





# Patient Blood Management

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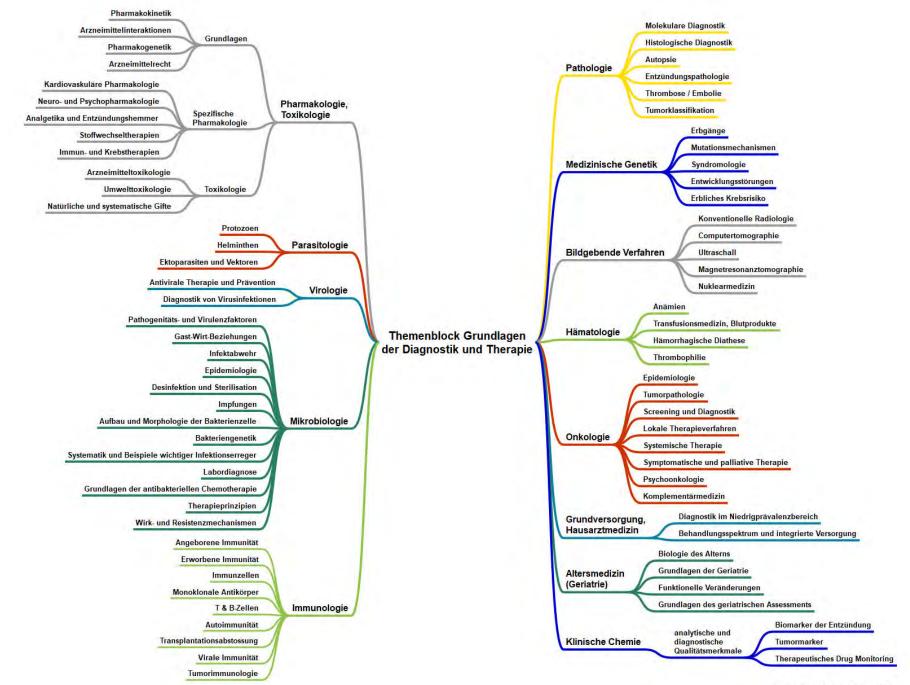
UZH - Themenblock Grundlagen der Diagnostik und Therapie, 13.11.2024

#### Interessenskonflikte

- AstraZeneca, Pharmacosmos: Advisory Board
- Bayer, CSL Behring, AstraZeneca: Honorare für Vorträge
- Baxter, Pharmacosmos, CSL Behring: Reise- und Fortbildungsgebühren



#### **Mindmap**





#### **Lernziele - Patient Blood Management**

- Sie können die 3 Säulen des Patient Blood Managements benennen.
- Sie können die Häufigkeit und die Bedeutung der präoperativen Anämie wiedergeben.
- Sie können die Behandlungsprinzipien der präoperativen Anämie-Behandlung beschreiben.
- Sie können die 4 wichtigsten Outcome-Benefits des Patient Blood Managements benennen.



## WHAT IS PATIENT BLOOD MANAGEMENT?

#### **Patient Blood Management**

(PBM) is the timely application of evidence-based medical and surgical concepts designed to maintain hemoglobin concentration, optimize hemostasis and minimize blood loss in an effort to improve patient outcomes.







## Die 3 Säulen des PBM

#### Optimierung des Erythrozytenvolumens

Unter anderem Diagnostik und Behandlung einer präoperativen Anämie, zeitliche Planung des operativen Eingriffs, ggf. postoperative Stimulation der Erythropoese

## Minimierung von Blutverlusten & Blutungen

Unter anderem Minimierung des diagnostischen und des interventionellen Blutverlusts, exakte Blutstillung

# Nutzung & Optimierung der physiologischen Anämie-Reserve

Unter anderem strenge Indikationsstellung zur Bluttransfusion (restriktives Transfusionsregime)





#### POLICY BRIEF

# THE URGENT NEED TO IMPLEMENT PATIENT BLOOD MANAGEMENT

#### This policy brief aims to:

- create awareness about the enormous, but greatly under-appreciated global disease burden of iron deficiency, anaemia, blood loss and bleeding disorders;
- create a sense of urgency for health care entities to implement PBM, a systematic, multidisciplinary, multiprofessional concept to routinely minimize these risk factors, and, in so doing, significantly and cost-effectively improve health and clinical outcomes for hundreds of millions of medical and surgical patients, pregnant women, neonates, children, adolescents, elderly people, and the population as a whole;
- announce the upcoming World Health Organization (WHO) initiative to develop PBM Implementation Guidelines that will serve as a framework for health care leaders of all Member States;

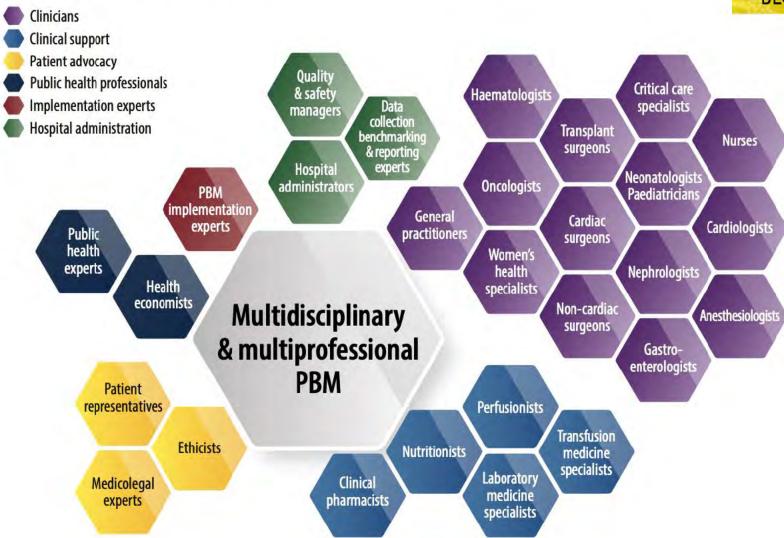




#### POLICY BRIEF

THE URGENT NEED TO IMPLEMENT PATIENT BLOOD MANAGEMENT

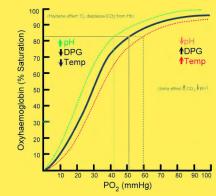


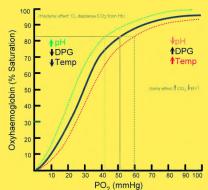


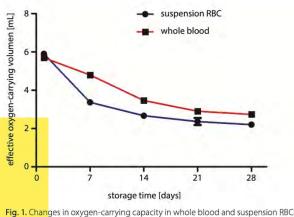


#### Probleme der EC-Transfusion

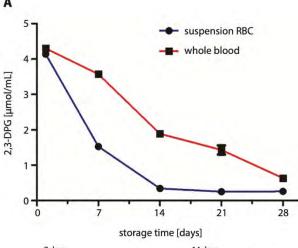
- Allergische Reaktionen
- Fieber und Schüttelfrost
- Infektionen
- TRALI
- **TACO**
- **Immunmodulation**
- Nierenschädigung (AKI)
- Hämolyse → freies Hämoglobin: irreversible Endothelschäden
- Geldrollenphänomen → gestörte Mikrozirkulation
- **Thromboembolien**











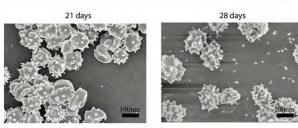
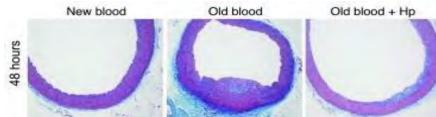


Fig. 3. Changes in erythrocyte morphology after different storage times. The scale bar was 100 nm



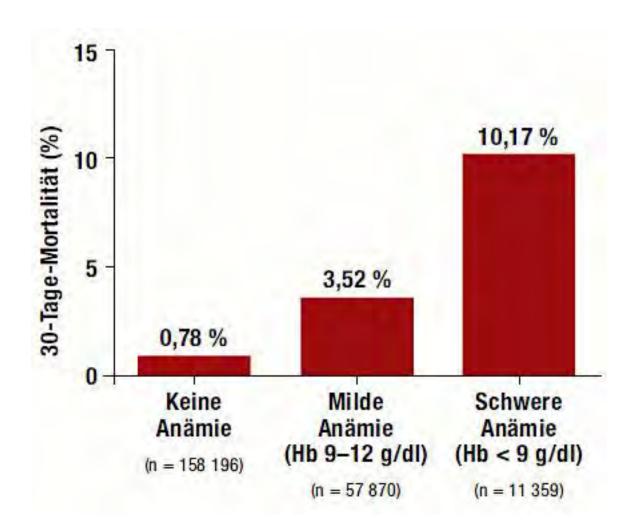




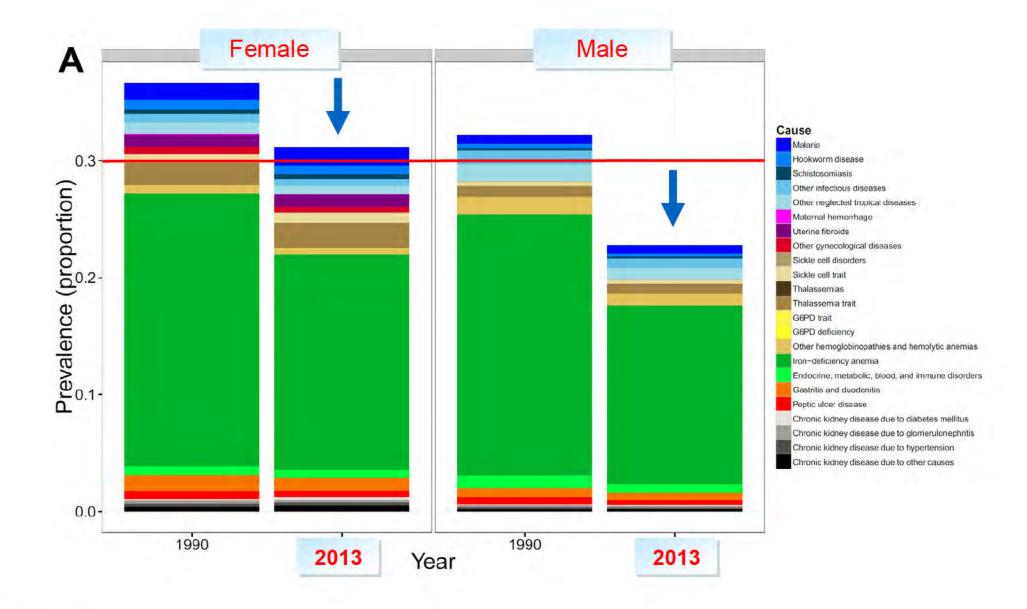
## 1. Säule: Präoperatives Anämie Screening und Behandlung



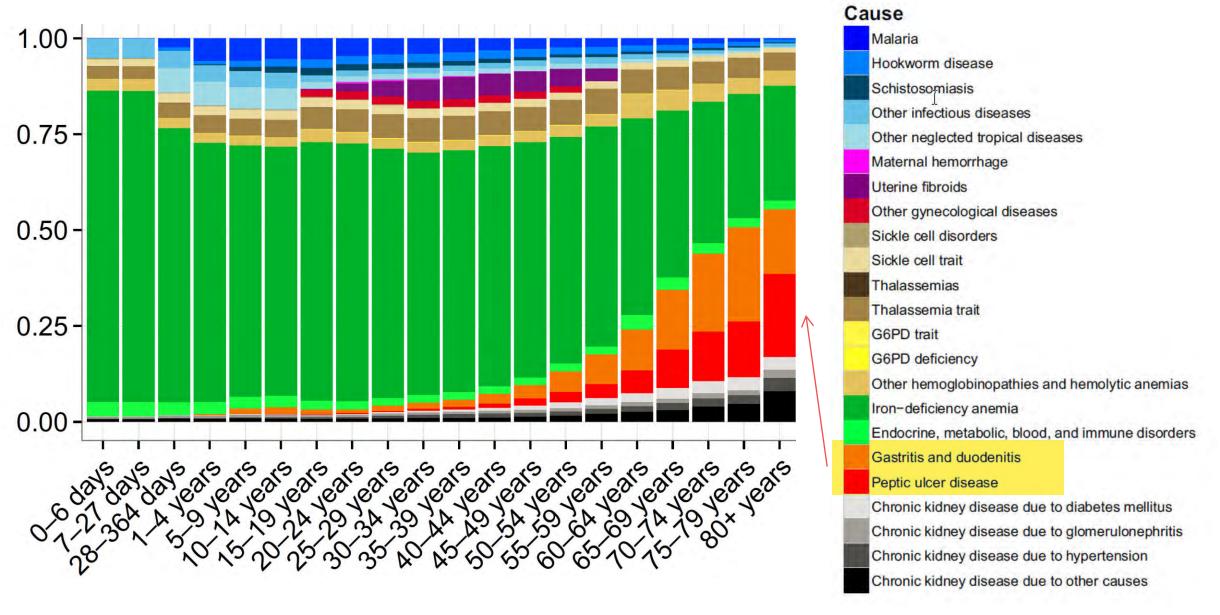
# Einfluss einer präoperativen Anämie auf die 30-Tage-Mortalität bei nicht herzchirurgischen Patienten











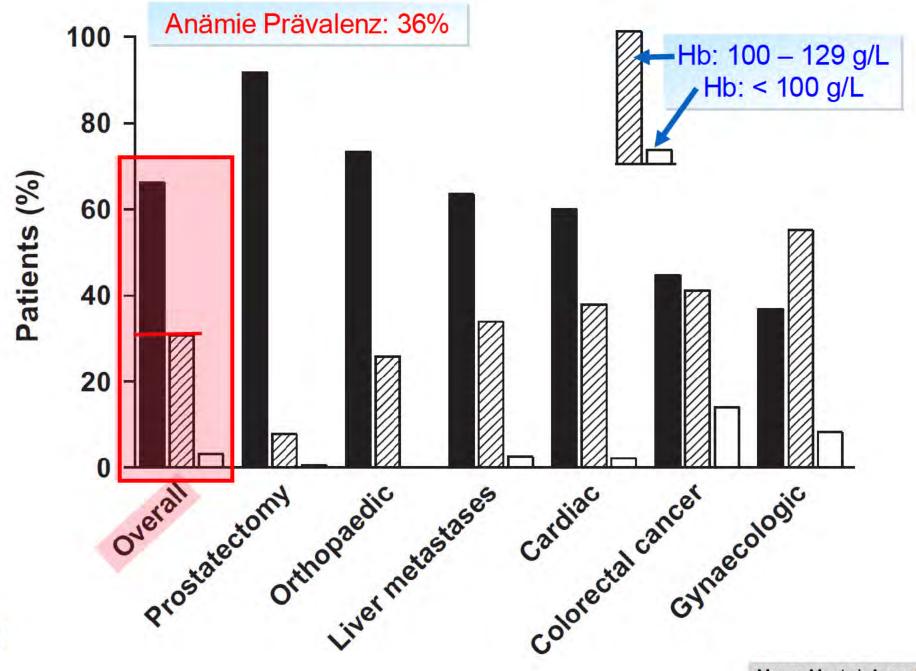


# Original Article

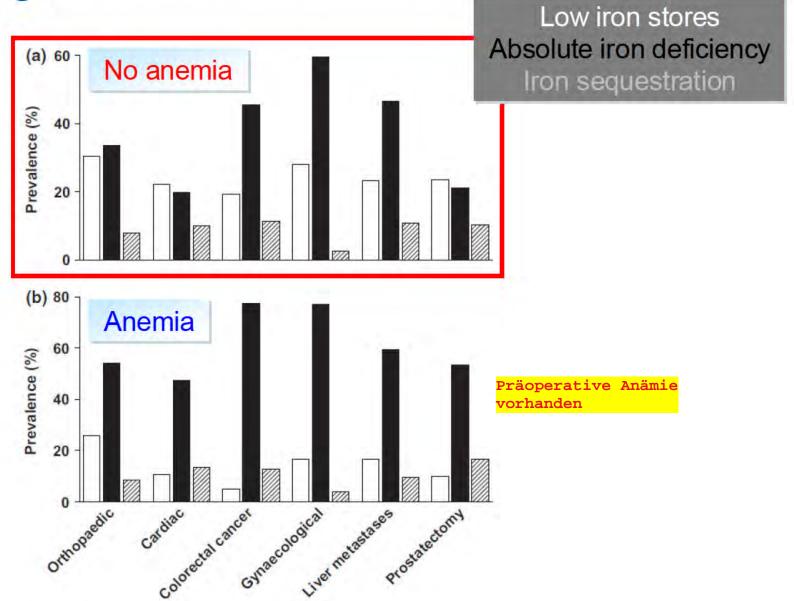
Pre-operative haemoglobin levels and iron status in a large multicentre cohort of patients undergoing major elective surgery\*

- 3'342 patients scheduled for orthopedic, cardiac, colon, prostatic, gynecological or liver (metastatic) surgery representing the focus group
  - ⇒ Expected RBC transfusion rate > 10%
  - ⇒ Expected blood loss > 500 ml
- Anemia definition: Hb < 130 g/L (male and female)</li>
- Iron deficiency (data available in 2'884 patients)
  - ⇒ Absolute iron deficiency: Ferritin < 30 ng/ml or < 100 ng/ml and (TSAT < 20% or CRP > 5 mg/l)
  - ⇒ Low iron store: Ferritin <100 ng/ml and TSAT > 20%
  - ⇒ Iron sequestration: Ferritin > 100 ng/ml and TSAT < 20%





#### **Eisenmangel**





# Iron deficiency is associated with higher mortality in patients undergoing cardiac surgery: a prospective study

- 730 patients undergoing cardiac surgery
- Iron deficiency = ferritin < 100 ng/ml</li>
- Anemia
  - ⇒ Hb < 130 g/l in man
  - ⇒ Hb < 120 g/l in women
- Outcome

  - ⇒ SAEs

  - ⇒ Transfusion of RBC, FFP, platelets
  - ⇒ LOS



#### 3-fach ★

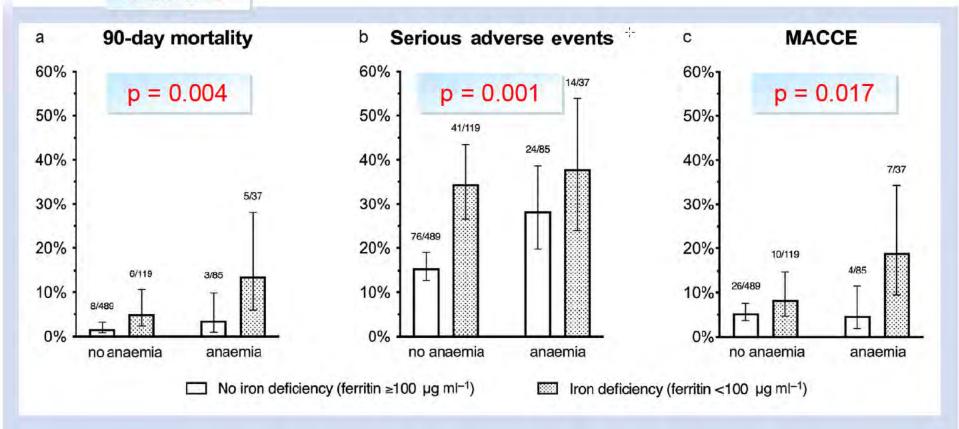
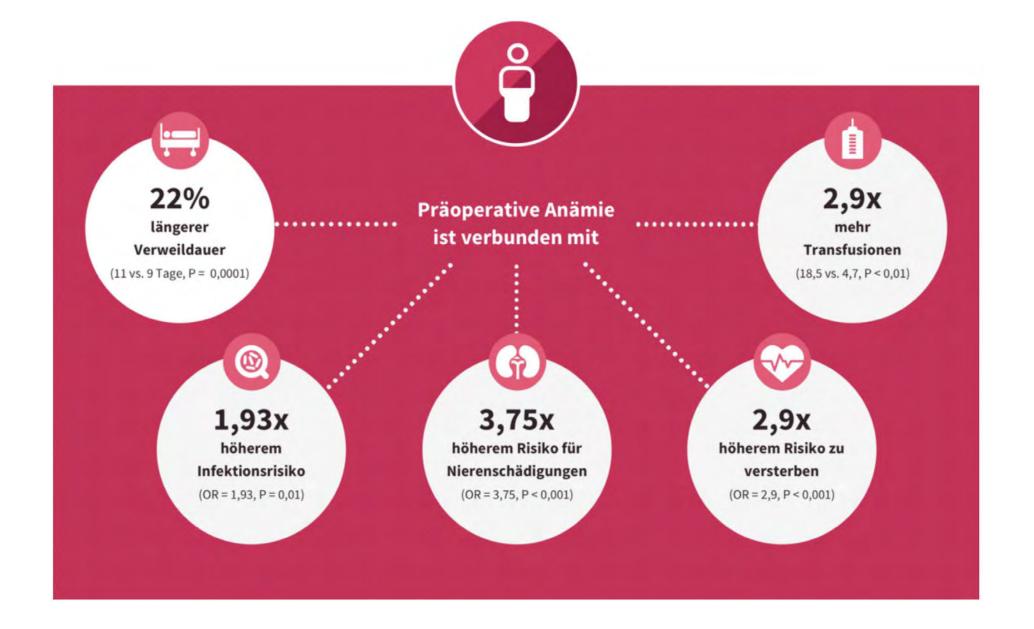


Fig 2. Frequency graphs. Bar graphs showing the frequency of (a) 90-day mortality, (b) serious adverse events, and (c) major adverse cardiac and cerebrovascular events (MACCE). The frequencies in the four groups are shown in percent with 95% Wilson confidence intervals. Group differences for iron deficiency adjusted for anaemia are significant: (a) P=0.004, (b) P=0.001, and (c) P=0.017.







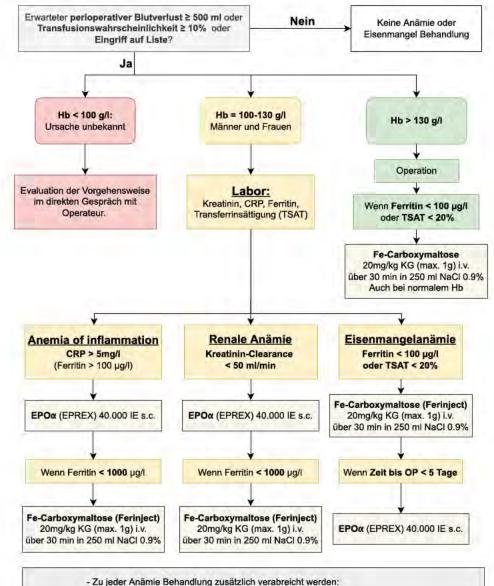
#### Anämietherapie





#### Behandlung präoperative Anämie (Hb < 130 g/l) und Eisenmangel (Ferritin < 100 μg/l oder TSAT < 20%)

Universitätsspital Zürich



- + Vitamin B12 (Vitarubin) 1 mg einmalig s.c.
- + Folsäure (Acidum folicum Streuli) 5mg einmal täglich p.o. bis zur Operation
- Bei Unklarheiten Patlent Blood Manager kontaktieren. Tel.: 043 253 90 36

#### Kosten





2x 177.40

Ferinject

**EPREX** 



383.60

Vitamin B12



1.90

Folsäure



5.25



# Effect of ultra-short-term treatment of patients with iron deficiency or anaemia undergoing cardiac surgery: a prospective randomised trial

- 505 herzchirurgische Patienten (CABG, valve and combination)
  - Anämie (n = 253)
  - Isolierter Eisenmangel (n = 252)
- Stratifizierung
  - Art der OP (CABG, valve or CABG-valve)
  - Primary vs. re-do surgery
  - On- vs. off-pump ACPB
  - Duale Plättchenhemmung
- Behandlung vs. Placebo
  - Erythropoitin alpha 40'000 units s.c.
  - 20 mg/kg ferric carboxymaltose (max. 1'000 mg) in 250 ml saline in 30 min iv
  - Vit B 12, 1 mg s.c.
  - Folic acid 5 mg po



# Effect of ultra-short-term treatment of patients with iron deficiency or anaemia undergoing cardiac surgery: a prospective randomised trial

	Treatment (n=243)	Placebo (n=241)	p value
RBC units transfused in first 7 days			0.036
Mean (SD)	1.5 (2.7)	1.9 (2.9)	40
Median (IQR)	0 (0-2)	1 (0-3)	
Distribution, n (%)			
0	135 (56%)	114 (47%)	#
1	31 (13%)	27 (11%)	
2	33 (14%)	38 (16%)	#
3	10 (4%)	23 (10%)	
4	12 (5%)	11 (5%)	#
≥5	22 (9%)	28 (12%)	

RBC units transfused in day 0		**	0.018
to POD 90			
Mean (SD)	1.7 (3.2)	2.3 (3.3)	**
Median (IQR)	0 (0-2)	1 (0-3)	
Distribution, n (%)			
0	129 (53%)	107 (44%)	-12
1	28 (12%)	23 (10%)	40
2	37 (15%)	39 (16%)	
3	13 (5%)	25 (10%)	**
4	12 (5%)	9 (4%)	- 11
≥5	24 (10%)	38 (16%)	12.

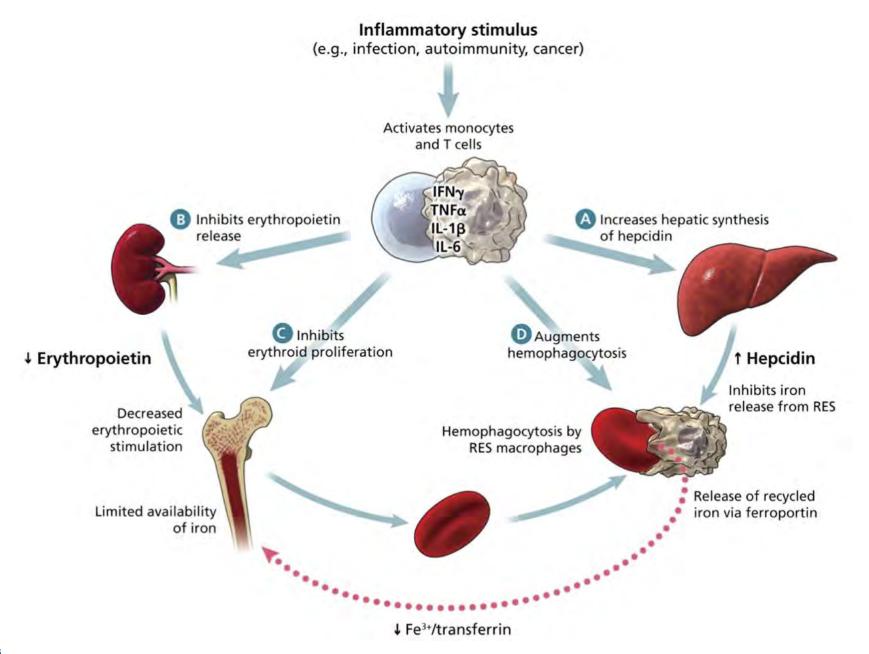
•	Treatment group (n=243)	Placebo group (n=241)	p value	
Patients transfused (≥1 RBC), first 7 days	108 (44%)	127 (53%)	0.084	
Combined allogeneic transfusions (≥1 RBC or ≥1 TC or ≥1 FFP), first 7 days	111 (46%)	129 (54%)	0.10	
Length of stay in ICU (days)	#	*	0.33	
Mean (SD)	3.5 (7.8)	2.7 (5.2)	×e	
Median (IQR)	1(0.9-2.0)	1 (0.9-2.0)	â	
Length of stay in hospital (days)	**	*	0.73	
Mean (SD)	12.0 (9.7)	12.3 (11.0)	10.	
Median (IQR)	8-8 (6-9-12-9)	8-9 (6-9-13-7)	2.0	
Duration of mechanical ventilation (h)			0.95	
Mean (SD)	28-1 (79-3)	20-8 (56-3)	-	
Median (IQR)	5.7 (3.7-9.2)	5.5 (3.9-10.6)		
MACCE	23 (10%)	21 (9%)	0.88	
Allergy	5 (2%)	2 (1%)	0.45	
Angina	4 (2%)	4 (2%)	1.00	
Myocardial infarction	1 (0%)	6 (3%)	0.068	
Maximum postoperative hs troponin until POD 2 (g/L)			0.40	
Mean (SD)	1259 (2654)	1259 (2383)		
Median (IQR)	571 (222-1110)	578 (293-1230)	0.	
Stroke	6 (3%)	6 (3%)	1.00	
Acute kidney injury	22 (9%)	18 (8%)	0.62	
Dialysis	15 (6%)	8 (3%)	0.20	
Atrial fibrillation	43 (18%)	52 (22%)	0.30	
Infection	88 (36%)	77 (32%)	0.34	

#### **Preoperative Inflammation**

Operation	C-reactive Protein threshold	Percent inflammation (CRP > x)
Colorectal cancer surgery	10 mg/L	34%
Colorectal cancer surgery	7 mg/L	50%
Colorectal cancer surgery	8 mg/L	31%
Esophageal cancer surgery	10 mg/L	37%
Pancreatic cancer surgery	3 mg/L	50%
Liver cancer surgery	10 mg/L	28%
Ovarian cancer surgery	10 mg/L	50%



Fuglestad A. J. et al. Acta Oncologic (2022) 61: 1248 Bath M. et al. PLoS ONE (2022) 17: e0269309 Kim W. R. et al. Ann Surg Oncol (2018) 25: 3898 Feng J-F. et al. OncoTargets Therap (2013) 6: 1147 Nurmi A. M. et al. Scinti Rep (2021) 11: 781 Frühling P. et aö. Br J Surg open (2021) 5: zrab104 Wesley E. et al. Gynecol Oncol (2021) 160: 193





## Eisendynamik bei Inflammation

		Entnahme Eingang Befund-Nr.	25.02.2022-05:00 25.02.2022-09:15 Q202250419	24.02.2022-05:00 24.02.2022-09:07 Q202240376	23.02.2022-08:30 23.02.2022-08:53 Q202230320
BLUTSTATUS					
Blutstatus					
Hämoglobin	117-153	g/l	126	124	143
Hämatokrit	0.350-0.460	1/1	0.375	0.379	0.414
Erythrozyten	3.9-5.2	(I) T/I	4.19	4.19	4.75
MCV	80-100	fl	89.5	90.5	87.2
MCH	26-34	pg	30.1	29.6	30.1
Entzündung					
CRP (C-reakt.Prot.)	< 5	mg/l	16 *	34 *	35 *
Eisenstoffwechsel					
Eisen	7.0 - 26.0	µmol/l	13.0 (2)	5.4 * (2)	3.4 * (2)
Ferritin (ECLIA)	(II) 13-300	µg/I	276 (2)	229 (2)	168 (2)
Ferritin bei Risiko (ECLIA)	> 100	µg/l	276 (2)	229 (2)	168 (2)
Transferrin	25 - 50	µmol/l			
Transferrin	25 - 50	µmol/l	24 * (2)	22 * (2)	29 (2)
Transferrin-Sättigung	15 - 50	%	27 (2)	12 * (2)	6 * (2)
TSAT bei Risiko	>20	%	27 (2)	12 * (2)	6 * (2)



#### **Review Article**

The role of iron in chronic inflammatory diseases: from mechanisms to treatment options in anemia of inflammation

Disease	Anemia prevalence	Al / Inflammation Hepcidin 🛨
CKD	21-62%	++
IBD	~67%	+++
Autoimmune disease	30-60%	+++
Cancer	40-80%	+++
Lung disease (COPD)	7-33%	+++
CHF	30-50%	++
Infectious disease		+++



I

#### IRON METABOLISM AND ITS DISORDERS

#### Anemia of inflammation

Guenter Weiss, 1,2 Tomas Ganz, 3 and Lawrence T. Goodnough 4,5

- Iron deficiency / reduced iron availability
- ◆ EPO response to anemia
- ◆ Erythroid cell differentiation
- RBC life span ▼

Table 1. Disease groups in which AI is common

Disease group	
Cancer and hematological malignancies	
Infections	
Immune-mediated diseases	
Inflammatory diseases	
Chronic kidney disease	
Congestive heart failure	
Chronic pulmonary disease	News
Obesity	20006
Anemia of the elderly	
Anemia of critical illness (accelerated course)	



### Erfolg der präoperativen Anämiebehandlung

Operation	Iron deficiency anemia	Benefit
Colorectal cancer surgery	F<30 or F<100+TSAT<20%	RBC♣, LOS♣, finance♠
Major abdominal surgery *	F<300 or TSAT<25%	RBC♣, LOS♣
Cardiac surgery * EPO+iv Iron	Anemia WHO	RBC♣, AKI♣
Cardiac surgery * EPO+iv Iron	F<100 or TSAT<30%	RBC♣
Cardiac surgery * EPO+iv Iron+VitB12+FA	Anemia WHO or F<100	RBC♣
Major orthopedic surgery * EPO +/- iv Iron	Anemia WHO	RBC-production <b></b>



Trentino K. M. et al. Anesth Analg (2021) 132: 344
Froessler B. et al. Ann Surg (2016) 264: 41
Yoo Y-C. et al. Anesthesiology (2011) 115: 926
Kong R. et al. Br J Anaesth (2022) 128: 796
Spahn D. R. et al. Lancet (2019) 393: 2201
Biboulet P. et al. Anesthesiology (2018) 129: 710

#### **Conclusion 1**

- Eine präoperative Anämie ist assoziiert mit:
  - Mortalität ★
  - Morbidität ★
  - RBC Transfusion •
  - Aufnahme IPS •
  - Länge des Spitalaufenthaltes •

• Eine präoperative Anämie muss und/oder ein isolierter präoperativer Eisenmangel (d. h. ohne Anämie) muss frühzeitig erkannt und präoperativ behandelt werden



#### Blutungsprävention

Petechiale Blutungen Schleimhautblutungen

Flächenhafte Blutungen Weichteilblutung Gelenkblutung

Nachbluten

thrombozytär

plasmatisch

von Willebrand, F XIII

Gefässfaktoren



#### **Strukturierte Gerinnungsanamnese**

Präoperative Bestimmung globaler Gerinnungstets schliessen Gerinnungsdefekte nicht sicher aus.

Sorgfältige, strukturierte Anamnese hat eine höhere Sensitivität und Spezifität für die Aufdeckung bis dahin unbekannter Störungen der Blutgerinnung als eine routinemäßig durchgeführte Bestimmung von Quick, PTT und Thrombozytenzahl.



Version 2.0 / 2018 28.08.18

	e Zutreffendes reuzen, unterstreichen, bzw. ergi	inzen:	N Nein	Ja		<u>satzfragen u.</u> tizen des Arztes:
1	Ist bei Ihnen jemals eine Blutung Thrombose festgestellt worden?		N	J	0	JA (2)
2	Beobachten Sie folgende Blutung auch ohne erkennbaren Grund	gsarten -			pos. Blutungs-Anamnese	
2a	Nasenbluten (ohne andere Ursachen wie Schnupft trockene Luft, starkes Nasenputzen e	en,	Z	J	0	immer (2) nur saisonal (0) bei Medikamenteneinnahme (1) arterielle Hypertonie? (4) HNO-Ursache? (3)
2b	blaue Flecken oder punktförmige (auch am Körperstamm; auch ohne s		N	IJ	0	unfallträchtige Tätigkeiten? (0)
2c	Gelenksblutungen, Blutungen in Weichteile oder M	uskel	N	J	0	JA (2)
3	Beobachten Sie bei Schnittwunde Schürfwunden ein längeres Nach		N	IJ	9	> 5 Minuten (2) bei Nassrasur etc.? (2) bei Medikamenteneinnahme (1)
	Gab es in der Vorgeschichte läng verstärktes Nachbluten beim Za		N	J	0	Nachbehandlung nötig? (5,2) bei Medikamenteneinnahme (1)
	Gab es in der Vorgeschichte eine Blutung während oder nach Ope		N	Ū	0	welche Operationen? (5) Blutung nicht chirurgisch (2)
3	Heilen Wunden schlecht ab?		N	J	0	lange nässend, klaffend? (2) Neigung zur Kelloidbildung? (2)
	Gab / gibt es in der Familie (Bluts Fälle von Blutungsneigung?	verwandtschaft)	N	J	9	
Ba Nehmen Sie Medikamente zur Blutverdünnung ein?  VKA: Sintrom®, Marcoumar®  XABANE: Eliquis®, Lixiana®, Xarelto®  DTI: Pradaxa®  APT: ASS, Plavix®, Efient®, Brilique®)  Nehmen Sie Schmerz- oder Rheumamittel ein? (z.B. Aspirin®, Thomapyrin®, Voltaren®, Proxen®, Seractil®, etc.)  Nehmen Sie pflanzliche Präparate oder Vitamin-Präparate ein? (z.B. Tebonin®, Tebofortan®, Ceremin®, Ginsana®, Geriatric-Pharmaton®, Supradyn vital plus®, Zintona®, Kwai® etc.)		Z	U	99	Angabe der Medikamente (1) Blutungsneigung bei Medikamenteneinnahme? (2, 4, 6)	
9 Zusatzfrage an Patientinnen: Sind Monatsblutungen verlängert (> 7 Tage) und/oder verstärkt (häufiger Wechsel von Tampons/Binden)?		N	IJ	Ø.	seit Menarche? (2)	
0) k	sequenzen teine; (1) Medikamentenanamnese sultation Internist; (5) Befundaushe					
Dati						Patient

# 2. Säule: Minimierung von Blutverlusten und Blutungen



### **Intraoperative Massnahmen**

- Blutungsarme Operationstechniken
- Cellsaver
- Gerinnungsalgorithmus











Individualisierte, faktorenbasierte, zielgerichtete Gerinnungstherapie



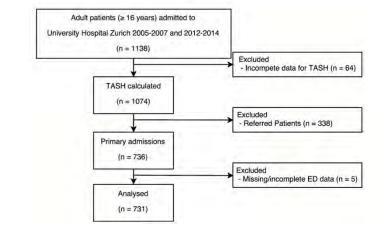
Schritte Diagnostik Intervention ABGA Blutverlust ≥ 50 % - ROTEM (HEPTEM bei Heparinisierung) Gerinnungsstatus\*: Hb, Tc, Quick, Fibrinogen, Faktor V, Faktor XIII diffuse Blutung (vor jeder Gabe von Faktor XIII) Aktive Erwärmung Korrigiere (Zielwert): - Hypothermie (≥ 35.0 °C) - Calcium i.v.-Gabe · Volumenersatz: Ringerfundin® u. Physiogel balanced® Hypokalziämie (Ca<sup>2+</sup> ≥ 1.15 mmol/l)<sup>1</sup> **Physiologie** Vasopressoren - Anämie (Hämoglobin ≥ 70 g/l) Erythrozytenkonzentrat-Transfusion Hypertension (MAP 55 – 65 mmHg) - Tranexamsäure (Cyklokapron<sup>®</sup>) 1g i.v. (15 mg/kg KG) empirisch (MAP 80 - 90 mmHg bei SHT) (zur Verhinderung einer lokalen Hyperfibrinolyse) INTEM: CT/CFT verlängert, HEPTEM normal Protaminsulfat (1:1 zur verabreichten Heparindosis) Heparin-Effekt - ACT pathologisch, Heparinase-ACT normal zur Heparin-Antagonisierung - Fibrinogen (Hämocomplettan®) 2 – 4g, max. 6g i.v. Fibrinogen - FIBTEM A5 ≤ 7mm Nach total 6g Fibrinogen: Faktor XIII (Fibrogammin<sup>®</sup>) 15 E/kg KG i.v. - Bei persistierendem FIBTEM ≤ 7 mm weitere Fibrinogen Substitution und Backup informieren Hyperfibrinolyse: - EXTEM / INTEM: Abfall der MCF nach Fibrinolyse - Tranexamsäure (Cyklokapron®): Bolus: 1 g i.v. (15 mg/kg KG) Erreichen des Maximums - APTEM: normal --> systemische Hyperfibrinolyse - Faktor XIII (Fibrogammin®) 15U/kg Faktor XIII < 60% empirisch nach 6g Fibrinogen Thrombozyten: Thrombozytenkonzentrat-Transfusion - EXTEM / INTEM MCF < 40 mm und FIBTEM > 7mm - ≤ 50 000/µl Anhaltende - ≤ 80 000/µl SHT und in der Herzchirurgie - Desmopressin (Minirin®)\_0.3 μg/kg (max. 16 μg) - Tc-Funktionsstörung (Tc-Hemmer) Blutung FFP-Transfusion (2 - 4 Beutel) Faktor V < 20% Zielbereich ≥ 20 %, insbesondere bei Leberinsuffizienz oder intra-abdominaler Sepsis Quick < 30% und Faktor V ≥ 20% Faktor II, VII, IX, X-Konzentrat (Beriplex®) 1000 - 2000 E. abhängig vom Patientengewicht, langsam i.v. EXTEM / INTEM: CT, CFT verlängert Bei Massivtransfusion Ziel Hämoglobin 70 g/l 35

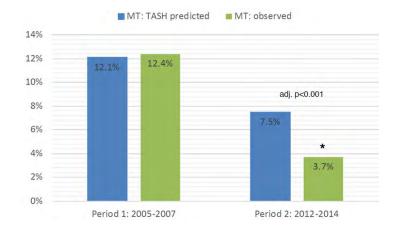
# Original Article

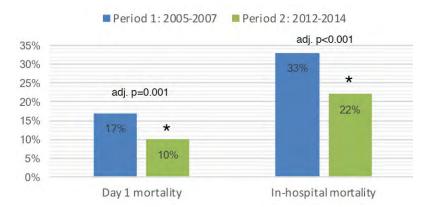
CPD available at http://www.learnataagbi.org/

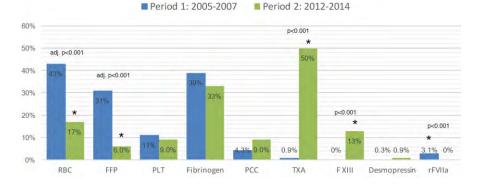
Change of transfusion and treatment paradigm in major trauma patients

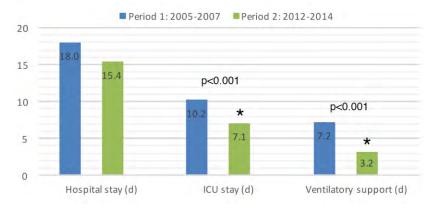
P. Stein, A. Kaserer, K. Sprengel, G. A. Wanner, B. Seifert, O. M. Theusinger and D. R. Spahn











#### **ScienceDirect**





#### Effect of a factor-based coagulation management on blood product use after major burn injury: A retrospective cohort study

Sebastian D. Sahli  $^a$ , Nadine Pedrazzi  $^b$ , Julia Braun  $^c$ , Donat R. Spahn  $^a$ , Alexander Kaserer  $^a$ , Jan A. Plock  $^d$ ,  $^e$ ,  $^e$ 

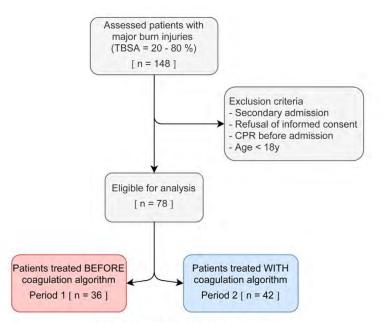


Fig. 2 - Flowchart of patient selection.

# Table 3 – Adjusted models for the comparison of transfused allogeneic blood products and administered coagulation factors between the periods.

	Coefficient	95% confidence interval	p-value
Allogenic transfusions		0.000	775.00
Red blood cells (units)	-33	-52.8 to -12.9	0.002
Fresh frozen plasma (units)	-9	-14.7 to -2.6	0.006
Platelet concentrate (units)	0	-0.7 to 0.2	0.300
Coagulation factors			
4-factor PCC (IU)	-61	-141.9 to 19.7	0.140
Coagulation factor XIII (IU)	-1211	-2443.7 to 20.9	0.054
Fibrinogen (g)	-1.4	-2.2 to -0.5	0.001

The coefficients represent the difference for the patients treated according to the coagulation algorithm (period 2) in comparison with patients treated before (period 1). The models are adjusted for age, sex, the Abbreviated Burn Severity Index (ABSI) and Charlson Comorbidity Index. Abbreviation: 4-factor PCC, 4-factor prothrombin complex concentrate.



Table 2. Differences in the number of patients receiving allogeneic blood products and coagulation factors between the periods during the length of hospital stay.

	Period 1 [n = 36]	Period 2 [n = 42]	Odds ratio	<i>p</i> -value
Allogenic transfusions				
Red blood cells	26 (72,2%)	23 (54.8%)	0.47 [0.18 to 1.19]	0,11
Fresh frozen plasma	14 (38.9%)	4 (9.5%)	0.17 [0.04 to 0.53]	<0.01
Platelet concentrate	2 (5.6%)	2 (4.8%)	0.85 [0.10 to 7.39]	0.87
Coagulation factors				
4-factor PCC	4 (11.1%)	2 (4.8%)	0.40 [0.05 to 2.18]	0.31
Coagulation factor XIII	17 (47.2%)	17 (40.5%)	0.76 [0.31 to 1.87]	0.55
Fibrinogen	11 (30.6%)	4 (9.5%)	0.24 [0.06 to 0.79]	0.03

Data reported as number and percentage (%). **Period 1** refers to the patient cohort before the introduction and **Period 2** to the cohort treated according to the coagulation algorithm.



Sahli et al. Burns 2021;47(7):1486-1494

Table 4. Descriptive of complications and outcome of patients treated before and according to the coagulation algorithm.

	4 m 14 m		
	Before algorithm	With algorithm	p-valu
	Period 1 [n = 36]	Period 2 [n = 42]	
Surgeries (frequency)	4.0 (2.0-9.5)	3.0 (2.0–6.0)	0.33
Multi-organ failure	4 (11.8%)	6 (14.6%)	1.00
Sepsis	15 (44.1%)	16 (39.0%)	0.81
Positive blood culture	17 (50.0%)	18 (43.9%)	0.65
Overall infection rate	21 (61.8%)	17 (41.5%)	0.11
Respiratory	17 (50.0%)	17 (41.5%)	0.49
Skin	18 (52.9%)	12 (29.3%)	0.06
Osteomyelitis	1 (2.8%)	1 (2.4%)	1.00
Abdominal	2 (5.9%)	2 (4.9%)	1.00
Urinary	14 (41.2%)	4 (9.8%)	<0.01
Antibiotic drugs	21 (61.8%)	22 (53.7%)	0.49
Antifungal drugs	14 (38.9%)	6 (14.3%)	0.02
ніт	0 (0%)	3 (7.3%)	0.25
Thrombo-embolic events	5 (14.7%)	7 (17.1%)	1.00
Duration			
Length ICU (days)	22.0 (5.0-48.2)	12.0 (3.0–30.0)	0.13
Length of stay (days)	29.0 (16.5–56.0)	23.0 (11.0-54.0)	0.49
In-hospital mortality	14 (38.9%)	11 (26.8%)	0.33

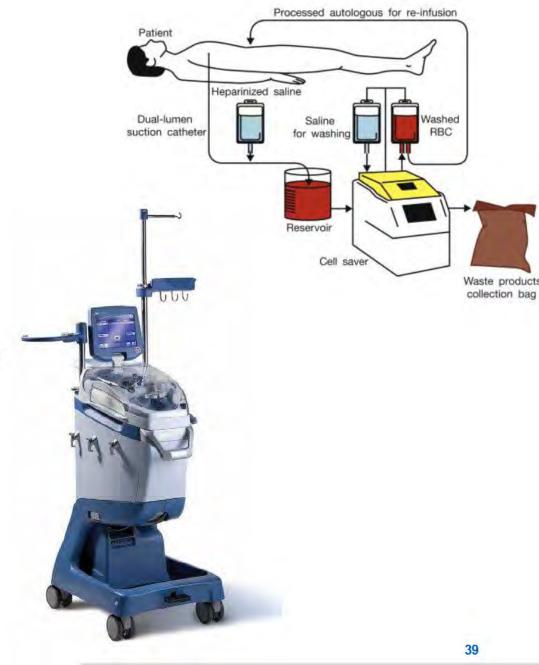
## Cellsaver

Insbesondere bei Blutungen im Bauch, Becken oder Thorax

Senkt Gabe von Fremdblut um bis zu 39%

- Verbesserte Immunfunktion nach Transfusion von patienteneigener gewaschenen Erythrozyten (ICS) im Vergleich zu allogenen Bluttransfusionen
  - · Mechanismus der Immunmodulation unklar
- Aufrüsten braucht Personal und Zeit → frühzeitig dran denken!





# Leukozytenfilter

- Filtration auch von Lipiden, Bakterien und Tumorzellen
   → durch Grösse (40 µm), Adhäsion, Oberflächenladung,
   Benetzbarkeit und Oberflächenstruktur
- 99.6 % 99.9 % Filtrierung von Tumorzellen ¹
   (quantitative Erschöpfung bei rupturierten Tumoren möglich → 2. Filter )
- Empfehlungsgrad IIb f
  ür Hochrisikopatienten in USA<sup>2</sup>



- Bestrahlung mit 50Gy Cäsium 137 für 15 Minuten
- Beseitigung von 99.86% aller Tumorzellen (max. 500ml) <sup>3</sup>
- Kein internationaler Standard für die Bestrahlungsdosis von Erythrozyten (USA/FDA: 15-25 Gy, UK: 25-50 Gy) <sup>4</sup>
- Sterilisation: Bestrahlung von Bakterien bis zu 50kGy (50'000 Gy)







<sup>2</sup> Ferraris VA et. al. Ann Thorac Surg. 2011 3 Hansen E et. al. Transfusion. 1999

<sup>4</sup> Chapman J et. al. Transfus Med. 1996

## Cellsaver

- Filtration-based autotransfusion device
- Retransfusion von Erythrozyten und Thrombozyten





Combined Platelet and Red Blood Cell Recovery during On-pump Cardiac Surgery using same™ by i-SEP Autotransfusion Device (i-TRANSEP Study)

First-in-human noncomparative study of a blood and platelet recovery device in 50 adult patients undergoing cardiac surgery with cardiopulmonary bypass



#### Hypothesis:

- Red blood cell recovery >80%
- Posttreatment hematocrit >40%
- Removal of >90% heparin
- Removal of >75% free hemoglobin

#### **Observed 50 patients:**

- 18 CABG
- 26 valve surgery
- 6 aortic root surgery



	Median (25-75th percentile)
RBC recovery/cycle (%)	86.1 (80.8 - 91.6)
Posttreatment Hct (%)	41.8 (39.7 - 44.2 )
Platelet recovery (%)	52.4 (44.2 - 60.1)
Posttreatment platelet concentration (10°/l)	116 (93 - 146)

In this first-in-human study, the blood and platelet recovery device was able to simultaneously recover and wash both platelets and red blood cells for transfusion during cardiac surgical procedures

Mansour A, et al. Anesthesiology, 2023.



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Mansour et al. Anesthesiology (2023) 139:287

# 3. Säule: Rationaler Einsatz von Blutprodukten



### EDITORIAL



# Transfusion Threshold of 7 g per Deciliter — The New Normal

Paul C. Hébert, M.D., and Jeffrey L. Carson, M.D.

We believe it has become abundantly clear that a transfusion threshold of 7 g per deciliter should become the new normal, recommended in all critically ill patients, including those with severe sepsis and septic shock. To speed up adoption, we should ensure that clinical practice



# Restrictive transfusion strategies are standard today



Tibi P. et al. Ann Thorac Surg (2021) 112: 981
Ferraris V. A. et al. Ann Thorac Surg (2011) 91: 944
Carson J. L. et al. Ann Intern Med (2012) 157: 49
Fleisher L. A. et al. Circulation (2014) 130: e278
ASA Practice Guidelines Anesthesiology (2015) 122: 241
Carson J. L. et al. JAMA (2016) 316: 2025
Mueller M. M. et al. JAMA (2019) 321: 983
Vlaar A. P. et al. Intensive Care Med (2020) 46:673

# Incidence and Impact of a Single-Unit Red Blood Cell Transfusion: Analysis of The Society of Thoracic Surgeons Database

- Database 2010 to 2019 (N=2'151'430)
  - Type of surgery (CABG, aortic valve replacement)
  - 0 RBC: 1'303'988 patients
  - 1 RBC: 206'555 patients
  - >1 RBC: 640'887 patients
- Comparison between patients with 0 vs. 1 RBC transfusions
  - Propensity matching between patients with 0 and 1 RBC transfusion

	No RBCs	1 Unit of RBCs		
Variable	(n = 206,555)	( n = 206,555)	P Value	
Operative mortality	2058 (1.0)	2990 (1.4)	<.001	
Stroke	2492 (1.2)	3458 (1.7)	<.001	
Sternal wound infection	1070 (0.5)	1343 (0.7)	<.001	
Prolonged ventilation	6990 (3.4)	13,305 (6.4)	<.001	
New hemodialysis	1816 (0.9)	3703 (1.8)	<.001	
Reoperation for bleeding	1064 (0.5)	2685 (1.3)	<.001	



# **Evidenzbasierte Transfusionstrigger**

Patientencharakteristika	Transfusion Trigger
Gesunde Gebärende	Hb < 60 g/L
Patienten mit: Schädelhirntrauma, Verbrennung, GI-Blutung, Sepsis, stabile IPS-Patienten, freie Lappenplastik, vv-ECMO	Hb < 70 g/L
Patienten bei komplexen herzchirurgischen Eingriffen	Hb < 75 g/L
Patienten mit: schwerer KHK, akutem Herzinfarkt, Herzinsuffizienz, Karotisstenose >70%, > 80 Jahre und kardiovaskulärer Krankheit, va-ECMO	Hb < 80 g/L

Tibi P. et al. Ann Thorac Surg (2021) 112: 981
Ferraris V.A. et al. Ann Thorac Surg (2011) 91: 944
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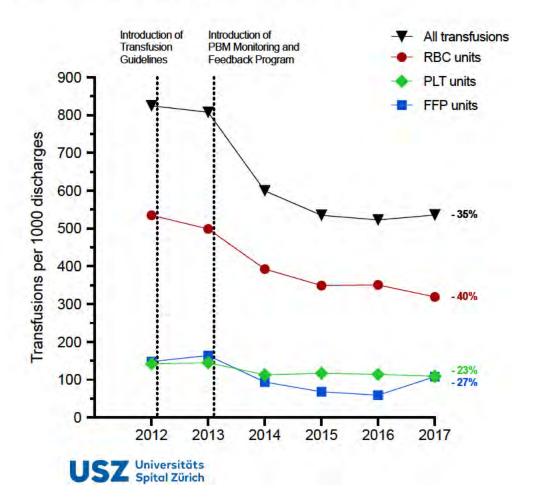
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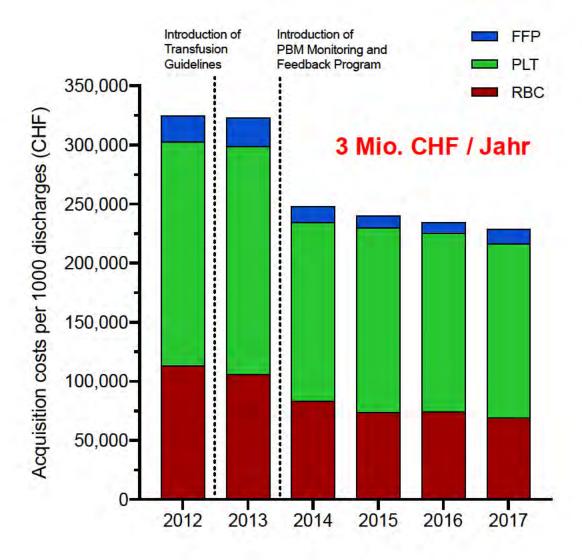
Anaesthesia 2019 doi:10.1111/anae.14816

#### Original Article

# Impact of a Patient Blood Management monitoring and feedback programme on allogeneic blood transfusions and related costs

A. Kaserer, <sup>1</sup> J. Rössler, <sup>1</sup> J. Braun, <sup>2</sup> F. Farokhzad, <sup>3</sup> H.-C. Pape, <sup>4</sup> P. Dutkowski, <sup>5</sup> A. Plass, <sup>6</sup> T. Horisberger, <sup>7</sup> J. Volbracht, <sup>8</sup> M. G. Manz <sup>9</sup> and D. R. Spahn <sup>10</sup>





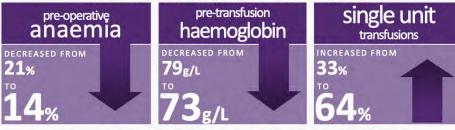
# PATIENT BLOOD MANAGEMENT PROGRAM

The Western Australian Patient Blood Management Program recently published the world's largest study on patient blood management outcomes. The study included over 600,000 patients admitted to Western Australia's four major adult hospitals between July 2008 and June 2014. Over the six-year study period, the program was associated with:

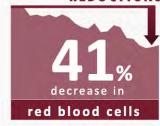
#### IMPROVED PATIENT OUTCOMES

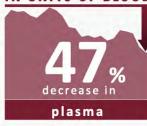


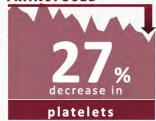
#### IMPROVED KEY PROGRAM INDICATORS



#### REDUCTIONS IN UNITS OF BLOOD TRANSFUSED







#### **PRODUCT COST SAVINGS**

Over the six-year study period blood product cost savings were:

\$18.5M

#### **ACTIVITY BASED COST SAVINGS**

...however with the hospital costs of administering a transfusion added, the gross savings are estimated to be between:

\$80M-\$100M

For more information see: Leahy MF et al. Improved outcomes and reduced costs associated with a health system-wide Patient Blood Management Program. *Transfusion*.



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

A Systematic Review and Meta-analysis

Meta-Analyse von 17 Studien mit 235,779 Patienten

⇒ Pre-PBM: 100,886 Patienten

⇒ PBM: 134,893 Patienten

- Outcomes:
  - ⇒ RBC transfusion rate
  - ⇒ RBC units transfused
  - ⇒ Hospital LOS
  - ⇒ Adverse outcomes
  - ⇒ Mortality

**TABLE 1.** Benefits of Patient Blood Management

	Change	P	Number of Patients
Transfusion rate	-39%	< 0.00001	207,006
RBC unit per patient	-0.43 unit	< 0.00001	216,657
Hospital LOS	-0.45  day	< 0.00001	219,850
Major complications	-20%	< 0.00001	214,298
Acute renal failure	-26%	< 0.00001	166,955
Infection rate	-9%	< 0.03	192,987
Thromboembolic events	-25%	< 0.00001	170,189
Mortality	-11%	< 0.02	221,528

LOS indicates length of stay.



# **Take Home Messages**

- Patient Blood Management (PBM) ist ein multimodales, hoch effizientes Konzept
- PBM muss implementiert werden, um die individuellen und kombinierten nachteiligen Auswirkungen von Anämie, Eisenmangel, Blutverlust und RBC-Transfusionen zu vermeiden
- Die prä- und postoperative Behandlung von Anämie und/oder Eisenmangel mit intravenösem Eisen und Epoetin alpha ist ein Schlüsselelement des PBM
- Präoperative Zielwerte der Patienten aus der Fokusgruppe (Transfusionsrate >10% oder erwarteter BV >500 ml): Hämoglobinwert von > 130 g/L (bei Männern und Frauen), Ferritin ≥ 100 mg/l und TSAT ≥ 20%



