0.001)
- Odds ratio of death for TACO cases compared with transfused controls of 3.8 (95% Cl, 2.2 to 6.7) (P<0.001)

Pre-transfusion RBC checklist

Red Blood Cell Pre-Transfusion Checklist

Alternatives failed or have been ordered?	 Anemia investigations completed (e.g., CBC, blood film, ferritin, iron saturation, vitamin B12, reticulocyte count) Iron (oral and IV), vitamin B12, erythropoietin ordered or not indicated
Consent?	Patient advised of risks of: TACO 1 in 100 Hemolytic reaction 1 in 7,000 TRALI 1 in 10,000 Major allergic reaction 1 in 40,000 Bacterial infection 1 in 250,000

Back of blood easy 5.1

Tseng, et al. 2016 Apr;26(2):104-10. doi: 10.1111/tme.12284. Epub 2016 Feb 10.

Pre-transfusion RBC Checklist

Female under 45?	Order Kell-negative units Rwanda: Kell+ donors deferred			
	Inform recipient of the potential risk of transfusion causing hemolytic disease of the newborn in future pregnancies			
At risk for FATAL	Order irradiated blood if patient has any history of the following:			
transfusion-	Hodgkin's lymphoma			
associated Graft vs. Host Disease?	Allogeneic or autologous stem cell transplant			
Or units > 14 days	Ever treated with fludarabine, cladribine, bendamustine, alemtuzumab, anti-thymocyte globulin (ATG)			
Or units >14 days	Congenital immunodeficiencies			

Pre-transfusion RBC Checklist





Science not the "Art of Medicine"

47 RCTs with 20,967 patients

restrictive (7.0-7.5-8.0) vs. liberal (9.0-9.5-10.0)

Clinical trials.gov – 14 ongoing studies that will add an additional 15,000 patients

Pre-TRICC

Hebert P, et al. Am J Resp CCM 1997; 155: 1618-23



$1400 \\ $	502 257 320 40 62 2427 29 1000 168 100 416 51 29 8321	1/5 25 14 0 87 1 19 15 3 34 16 2 689	496 382 317 40 65 2429 29 1003 177 100 417 49 31 8408	13% 4.0% 3.2% 9.6% 0.3% 4.5% 3.6% 1.0% 5.0% 3.9% 0.4% 100.0%	0.95 [0.80 , 1.13] 0.83 [0.44 , 1.57] 0.99 [0.48 , 2.04] Not estimable 0.85 [0.63 , 1.15] 1.00 [0.07 , 15.24] 1.37 [0.76 , 2.46] 1.12 [0.57 , 2.20] 1.67 [0.41 , 6.79] 0.56 [0.32 , 0.97] 0.72 [0.38 , 1.36] 0.53 [0.05 , 5.58] 0.99 [0.86 , 1.15]		••••••••••••••••••••••••••••••••••••
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14 14 14 0 74 1 26 16 5 3 19 2 2	502 257 320 40 62 2427 29 1000 168 100 416 51	1/5 25 14 0 0 87 1 19 15 3 34 16	496 382 317 40 65 2429 29 1003 177 100 417 49	13.3% 4.0% 3.2% 9.6% 0.3% 4.5% 3.6% 1.0% 5.0% 3.9%	0.95 [0.80 , 1.13] 0.83 [0.44 , 1.57] 0.99 [0.48 , 2.04] Not estimable 0.85 [0.63 , 1.15] 1.00 [0.07 , 15.24] 1.37 [0.76 , 2.46] 1.12 [0.57 , 2.20] 1.67 [0.41 , 6.79] 0.56 [0.32 , 0.97] 0.72 [0.38 , 1.36]		
14 14 14 0 0 74 1 26 16 5 3 19	502 257 320 40 62 2427 29 1000 168 100 416	1/5 25 14 0 0 87 1 19 15 3 34	496 382 317 40 65 2429 29 1003 177 100 417	13.3% 4.0% 3.2% 9.6% 0.3% 4.5% 3.6% 1.0% 5.0%	0.95 [0.80 , 1.13] 0.83 [0.44 , 1.57] 0.99 [0.48 , 2.04] Not estimable 0.85 [0.63 , 1.15] 1.00 [0.07 , 15.24] 1.37 [0.76 , 2.46] 1.12 [0.57 , 2.20] 1.67 [0.41 , 6.79] 0.56 [0.32 , 0.97]		
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14 14 0 74 1 26	502 257 320 40 62 2427 29 1000	1/5 25 14 0 0 87 1 19	496 382 317 40 65 2429 29 1003	9.6% 0.3%	0.95 [0.80 , 1.13] 0.83 [0.44 , 1.57] 0.99 [0.48 , 2.04] Not estimable 0.85 [0.63 , 1.15] 1.00 [0.07 , 15.24] 1.37 [0.76 , 2.46]		
14 14 14 0 0 74 1	502 257 320 40 62 2427 29	175 25 14 0 0 87 1	496 382 317 40 65 2429 29	4.0% 3.2% 9.6% 0.3%	0.95 [0.80 , 1.13] 0.83 [0.44 , 1.57] 0.99 [0.48 , 2.04] Not estimable 0.85 [0.63 , 1.15] 1.00 [0.07 , 15.24]		
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14 14	502 257 320	175 25 14	496 382 317	4.0% 3.2%	0.95 [0.80 , 1.13] 0.83 [0.44 , 1.57] 0.99 [0.48 , 2.04]	- -	
162	502 257	25	496 382	4.0%	0.83 [0.44 , 1.57]	.	
102	502	1/5	496	13,3%	0.95[0.80,1.13]	•	
1.00		4 7 7	100	10 504	0.05 [0.00 1.10]		
78	418	98	420	10.7%	0.80 [0.61 , 1.04]	-	• • • • • ? •
8	33	9	36	2.6%	0.97 [0.42 , 2.22]	+	• ? • • • ? •
15	249	13	253	3.2%	1.17 [0.57 , 2.41]	- - -	$\bullet \bullet \bullet \bullet \bullet \bullet \bullet$
C	109	1	109	0.2%	0.33 [0.01 , 8.09]		🔁 🖶 🖶 🖶 📍 ? ?
i 21	144	12	140	3.6%	1.70 [0.87 , 3.32]	+- -	
7	23	1	21	0.5%	6.39 [0.86 , 47.70]	↓ • − −	• • • • • • • ?
2	26	1	36	0.4%	2.77 [0.26, 28.95]	_ 	$\bullet \bullet \bullet \bullet \bullet \bullet \bullet$
5	60	0	60	0.2%	11.00 [0.62 , 194.63]		• • • • ? • •
19	342	25	324	4.6%	0.72 [0.40 , 1.28]	-	
1	59	2	30	0.4%	0.25 [0.02 , 2.69]		
15 23	101	8	97	3.0%	2.76 [1.30 , 5.87]		
2	23	1	21	0.4%	1.83 [0.18 , 18.70]		
7	55	1	55	0.5%	7.00 [0.89 , 55.01]		
43	1009	52	1007	7.4%	0.83 [0.56 , 1.22]	-	
1	42	1	42	0.3%	1.00 [0.06 , 15.47]		$\bullet \bullet \bullet \bullet \bullet \bullet ? \bullet$
4	50	4	49	1.1%	0.98 [0.26 , 3.70]		• • • • • • ? •
C	26	2	24	0.2%	0.19 [0.01 , 3.67]		? ?
l	84 0 4 1 43 7 2 5 23 .5 1 1 9	84 151 0 26 4 50 1 42 43 1009 7 55 2 23 .5 23 101 1 59 19 342	84 151 67 0 26 2 4 50 4 1 42 1 43 1009 52 7 55 1 2 23 1 .5 23 101 8 1 59 2 19 342 25	84 151 67 149 0 26 2 24 4 50 4 49 1 42 1 42 43 1009 52 1007 7 55 1 55 2 23 1 21 .5 23 101 8 97 1 59 2 30 19 342 25 324	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	84 151 67 149 11.8% 1.24 [0.99, 1.55] 0 26 2 24 0.2% 0.19 [0.01, 3.67] 4 50 4 49 1.1% 0.98 [0.26, 3.70] 1 42 1 42 0.3% 1.00 [0.06, 15.47] 43 1009 52 1007 7.4% 0.83 [0.56, 1.22] 7 55 1 55 0.5% 7.00 [0.89, 55.01] 2 23 1 21 0.4% 1.83 [0.18, 18.70] .5 23 101 8 97 3.0% 2.76 [1.30, 5.87] 1 59 2 30 0.4% 0.25 [0.02, 2.69] 19 342 25 324 4.6% 0.72 [0.40, 1.28]	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$

Reduces the risk of transfusion: 0.54 (0.47-0.63; P<0.001)

And the number of units transfused (mean difference -1.43 unit, (-2.01 to -0.86; P<0.001)

Cost to put a single RBC unit into a patient US\$741 in 2010

Holst L, et al. BMJ 2015;350:h1354

RBC - TRICC Study

- NEJM 1999; 340:409-17 Hebert et al
 - n=838 non-bleeding, ICU patients, Hb <9 g/dL</p>
 - RCT transfusion Hb <7 vs <10</p>
 - Non-leukoreduced RBCs
 - Stratified by APACHE 2 score
 - Groups equal with respect to baseline characteristics
 - Average patient: 58 year old male, with I-2 organ failure, mechanically vented, admitted to the ICU from the OR

Outcome - Mortality

Outcome	<7 g/dL	<10 g/dL	P value
30-day	18.7%	23.3%	P=0.11
Hospital	22.2%	28.1%	P=0.05

NNT = 17 patients to prevent one in-hospital death

Morbidity Outcomes in TRICC

	Restrictive N (%)	Liberal N (%)	P Value
МІ	3 (0.7)	12 (2.9)	0.02
Pulmonary Edema	22 (5.3)	45 (10.7)	0.01
ARDS	32 (7.7)	48 (11.4)	0.06

Key RBC Trials



BMJ 2016;352:i1351

No benefit in CVD patients

	No of events/ total No of patients				
Study	Restrictive	Liberal	Risk ratio MH random	Weight	Risk ratio MH random
All studies	transfusion	transfusion	effect (95% CI)	(%)	effect (95% CI)
Almeida 2015	7/22	0/12		0.9	8.48 (0.53 to 136.76)
Bush 1997	4/49	4/50		3.8	1.02 (0.27 to 3.85)
Carson 2011	43/1008	52/995		27.7	0.82 (0.55 to 1.21)
Carson 2013	7/55	1/55		1.6	7.00 (0.89 to 55.01)
Cooper 2011	2/24	1/21		1.3	1.75 (0.17 to 17.95)
Gregersen 2015	6/34	3/25		4.0	1.47 (0.41 to 5.32)
Hebert 1999	29/111	31/146	- - -	23.9	1.23 (0.79 to 1.91)
Holst 2014	33/75	24/66	- 	26.5	1.21 (0.80 to 1.82)
Jairath 2015*	6/49	2/67		2.8	4.10 (0.86 to 19.47)
Parker 2013	4/70	4/67		3.7	0.96 (0.25 to 3.67)
Walsh 2013	3/17	4/15		3.8	0.66 (0.18 to 1.50)
Total	144/1514	126/1519	↓	100.0	1.15 (0.88 to 1.50)
Test for heterogeneity: τ^2 =0.03,	χ ² =11.58, df=10,	P=0.31, I ² =14%			
Test for overall effect: z=1.04, P=	=0.30				
Studies randomised by CVD					
Bush 1997	4/49	4/50		3.8	1.02 (0.27 to 3.85)
Carson 2011	43/1008	52/995		27.7	0.82 (0.55 to 1.21)
Carson 2013	7/55	1/55		1.6	7.00 (0.89 to 55.01)
Cooper 2011	2/24	1/21		1.3	1.75 (0.17 to 17.95)
Walsh 2013	3/17	4/15		3.8	0.66 (0.18 to 1.50)
Total	59/1153	62/1136	+	100.0	0.96 (0.58 to 1.59)
Test for heterogeneity: τ^2 =0.06,	χ ² =4.67, df=4, P=	=0.32, I ² =14%	0.01 0.1 1 10 10	00	
Test for overall effect: z=0.17, P=	=0.87		Favours restrictive Favours liber	al	

transfusion

transfusion

No benefit for cardiac surgery patients

F	Restric	tive		Liber	al		Risk Ratio	Risk Ratio
Events	Total	Percent	Events	Total	Percent	Weight	IV, Fixed, 95% CI	IV, Fixed, 95% CI
ion Thre	shold	Trials						
3	215	1.4%	6	222	2.7%	3.0%	0.52 [0.13, 2.04]	
15	249	6.0%	12	253	4.7%	10.2%	1.27 [0.61, 2.66]	
4	25	16.0%	1	25	4.0%	1.2%	4.00 [0.48, 33.33]	
0	38	0.0%	0	35	0.0%		Not estimable	
26	1000	2.6%	19	1003	1.9%	16.3%	1.37 [0.76, 2.46]	
3	363	0.8%	6	354	1.7%	2.9%	0.49 [0.12, 1.93]	
74	2427	3.0%	87	2429	3.6%	60.0%	0.85 [0.63, 1.15]	-
125	4317	2.9%	131	4321	3.0%	93.7%	0.95 [0.75, 1.22]	•
6.03, df	= 5 (P	= 0.30); 12	= 17%				CONTRACTOR PORT AND ADDRESS	1
Z = 0.38	3 (P = 0	0.71)						
fusion 1	63 30	3.2%	2	62 30	3.2%	1.5%	0.98 [0.14, 6.77]	
0	50	0.0%		50	0.0%	0.0%	0.33 [0.01, 7.87]	· · · · ·
6	82	7 3%	5	80	6.3%	4 2%	1 17 10 37 3 681	
	228	2 50/		226	3 5%	6 3%	1.01 [0.37, 3.00]	
0.54 df	= 2 /P	= 0.76) - 12	= 0%	220	0.0 %	0.370	1.01 [0.38, 2.07]	
7=0.0	- 2 (F	0.10), 1	- 0 /0					
2 - 0.01	. (
133	4545	2.9%	139	4547	3.1%	100.0%	0.96 [0.76, 1.21]	+
6.58, df	= 8 (P	= 0.58); l ²	= 0%					
Z = 0.36	5 (P = 0).72)						
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Figure 4 Mortality within 30 days of surgery in randomized controlled trials of adult and paediatric cardiac surgery patients. Fixed-effects metaanalysis.

OR 0.96 (0.76-1.21)

Shehata et al. Eur Heart J 2019; 1: 1081-1088

REALITY Trial – RCT 80 vs. 100 g/L in patients with acute myocardial infarction

Table 3. Primary and Secondary Outcomes at 30 Days Among the As-Randomized Population in a Study of the Effect of a Restrictive vs Liberal Blood Transfusion Strategy on Patients With Acute Myocardial Infarction and Anemia

	No. (%)		Difference	Relative risk	
Outcome	Restrictive	Liberal	(95% CI), %	(1-sided 97.5% CI)	
Primary (major adverse cardiovascular events) No./total No. (%) [95% CI] ^a),				
As-treated population	36/327 (11.0) [7.5 to 14.6]	45/322 (14.0) [10.0 to 17.9]	-3.0 (-8.4 to 2.4)	0.79 (0.00 to 1.19)	
As-randomized population	38/342 (11.1) [7.6 to 14.6]	46/324 (14.2) [10.2 to 18.2]	-3.1 (-8.4 to 2.3)	0.78 (0.00 to 1.17)	

MINT Trial 70-80 vs 100 g/L for patients with acute myocardial infarction



No difference in primary Outcome of recurrent MI Or death

RR 1.15 (0.99-1.34)

2 major problems:

- (1) one-third transfused before randomization
- (2) Recurrent MI rate was 2x the rate of all other MI trial (raising concerns for validity of the primary outcome)



Villanueva et al. NEJM Jan 2013;368:11-21

PPH – WOMB Trial

- > 37 Dutch hospitals, 521 women randomized
- PPH with >1000 ml, Hb drop of 19+ points, and hemoglobin between 4.8-7.9 g/L, no severe symptoms of anemia (dyspnea, syncope, HR>100)

transfusion

Randomized to transfusion or no transfusion



Variable	Transfusion (n = 258)	Non-intervention (n = 261)	Р
RBC transfusion			
Units per woman	2 (2–2)	0 (0–0)	<0.001
Total units*	517	88	<0.001
Hb concentration after transfusion, g/dl)**	9.0 (8.5–9.6)	8.9 (8.2–9.7)	0.56
Hb concentration at discharge (g/dl)***	9.0 (8.5–9.5)	7.4 (6.8–7.7)	<0.001
Hb concentration at 6 weeks (g/dl)****	12.1 (11.3–12.6)	11.9 (10.9–12.6)	0.18

Table 2. Blood loss, haemoglobin concentration, and RBC



ORACL Trial

- Patients: Ortho trauma past initial resuscitation phase, hemodynamically stable, aged 18-50, Hb<90 g/L
- ▶ N=65
- Multicentre trial
- Intervention: Restrictive threshold 55 g/L
- Control: Liberal threshold 70 g/L
- Time: I year follow-up
- Outcome: Infection

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

Mullis BH, et al. J Orthop Trauma. 2024 Jan 1;38(1):18-24

AABB RBC Guideline 2023

Recommendations for Adults

Recommendation 1

For hospitalized adult patients who are hemodynamically stable, the international panel recommends a restrictive RBC transfusion strategy in which the transfusion is considered when the hemoglobin concentration is less than 7 g/dL (strong recommendation, moderate certainty evidence).

Remark: in accordance with the restrictive strategy threshold used in most of the trials for subgroups of patients, clinicians may choose a threshold of 7.5 g/dL for patients undergoing cardiac surgery and 8 g/dL for patients undergoing orthopedic surgery or those with preexisting cardiovascular disease.

Recommendation 2

D

For hospitalized adult patients, the panel suggests a restrictive RBC transfusion strategy in which transfusion is considered when the hemoglobin concentration is less than 7 g/dL in those with hematologic and oncologic disorders (conditional recommendation, low certainty evidence).

Recommendations for Children

Recommendation 3

For critically ill children and hospitalized children at risk of critical illness who are hemodynamically stable and without a transfusion-dependent hemoglobinopathy, cyanotic cardiac condition, or severe hypoxemia, the international panel recommends a restrictive transfusion strategy in which a transfusion is considered when the hemoglobin level is less than 7 g/dL compared with one of less than 9.5 g/dL (strong recommendation, moderate certainty evidence).

Recommendation 4

The international panel suggests considering a transfusion threshold for hemodynamically stable children with congenital heart disease that is based on the cardiac abnormality and stage of surgical repair: 7 g/dL (biventricular repair), 9 g/dL (single-ventricle palliation), or 7 to 9 g/dL (uncorrected congenital heart disease) (conditional recommendation, low certainty evidence).

Carson JL, et al. JAMA. 2023 Nov 21;330(19):1892-1902

2018 Frankfurt Guidelines

Newer but same as AABB plus:

- The panel recommended a restrictive RBC transfusion threshold (hemoglobin concentration <7.5 g/dL) in patients undergoing cardiovascular surgery
- The panel recommended a restrictive transfusion threshold (hemoglobin concentration 7-8 g/dL) in hemodynamically stable patients with acute gastrointestinal bleeding

Reasonable approach for inpatients Remember not to transfuse for pallor/fatigue!

Patient scenario	Hemoglobin threshold	Transfusion approach
Young patient with severe iron or B12 deficiency anemia with only fatigue and pallor	Any	lv iron (or B12 im/po)
Young patient with reversible asymptomatic anemia (eg. Postpartum, recovering young trauma)	<5 g/dL	l unit
Average patient without symptoms or cardiac history (eg. ICU, CVICU, hem-onc)	<7 g/dL	l unit
Cardiac history without symptoms	<7-8 g/dL	l unit
Hemodynamic symptoms (tachycardia, pre-syncope, etc)	<9 g/dL	l unit
Myocardial infarction with only fatigue and pallor	<8 g/dL	l unit GO SLOW
Slow bleeding and asymptomatic anemia	<7 g/dL	I-2 units
Rapid hemorrhage (eg. stabbing, gunshot, varices)	Keep 6-11 g/dL	As many as you need! Don't forget to use the term uncrossmatched!



Case 1

- 27 year old female, 2 days post C-section complicated by moderate post-partum hemorrhage
- Pre-delivery hemoglobin was 10.5 g/dL
- Hemoglobin this morning is 5.7 g/dL
- Heart rate 87, blood pressure 102/56.
- She is pale and tired but no pre-syncope or lightheadedness
- You have ordered a dose of intravenous iron
- Plan is for discharge today
- Should you transfuse I unit of RBCs before sending her home?

Case 2

- 76 year old woman 4 days post-op from hip fracture surgery (after fracture from a fall)
- Plan to discharge tomorrow
- Hemoglobin 7.1 g/dL
- Asymptomatic, vitals stable
- No cardiac history
- Should you transfuse red blood cells?

Summary

- Use these trials to help set your 'general' transfusion trigger where you might <u>consider</u> a transfusion
- Don't be overly prescriptive just because the hemoglobin is
 6.9 g/dL you have to transfuse...or 7.1 g/dL and you hold off
- Look at your patient Are they symptomatic? Adjust the trigger to your patient's co-morbidities
- Unless rapid bleeding I unit at a time
- Write a rate
- Anticipate and prevent TACO





023: RBC Transfusion Guidelines with Jeff Carson

Whither RBCs? There's no one better than lead author Dr. Jeff Carson to discuss the 2016 AABB RBC transfusion threshold recommendations!



035: Why Give Platelets? with Rick Kaufman

Platelets are tiny, but they can be a big issue! Dr. Rick Kaufman magnifies what the evidence shows about platelet transfusion.

Listen to This Episode!



016: Plasma Transfusion with Jeannie Callum

As many as 50% of plasma transfusions are unnecessary or inappropriate! You need to know why, and Dr. Jeannie Callum explains it SO well!

