

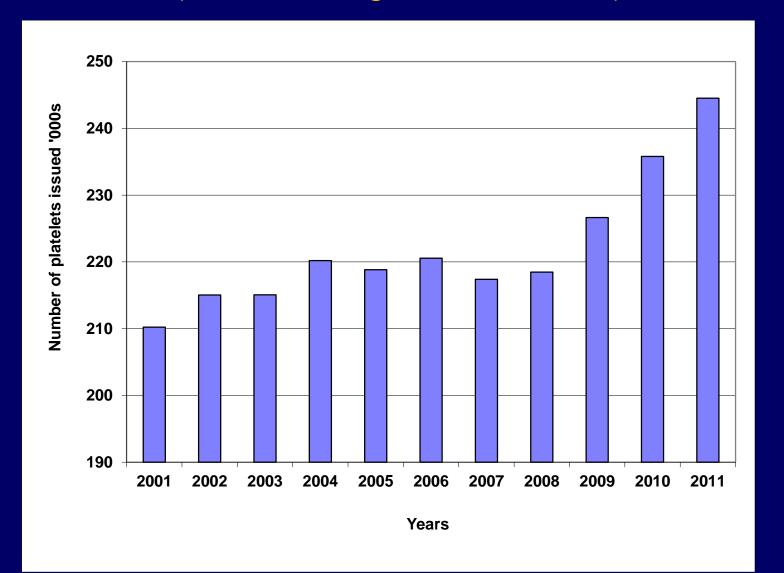


### Using platelets in Haematology Patients New Studies

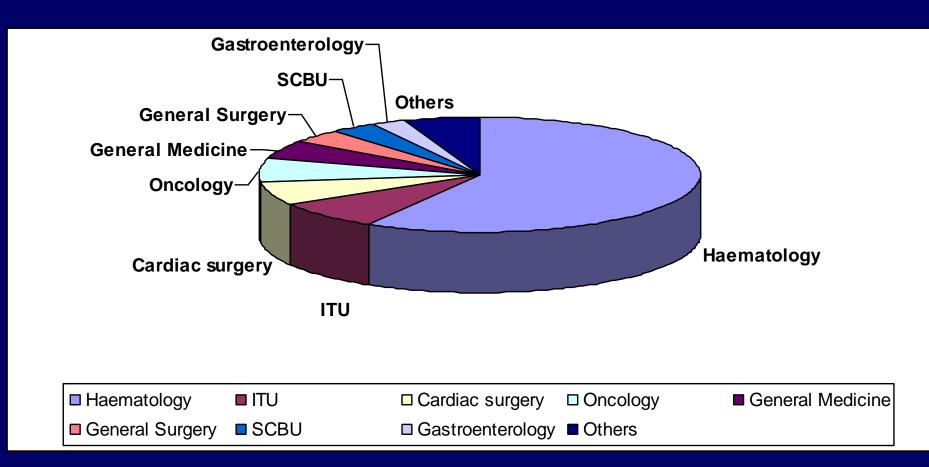
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### Platelet usage increasing

(data from England & N. Wales)



### Haematology patients use majority of platelet transfusions issued



Data from North West of England and Wales Audit of platelet use and wastage.

## Majority of platelet transfusions are prophylactic

Reason for Transfusion	Audited episodes in each category	Appropriate	Indeterminate	Outside guidelines
Prophylactic	69%	60%	6%	34%
Pre -procedure	15%	64%	13%	23%
Therapeutic	13%	84%	12%	5%
Unclear	3%	0%	100%	0%

### Current Issues in Prophylactic Platelet Transfusion Studies

Platelet dose

Platelet threshold

Therapeutic versus prophylactic

Type of platelet component

### Systematic Review of Prophylactic Platelet Transfusions

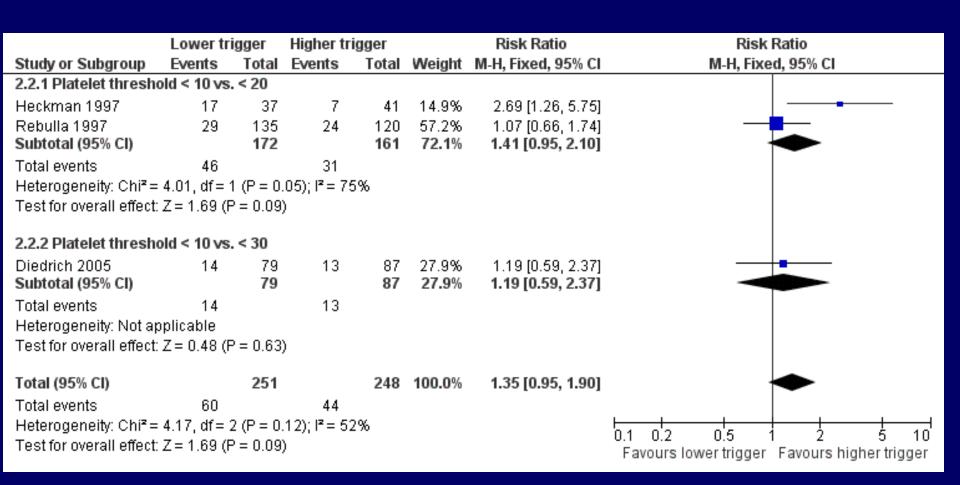
- Update of Cochrane Review
- 18 studies
  - 13 included in analysis
  - 3 completed not published
  - 1 actively recruiting & 1 not actively recruiting
- 3 main questions
  - Platelet Dose
  - Platelet Threshold
  - Prophylactic vs. Therapeutic-only Platelets

# Platelet Dose Number of Patients with bleeding of WHO grade 2 or above

	Study dose		Standard dose		Risk Ratio	Risk Ratio		
Study or Subgroup	Events				Weight	M-H, Fixed, 95% CI	M-H, Fixed, 95% CI	
3.2.1 Low dosage platelet transfusions versus standard dosage platelet transfusions								
Heddle 2009	30	58	30	61	9.0%	1.05 [0.74, 1.50]	<del>-</del>	
Slichter 2010	296	417	292	423	89.7%	1.03 [0.94, 1.12]		
Tinmouth 2004 Subtotal (95% CI)	6	56 <b>531</b>	4	55 <b>539</b>	1.2% <b>100.0</b> %	1.47 [0.44, 4.94] 1.04 [0.95, 1.13]	•	
Total events	332		326					
	Heterogeneity: Chi² = 0.36, df = 2 (P = 0.84); l² = 0%							
Test for overall effect:	Z = 0.79 (	P = 0.4	3)					
3.2.2 High dosage pla	atelet tran	sfusio	ns versus	standar	d dosage	platelet transfusions		
Sensebe 2004	3	48	2	48	0.7%	1.50 [0.26, 8.58]		
Slichter 2010	302	432	292	423	99.3%	1.01 [0.93, 1.11]	· ·	
Subtotal (95% CI)		480		471	100.0%	1.02 [0.93, 1.11]	•	
Total events	305		294					
Heterogeneity: Chi² = 0.20, df = 1 (P = 0.66); l² = 0%								
Test for overall effect: Z = 0.35 (P = 0.73)								
							0.1 0.2 0.5 1 2 5 10 Study dose Standard dose	

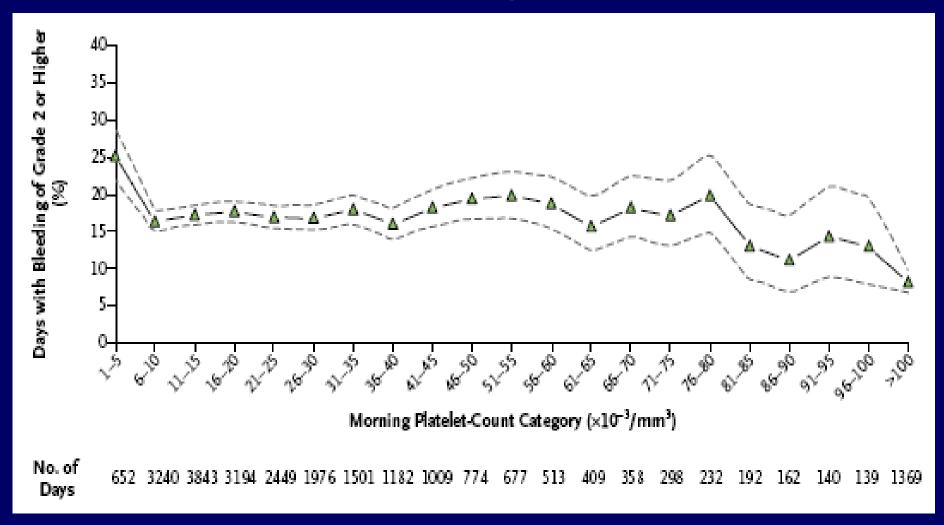
Prophylactic platelet transfusion for prevention of bleeding in patients with haematological disorders after chemotherapy and stem cell transplantation Estcourt *et al* 2012. Cochrane Database of Systematic Reviews

## Platelet Threshold Number of Patients with bleeding of WHO grade 2 or above

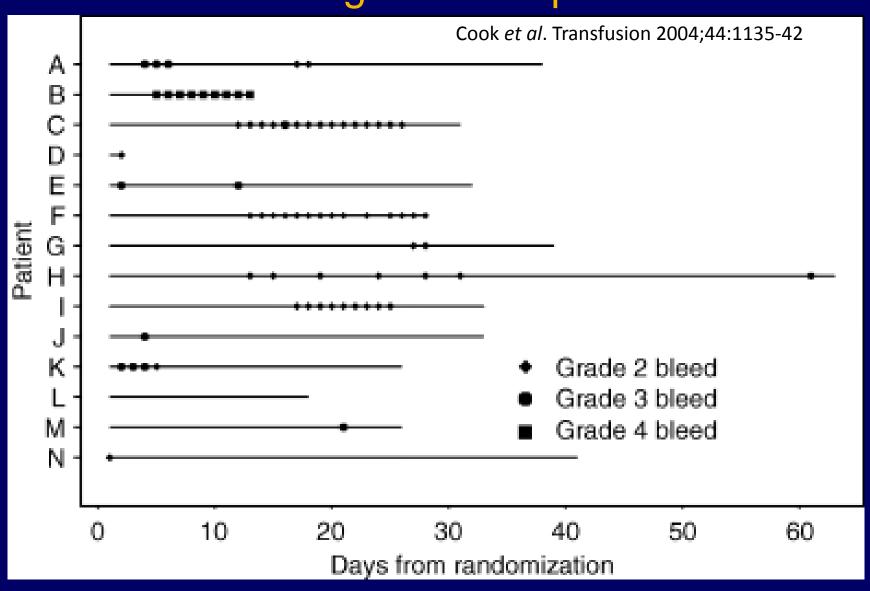


Prophylactic platelet transfusion for prevention of bleeding in patients with haematological disorders after chemotherapy and stem cell transplantation Estcourt *et al* 2012. Cochrane Database of Systematic Reviews

### Morning platelet count is a poor predictor of bleeding risk



### Variability in frequency and severity of bleeding between patients



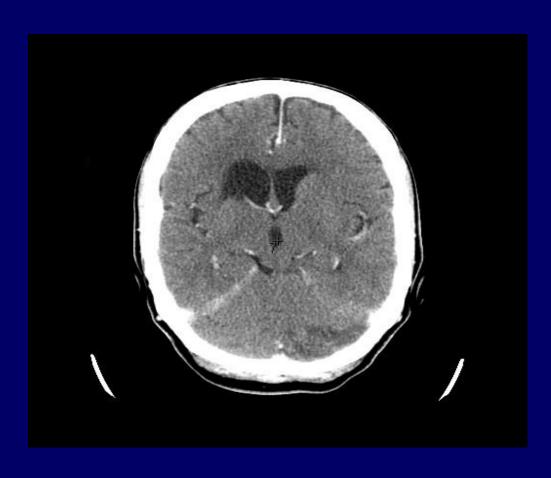
#### Consequences

 Exposing patients at low risk of bleeding to unnecessary blood products

May be undertreating patients at high risk of bleeding

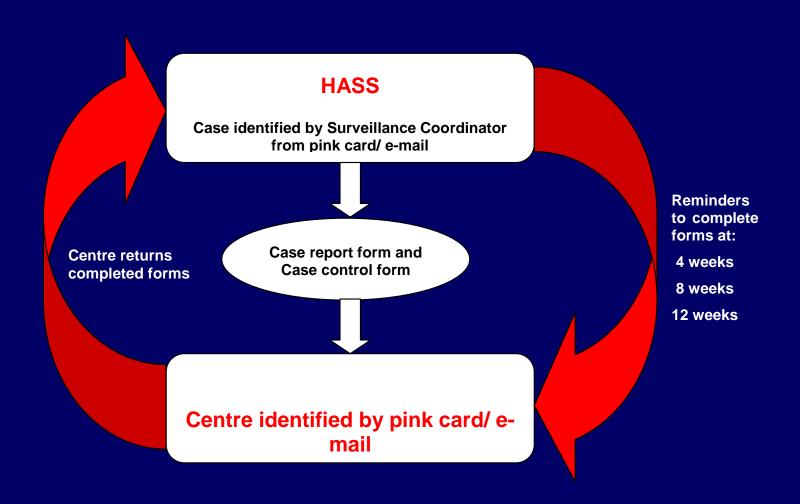
Not using limited platelet supply effectively

### Case-control study of ICH in thrombocytopenic patients with haematological malignancies



- What factors (e.g. age, haematological disease, treatment, infection) predispose patients to ICH?
- What is frequency of ICH in these patients?
- What are short-term outcomes? (e.g. death or persistent neurological deficit)

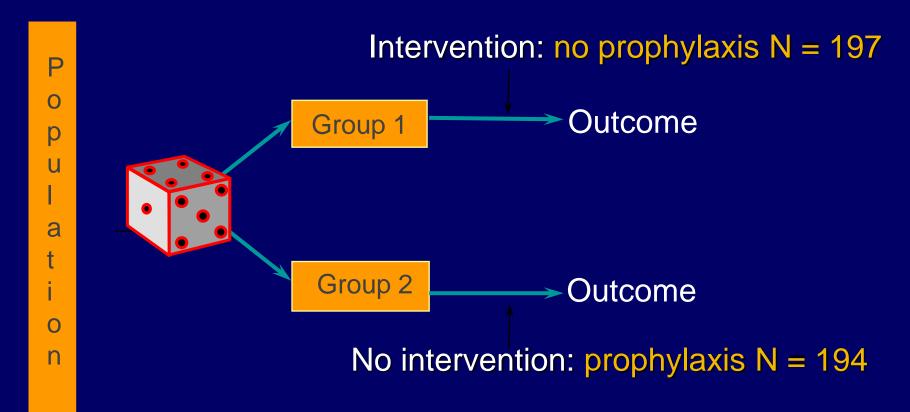
#### **INCITE STUDY**



### Pre-procedure platelet transfusions Central Lines

- 2 retrospective observational studies
  - Ziedler et al 2011–173 patients (acute leukaemia)
    - No increase in bleeding if plts > 20
  - Hass et al 2010 344 patients (majority haematology)
    - No significant bleeding complications
- Prospective observational study
  - Barrera et al 1996 108 patients
    - No difference in platelet count between patients with and without bleeding complications

### German Multicentre trial of prophylaxis vs. no prophylaxis



Primary outcome

Number of platelet transfusions given during 14 day observation period

Therapeutic platelet transfusion versus routine prophylactic transfusion in patients with haematological malignancies: an open-label, multicentre, randomised study Wandt et al 2012. Lancet

#### Primary end-point. Platelet transfusions per patient

	Prophylactic	Therapeutic	Reduction (%)	P value
All patients No. (treatment cycles)	194 (343 Rx cycles)	197 (301 Rx cycles)		
	2.44 (2.22 to 2.67)	1.63 (1.42 to 1.83)	33.5% (22.2 to 43.1)	< 0.0001
AML No. (treatment cycles)	96 (245 Rx cycles)	94 (198 Rx cycles)		
	2.68 (2.35 to 3.01)	1.83 (1.58 to 2.10)	31.6% (18.6 to 42.6)	< 0.0001
Autologous SCT No. (treatment cycles)	98 (98 Rx cycles)	103 (103 Rx cycles)		
	1.80 (1.45 to 2.15)	1.18 (0.82 to 1.55)	34.2% (6.6 to 53.7)	0.0193

Therapeutic platelet transfusion versus routine prophylactic transfusion in patients with haematological malignancies: an open-label, multicentre, randomised study Wandt et al 2012. Lancet

#### Secondary outcome. Bleeding per treatment cycle

	All patients			AML Patients			Autologous SCT Patients		
	Prophylactic (343 Rx cycles)	Therapeutic (301 Rx cycles)	P value	Prophylactic (245 Rx cycles)	Therapeutic (198 Rx cycles)	P value	Prophylactic (98 Rx cycles)	Therapeutic (103 Rx cycles)	P value
Grade 2 or higher	65 19% (14 to 23)	127 42% (36 to 48)	< 0.001	57 24% (18 to 30)	98 51% (43 to 59)	<0.001	8 8% (3 to 14)	29 28% (19 to 37)	<0.001
Grade 3	3 1% (0 to 2)	7 2% (0 to 4)	0.21	3 1% ( 0 to 4)	6 3% (1 to 7)	0.32	0 0% (0 to 5)	1 1% (0 to 6)	1
Grade 4	4 1% (0 to 2)	14 5% (2 to 7)	0.0159	4 2% (0 to 3)	13 7% (3 to 11)	0.0095	0	0	-

#### Number of days with thrombocytopenia (<20)

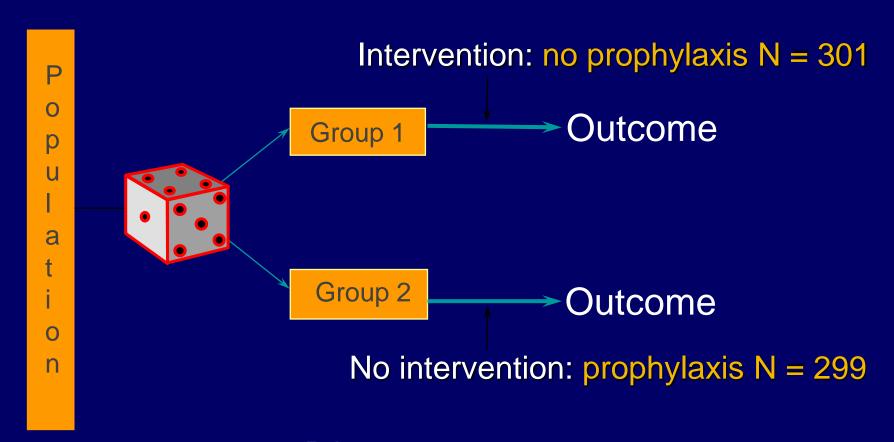
- No difference between study arms prophylactic vs. therapeutic
- Higher in AML group (12.68) vs. Autologous SCT (6.36 to 7.88).

Therapeutic platelet transfusion versus routine prophylactic transfusion in patients with haematological malignancies: an open-label, multicentre, randomised study Wandt *et al* 2012. Lancet

#### Other secondary outcomes

- No difference between prophylactic and therapeutic groups
  - Red blood cell transfusions per patient
  - Days in hospital
  - Side effects of transfusions
  - Overall survival

### TOPPS: Multinational trial of prophylaxis vs. no prophylaxis



Primary outcome
Proportion of patients who have ≥ WHO grade 2 bleeding

#### Review of current Issues in Prophylactic Platelet Transfusion Studies

Platelet dose

Platelet threshold

Therapeutic versus prophylactic

#### Future research

 More evidence required for use of platelet transfusions prior to procedures

 A better understanding of all haemostatic changes in patients with haematological disorders

 A more patient-centred approach to platelet transfusions required





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