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# Intraoperative transfusion practices in Europe

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### Abstract

**Background:** Transfusion of allogeneic blood influences outcome after surgery. Despite widespread availability of transfusion guidelines, transfusion practices might vary among physicians, departments, hospitals and countries. Our aim was to determine the amount of packed red blood cells (pRBC) and blood products transfused intraoperatively, and to describe factors determining transfusion throughout Europe.

**Methods:** We did a prospective observational cohort study enrolling 5803 patients in 126 European centres that received at least one pRBC unit intraoperatively, during a continuous three month period in 2013.

**Results:** The overall intraoperative transfusion rate was 1.8%; 59% of transfusions were at least partially initiated as a result of a physiological transfusion trigger- mostly because of hypotension (55.4%) and/or tachycardia (30.7%). Haemoglobin (Hb)- based transfusion trigger alone initiated only 8.5% of transfusions. The Hb concentration [mean (sD)] just before transfusion was 8.1 (1.7) g dl<sup>-1</sup> and increased to 9.8 (1.8) g dl<sup>-1</sup> after transfusion. The mean number of intraoperatively transfused pRBC units was 2.5 (2.7) units (median 2).

**Conclusions:** Although European Society of Anaesthesiology transfusion guidelines are moderately implemented in Europe with respect to Hb threshold for transfusion (7–9 g dl<sup>-1</sup>), there is still an urgent need for further educational efforts that focus on the number of pRBC units to be transfused at this threshold.

Clinical trial registration: NCT 01604083.

Key words: anaemia; anesthesia; blood transfusion; surgery; transfusion trigger

It is now considered good clinical practice to use a restrictive transfusion regimen into clinical pathways in order to minimize unnecessary use of allogeneic blood.<sup>1–3</sup> This is consistent with the current 'patient blood management' (PBM) paradigm, a

multidisciplinary, multimodal approach to best transfusion practice.<sup>4–6</sup> However, despite the general belief that PBM is useful and improves outcome, implementation of all measures of this package is difficult, time consuming, and as a consequence

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Editor's key points

- There is marked variation in blood transfusion practices.
- Most transfusions given intraoperatively are for hypotension or other indicators of tissue hypoperfusion.
- A transfusion threshold or trigger is less relevant in the intraoperative setting.
- Single unit red cell transfusions should be used more often.

is seldom performed completely.<sup>6–8</sup> The degree to which guidelines have resulted in consensus in community transfusion practice is unknown. Previous European studies evaluating transfusion practices were limited in size and pertained to a single country.<sup>9–11</sup> Little is known about the intraoperative transfusion triggers used and the transfusion strategy applied in clinical practice across Europe.<sup>12</sup> Therefore, the primary goal of our study was to assess transfusion triggers and existing transfusion practices and the use of packed red blood cell (pRBC), fresh frozen plasma (FFP), and platelet transfusions in a large sample of European centres.

## Methods

## Study design and participants

The ETPOS (European Transfusion Practice and Outcome Study) protocol was designed as a prospective, observational, multicentre European study. Centres volunteered to participate in the study via the homepage of the European Society of Anaesthesiology (ESA). Each centre had to contact the local ethics committee and/or the relevant regulatory approving body in order to determine whether obtaining informed consent was necessary or could be waived. If informed consent was required, only patients that agreed to participate and signed a consent form were included in the study. The inclusion procedure stipulated postoperative informed consent for the usage of data obtained. If a patient died before consent could be obtained, data were not used. No centre started without all mandatory local ethical or regulatory requirements being fulfilled.

All patients undergoing an elective noncardiac surgical procedure at each of the participating hospitals were screened for inclusion. Only patients that received intraoperatively at least one pRBC unit during the study period (April 1st – December 31st, 2013) were included in the analysis. Data acquisition time was three consecutive months for each centre. There were no further specific inclusion criteria. The only exclusion criteria was age <18 yr and cardiothoracic, emergency, or trauma surgery.

#### Pre-study survey

Each centre participating in the study was asked to fill in a prestudy survey. The main characteristics requested were the size of the hospital (beds), specialities provided, and several aspects of the transfusion regimen and blood management of the hospital.

#### Data collection

Data were collected on patient characteristics (sex, age, body weight and height); ASA physical status; duration of anaesthesia; type of surgery; usage of point of care coagulation monitoring; laboratory values (Hb, INR, aPTT, platelets, fibrinogen) at beginning of surgery, just before transfusion of first pRBC, and at the end of surgery; reason for the transfusion of the first pRBC (Hb threshold, physiological transfusion triggers, transfusion relevant comorbidities, massive acute bleeding); fluid volume, blood products, and procoagulant drugs administered until end of surgery [pRBC, crystalloids, colloids, cell saver, FFP, platelet concentrates, tranexamic acid, prothrombin complex concentrate (PCC), fibrinogen, recombinant factor VIIa, cryopreciptate, factor XIII]; estimated volume loss until end of surgery (blood loss, urine output, other fluids). An overview of the protocol is provided in the appendix.

Primary endpoints were: (i) amount of pRBC and blood products and coagulation factors transfused, and (ii) factors determining transfusion of pRBC and blood products in different regions of Europe.

#### Data acquisition and quality management

The data collection and management was done using the Open-Clinica open source software, version 3.2. (Copyright<sup>®</sup> OpenClinica LLC and collaborators, Waltham, MA, USA, www.OpenClinica. com). Data were collected on paper by the physician providing anaesthesia and recorded afterwards into the study database. The login names and passwords were provided for registration of patients, monitoring of recruiting progress, query management, and source data verification and an internal communication platform. Automatic data entry plausibility checks and mandatory data items enforced high data quality. Furthermore three different data cleaning runs were done with several checks ensuring high data quality.

#### Statistical analysis

Our aim was to recruit as many participating hospitals as possible and to recruit every eligible patient receiving at least one pRBC during surgery in those hospitals. We anticipated that a minimum sample size of 10 000 patients would enable a precise estimate of current transfusion triggers throughout Europe. Therefore we aimed at 100 participating centres providing 100 patients each. This sample size was also expected to provide sufficient data to be able to describe transfusion practices in different settings.

We used SPSS (version 21.0) for data analysis. Categorical variables are presented as number (%) and continuous variables as mean (SD) when normally distributed or median (IQR) when not.

This study is registered with ClinicalTrials.gov, number NCT01604083.

#### Role of the funding source

The study was funded by a grant of the ESA Clinical Trial Network. An independent steering committee was responsible for study design, conduct, and data analysis. Members of the steering committee had full access to the study data and were solely responsible for interpretation of the data, drafting and critical revision of the report, and the decision to submit for publication.

## Results

One hundred and twenty six hospitals of 30 European countries participated in the study (overview is given in the appendix).

#### Pre-study survey

An overview of the parameters included in the pre-study survey is given in Table 1.

Of the 126 centres with valid patients all 126 filled in the prestudy survey (with one being incomplete). Approximately a third of the centres (n=40) had less than 500 beds, 50 (40%) had between 500 and 1000 beds and 36 (28%) had more than 1000 beds.

liberal: 12%

liberal: 13%

restrictive: 82%

restrictive: 80%

Table 1 Prestudy survey of participating hospitals (n=126): casemix, haemostasis and transfusion practices. Values are given as median (IQR) or %. INR, international normalized ratio, apTT, partial thromboplastin time, ACT, activated clotting time, TEG, thrombelastography, TEM, thrombelastometry; percentage values do not necessarily add up to 100% as multiple selections were possible

Hospital characteristics	
Beds	40
<500	40
501–999	50
>1000	36
no. of operating theatres	18 (10–26)
no. of surgical procedures per yr	15 000 (7050–25 000)
(estimation)	15 000 (7 050 25 000)
no. of board certified anaesthetists	26 (13–47)
no. of anaesthetists in training	17 (7–30)
Hospital specialties	
Orthopaedics	88%
Visceral surgery	89%
Urology	86%
Gynaecology	90%
Vascular Surgery	79%
Trauma	80%
Ear Nose Throat	84%
Thoracic Surgery	63%
Cardiac Surgery	51%
Neurosurgery	67%
Hepato – biliary surgery	83%
Dental surgery	64%
Plastics	74%
Ophthalmology	73%
Estimated % of patients prepared with	
Oral iron	1 (0–5)
i.v. iron	1 (0-2)
Erythropoietin	0 (0-1)
Perioperative haemostasis monitoring	- ()
- conventional monitoring	90%
- INR	90%
- apTT	85%
- fibrinogen	80%
- platelets	90%
- ACT	24%
Point of care monitoring	2170
- INR/apTT	18%
- TEG	16%
- ROTEM	30%
- Multiplate, Verify Now, Platelet	5078
Mapping	14%
Mapping Massive transfusion	
% of hospitals with transfusion	
protocols	57%
- based on packages	21%
- based on ratios	21%
- based on conventional coagulation	2,70
tests	50%
- based on POC testing	31%
- based on experience	37%
Transfusion regimen	5/ /0
- for pRBCs	liberal: 30%
- 101 httpcs	restrictive: 63%
- for FFPs	liberal: 31%
1011113	restrictive: 63%
	103010076. 03/0
	Continued

Table 1 Continued	
- for coagulation factors	
- for platelets	

Transfusion practice		
<ul> <li>leukocyte reduced pRBCs used</li> </ul>	54%	
<ul> <li>non-leukocyte reduced pRBCs used</li> </ul>	46%	
- cryoprecitates used	48%	

Table 2 Patient and perioperative characteristics (n=5803)

Variable	
Age (yr)	63.7 (16.1)
Male	49%
BMI	26.2 (5.3)
ASA physical status	
Ι	5.5%
П	31.0%
III	45.9%
IV	16.5%
V	1.1%
Type of surgery	
Orthopaedic	21.6%
Lower gastrointestinal	12.4%
Vascular	9.9%
Upper gastrointestinal	9.3%
Hepatobiliary	7.9%
Gynaecological	7.5%
Urological	6.7%
Others	24.7%
Cancer Surgery	36.8%
Duration of Surgery (h)	4.1 (2.7)
Point of Care Device used	15.6%

The median number of operating theatres in each hospital was 18 (10-20). Most centres reported a higher number of board certified anaesthetists than anaesthetists in training. All relevant specialities were covered by the participating hospitals. A negligible amount of centres reported that they used measures to pre-optimize their patients before surgery with either iron or erythropoietin. Conventional coagulation tests were used by more than 90% of the centres, whereas point of care monitoring systems were implemented only by 14-30% of the centres depending on the specific monitoring device. Fifty seven percent of the hospitals use transfusion protocols, the majority of them based on conventional (laboratory) coagulation tests. Sixty three percent of the hospitals reported having a restrictive transfusion protocol implemented for pRBC, 63% for FFP, 82% for platelets and 80% for coagulation factors.

#### General data

In total 373 732 patients were screened (Table 2). The first patient was enrolled on April 1st, 2013, and the last patient December 31st, 2013. Data were obtained for 5929 patients for whom informed consent was obtained or waived, of which 126 were excluded, having been identified as duplicates or having missing transfusion data, leaving 5803 subjects for analysis (Table 1). A mean number of 45 (range: 1-165) patients were included per Table 3 Transfusion data (n=5803). Values are reported as mean (sD). Hb, haemoglobin. INR, international normalized ratio, aPTT, activated partial thromboplastin time

Variable	No.	Value
Estimated blood loss (ml)		1392 (2040)
Begin of surgery		
Hb (g dl <sup><math>-1</math></sup> )	5674	10.6 (2.4)
INR (%)	4500	1.2 (0.5)
aPTT (sec)	4004	33 (17)
Platelets (n microl <sup>-1</sup> )	5180	265 (132)
Fibrinogen (mg dl <sup>-1</sup> )	1838	400 (177)
Just before transfusion		
Hb (g dl <sup><math>-1</math></sup> )	3977	8.1 (1.7)
INR (%)	630	1.4 (0.9)
aPTT (s)	570	43 (38)
Platelets (n microl <sup>-1</sup> )	844	209 (127)
Fibrinogen (mg dl <sup>-1</sup> )	481	297 (174)
End of surgery		
Hb (g dl <sup><math>-1</math></sup> )	4352	9.8 (1.8)
INR (%)	1985	1.4 (4.7)
aPTT (s)	1944	38 (27)
Platelets (n microl <sup>-1</sup> )	2639	202 (119)
Fibrinogen (mg dl <sup>-1</sup> )	1402	307 (163)

Table 4 Blood product volume administered. pRBC, packed red blood cells, FFP, fresh frozen plasma, PCC, prothrombin complex concentrates

Blood product	Patients	Amount administered
pRBC –intraoperative	100%	2.5 (2.7) units
Crystalloids	99%	2434 (1784) ml
Colloids	65%	907 (711) ml
Cell Saver blood	6%	1116 (1388) ml
FFP	31%	4.5 (5.9) units
Platelet concentrate	7%	3.0 (4.1) units
Tranexamic acid	13%	1.4 (1.0) g
PCC	2%	1846 (1476) I.U.
Fibrinogen concentrate	5%	3.2 (2.4) g
Cryoprecipitate	1%	7.2 (5.8) units
Factor XIII	<1%	2119 (1338) I.U.
pRBC –postoperative	56%	3.7 (4.1) units

hospital and 193 (range: 5–783) per country. The study included almost equal numbers of females and males, with a mean age of 64 (13; range 18–100) yr, approximately half of whom were categorized into ASA class III. Orthopaedic and lower gastrointestinal surgery were performed most frequently and around third of the operations were for cancer. A point of care device for monitoring of blood coagulation was used in 15.6% of patients.

#### Course of transfusion

The mean reported blood loss was 1392 (2040) ml (Table 3). In general, 82% of the patients were anaemic as defined by the WHO criteria (male<13 g dl<sup>-1</sup>, female<12 g dl<sup>-1</sup>) preoperatively with baseline (pre-surgery) Hb concentration of 10.6 (2.4) g dl<sup>-1</sup>. Before transfusion of the first pRBC, measured Hb concentration was 8.1 (1.7) g dl<sup>-1</sup> and increased to 9.8 (1.8) g dl<sup>-1</sup> at the end of surgery. The mean number of transfused pRBC was 2.5 (2.7) units (Median

Table 5 Reasons for transfusion of first packed red blood cell unit. Hb, haemoglobin;  $Sv_{O_2}$ , mixed venous oxygen saturation,  $ScvO_2$ , central venous oxygen saturation

Hb alone	8.5%
Hb & physiological trigger	8.2%
Hb & comorbidity	8.7%
Hb & physiological trigger & comorbidity	11.4%
physiological trigger irrespective of Hb	14.3%
physiological trigger & comorbidity	12.5%
physiological trigger & blood loss	7.1%
physiological trigger & blood loss & comorbidity	5.4%
Comorbidity alone	5.3%
Hb & physiological trigger & blood loss & comorbidity	4.6%
other combinations	14%
Physiological transfusion triggers	
Hypotension	55.4%
Tachycardia	30.7%
Acidosis	7.8%
Lactate	7.3%
Arrhythmia	5.1%
ECG	2.7%
ScvO <sub>2</sub> or Sv <sub>O2</sub>	3.4%
Other	10.2%
Comorbidity	
Cardiovascular	35.7%
Renal	8.1%
Pulmonary	7.7%
Haematological	6.9%
Gastrointestinal	6.4%
Others	8.6%

2 [IQR 2-2]). One third of patients received 1 pRBC and 41% of patients received 2 pRBC. There was no correlation between the Hb concentration before transfusion or at the end of surgery and the amount of pRBC units administered (correlation coefficient -0.1 and -0.2, respectively). Conventional coagulation tests varied only slightly over time.

Thirty one percent of patients received FFP and 7% of patients received platelets. In patients receiving FFP the mean amount administered was 4.5 (5.9) units and with platelets the mean amount was 3.0 (4.1) units. An overview for all other fluids administered is given in Table 4.

#### Reason for transfusion

An overview of the rationales for transfusion is given in Table 5. The most prevalent basis for transfusion of the first pRBC was the occurrence of a physiological transfusion trigger (14.3%). The second most important reason was the combination of a physiological transfusion trigger with a suspected or known comorbidity (12.5%). In total 58.9% were transfused at least in part as a result of a physiological transfusion trigger. Only 8.5% of transfusions have been administered solely because of an Hbbased transfusion trigger. If an Hb based transfusion trigger was used, the physician chose an Hb value of 8.1 (1.3) g  $dl^{-1}$ . Notably, most of the physiological transfusion triggers were not lactic acidosis (7.3%) or a decline of  $ScvO_2$  (1.3%) but hypotension (55.4%) and tachycardia (30.7%). The most important comorbidity resulting in transfusion was cardiovascular (35.7%), whereas all other comorbidities were uncommon (<9%). Regression analysis found that the amount of pRBC administered decreased with age of patients, and was also not affected by the type of surgery.

	Restrictive (n=3738)	Liberal (n=2065)	P value
Haemoglobin value (g dl <sup>-1</sup> )			
Beginning of surgery	10.6 (2.5)	10.8 (2.4)	0.0004
Just before transfusion	8.0 (1.7)	8.4 (1.6)	0.0000
End of surgery	9.6 (1.7)	10.1 (1.8)	0.0000
Units administered			
pRBC –intraoperative	2.4 (2.5)	2.7 (2.5)	0.0000
FFP	4.1 (5.3)	5.2 (6.7)	0.0003
Platelet concentrate	3.1 (4.4)	2.9 (3.7)	0.89
pRBC –postoperative	1.9 (3.4)	2.3 (3.9)	0.0008
Estimated blood loss	1382 (2060) ml	1408 (2006) ml	0.22

Table 6 Perioperative characteristics according to self-reported restrictive or liberal red cell transfusion strategy. pRBC, packed red blood cells, FFP, fresh frozen plasma

Furthermore, the country had a minimal role, with only two pairwise comparisons between countries being statistically significant although of arguable clinical relevance. The size of the hospital and academic degree did not influence transfusion practice. Differences between single hospitals have were not analysed because of insufficient sample sizes.

#### Hospitals with different transfusion strategies

Sixty three percent of all hospitals stated to have a restrictive transfusion strategy in the pre-study survey, in contrast to the 37% with a liberal transfusion strategy. Hospitals with a restrictive strategy treated 64% of patients. Indeed, hospitals claiming a more restrictive strategy tended to transfuse at lower Hb concentrations and accept a lower Hb value post-transfusion. As a consequence, in these hospitals significantly less transfusions were needed despite comparable blood loss in both groups. Furthermore less postoperative transfusions and less substitution of FFP coagulation factors were necessary in the restrictive group (Table 6).

## Discussion

It has been previously demonstrated that transfusion practice differs between different physicians, centres, and countries.<sup>9–11 13</sup> For elective surgery, the hospital might therefore be the most important determinant of the number of administered transfusions, with some adopting programmes to reduce transfusions, while others negating the importance of PBM measures on perioperative outcome.<sup>14</sup> There is growing evidence that a restrictive transfusion strategy can be used in most clinical settings, with published guide-lines focusing on PBM of the surgical patient.<sup>24</sup>

We found that blood transfusions are rarely used in most surgical procedures, with only 1.8% transfused intraoperatively with pRBC. The intraoperative transfusion rate will depend, among other things, on a hospital's surgical casemix and transfusion practices. In the current study the pre-transfusion Hb concentration averaged 8.1 (1.7) g dl<sup>-1</sup>. A Hb trigger of 7–9 g dl<sup>-1</sup> has been recommended during active bleeding by ESA guidelines.<sup>15</sup> Thus, the old commonly used Hb threshold of 10 g dl<sup>-1</sup> is no longer the 'magic number' for transfusion, with European centres now practicing according to current transfusion guidelines, at least in the intraoperative period. The post-transfusion Hb, though, was unnecessarily high (9.8 (1.8) g dl<sup>-1</sup>), suggesting that the decision to transfuse led to more than 1 pRBC unit at a time. Some hospitals that claimed to have a restrictive transfusion strategy transfused at rather high Hb concentrations. There is thus still a need for further educational efforts that focus restrictive transfusion approach and on the number of pRBC units to be transfused at this threshold.

The primary rationale for pRBC transfusion has not been previously studied in depth. The current study found that the most frequent triggers for pRBC administration are physiological parameters; 8.5% of transfusions were initiated based only on Hb value. It is worth noticing that if a physiological transfusion trigger had been used, the Hb value chosen for transfusion was 8.1 (1.3) g dl<sup>-1</sup>, a level very close to the Hb value just before transfusion of the first pRBC in all patients. This fact suggests that in reality many clinicians use the Hb value as an adjunctive ('hidden') transfusion trigger, even if physiological triggers are believed to be the primary trigger. The physiological parameters used to trigger transfusion in the majority of patients were hypotension and tachycardia. The triggers with the highest discriminatory power to manifest tissue hypoxia (e.g. lactate and mixed venous central or central venous saturation) played only a secondary role. This may reflect the tendency of many anaesthetists to use the most easily accessible parameters (heart rate and bp) rather than those that require additional invasive catheters. It may also reflect the emphasis anaesthetists give to maintenance of haemodynamic stability, with less focus on tissue oxygenation.

Preoperative anaemia is one of the most important determinants of intraoperative transfusion, and therefore it had been recommended that elective surgery of anaemic patients should be postponed to enable adequate patient preparation.<sup>16</sup> Furthermore, it has been demonstrated, that even mild or moderate preoperative anaemia is associated with significant morbidity and mortality.<sup>17</sup> Therefore, PBM guidelines state that preoperative optimization of Hb is recommended to avoid unnecessary transfusions.<sup>1 2 4 6 18 19</sup> The present study found that the majority of centres have not adopted this recommendation. Only 1% of centres prepared their patients with oral/i.v. iron and/or erythropoietin. As 82% of transfused patients in the current study were anaemic at the beginning of the surgery, it may be hypothesized that preoperative Hb optimization could have avoided transfusion to a certain extent. Because of the small number of patients that were pre-treated for their anaemia, we cannot determine whether those patients were less likely to be transfused.

Previous studies suggest that often more than 1 unit of pRBC are transfused once a decision to administer blood has been made, in part because of blood allocation strategies.<sup>20</sup> In our study more than 40% of patients had two pRBC units transfused; as less than 25% of patients received three or more units of pRBC it can be speculated that because two pRBC units is often ordered

initially, they were given as package. To overcome this problem Australia initiated a campaign, known as the 'one unit policy' where Hb levels are measured after each pRBC administered to determine the necessity for further transfusion. The results of the present study suggest that a similar campaign should be implemented across Europe.

In our study the mean estimated blood loss was 1392 (2040) ml, reflecting a high variability in blood loss, which warrants cautious interpretation of our data. Unlike in trauma, where massive transfusion protocols have been developed and shown to improve outcome,<sup>21</sup> protocols for the administration of blood products in the actively bleeding patient in the operating room are missing. Modern guidelines promote use of a point of care monitoring and coagulation factor-based bleeding management.<sup>15</sup> This was not the case in the present study. Almost half of the patients received FFP, platelet concentrates or coagulation factor concentrates. FFP, despite several adverse effects and poor efficacy<sup>22</sup> is still the most commonly used agent during intraoperative bleeding, whereas in five percent and two percent of patients fibrinogen concentrate and PCC were used, respectively. Parameters of routine coagulation tests did not reveal coagulopathy either preoperatively and at the end of surgery. However, as immediate pre-transfusion coagulation tests were not documented for participating patients, conclusion regarding the appropriateness of the administration of FFP, platelets or coagulation factors cannot be made.

Our study has some limitations. We recorded selected data and might therefore miss some of the factors that might influence transfusion habits. We did not collect data on those not transfused intraoperatively, not in those transfused before or after surgery. We did not find clinically relevant differences between countries. As centres participating and procedures included varied essentially from country to country, differences in transfusion habits might be more centre-specific than country-specific.

## Conclusion

Across Europe, the vast majority of elective surgical patients receiving one or more units of RBC intraoperatively are anaemic at the commencement of surgery (Hb 10.6 g dl<sup>-1</sup>), and correction of anaemia before surgery deserves further study. Although the transfusion trigger (Hb 8.1 g dl<sup>-1</sup>) is probably appropriate intraoperatively, post-transfusion Hb values were uniformly high (Hb 9.8 g dl<sup>-1</sup>) suggesting opportunity to use a single unit transfusion stratagem. Physiological transfusion triggers seem to be the most important catalyst for transfusion intraoperatively. But the physiological triggers mainly used (hypotension and tachycardia) might have a low discriminative power for tissue hypoperfusion and often occur at Hb values that are considered safe for tissue oxygenation in most patients.

## Authors' contributions

Study design/planning: all authors Study conduct: all authors and collaborators Data analysis: all authors Writing paper: all authors Revising paper: all authors

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## **Declaration of interest**

None declared.

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Schuster	Michael	University Medical Center Johannes Gutenberg Mainz	Germany
Seco	Carlos	Hospital da Universidade de Coimbra	Portugal
Selvi	Onur	Maltepe University	Turkey
Senard	Marc	Centre hospitalier Universitaire de Liège	Belgium
Serra	Sofia	Instituto Português Oncologia	Portugal
Serrano	Helena	Hospital Vall d Hebron	Spain
Shmigelsky	Alexander	Burdenko Neurosurgery Institute	Russia
Silva	Luisa	Hospital da Universidade de Coimbra	Portugal
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Singh	Rita	Newcastle Upon Tyne Hospitals NHS Trust The Freeman Hospital High Heaton	United Kingdom
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Skitek	Kornel	Carl – Thiem Klinikum Cottbus	Germany
Skok	Ira	University Hospital "Sveti Duh"	Croatia
Smékalová	Olga	Faculty Hospital Brno	Czech Republic
Smirnova	Nadezda	North Estonian Center	Estonia
Sofia	Machado	Hospital Clínico Universitario de Valencia	Spain
Soler Pedrola:	Maria	Hospital Son Llatzer	Spain
Söndergaard	Sören	Sahlgrenska University Hospital	Sweden
Sõrmus	Alar	Tartu University Hospital	Estonia
Sørvoll	Ingvild	University Hospital of North Norway, Tromsø	Norway
	Hausberg	·····, ····, ····	
Soumelidis	Christos	Ippokrateio Hippokrateion General Hospital of Thessaloniki	Greece
Spindler Yesel	Alenka	University Medicine Centre Ljubljana	Slovenia
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Stevanovic	Ana	University Hospital Aachen	Germany
Stevikova	Jordana	Faculty hospital, Nitra , Slovak republic	Slovakia
Stivan	Sabina	University Medicine Centre Ljubljana	Slovenia
Štourač	Petr	Faculty Hospital Brno	Czech Republic
Striteska	Jana	University Hospital Hradec Kralove	Czech Republic
Strys	Lydia	University Medical Center Johannes Gutenberg Mainz	Germany
Suljevic	Ismet	University Clinical Center Sarajevo	Bosnia and
Bullevie	isince	Shiversity Gimical Genter Satajevo	Herzegovina
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Tena	Beatriz	Hospital Clinic Barcelona	0
Theodoraki	Kassiani	Aretaieion University Hospital	Spain Greece
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Tikuisis	Popatas		Lithuania
Tolós	Renatas	Vilnius University Hospital - Institute of Oncology	Lithuania
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Tomasi Tomasay	Roland	University Hospital Munich	Germany
Tomescu Tomkute	Dana	Fundeni Clinical institute - Intensive Care Unit	Romania Lithuania
	Gabija Dilor	Vilnius University Hospital - Santariskiu Clinics	Lithuania
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Trepenaitis	Darius	Kaunas Medical University Hospital, Hospital of Lithuanian University of Health Sciences	Lithuania
Troyan	Galina	Zaporizhzhia State Medical University	Ukraine
Unic-Stojanovic	Dragana	Cardiovascular Institute Dedinje Belgrade	Serbia
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Uranjek	Jasna	General Hospital Slovenj Gradec	Slovenia
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van Dasselaar	Nick	Reinier De Graaf Gasthuis Delft	Netherlands
Van Limmen	Jurgen	Ghent University Hospital	Belgium

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Veljovic	Milic	Military Medical Academy	Serbia
Vera Bella	Jorge	Hospital San Jorge	Spain
Vercauteren	Marcel	UZA	Belgium
Verdouw	Bas	Reinier De Graaf Gasthuis Delft	Netherlands
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Vieira	Helena	Hospital da Universidade de Coimbra	Portugal
Villar	Tania	Hospital Mar-Esperança. Parc de Salut Mar	Spain
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Voje	Minca	University Medicine Centre Ljubljana	Slovenia
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Von Langen	Daniel	Medical University Hospital Innsbruck	Austria
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Werner	Eva	University Hospital Regensburg	Germany
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Wigmore	Thimothy	Royal Marsden Hospital	United Kingdom
Wittmann	Maria	University Hospital Bonn	Germany
Wright	Margaret	Hospital James Paget University Hospital NHS Foundation Trust	United Kingdom
Wunder	Christian	University Hospital of Wuerzburg	Germany
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