

Периоперативна Анемија

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Anemia



Talking anemia is offensive???

Paradigm

- Started a study for Anemia and PBM issues
- In 80 primary care settings
- Not Published, GOING ON
- DEVESTATING

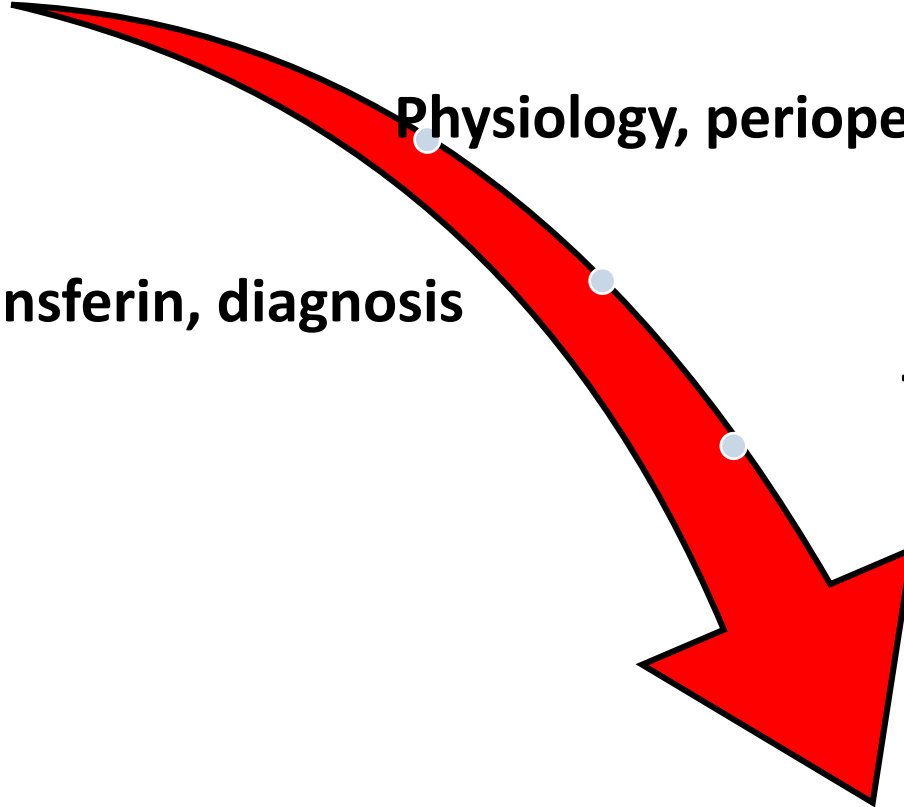
Anemia , perioperative risks

Physiology, perioperative implications

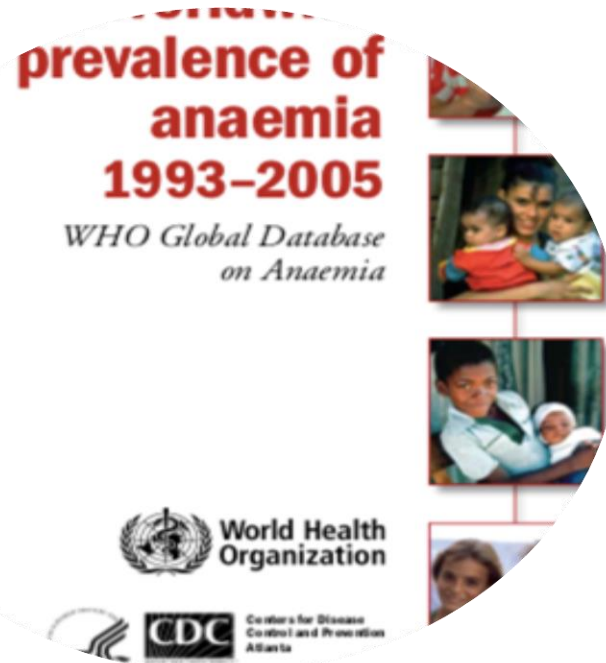
Iron, ferritin, transferin, diagnosis

Treatment modalities

PBM -Recommendations



Facts



- prevalence 5-75%(*Shander et al.2004*)

- 50% is iron deficiency anemia

- *de Benoist B et al., eds. Worldwide prevalence of anaemia 1993-2005. WHO Global Database on Anaemia Geneva, World Health Organization, 2008*

- *[the global prevalence of anaemia in 2011 - World Health ...](#)*

https://apps.who.int/iris/bitstream/handle/9789241564960_eng



Anaemia in the older surgical patient: a review of prevalence, causes, implications and management

International Surgery Journal
Somani B et al. Int Surg J. 2016 Feb;3(1):71-76
<http://www.ijurgery.com>

pISSN 2349-3305 | eISSN 2349-2902

Research Article

DOI: <http://dx.doi.org/10.18203/2349-2902.ij20151535>

Anaemia in surgical patients and its effect on recovery of patients

J R Soc Med 2013; 106: 269-277. DOI: 10.1177/0141076813479580

Facts -surgery

- >30% have anemia on the day of surgery
- 48 % (>65y) have unexplained anemia
- cardiac surgery (40,4 %), Emergency hip(>45 %)
- **Surgery, age, gender and status**

Steinberg A et al. Anesth Intensivmed 2015;56:64-74

Anemia is harmless- we have weapon for that?



- MOST CLINICANS THINK TRANSFUSION IS THE WEAPON

- ARE WE SURE?

Eur. J Anaesthesiology 2017; 34: 332-395

- Preoperative anemia in adults and children appears to be a **strong predictor** for **perioperative blood transfusion** across various types of conditions and surgeries and may be **associated with adverse events. B**

Anemia as a risk

- Anemia is – RISK FACTOR –multi dimensional
 - *Fowler BJM 2015*
- Mild anemia- (colorectal surgery >20 000) (AMI, stroke, death)
 - *Leichete J. Am Coll Surg 2011*
- Independent predictive risk - mortality and death
 - Beatie et al, 2009;Spahn D, 2010;Musllam et al, 2011*

- Preop anemia - Co existing anemia increases health care costs

Dowling, 2007;

Kaser et al 2019

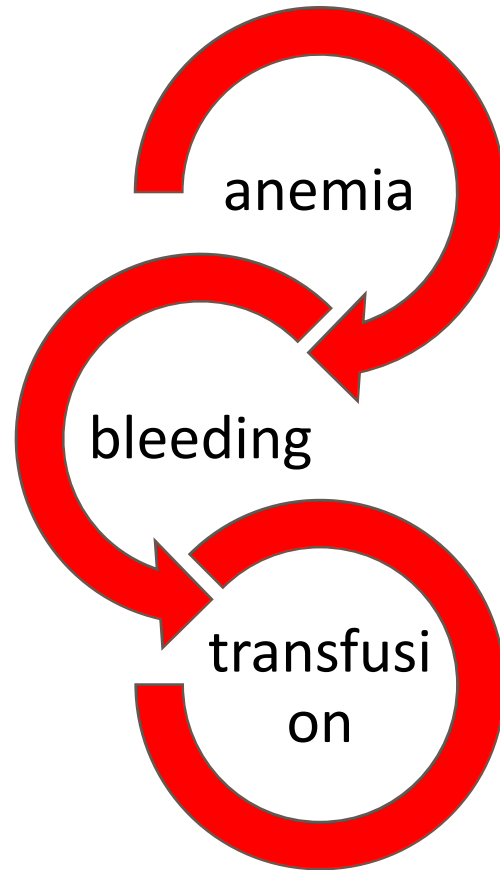
- Increases the adverse events

Fredericke C et al 2019

- Predictor for transfusion, and transfusion costs

- *Munoz, BJA 2015*

- Kaser et al 2019



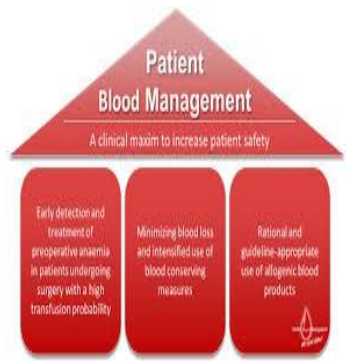
>5 mil pts (24 mil of units blood) yearly

(ANNUAL SUMMARY OF THE REPORTING ADVERSE REACTIONS AND EVENTS 2015, EUROPEAN COMMISSION).



- With such numbers, WE HAVE TO CHANGE OUR ATTITUDES
- AWARENESS YES, BUT ADDRESSABLENESS ALSO is a matter of thing
- DOING SMTH.
- Anemia has influence on every part of the perioperative part

	DETECTION AND MANEGMENT ANEMIA	MINIMISATION OF BLOOD LOOS AND BLEEDING	MANEGMENT ANEMIA AND IMPROVE TOLERANCE
PRE-OP	• Aim for assessment of anaemia 4–6 weeks before surgery • Identify, evaluate and treat anaemia • Treat absolute or functional iron deficiency with oral or i.v. iron • Consider erythropoiesis stimulating agents if nutritional anaemia is ruled out/treated • Refer for further evaluation as necessary	• Identify and manage bleeding risk (past medical and family history) • Review medications (antiplatelet, anticoagulation therapy) • Minimize iatrogenic blood loss • Procedure planning and rehearsal	• Compare estimated blood loss with patient specific tolerable blood loss • Assess and optimize patient's physiologic reserve e.g. pulmonary and cardiac function • Formulate patient-specific management plan using appropriate blood conservation modalities
INTRA-OP	• Schedule surgery with optimization of red cell mass	• Meticulous haemostasis and surgical techniques • Anaesthetic blood sparing techniques e.g. central neuraxial blockade • Balanced physiology to aid optimal coagulation • Patient positioning • Goal directed management using point of care and VET • Antifibrinolysis and cell salvage • Monitor and manage bleeding • Avoid secondary haemorrhage • Maintain normothermia (unless specifically indicated) • Autologous blood salvage • Minimize iatrogenic blood sampling loss • Haemostasis/anticoagulation management	• Optimize cardiac output • Optimize oxygenation and ventilation • Evidence based transfusion thresholds
POST-OP	• Stimulate erythropoiesis • Manage nutrition and correctable anaemia (e.g. avoid folate deficiency, iron restricted erythropoiesis)		• Maximize oxygen delivery • Minimize oxygen consumption • Avoid/treat infections promptly • Evidence based transfusion thresholds



The timely application of EBM concepts designed to maintain HB concentrations, optimize hemostasis minimize blood loss in a effort to improve patient outcome

WHO ASSEMBLY – PBM – PATIENT SAFETY
WHA 63.12 RESOLUTION WHO MEMBERS SHOULD IMPLEMENT



PBM organization chart per the Society for the advanced Blood Management



Multimodal Patient Blood Management Program Based on a Three-pillar Strategy

A Systematic Review and Meta-analysis

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Objectives: To determine whether a multidisciplinary, multimodal Patient Blood Management (PBM) program for patients undergoing surgery is effective in reducing perioperative complication rate, and thereby is effective in improving clinical outcome.

Background: PBM is a medical concept with the focus on a comprehensive anemia management, to minimize iatrogenic (unnecessary) blood loss, and to harness and optimize patient-specific physiological tolerance of anemia.

Methods: A systematic review and meta-analysis was performed. Eligible studies had to address each of the 3 PBM pillars with at least 1 measure per pillar, for example, preoperative anemia management plus cell salvage plus

rational transfusion strategy. The study protocol has been registered with PROSPERO (CRD42017079217).

Results: Seventeen studies comprising 235,779 surgical patients were included in this meta-analysis (100,886 pre-PBM group and 134,893 PBM group). Implementation of PBM significantly reduced transfusion rates by 39% [risk ratio (RR) 0.61, 95% confidence interval (CI) 0.55–0.68, $P < 0.00001$], 0.43 red blood cell units per patient (mean difference -0.43 , 95% CI -0.54 to -0.31 , $P < 0.00001$), hospital length of stay (mean difference -0.45 , 95% CI -0.65 to -0.25 , $P < 0.00001$), total number of complications (RR 0.80, 95% CI 0.74–0.88, $P < 0.00001$), and mortality rate (RR 0.89, 95% CI 0.80–0.98, $P = 0.02$).

Conclusions: Overall, a comprehensive PBM program addressing all 3 PBM pillars is associated with reduced transfusion need of red blood cell units, lower complication and mortality rate, and thereby improving clinical outcome. Thus, this first meta-analysis investigating a multimodal approach should motivate all executives and health care providers to support further PBM activities.

Keywords: blood transfusion, complication rate, effectiveness, mortality, Patient Blood Management

(Ann Surg 2019;269:794–804)

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- 200 000 patients with or without PBM
- Different types of surgery
- Risk for transfusion
- Transfusion rates per patients
- LOHS
- Mortality rate

No of patients exposed to allogenic RBC

orthopedic

Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI
Orthopedic surgery						
So-Osmann 2014	32	190	29	127	3.0%	0.74 [0.47, 1.16]
Rineau 2016	5	183	24	184	1.1%	0.21 [0.08, 0.54]
Kopanidis 2016	6	100	10	100	1.0%	0.60 [0.23, 1.59]
Loftus 2016	771	6593	1253	5997	5.8%	0.56 [0.52, 0.61]
Holt 2016	14	1010	325	1814	2.5%	0.08 [0.05, 0.13]
Freedman 2008	200	1127	266	1089	5.3%	0.73 [0.62, 0.86]
Ma 2014	3	33	12	37	0.8%	0.28 [0.09, 0.91]
Morais 2014	0	71	95	347	0.2%	0.03 [0.00, 0.40]
Albinarrate 2015	81	357	111	271	4.7%	0.55 [0.44, 0.70]
Frew 2016	9	406	107	717	1.8%	0.15 [0.08, 0.29]
Theusinger 2014	656	6721	431	2150	5.7%	0.49 [0.44, 0.54]
Meybohm 2016	2852	16298	2407	12633	5.9%	0.92 [0.87, 0.96]
Leahy 2014	63	2343	96	2669	4.0%	0.75 [0.55, 1.02]
Subtotal (95% CI)		35432		28135	41.7%	0.45 [0.35, 0.59]

Total events 4692 5166
Heterogeneity: $\tau^2 = 0.16$; $\chi^2 = 290.05$, $df = 12$ ($P < 0.00001$); $I^2 = 96\%$
Test for overall effect: $Z = 5.87$ ($P < 0.00001$)

cardio

Cardiac surgery						
Freedman 2008	119	275	165	274	5.3%	0.72 [0.61, 0.85]
Brevig 2009	86	479	229	530	4.9%	0.42 [0.34, 0.52]
Leahy 2014	34	266	81	295	3.6%	0.47 [0.32, 0.67]
Gross 2015	473	2275	152	387	5.4%	0.53 [0.46, 0.61]
Moskowitz 2010	62	586	249	586	4.5%	0.25 [0.19, 0.32]
Meybohm 2016	3837	7904	3378	5630	6.0%	0.81 [0.78, 0.83]
Subtotal (95% CI)		11785		7702	29.7%	0.50 [0.36, 0.70]

Total events 4611 4254
Heterogeneity: $\tau^2 = 0.17$; $\chi^2 = 156.24$, $df = 5$ ($P < 0.00001$); $I^2 = 97\%$
Test for overall effect: $Z = 4.00$ ($P < 0.0001$)

vascular

Vascular surgery						
Freedman 2008	105	232	143	287	5.1%	0.91 [0.76, 1.09]
Leahy 2014	17	406	20	471	2.0%	0.99 [0.52, 1.86]
Meybohm 2016	2119	5823	1731	4377	5.9%	0.92 [0.88, 0.97]
Subtotal (95% CI)		6461		5135	13.1%	0.92 [0.88, 0.96]

Total events 2241 1894
Heterogeneity: $\tau^2 = 0.00$; $\chi^2 = 0.06$, $df = 2$ ($P = 0.97$); $I^2 = 0\%$
Test for overall effect: $Z = 3.41$ ($P = 0.0006$)

general

General surgery						
Leahy 2014	56	1276	61	1853	3.7%	1.33 [0.93, 1.90]
Meybohm 2016	2772	13694	2051	9164	5.9%	0.90 [0.86, 0.95]
Subtotal (95% CI)		14970		11017	9.6%	1.05 [0.73, 1.53]

Total events 2828 2112
Heterogeneity: $\tau^2 = 0.06$; $\chi^2 = 4.49$, $df = 1$ ($P = 0.03$); $I^2 = 78\%$
Test for overall effect: $Z = 0.27$ ($P = 0.78$)

other

Other field						
Meybohm 2016	6259	49856	4818	36513	6.0%	0.95 [0.92, 0.99]
Subtotal (95% CI)		49856		36513	6.0%	0.95 [0.92, 0.99]

Total events 6259 4818
Heterogeneity: Not applicable
Test for overall effect: $Z = 2.78$ ($P = 0.005$)

Total (95% CI)		118504		88502	100.0%	0.61 [0.55, 0.68]
Total events	20631		18244			
Heterogeneity: $\tau^2 = 0.05$; $\chi^2 = 598.23$, $df = 24$ ($P < 0.00001$); $I^2 = 96\%$						
Test for overall effect: $Z = 8.75$ ($P < 0.00001$)						
Test for subgroup differences: $\chi^2 = 43.51$, $df = 4$ ($P < 0.00001$), $I^2 = 90.8\%$						

pre PBM

post PBM

Abdominal/
Pediatric,
obstetrics

Total no. of complications

orthopedic

cardio

vascular

general

other

Study or Subgroup	PBM		Control (pre-PBM)		Weight	Risk Ratio M-H, Random, 95% CI	Risk Ratio M-H, Random, 95% CI
	Events	Total	Events	Total			
orthopedic surgery							
Leahy 2017	12	214	5	138	0.7%	1.55 [0.56, 4.30]	
Rineau 2011	14	183	11	184	1.2%	1.28 [0.60, 2.74]	
Kopanidis 2016	17	100	16	100	1.7%	1.06 [0.57, 1.98]	
Loftus 2016	2954	6593	3694	5997	8.8%	0.73 [0.70, 0.75]	*
Holt 2016	5	1010	13	1814	0.7%	0.69 [0.25, 1.93]	
Freedman 2008	23	1127	41	1089	2.3%	0.54 [0.33, 0.90]	
Ma 2014	5	33	3	37	0.4%	1.87 [0.48, 7.22]	
Albinarrate 2015	32	357	39	271	2.8%	0.62 [0.40, 0.97]	
Frew 2016	36	406	97	717	3.6%	0.66 [0.46, 0.94]	
Meybohm 2016	599	16298	464	12633	7.7%	1.00 [0.89, 1.13]	
Leahy 2017	204	8413	266	7282	6.6%	0.66 [0.55, 0.79]	
Subtotal (95% CI)	34734	30262	36.5%	0.78 [0.66, 0.92]			
Total events	3901		4649				
Heterogeneity: $\tau^2 = 0.03$; $\chi^2 = 37.62$, $df = 10$ ($P < 0.0001$); $I^2 = 73\%$							
Test for overall effect: $Z = 2.96$ ($P = 0.003$)							
cardiovascular surgery							
Freedman 2008	16	275	30	274	1.9%	0.53 [0.30, 0.95]	
Brevig 2009	54	479	77	530	4.1%	0.78 [0.56, 1.07]	
Moskowitz 2010	91	586	178	586	5.7%	0.51 [0.41, 0.64]	
Gross 2015	164	2275	43	387	4.2%	0.65 [0.47, 0.89]	
Meybohm 2016	813	7904	618	5630	8.0%	0.94 [0.85, 1.03]	
Leahy 2017	86	949	69	750	4.4%	0.99 [0.73, 1.33]	
Subtotal (95% CI)	12468	8157	28.3%	0.73 [0.56, 0.94]			
Total events	1224		1015				
Heterogeneity: $\tau^2 = 0.08$; $\chi^2 = 29.54$, $df = 5$ ($P < 0.0001$); $I^2 = 83\%$							
Test for overall effect: $Z = 2.46$ ($P = 0.01$)							
vascular surgery							
Meybohm 2016	541	5828	494	4377	7.7%	0.82 [0.73, 0.92]	
Leahy 2017	72	1155	70	986	4.2%	0.88 [0.64, 1.21]	
Subtotal (95% CI)	6983	5363	11.9%	0.83 [0.74, 0.92]			
Total events	613		564				
Heterogeneity: $\tau^2 = 0.00$; $\chi^2 = 0.14$, $df = 1$ ($P = 0.70$); $I^2 = 0\%$							
Test for overall effect: $Z = 3.39$ ($P = 0.0007$)							
general surgery							
Meybohm 2016	748	13694	597	9164	7.9%	0.84 [0.76, 0.93]	
Leahy 2017	235	3736	213	3368	6.6%	0.99 [0.83, 1.19]	
Subtotal (95% CI)	17430	12532	14.5%	0.90 [0.76, 1.06]			
Total events	983		810				
Heterogeneity: $\tau^2 = 0.01$; $\chi^2 = 2.60$, $df = 1$ ($P = 0.11$); $I^2 = 62\%$							
Test for overall effect: $Z = 1.28$ ($P = 0.20$)							
other fields							
Meybohm 2016	4348	49856	3465	36513	8.7%	0.92 [0.88, 0.96]	
Subtotal (95% CI)	49856	36513	8.7%	0.92 [0.88, 0.96]			
Total events	4348		3465				
Heterogeneity: Not applicable							
Test for overall effect: $Z = 3.89$ ($P < 0.0001$)							
Total (95% CI)	121471	92827	100.0%	0.80 [0.74, 0.88]			
Total events	11069		10503				
Heterogeneity: $\tau^2 = 0.02$; $\chi^2 = 143.08$, $df = 21$ ($P < 0.00001$); $I^2 = 87\%$							
Test for overall effect: $Z = 4.75$ ($P < 0.00001$)							
Test for subgroup differences: $\chi^2 = 8.64$, $df = 4$ ($P = 0.07$), $I^2 = 53\%$							
						pre PBM	post

pre PBM

post PBM

Number of units per patients

orthopedic

cardio

vascular

general

other

Study or Subgroup	PBM		Control (pre-PBM)		Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI	
	Mean	SD	Mean	SD				
orthopedic								
Kopanidis 2016	1.1	1.1	0.61	1.6	4.6%	-0.25 [-0.57, 0.07]		
Freedman 2008	3.16	2.04	100	2.05	1.1	100	3.4%	1.11 [0.66, 1.56]
Freedman 2008	0.34	0.83	1127	0.51	1.04	1089	6.7%	-0.17 [-0.25, -0.09]
Ma 2014	0.24	0.83	33	0.81	1.28	37	3.1%	-0.57 [-1.07, -0.07]
Theusinger 2014	0.34	1.25	6721	0.68	1.68	2150	6.7%	-0.34 [-0.42, -0.26]
Meybohm 2016	1.24	0.16	16298	1.46	0.16	12633	6.9%	-0.22 [-0.22, -0.22]
Leahy 2017	0.16	0.75	11942	0.27	1.02	10304	6.9%	-0.11 [-0.13, -0.09]
Subtotal (95% CI)		36411		26440	38.3%	-0.18 [-0.26, -0.09]		
Heterogeneity: $\tau^2 = 0.01$; $\chi^2 = 125.65$, $df = 6$ ($P < 0.00001$); $I^2 = 95\%$								
Test for overall effect: $Z = 4.14$ ($P < 0.0001$)								
cardio								
Freedman 2008	1.16	2.03	275	2.01	2.7	274	3.8%	-0.85 [-1.25, -0.45]
Brevig 2009	0.53	1.82	479	1.43	3.08	530	4.7%	-0.90 [-1.21, -0.59]
Gross 2015	0.61	1.57	2275	1.28	2.34	387	5.4%	-0.67 [-0.91, -0.43]
Meybohm 2016	3.68	0.46	7904	4.65	0.46	5630	6.9%	-0.97 [-0.99, -0.95]
Leahy 2017	0.65	2.15	1431	1.46	3.67	1096	5.3%	-0.81 [-1.05, -0.57]
Subtotal (95% CI)		12364		7917	26.1%	-0.87 [-1.00, -0.74]		
Heterogeneity: $\tau^2 = 0.01$; $\chi^2 = 8.02$, $df = 4$ ($P = 0.09$); $I^2 = 50\%$								
Test for overall effect: $Z = 12.91$ ($P < 0.00001$)								
vascular								
Freedman 2008	1.75	3.82	232	2.12	4.39	287	2.0%	-0.37 [-1.08, 0.34]
Meybohm 2016	3.77	0.52	5823	4.9	0.53	4377	6.9%	-1.13 [-1.15, -1.11]
Leahy 2017	0.38	1.51	1870	0.77	2.43	1760	6.4%	-0.39 [-0.52, -0.26]
Subtotal (95% CI)		7925		6424	15.2%	-0.66 [-1.29, -0.04]		
Heterogeneity: $\tau^2 = 0.27$; $\chi^2 = 121.12$, $df = 2$ ($P < 0.00001$); $I^2 = 98\%$								
Test for overall effect: $Z = 2.08$ ($P = 0.04$)								
general								
Meybohm 2016	2.09	0.16	13694	2.39	0.16	9164	6.9%	-0.30 [-0.30, -0.30]
Leahy 2017	0.31	1.59	5437	0.49	2.17	4512	6.7%	-0.18 [-0.26, -0.10]
Subtotal (95% CI)		19131		13676	13.6%	-0.25 [-0.36, -0.13]		
Heterogeneity: $\tau^2 = 0.01$; $\chi^2 = 9.52$, $df = 1$ ($P = 0.002$); $I^2 = 89\%$								
Test for overall effect: $Z = 4.13$ ($P < 0.0001$)								
other								
Meybohm 2016	1.4	4.8	49856	1.7	5.6	36513	6.7%	-0.20 [-0.27, -0.13]
Subtotal (95% CI)		49856		36513	6.7%	-0.20 [-0.27, -0.13]		
Heterogeneity: Not applicable								
Test for overall effect: $Z = 5.50$ ($P < 0.00001$)								
Total (95% CI)		125687		90970	100.0%	-0.43 [-0.54, -0.31]		
Heterogeneity: $\tau^2 = 0.05$; $\chi^2 = 15280.28$, $df = 17$ ($P < 0.00001$); $I^2 = 100\%$								
Test for overall effect: $Z = 7.08$ ($P < 0.00001$)								
Test for subgroup differences: $\chi^2 = 88.79$, $df = 4$ ($P < 0.00001$), $I^2 = 95.5\%$								

pre PBM

post PBM

pre PBM

post PBM

Length of hospital stay

orthopedic

Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
Orthopedic surgery									
Rineau 2016	9.28	2.7	183	9.35	2.5	184	6.7%	-0.07 [-0.60, 0.46]	
Loftus 2016	3.02	1.76	6593	3.32	1.91	5997	12.7%	-0.30 [-0.36, -0.24]	
Freedman 2008	6.25	18.2	1127	7.16	7.51	1089	2.4%	-0.91 [-2.06, 0.24]	
Ma 2014	12.4	1.2	33	13.2	1.5	37	5.6%	-0.80 [-1.43, -0.17]	
Albinarrate 2015	9.1	3.8	357	10.1	3.8	271	5.9%	-1.00 [-1.60, -0.40]	
Meybohm 2016	12.85	16.3	16298	12.98	16.03	12633	8.8%	-0.13 [-0.51, 0.25]	
Leahy 2017	5.89	8.74	11942	6.39	8.86	10304	10.9%	-0.50 [-0.73, -0.27]	
Subtotal (95% CI)			36533			30515	52.9%	-0.41 [-0.60, -0.22]	
Heterogeneity: $\tau^2 = 0.03$; $\chi^2 = 12.66$, $df = 6$ ($P = 0.05$); $I^2 = 53\%$									
Test for overall effect: $Z = 4.16$ ($P < 0.0001$)									

cardio

Cardiac surgery									
Freedman 2008	7.81	7.55	275	10.78	10.4	274	1.5%	-2.97 [-4.49, -1.45]	
Gross 2015	10.4	8	2275	12.2	9.6	387	3.0%	-1.80 [-2.81, -0.79]	
Meybohm 2016	16.6	19.4	7904	16.95	19.8	5630	5.2%	-0.35 [-1.02, 0.32]	
Leahy 2017	11.07	11.64	1431	11.9	11.74	1096	3.4%	-0.83 [-1.75, 0.09]	
Subtotal (95% CI)			11885			7387	13.1%	-1.34 [-2.34, -0.34]	
Heterogeneity: $\tau^2 = 0.77$; $\chi^2 = 12.58$, $df = 3$ ($P = 0.006$); $I^2 = 76\%$									
Test for overall effect: $Z = 2.63$ ($P = 0.009$)									

vascular

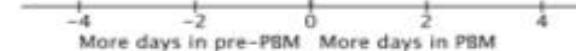
Vascular surgery									
Freedman 2008	8.07	23.4	232	12.91	27.5	287	0.2%	-4.84 [-9.22, -0.46]	
Meybohm 2016	18.13	25.1	5828	18.68	24.98	4377	3.1%	-0.55 [-1.53, 0.43]	
Leahy 2017	9.16	12.91	1870	9.9	13.02	1760	3.9%	-0.74 [-1.58, 0.10]	
Subtotal (95% CI)			7930			6424	7.2%	-0.85 [-1.83, 0.13]	
Heterogeneity: $\tau^2 = 0.31$; $\chi^2 = 3.51$, $df = 2$ ($P = 0.17$); $I^2 = 43\%$									
Test for overall effect: $Z = 1.70$ ($P = 0.09$)									

general

General surgery									
Meybohm 2016	14.38	21.33	13694	14.58	19.58	9164	6.6%	-0.20 [-0.74, 0.34]	
Leahy 2017	6.97	10.06	5437	6.49	8.79	4512	8.9%	0.48 [0.11, 0.85]	
Subtotal (95% CI)			19131			13676	15.5%	0.17 [-0.50, 0.83]	
Heterogeneity: $\tau^2 = 0.18$; $\chi^2 = 4.17$, $df = 1$ ($P = 0.04$); $I^2 = 76\%$									
Test for overall effect: $Z = 0.50$ ($P = 0.62$)									

other

Other fields									
Meybohm 2016	10.7	14.1	49856	11.1	14.6	36513	11.4%	-0.40 [-0.59, -0.21]	
Subtotal (95% CI)			49856			36513	11.4%	-0.40 [-0.59, -0.21]	
Heterogeneity: Not applicable									
Test for overall effect: $Z = 4.04$ ($P < 0.0001$)									
Total (95% CI)			125335			94515	100.0%	-0.45 [-0.65, -0.25]	
Heterogeneity: $\tau^2 = 0.08$; $\chi^2 = 57.74$, $df = 16$ ($P < 0.00001$); $I^2 = 72\%$									
Test for overall effect: $Z = 4.45$ ($P < 0.00001$)									
Test for subgroup differences: $\chi^2 = 7.02$, $df = 4$ ($P = 0.13$), $I^2 = 43.0\%$									



What is anemia?

- Hgb < 130g/L m
- Hgb <120 g/L F*
- Mild to moderate: 100-129 g/L
- Moderate to severe: <100 g/L
- **ANEMIA means – low O₂ carrying capacity AND inability to meet body physiological needs for quantity of O₂ of the tissues**
- WHO (1968-2011)



Eur. J Anesthesiology 2017; 34: 332-395

- If anemia is present, we recommend identifying the cause. **1C**

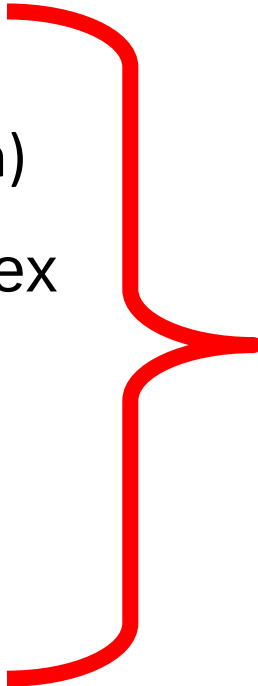
holly trinity of anemia

- **Iron deficiency**
- **Renal disease**
- **Chronic inflammation**

PREOPERATIVE : SCREENING OF ANEMIA

Laboratory Tests

- Complete Blood Cell Count
- Iron Status (Iron, **Ferritin**, Transferrin saturation)
- Serum sol transferrin receptors –ferritin index (<1>2)
- Inflammation Marker
- Vitamin B12, Folic Acid (>65 years old)



**30 days
before
scheduled
operation**

Review Article

Pre-operative haematological assessment in patients scheduled for major surgery

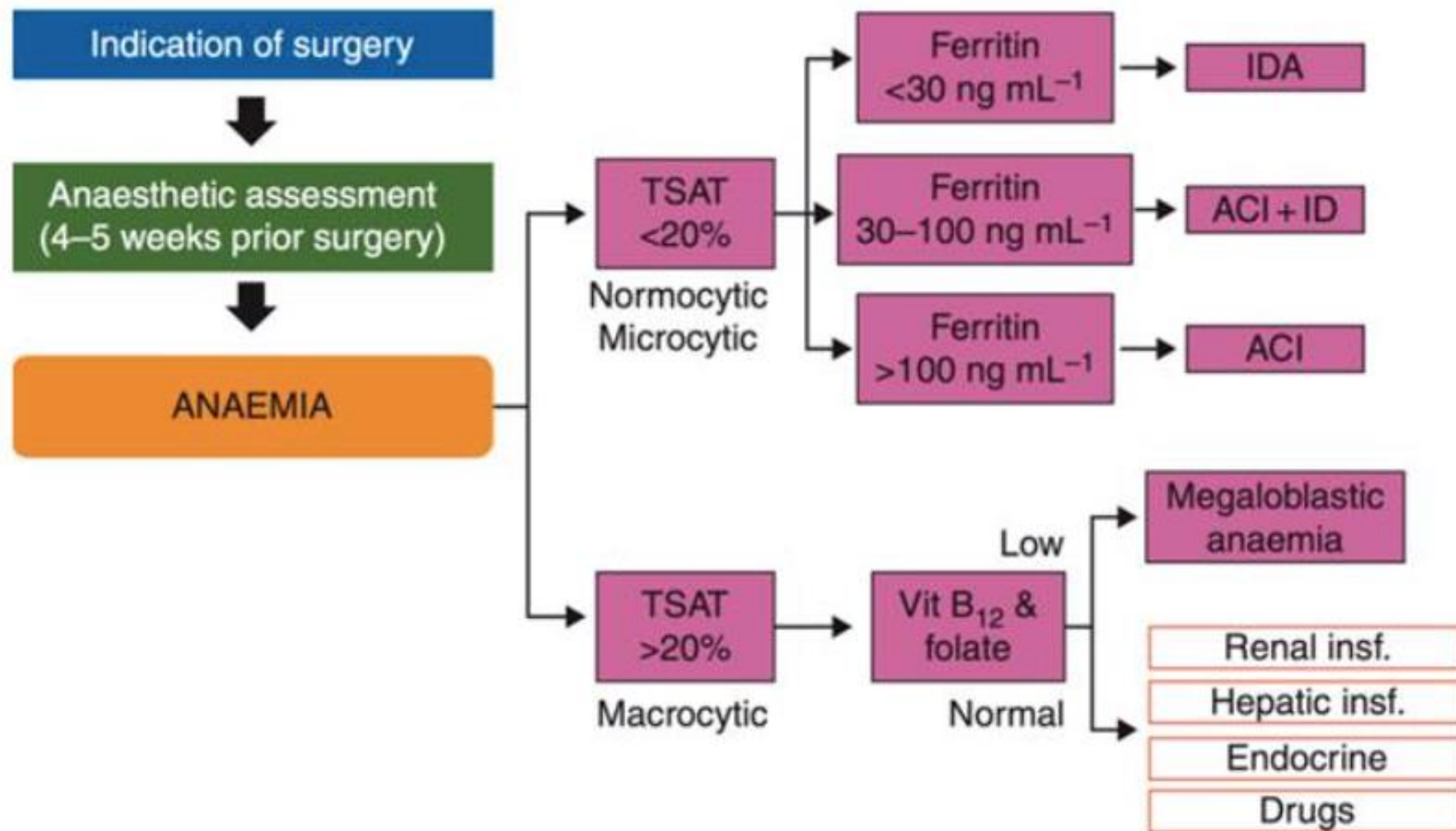
M. Muñoz,¹ S. Gómez-Ramírez² and S. Kozek-Langeneker³¹ Professor, Peri-operative Transfusion Medicine, School of Medicine, University of Málaga, Málaga, Spain² Consultant, Department of Internal Medicine, Xanit International Hospital, Benalmádena, Spain³ Professor, Department of Anaesthesia and Intensive Care, Evangelical Hospital, Vienna, Austria

Summary

Peri-operative anaemia, blood loss and allogeneic blood transfusion are associated with increased postoperative morbidity and mortality, and prolonged hospital stay. A multidisciplinary, multimodal, individualised strategy, collectively termed 'patient blood management', may reduce or eliminate allogeneic blood transfusion and improve outcomes. This approach has three objectives: the detection and treatment of peri-operative anaemia; the reduction of peri-operative bleeding and coagulopathy; and harnessing and optimising the physiological tolerance of anaemia. This review focuses on the pre-operative evaluation of erythropoiesis, coagulation status and platelet function. Where possible, evidence is graded systematically and recommended therapies follow recently published consensus guidance.

Diagnosis

- HgB, MCV
- Serum iron-TIBC
- **Serum ferritin (30-100micg/L)- most specific**
- Transferrin saturation(TSAT) (20 %)
- Serum soluble transferrin receptors (sTRF)(<2,4,5)
- (not affected by inflammation)
- Serum sol transferrin receptpors –ferritin index (<1>2)
- Ferritin (Inflamat <100; CHF <300; CKD<500, cancer <800) Tsat <20%



FE Metabolism basics

Iron, transferrin should be bounded

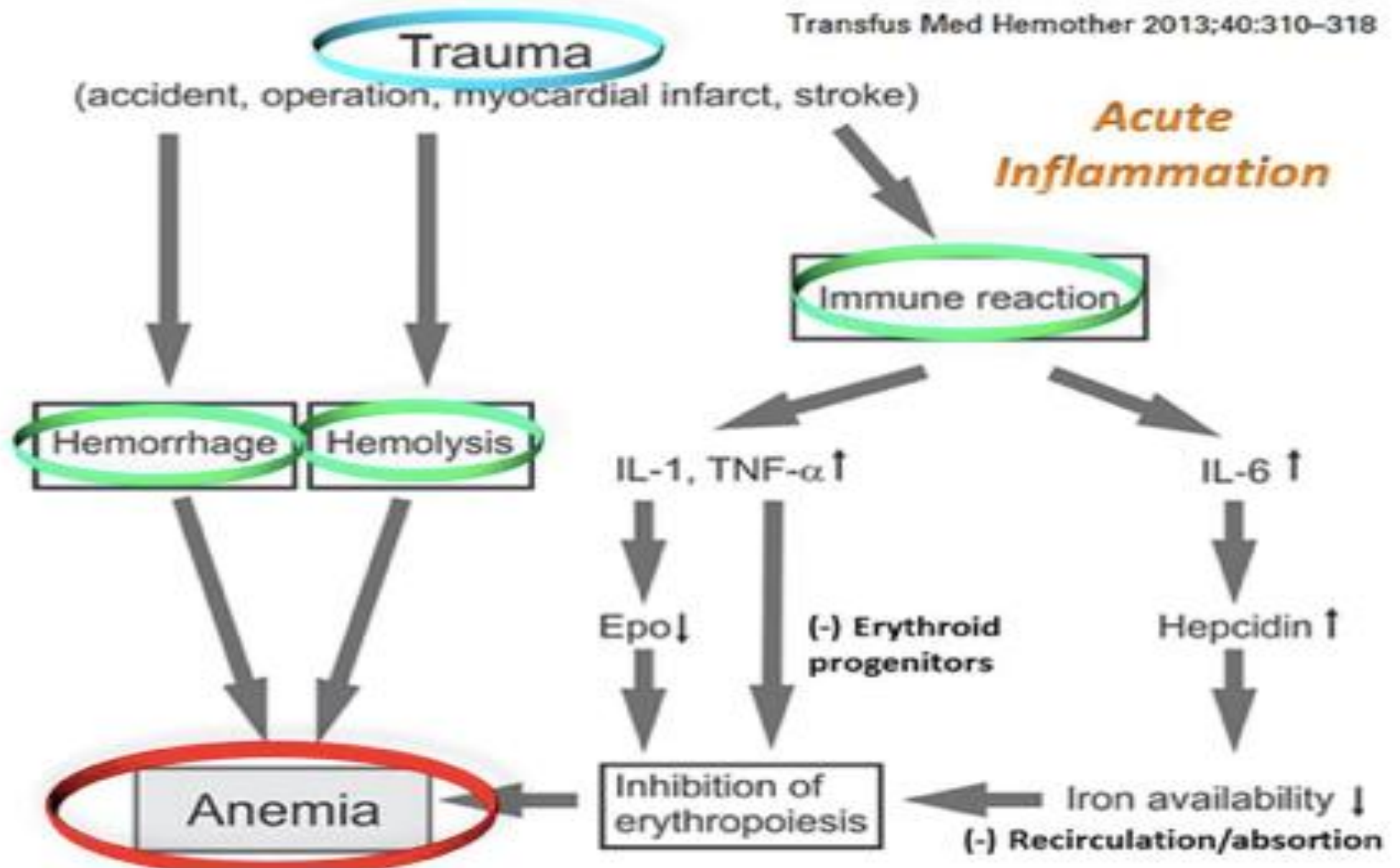
Fe, from nutrition or recycled

functional pool (heart, muscles- non erythroid, bone marrow, RBC);storage pool (liver)

Fe, needed for RBC renewal 20-30 mg,1-2 mg absorbed- lost

Fe, Ferrous form transported through FERROPORTIN

HEPCIDIN- PEPTIDE (HEPATOCITES – IN RESPONSE OF INLTAMATION) DYSREGULATES IRON



Challenge is to implement anemia management pathways

BJA

British Journal of Anaesthesia 115 (1): 15–24 (2015)

doi: 10.1093/bja/aev165
Review Articles

REVIEW ARTICLES



‘Fit to fly’: overcoming barriers to preoperative haemoglobin optimization in surgical patients[†]

M. Muñoz^{1,*}, S. Gómez-Ramírez³, S. Kozek-Langeneker⁴, A. Shander⁵, T. Richards⁶, J. Pavía², H. Kehlet⁷, A. G. Acheson⁸, C. Evans⁹, R. Raobaikady¹⁰, M. Iavidroozi⁵ and M. Auerbach¹¹

BJA

British Journal of Anaesthesia 115 (1): 15–24 (2015)

doi: 10.1093/bja/aev165
Review Articles

REVIEW ARTICLES



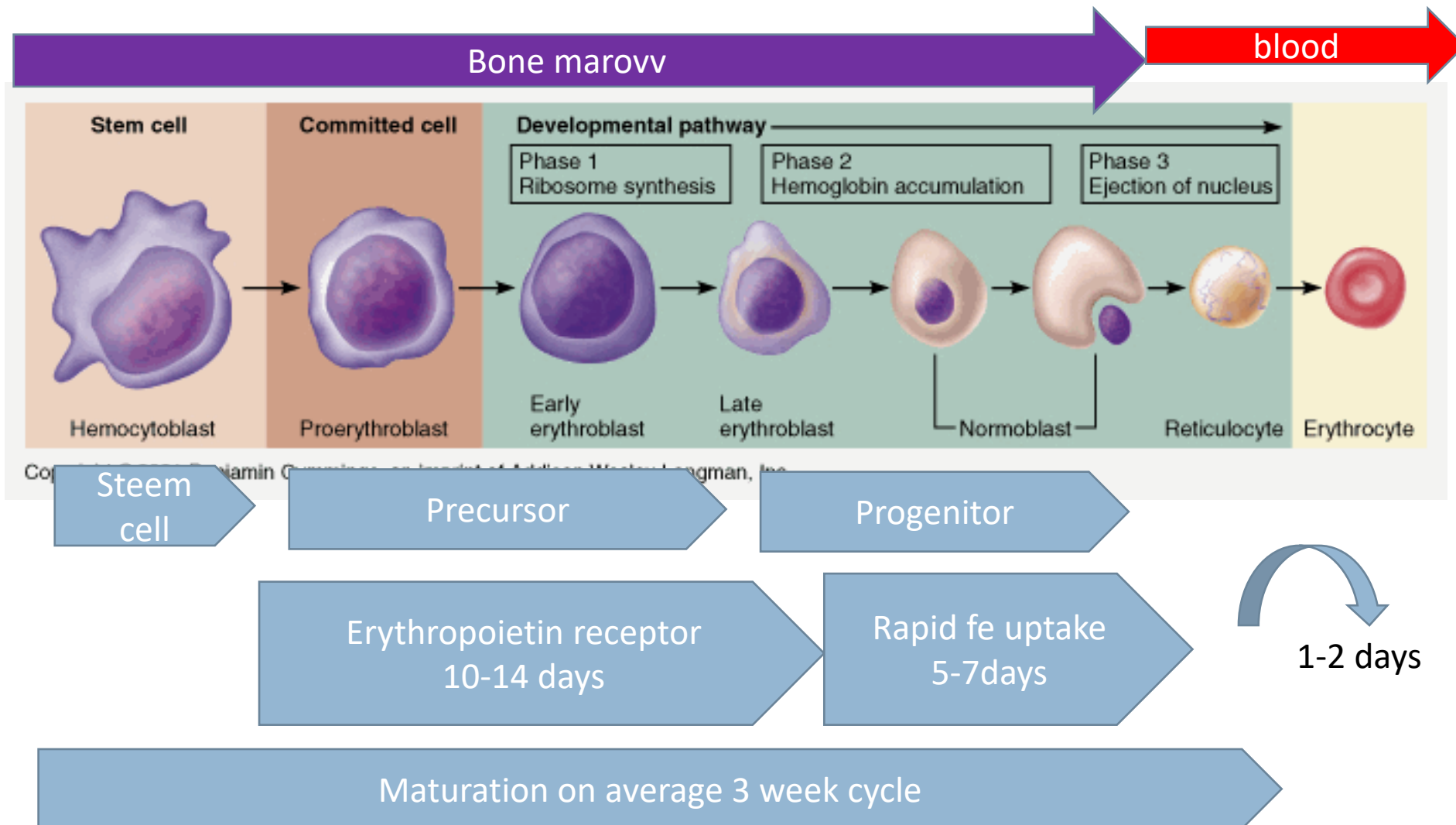
‘Fit to fly’: overcoming barriers to preoperative haemoglobin optimization in surgical patients[†]

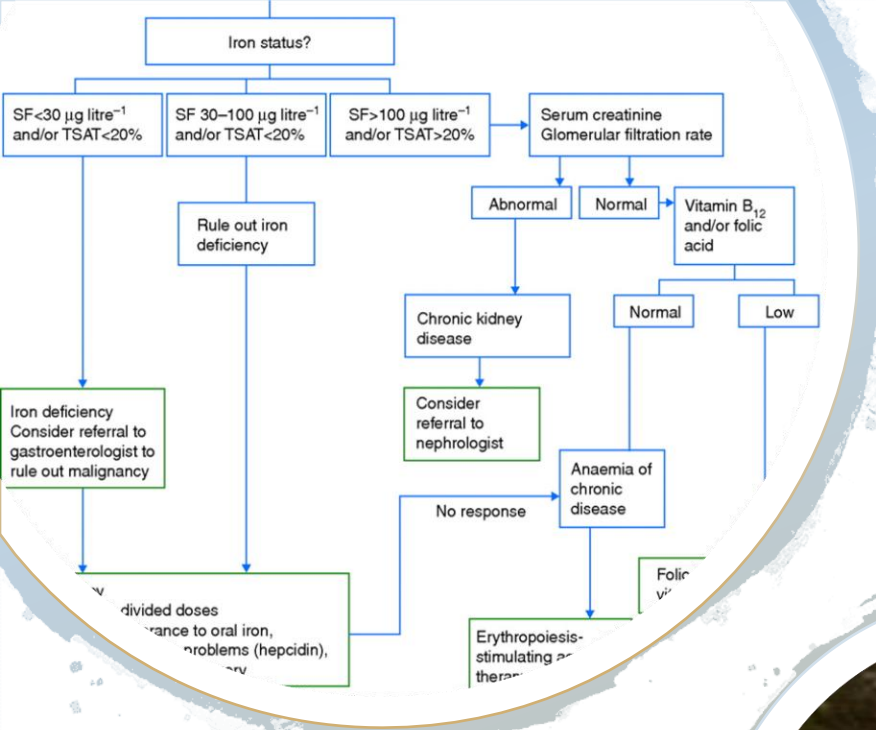
M. Muñoz^{1,*}, S. Gómez-Ramírez³, S. Kozek-Langeneker⁴, A. Shander⁵, T. Richards⁶, J. Pavía², H. Kehlet⁷, A. G. Acheson⁸, C. Evans⁹, R. Raobaikady¹⁰, M. Iavidroozi⁵ and M. Auerbach¹¹

Eur. J Anesthesiology 2017; 34: 332-395

- We recommend that patients at risk of bleeding are assessed for anemia 3 to 8 weeks before surgery. **1C**
- In non-cancer patients with preoperative anemia scheduled for elective major surgery, we recommend **postponing surgery until anemia has been corrected. 1C**
- **Time Supporting hematopoiesis (iron, folic ESAS)**

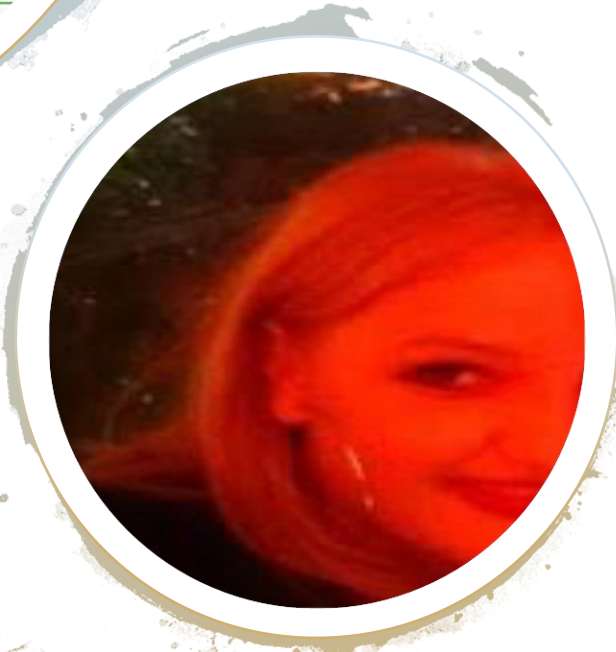
Red cell maturation





HELP us Celebrities

- Venus williams
- Selena Gomez
- Angelina Jolie
- Brittany Murphy





- Esa is in line with NATA

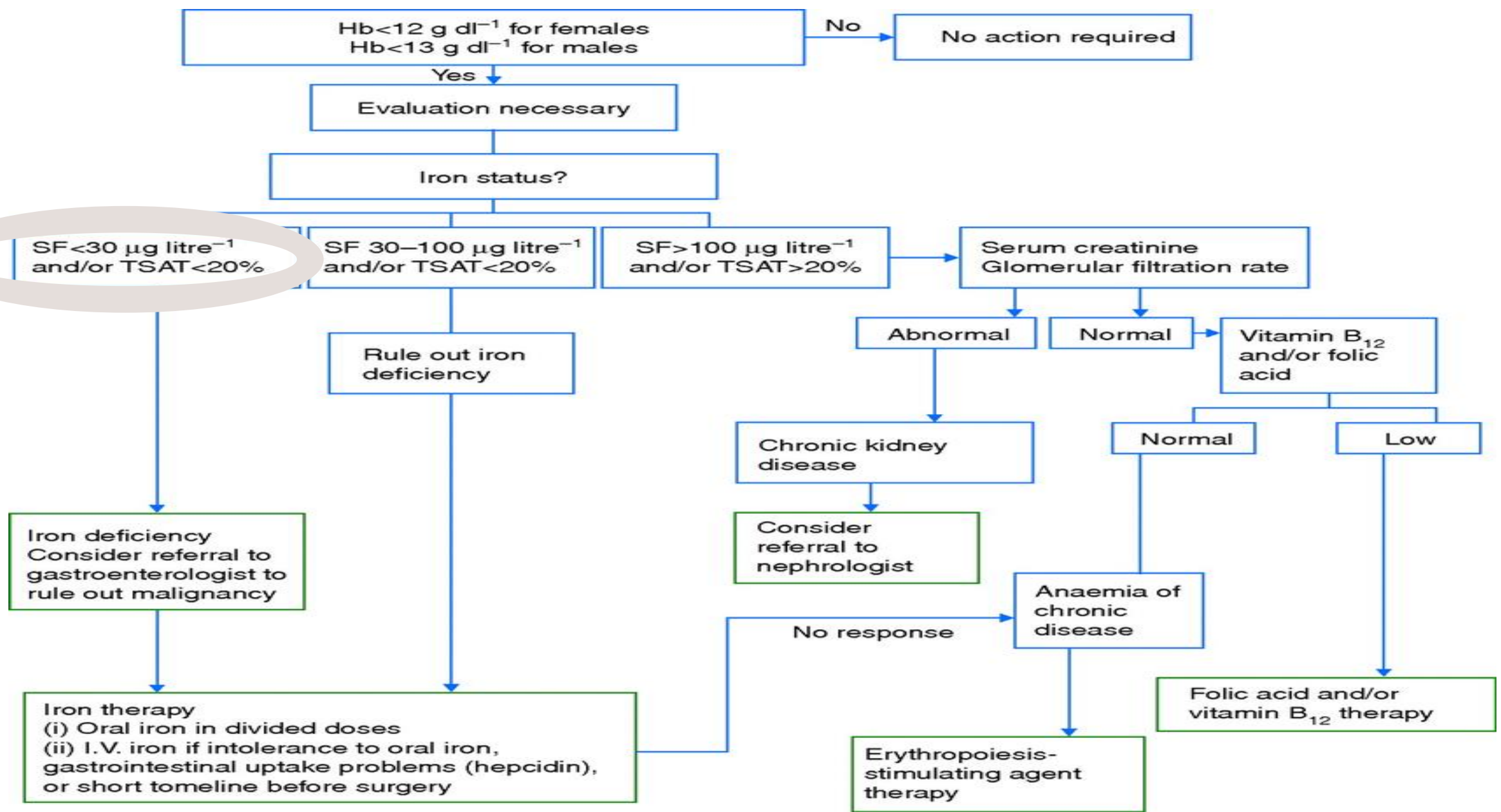
Eur. J Anesthesiology 2017; 34: 332-395

- We recommend treating iron deficiency with iron supplementation. **1B-**
- We recommend the use of intravenous iron in preference to oral iron. **1C-**

Munoz M Management of Preoperative anemia :NATA consensus statement ISBP Sci Ser 2012;7:283-287

Bruce et al Practical recommendation for patient blood management to reduce of perioperative transfusion in joint replacement

SF < 30 $\mu\text{g litre}^{-1}$
and/or TSAT < 20%



	Ferumoxyl	Iron Carboxymaltose	Iron Isomaltoside 1000	Low Molecular Weight Iron Dextran	Iron Succinate	Iron Glucamate
Brand name	Feraheme [®]	Deriject [®]	Monoferr [®]	Conosfer [®]	Venosfer [®]	Derlijet [®]
Maximum single dose	510 mg	1000 mg	20 mg/kg	20 mg/kg	200 mg	125 mg
Minimum administration time (minutes)	15	15	15	60	30	30-60
Replacement dose possible in a single infusion	No	Yes	Yes	Yes	No	No

Oral Iron therapy

Is cheap, takes >8 weeks

Pitfalls: not predictable, GI side effects, bioavailability is changed (H. Pylori)

Doses 40-60 mg /day, without food, interfered with drugs and minerals

Fe, Ferrous form- HAEM form is better choice

Hepcidin levels block it, controversies : does it rise its levels

Blood 2015: 126:1981-1989
Cochrane Library: database of systematic reviews 2016

IV IRON THERAPY

EXPENSIVE , BUT MORE EFFECTIVE

Decrease the need for ABT and mortality rates
increases Hbg, decreases postop anemia

Novel preparations -without high molecular dextran's ,less- adverse effects

Benefits overcome risks

Recommended where time to surgery is short, no other treatment has
results 1 B

Lancet Haematol 2018;5:310-20

PLOS ONE 2015: 10:E011783

Vox Sanguinis 2015;257-266

Transfusion 2014;54:113-1145

Pragmatic approach „global correction“

- 1 g isomaltosid IV
- + 40.000 U erythropoietin alpha s.c.
- + 1 mg/day vitamin B12 p.o.
- + 5 mg/day folate p.o.



pharmaceuticals



Review

Intravenous Irons: From Basic Science to Clinical Practice

Sunil Bhandari ¹ , Dora I. A. Pereira ^{2,3}, Helen F. Chappell ⁴ and Hal Drakesmith ^{5,6,*}

Eur. J Anesthesiology 2017; 34: 332-395

- If other causes of anemia have been excluded or treated
we suggest erythropoiesis stimulating agents .

2C

Erythropoiesis stimulating agents

WHO list of essential medicines, in patient ACD (>500 ml blood lost),
Hgb <13

Epoetin alpha, Darbeopetin, Erythropoietin biosimilars

Increases red cell mass

Reduces ABT, not in hypertensive , increases the risk for DVT, cancer

Controversies for to much still going on

Lancet Haematol 2018;5:310-20
Vox Sanguinis 2015;257-266

Eur. J Anesthesiology 2017; 34: 332-395

- In patients with preoperative anemia, **we recommend the use of combined therapy with intravenous iron and erythropoietin along with a restrictive transfusion policy. 1C**
- **If autologous blood donation is performed suggest treatment with Iv iron, ESA to avoid preoperative anemia and transfusion rates. 2C**

Anemia Treatment

Safely increase Hgb 0.5-1 gm/week

- Treatment Options: IV iron, enteric iron, Erythropoietin stimulating agents (ESA's), Vitamin B-12, folate, treat co-morbidities, delay and defer
- IV iron, effect at 1 week, max effect at 2 weeks,
- several doses needed (1 gram divided dose common)
- ESA action onset 4-6 days, max at 10 days, must
- give with iron
- Pre- OP-One week treatment “lead time” needed, Post-OP start ASAP.

Ultra short combination treatment, IV iron, B12, ESA

- Single dose intravenous iron for iron deficiency: new paradigm

Haemathology 2016

Fast track anemia clinic in the Emergency department

Blood Transfusions 2016;14;126-133

Effect of ultra short term treatment of patients with iron deficiency anemia or anemia undergoing cardiac surgery a prospective trial

Lancet 2019 393;2201-2212

- Perioperative

-
- *Fluid management*
 - *T*
 - *Ph*
 - *Prolene*
 - *Cell salvage*
 - Antifibrinolytics

- Postoperative anemia



Eur. J Anesthesiology 2017; 34: 332-395

- If patients are anemic following surgery we suggest use of intravenous iron post op. **2C**

- Postoperative anemia

Eur. J Anesthesiology 2017; 34: 332-395

- Hemolysis,
- Blood loss, low EPO stores,
- Hepcidin levels,
- Nutritional or
- Hemodilution

- Postoperative anemia

EBM recommendation

- Ferritin levels
- Insufficient Iron stores and Ferritin <100
- We recommend a target Hb of 7-9 g/dl during active bleeding.**1C**

Restrictive transfusion criteria's for postoperative bleeding

Hb level (g/dl)	Clinical criteria: • ability to compensate anaemia? signs of hypoxia? • risk factors: comorbidity? • relevant postOP bleeding?	Decision for pRBC transfusion
< 6		yes (1-2 pRBCs)
6 - 8	compensation adequate no risk factors no relevant postOP bleeding	no
	compensation recuded e.g. ST-segment dynamics, tachycardia > 80 bpm, hypotension, lactate acidosis risk factors present: e.g.. CHD, cardiac failure, stroke, renal dysfunction	yes
8 - 10	compensation adequate	no
	compensation reduced: e.g. ST-segment dynamics, tachycardia > 80 bpm, hypotension, lactate acidosis	yes
> 10		no



Proposed mechanism of action of the erythroid regulator ERFE

BLOOD, 24 JULY 2014 • VOLUME 124, NUMBER 4



Hemodialysis International

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Supplement Article

Role of hepcidin-ferroportin axis in the pathophysiology, diagnosis, and treatment in anemia of chronic inflammation

Arielle L. Langer, Yelena Z. Ginzburg

First published: 22 March 2017 Full publication history

DOI: 10.1111/hdi.12543 View/Save Citation

Early View



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Online Version of Record
published before inclusion
in an issue

Iron man could cure
anemia.



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- **Total iron-binding capacity.** The transferrin molecule binds free **iron** and transports it in the blood. **Total iron-binding capacity** (TIBC) or sometimes transferrin **iron-binding capacity** is a medical laboratory test that measures the blood's **capacity** to **bind iron** with transferrin.

- **Transferrin saturation**, measured as a percentage, is a medical laboratory value. It is the value of serum **iron** divided by the total **iron**-binding capacity. ... For instance, a value of 15% means that 15% of **iron**-binding sites of **transferrin** are being occupied by **iron**. The three results are usually reported together.

Blood transfusion overuse

- World health Organisation resolution in 2012 identified patient blood management as one of the most important issues in the care of surgical patients worldwide
- WHO 2010 RECOGNIZED PBM AS A MEANS TO promote the availability of transfusion alternatives
- [Http://apps.who.int/ebwha](http://apps.who.int/ebwha)