ORIGINAL ARTICLE - BRAIN TRAUMA



Variation in neurosurgical management of traumatic brain injury: a survey in 68 centers participating in the CENTER-TBI study

Thomas A. van Essen ^{1,2} • Hugo F. den Boogert ³ • Maryse C. Cnossen ⁴ • Godard C. W. de Ruiter ² • Iain Haitsma ⁵ • Suzanne Polinder ⁴ • Ewout W. Steyerberg ^{4,6} • David Menon ⁷ • Andrew I. R. Maas ⁸ • Hester F. Lingsma ⁴ • Wilco C. Peul ^{1,2} • on behalf of the CENTER-TBI Investigators and Participants

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Abstract

Background Neurosurgical management of traumatic brain injury (TBI) is challenging, with only low-quality evidence. We aimed to explore differences in neurosurgical strategies for TBI across Europe.

Methods A survey was sent to 68 centers participating in the Collaborative European Neurotrauma Effectiveness Research in Traumatic Brain Injury (CENTER-TBI) study. The questionnaire contained 21 questions, including the decision when to operate (or not) on traumatic acute subdural hematoma (ASDH) and intracerebral hematoma (ICH), and when to perform a decompressive craniectomy (DC) in raised intracranial pressure (ICP).

Results The survey was completed by 68 centers (100%). On average, 10 neurosurgeons work in each trauma center. In all centers, a neurosurgeon was available within 30 min. Forty percent of responders reported a thickness or volume threshold for evacuation of an ASDH. Most responders (78%) decide on a primary DC in evacuating an ASDH during the operation, when swelling is present. For ICH, 3% would perform an evacuation directly to prevent secondary deterioration and 66% only in case of clinical deterioration. Most respondents (91%) reported to consider a DC for refractory high ICP. The reported cut-off ICP for DC in refractory high ICP, however, differed: 60% uses 25 mmHg, 18% 30 mmHg, and 17% 20 mmHg. Treatment strategies varied substantially between regions, specifically for the threshold for ASDH surgery and DC for refractory raised ICP. Also within center variation was present: 31% reported variation within the hospital for inserting an ICP monitor and 43% for evacuating mass lesions.

This article is part of the Topical Collection on Brain trauma

The CENTER-TBI Investigators and Participants and their affiliations are listed at the end of the manuscript in the *Appendix*.

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- ☐ Thomas A. van Essen essen@lumc.nl
- Department of Neurosurgery, Leiden University Medical Center, University Neurosurgical Center Holland (UNCH), Leiden, The Netherlands
- Department of Neurosurgery, Haaglanden Medical Center, University Neurosurgical Center Holland (UNCH), The Hague, The Netherlands
- Department of Neurosurgery, Radboud University Medical Center, Nijmegen, The Netherlands

- Center for Medical Decision Sciences, Department of Public Health, Erasmus Medical Center, Rotterdam, The Netherlands
- Department of Neurosurgery, Erasmus Medical Center, Rotterdam, The Netherlands
- Department of Biomedical Data Sciences, Leiden University Medical Center, Leiden, The Netherlands
- Division of Anaesthesia, Addenbrooke's Hospital, University of Cambridge, Cambridge, UK
- Department of Neurosurgery, Antwerp University Hospital and University of Antwerp, Edegem, Belgium



Conclusion Despite a homogeneous organization, considerable practice variation exists of neurosurgical strategies for TBI in Europe. These results provide an incentive for comparative effectiveness research to determine elements of effective neurosurgical care.

Keywords Traumatic brain injury · Neurosurgery · Practice variation · Acute subdural hematoma

Neurosurgical decision-making in patients with traumatic brain injury (TBI) is often challenging for several reasons. First, no two TBI patients are identical—clinical and radiological findings may differ greatly [26]. Second, there is no high-quality evidence to support the range of possible neurosurgical procedures in TBI. Indications for surgical management are summarized in the Brain Trauma Foundation guidelines, [5] but are merely based on retrospective studies of small groups of selected patients. These guidelines provide general advice on surgical indications for evacuation of acute epidural (EDH), acute subdural (ASDH), and contusions/ intracerebral hematomas (ICH) based on the size of the hematoma and midline shift. The guidance for decompressive surgery is even less clear. It is mostly performed to decrease raised intracranial pressure (ICP), either as a primary procedure in an acute setting, or as a secondary procedure to deal with diffuse edema or peri-contusional swelling. The guidelines state that this latter use of secondary decompression can reduce ICP, but does not necessarily improve outcome [6]. More fundamentally, the rationale for ICP monitoring has been challenged by the BEST TRIP randomized controlled trial (RCT), which found no benefit of a management protocol based on intracranial pressure monitoring, compared to one based on serial imaging and clinical examination. These results have generated doubts regarding ICP monitoring [1, 7, 15, 20, 28]. Overall, there is no clear consensus on the indications, extent, and timing of surgery [32].

This limited high-quality evidence for surgical management in TBI arises from a lack of RCTs, which may be difficult to conduct due to pragmatic, ethical, and methodological barriers [3]; however, observational studies to determine effectiveness are more prone for bias [2]. A promising alternative approach could be comparative effectiveness research (CER) [24, 33]. In this design, the heterogeneity and variability, that trouble RCTs in TBI, are accepted and exploited to study effectiveness of treatments as they occur in real-life practice. The current Collaborative European Neurotrauma Effectiveness Research in Traumatic Brain Injury (CENTER-TBI) study aims to use CER methodology to study treatment effectiveness of several neurosurgical interventions [25].

The aim of this study was to explore differences in neurosurgical strategies for TBI across Europe to provide a context for CENTER-TBI, an up-to-date insight into European neurosurgical management of TBI, and to identify naturally occurring variation between trauma centers in order to identify substrates for neurosurgical research questions that might be answered using CER in the study.

Materials and methods

This study was conducted within the setting of the international observational study CENTER-TBI [25]. Between 2014 and 2015, all centers participating in the international multicenter observational study CENTER-TBI (www.CENTER-TBI.eu) were asked to complete a questionnaire on neurosurgical management of TBI (Supplementary file 1) [9]. The questionnaire was sent to 71 centers (Fig. 1), of which five centers dropped out and two joined in, resulting in 68 eligible centers from Austria (n = 2), Belgium (n = 4), Bosnia Herzegovina (n = 2), Denmark (n = 2), Finland (n = 2), France (n = 7), Germany (n = 4), Hungary (n = 3), Israel (n = 2), Italy (n = 10), Latvia (n = 3), Lithuania (n = 2), Norway (n = 3), Romania (n = 1), Serbia (n = 1), Spain (n = 4), Sweden (n = 2), Switzerland (n = 1), The Netherlands (n = 6), and The United Kingdom (n = 7).

Questionnaire development and administration

We developed a set of questionnaires based on available literature and experts to measure the structure and processes of TBI care in individual centers. Details regarding this process and the questionnaires used are described in a separate paper [9]. Pilot testing was undertaken in 16 of the participating centers and feedback was incorporated into the final design.

One of the questionnaires was on neurosurgical standard practice. This survey contained 21 questions which could broadly be divided into 3 categories: (1) center characteristics and internal structure; (2) general (neuro) surgical trauma care and processes; and (3) site specific neurosurgical management for treating ASDH, EDH, ICH, the use of DC, and policy with regard to orthopedic injuries in the context of patients who had suffered a TBI.

Questions either sought quantitative estimates of key metrics (e.g., annual surgical volume, staff size, ASDH thickness, or ICP thresholds for surgery) or attempted to elicit the "general policy" of the center. To capture the latter, these questions were formulated in two ways: respondents were asked to



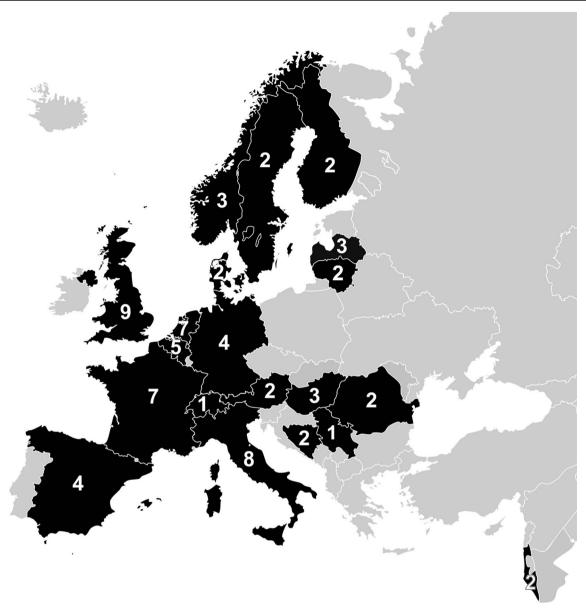


Fig. 1 Centers and countries included in the Collaborative European NeuroTrauma Effectiveness Research in Traumatic Brain Injury (CENTER-TBI) study. Reprinted and updated from Cnossen et al. (2016) with permission from Dr. Cnossen and Maas et al. (2015).

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estimate what the management strategy is in more than three quarters of patients in their center in a given context; or respondents were asked to indicate how often they used a particular surgical technique or how often specific factors influence their decision-making (never = 0-10%, rarely = 10-30%, sometimes = 30-70%, frequently = 70-90%, and always 90-100%). The options "frequently" and "always" were interpreted as "general policy", in line with a previous report [17] and similar to previous publications on other questionnaires [8, 9].

The reliability of the surveys was tested by calculation of concordance in a previous publication [9]. Overall, the median

concordance rates between duplicate questions were 0.81 (range 0.44–0.97) and specifically for the "Neurosurgery" survey 0.78 (range 0.68–0.86).

Analyses

The median and interquartile range (IQR) were calculated for continuous variables, and frequencies were reported along with percentages for categorical variables. Countries were divided into seven geographic regions: Northern Europe (Norway 3, Sweden 2, Finland 2 and Denmark 2 centers), Western Europe (Austria 2, Belgium 4, France 7, Germany



4, Switzerland 1 and The Netherlands 6 centers), The United Kingdom (7 centers), Southern Europe (Italy 10 and Spain 4 centers), Eastern Europe (Hungary 3, Romania 1, Serbia 1 and Bosnia Herzegovina 2 centers), Baltic States (Latvia 3 and Lithuania 2 centers), and Israel (2 centers).

For the following neurosurgical treatment strategies, we quantified regional differences: an absolute cutoff of hematoma thickness as an indication for surgery for ASDH, DC in the primary evacuation of an ASDH, early/pre-emptive surgical evacuation for ICH, and DC as a general policy in case of refractory raised ICP.

To assess the association of region with one of these treatment choices, a logistic regression was performed with treatment choice (general policy or "yes/no") as a dependent variable and the region (categorical) as independent variable. Nagelkerke R2 indicated the variance explained by geographic region. Analyses were done in IBM SPSS Statistics version 20 (IBM, Chicago, IL, USA).

Results

Center characteristics

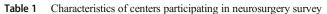
All 68 eligible centers completed the questionnaire on neurosurgery (response rate 100%). Questionnaires were mainly completed by neurosurgeons (n = 53, 78%), followed by local CENTER-TBI investigators (mainly research physicians or nurses: 19%). On average, 10 neurosurgeons (IQR 8–13) and four trauma surgeons (IQR 0–12) worked in each center. All centers reported that neurosurgical coverage was available 24 h a day/7 days a week, either by way of in-house availability of a qualified neurosurgeon (47%), or the availability of such an individual in less than 30 min (53%) (Table 1).

General (neuro) surgical care and processes

Treatment decisions regarding cranial surgical interventions in TBI patients within the critical care ER and ICU period are in most centers determined by the neurosurgeon (n = 65, 96%), followed by the orthopedic surgeons and neuro-intensivist in respectively 3% (n = 2) and 1% (n = 1). Urgent neurosurgical interventions (ICP monitor device insertion not included) for life-threatening traumatic intracranial lesions, are made by the neurosurgeon in 98.5% and trauma surgeons in 1.5% of the centers. Raised ICP will almost always be incorporated in decision-making, the time of day almost never (Fig. 2).

With regard to extremity fractures, the general policy in 59 (87%) centers was so-called damage control with priority for TBI and delayed definitive treatment of the limb fractures (Table 2). This policy is protocolized in 21 centers (22%).

Of all centers, 58 (85%) estimated the space-occupying effect of traumatic lesions on the surrounding tissue by



Characteristic	N completed	No. (%) or median (IQR)
Profession of respondent	68	
Neurologist		3 (4)
Neurosurgeon		53 (78)
Trauma surgeon		3 (4)
ED physician		1 (2)
Intensivist		$(1)^{a}$
Administrative staff member		11 (16) ^a
CENTER-TBI local investigator		13 (19) ^a
Volume of surgeries in 2013 ^c		
ASDH	59	25 (15–49)
ICH/contusion	58	10 (5–21)
EDH	59	10 (5–19)
DC		
Hemicraniectomy	57	10 (5–16)
Bifrontal	57	0 (0–2)
Removal bone flap	55	1 (0–3)
Ventriculostomy	57	7 (2–21)
Cranioplasty	56	10 (6–14)
Depressed skull fracture	57	5 (2–12)
Staffing (FTE)		
Neurosurgeons	66	10 (8–13)
Residents in training	65	5 (3–8)
Residents not in training	61	0 (0–3)
Trauma surgeons	64	4 (0–12)
Organization of care		
Neurosurgical decision making in ICU	68	
Neurosurgeon		65 (96)
Trauma surgeon		1 (3)
Neurologist		0
Neurointensivist or general intensivist		1 (2)
24/7 neurosurgical coverage ^b	68	
Qualified neurosurgeon in-house		32 (47)
Resident neurosurgery in-house		30 (44)
Neurosurgeon within 30 min		36 (53)
Neurosurgical resident within 30 min		11 (16)
Neurosurgeon more than 30 min		0 (0)

ASDH acute subdural hematoma, EDH epidural hematoma, ICH intracerebral hematoma, DC decompressive craniectomy, FTE full time equivalent, ICU intensive care unit



^a Numbers do not add up because the local investigators also depicted their profession and one responder declared to be an intensivist as well as an administrative staff member

^b Multiple options possible

^c Head trauma-related surgeries

calculation of the thickness of the hematoma and midline shift on CT. A quarter of centers used actual volume measurement to make surgical decisions (Table 2).

Neurosurgical management of ASDH, EDH, ICH, and the use of decompressive craniectomy

ASDH provided the highest volume of neurosurgical TBI cases, on average 25 cases per year. When performing a DC (for any indication), hemicraniectomy was the preferential technique, and bifrontal craniectomy was rarely performed (Table 1). Less than half of the centers (n = 27, 40%) reported an absolute threshold for evacuating an ASDH. Four out of 10 centers generally incorporate age in their decision for evacuating an ASDH (Table 2 and Fig. 2).

ICH were seldom operated upon pre-emptively, but 67% of centers reported undertaking delayed surgery in the event of deterioration. Almost a third of centers reported within-center variations between individual neurosurgeons in decisions regarding surgical evacuation of contusions or traumatic ICH.

Only a very low proportion of centers would routinely perform a DC at the time of evacuation of either ASDH or ICH (respectively 6% and 1.5% of the centers). For refractory raised ICP, most centers (n = 64, 91%) would consider a decompressive craniectomy, while 32 (47%) see this as a general policy in their center (Fig. 3, Table 2 and figure in supplementary file 2). Ninety-six percent (n = 65) reported to have a specific threshold for DC in refractory raised ICP. This was most commonly specified as 25 mmHg (n = 39, 58%), followed by 30 mmHg (n = 12, 18%) and 20 mmHg (n = 11, 17%).

Guidelines and practice variation

Overall, the reported adherence to the BTF guidelines was high (Fig. 4). The use of surgical interventions and specific indications for these interventions varied substantially within and between regions (Table 3). Surgical evacuation of ICH was only performed in the Baltic States and Southern Europe and geographic region explained 35% of the variance in use of the intervention. Having a specific threshold for ASDH surgery and employing a DC for refractory-raised ICP showed the largest within-region and also between-region variation. Lastly, when directly asked whether variation in specific management strategies exist, respectively 31% and 43% indicated to have a structural variation within their center staff with regard to ICP sensor insertion and mass lesion evacuation (Table 4).

Discussion

The aim of this study was to explore differences in neurosurgical strategies for TBI across Europe. We found substantial variability in practice and thereby provide useful indications regarding potential substrates for CER in CENTER-TBI. The structures and processes of neurosurgical care are generally homogeneous across centers with a comparable number of neurosurgeons, similar organization of neurosurgical coverage and uniform organization of responsibility for most surgical decisions on the ER and ICU. The indications for surgery, however, differ substantially with high within-region and between-region practice variations.

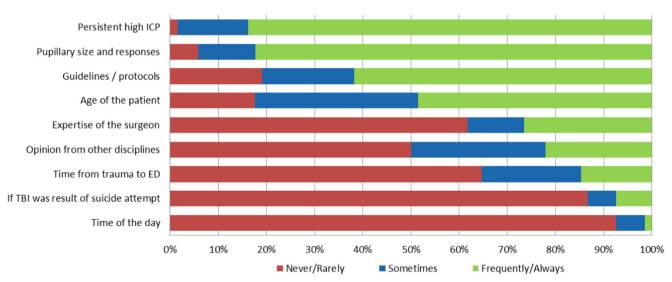


Fig. 2 Factors of influence on neurosurgical decision-making. Shown are the percentages of centers that would be never/rarely, sometimes or frequently/always influenced by the described factors in the decision to

perform neurosurgical procedures. Question was completed by all 68 centers. ICP: intracranial pressure; ED: Emergency Department ^B Other factors were not predetermined but were specified by responders



 Table 2
 Neurosurgical treatment policy of traumatic brain injury

Characteristic	N completed	No. (%) or mean (sd)
Structural estimation of mass lesions on CT ^a	68	
Visual intuition (e.g., no actual measurement)		27 (40)
Width, diameter and/or amount of MLS of the mass lesion		58 (85)
Volume measurements with imaging software		11 (16)
Volume measurements with direct calculation		17 (25)
Other		1 (2)
ASDH operation determinants		
Age considered important in surgery decision ^d	68	26 (42)
Size (volume or thickness) threshold for surgery	68	27 (40)
Minimum volume or thickness:	28 ^b	
15 mm		2 (3)
10 mm		16 (24)
10 mm and/or > 5 mm MLS		2 (3)
5 mm		3 (4)
ASDH thickness > width of cranium		3 (4)
Midline shift > thickness ASDH		2 (3)
DC indications	68	
Routine		4 (6)
Intra-operative brain swelling		59 (86)
Sometimes as a second procedure in case of uncontrollable ICP		5 (7)
Never		0 (0)
ICH/contusion operation determinants		
General policy	68	
Pre-emptive (to prevent deterioration)		2 (3)
Delayed (after deterioration)		45 (66)
Variable (depends on surgeon)		18 (27)
Other		3 (4)
DC indications	68	
Routine		1 (2)
Intra-operative brain swelling		55 (81)
Sometimes as a delayed procedure in case of uncontrollable ICP		10 (15)
Never		2 (3)
Raised ICP determinants		
DC employed > 70% of refractory high ICP cases	68	32 (46)
Mostly early DC (within 6–12 h of refractory ICP)	64	32 (47)
Mostly late DC (as last resort to control ICP)	64	32 (47)
ICP threshold for DC	68	65 (96)
Raised ICP threshold for DC (mmHg):	64 ^c	
30		12 (18)
25		39 (60)
20		11 (17)
15		1 (2)
Not standardized		1 (2)
DC indications considered ^a		
Pre-emptive in raised ICP (not last resort)		7 (10)
Refractory raised ICP (last resort)	68	64 (91)
CT evidence of raised ICP		9 (13)
Intra-operative brain swelling		45 (66)
Routine with every ASDH or ICH evacuation		2 (3)



Table 2 (continued)

Characteristic	N completed	No. (%) or mean (sd)
Policy towards extremity limb fractures ^e		
Damage control		59 (87)
Definitive care	68	9 (13)

MLS midline shift, BTF Brain Trauma Foundation, ICP intracranial pressure, hrs hours

Contemporary neurosurgical care

There are no recent comparable studies providing an overview of neurosurgical management on this scale. Two recent national surveys, in The United Kingdom and the Republic of Ireland and The Netherlands, have shown a comparable variability among neurosurgeons regarding the decision to evacuate an ASDH or to perform a primary DC [21, 34].

When comparing our results to existing—much older—surveys, evacuation of a traumatic ICH seems to be less often considered than in the past [11, 30]. Our results are concordant with older surveys in reporting variable use of DC for refractory raised ICP, despite the DECRA trial (the RECUEicp was not published

yet) [12, 19]. Interestingly, although the mostly applied cutoff for DC in refractory is reported to be 25 mmHg (60%), a lower value, 20 mmHg, and a higher value, 30 mmHg, are both reported to be used in almost 20% of centers.

More broadly, our results replicate past data that suggest poor guideline adherence and practice variability. Rayan et al. showed that in only 17% of a random sample of (brain) trauma patients care was delivered according to the BTF guidelines [31]. Of note, in the current study, surveys were sent to the centers between 2014 and 2015, so the more recent, updated BTF guidelines were not published yet, although the update was for medical management mainly (except DC in refractory IC) [6].

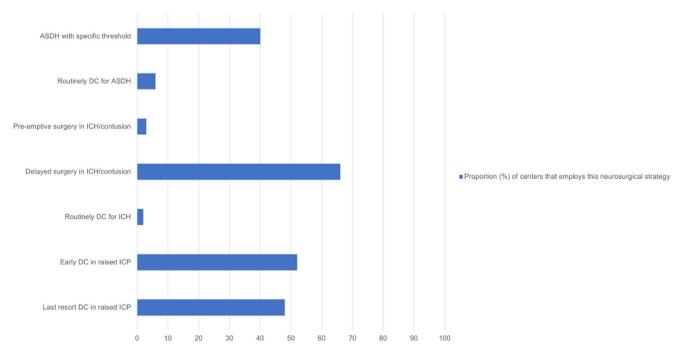


Fig. 3 Treatment indications for neurosurgical interventions. Shown are the proportions of centers that generally have these specific preferences with regard to operating or not in ASDH, ICH, and raised intracranial

pressure, respectively. ASDH: acute subdural hematoma; DC: decompressive craniectomy; ICH: intracerebral hematoma; ICP: intracranial pressure



^a Multiple options possible

^b One responder did not report a threshold for surgery while answering a specific threshold (10 mm)

^c One responder reported to employ a threshold for DC in raised ICP while not giving their specific threshold

^d The question was whether the responder considers if the decision on surgery in acute SDH is influenced by age (based on a general consensus in their respective center)

^e Damage control is focused on the TBI. All extremity fractures are stabilized, but definitive treatment delayed. Definitive care: the extremity fractures are operated as soon as possible

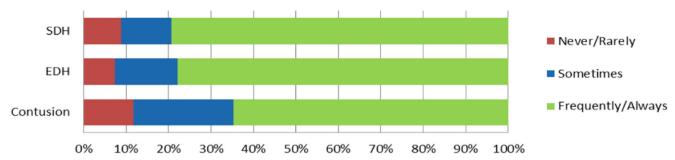


Fig. 4 BTF guideline adherence. Shown are the percentages of centers that reported to never/rarely, sometimes or frequently/always follow the Brain Trauma Foundation guidelines for the management of SDH, EDH,

or contusions. Question was completed by 68 of the 68 centers. TBI: traumatic brain injury; SDH: subdural hematoma; EDH: epidural hematoma

Comparable questionnaires on other aspects of TBI care have recently been published for ER and ICU management that, without exception, show practice variation [8, 9, 14, 18]. Practice variation has also been reported for other life-threatening or emergency disorders including ruptured abdominal aneurysm [4] and the spontaneous intracerebral hemorrhage [16].

Strengths and limitations

A strength of the current study is the methodology that we used to investigate practice variation. First, detailed questions were posed to shed light on specific clinical decisions with regard to neurosurgical interventions. Subsequently, (objective) answers on amounts (volume load, mostly from in-hospital registries) were combined with qualitative information (estimations of general policies, using two approaches). When integrated with the high response rate and low amount of missing data in 68 centers, this overview provides a complete picture of reported neurosurgical care across Europe.

This study also had weaknesses. First, responses to the questionnaire may have been biased by the abstract nature of the questions posed, which neglected to provide a more concrete clinical context for judgments about reported practice.

Although the respondents were experienced neurosurgeons with a scientific background, the difficulty of weighing individual patient characteristics with potentially fatal consequences can never be fully captured by a theoretical survey. In particular, the rational decision-making can obviously be completely different due to the cognitive biases of neurosurgeons in the acute critical care period.

Second, there might be a concern as to how well the individual neurosurgeon respondent can represent the general center neurosurgical policy. Although we urged the respondent to report the general consensus on treatment at their center rather than individual management preferences (see Supplementary file 1), neurosurgical strategies may still be variable within centers between neurosurgeons; however, we did capture a qualitative assessment of this intra-center variability (Table 4). Third, we did not fully account for inherent regional variations such as evidence knowledge, caseload, and casemix due to referral patterns or admission policies, as a potential explanation for differences in neurosurgery policies. Variations in evidence knowledge for some questions, such as those on guidelines, are important. Moreover, while we did asses the center's caseload and casemix, the caseload and casemix of the (individual) respondent was not specifically asked. Fourth, the questions dealt with individual decisions in isolation, rather than the more complex real-life situation

 Table 3
 Within- and between-region variation in surgical management

Decision	Northern Europe	Western Europe	United Kingdom	Southern Europe	Eastern Europe	Baltic States	Israel	Nagelkerke R ² value
ASDH								
- Size threshold for evacuation	56	29	0	29	71	80	100	0.34
- Routine or intraoperative DC <i>ICH/contusion</i>	89	92	100	100	86	80	100	0.17
- Pre-emptive surgery <i>Refractory</i> raised <i>ICP</i>	0	0	0	7	0	20	0	0.35
- DC	44	37	29	57	43	80	100	0.15

ASDH acute subdural hematoma, ICH intracerebral hematoma, DC decompressive craniectomy, ICP intracranial pressure

Table presents the proportion (%) of respondent within each region that indicated that they used the described strategy as their general policy for patients with respectively ASDH, ICH, or refractory raised ICP. The Nagelkerke R² value represents the variation in treatment that can be explained by the region



Table 4 Neurosurgical decision making

Characteristic	N completed	No (%)	
Structural variation ^a ICP monitor insertion	68		
No		47 (69)	
Yes		21 (31)	
Structural variation ^a mass lesion evacuation	65		
No		29 (43)	
Yes		29 (43)	
Depending on lesion type		7 (10)	

ED emergency department, GCS Glasgow Coma Scale

where several competing priorities need to be addressed. Fifth, the reports may have been biased (in varying extents) towards how centers would have been liked to be perceived, rather than a faithful report of actual clinical policy and practice. This issue will be addressed by a planned comparison of these Provider Profiling responses with actual treatment strategies employed in patient-level data from these centers in the CENTER-TBI Core study.

Finally, our study sample represents centers participating in TBI-research which are likely specialized neurotrauma centers with a tendency to have practice that is skewed towards up-to-date knowledge. An example is the fact that almost half of all centers stated to have a neurosurgeon in house 24 h a day. When studying all centers in Europe providing care to TBI patients, variability might be even larger.

Implications

Our results should be interpreted in combination with the current evidence on the effectiveness of different surgical strategies. For the use of DC in refractory raised ICP due to diffuse swelling, two RCTs have provided useful guidance. The DECRA trial showed that early use of DC for modest rises in ICP was associated with worse outcomes [12]. More recently however, after the conduct of this survey, the RESCUEicp trial showed that, when used for refractory severe intracranial hypertension, DC can save lives, but results in an excess of severely disabled survivors [19]. It is clear that the intervention is not uniformly beneficial: while some functional improvements occur by 12 months, many survivors remain severely disabled. Rescue-ICP was not published yet at the conduct of this study. In our study, the majority of centers indicated that DC is often employed for both indications (preemptive and last resort).

With regard to focal lesions, a recent study suggested that in patients with an ASDH an aggressive approach towards evacuation is associated with better outcome compared to a conservative approach [35]. Similar trends were noted in the STITCH-trauma trial, which suggested better outcome with early surgical management of ICH [29]. In our study, a minority of centers considers an early strategy for ICH evacuation.

Lastly, DC in the primary evacuation of an ASDH seems to be associated with more favorable outcomes [22]. There is no class 1 evidence, although the research question is currently being challenged in an RCT (Rescue-ASDH; ISRCT87370545). In the current survey standard (in some cases preventive) DC in ASDH evacuation is rarely employed but mostly done in case of intraoperative swelling.

There may be several explanations for the practice variation that we observed. Although high practice variation rates can be a sign of poor implementation of evidence-based care, in this context it probably reflects the lack of strong evidence to underpin practice. In such a low evidence context, clinical decisions are not driven by careful consideration or penetration of the evidence, but by local customs and surgical training, handed down over the years from one surgeon to the other in a given center (or country). The professional cultural drivers that underpin such learned treatment preferences are resistant to change, and provide an important hurdle to the design and conduct of randomized studies for neurosurgical interventions in TBI [27].

Additionally, even where the results of RCTs are available, it is possible that many neurosurgeons do not think the RCT results applicable to their (individual) patients, or restrict their focus to short term clinical outcomes such as mortality and complication rates (instead of long-term clinical or patient reported outcomes). [13]

The results of the questionnaire point out burning clinical questions for neurosurgery in TBI. For ASDH and ICH, important questions include whether to operate or not, the timing of operative evacuation, and whether or not a primary DC should be undertaken. Future studies should address these questions. For DC, the variation should lead to studies exploring the lack of evidence penetration, in addition to studying effectiveness of DC in refractory raised ICP.

While RCTs may provide the security of randomization as a basis for examining answering these questions, RCTs have no successful history in TBI due to various reasons [24]. The CENTER-TBI Provider Profiling exercise has revealed large practice variation that can be related to variation in patient outcome [23]. Such a CER approach may be a pragmatic alternative to RCTs.

Therefore, different steps are required. Firstly, to specify, ideally a-priori, how and where treatment variation occurs. This was one of the goals of this provider profiling. Secondly, the CENTER-TBI Core Study will need to collect patient-level data from a large variety of centers, capturing the range of treatment variation and relate it to outcome. The main challenge is to disentangle the effect of specific surgical strategies in a center from other regional care variation that might



^a Structural variation refers to a situation in which one or more of the clinicians are generally more likely to perform the (diagnostic) intervention than others

affect outcome. To do so, we propose random-effect models in which the effect of "surgical strategy" on outcome is estimated with adjustment for other between-hospital differences in a random effect for hospital [10, 34, 35].

Conclusions

This survey study explored differences in neurosurgical strategies for TBI. Current neurosurgical care differs within Europe (and Israel), while the organization of trauma centers does not. This variation in practice likely reflects the lack of high-quality evidence for these important, potentially life-saving, emergency neurosurgical interventions. In addition, local professional culture may drive practice in ways that are not dependent on the availability or penetration of evidence. The resulting entrenched practice variation does not facilitate equipoise that makes RCTs easy to deliver. CER may provide a pragmatic approach to generate evidence on optimal neurosurgical strategies for TBI patients.

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Compliance with ethical standards

Conflict of interest The authors declare that they have no conflicts of interest

All procedures performed in studies involving human participants were in accordance with ethical standards of the institutional and national research committee and with the 1964 Helsinki declaration and its later amendments.

Appendix

The CENTER-TBI Investigators and Participants and their affiliations:

Ackerlund Cecilia¹, Adams Hadie ², Agnoletti Vanni ³, Allanson Judith ⁴, Amrein Krisztina ⁵, Andaluz Norberto ⁶, Andelic Nada ⁷, Andreassen Lasse ⁸, Antun Azasevac ⁹, Anke Audny ¹⁰, Antoni Anna ¹¹, Ardon Hilko ¹², Audibert Gérard ¹³, Auslands Kaspars ¹⁴, Azouvi Philippe ¹⁵, Azzolini Maria Luisa ¹⁶, Baciu Camelia ¹⁷, Badenes Rafael ¹⁸, Bartels Ronald ¹⁹, Barzó Pál ²⁰, Bauerfeind Ursula ²¹, Beauvais Romuald ²², Beer Ronny ²³, Belda Francisco Javier ¹⁸, Bellander Bo-Michael ²⁴, Belli Antonio²⁵, Bellier Rémy ²⁶, Benali Habib ²⁷, Benard Thierry ²⁶, Berardino Maurizio ²⁸, Beretta Luigi ¹⁶, Beynon Christopher ²⁹, Bilotta Federico ¹⁸, Binder Harald ¹¹, Biqiri Erta ¹⁷, Blaabjerg Morten ³⁰, den Boogert Hugo ¹⁹,

Bouzat Pierre 31, Bragge Peter 32, Brazinova Alexandra 33, Brinck Vibeke ³⁴, Brooker Joanne ³⁵, Brorsson Camilla ³⁶, Buki Andras ³⁷, Bullinger Monika ³⁸, Calappi Emiliana ³⁹, Calvi Maria Rosa ¹⁶, Cameron Peter ⁴⁰, Carbayo Lozano Guillermo ⁴¹, Carbonara Marco³⁹, Carise Elsa ²⁶, Carpenter K. ⁴², Castaño-León Ana M. ⁴³, Causin Francesco ⁴⁴, Chevallard Giorgio ¹⁷, Chieregato Arturo ¹⁷, Citerio Giuseppe 45, 46, Cnossen Maryse 47, Coburn Mark 48, Coles Jonathan ⁴⁹, Coles-Kemp Lizzie ⁵⁰, Collett Johnny ⁵⁰, Cooper Jamie D. 51, Correia Marta 52, Covic Amra 53, Curry Nicola 54, Czeiter Endre ⁵⁵, Czosnyka Marek ⁵⁶, Dahyot-Fizelier Claire ²⁶, Damas François ⁵⁷, Damas Pierre ⁵⁸, Dawes Helen ⁵⁹, De Keyser Véronique ⁶⁰, Della Corte Francesco ⁶¹, Depreitere Bart ⁶², de Ruiter Godard C.W. ⁶³, Dilvesi Dula ⁹, Ding Shenghao ⁶⁴, Dippel Diederik ⁶⁵, Dixit Abhishek ⁶⁶, Donoghue Emma ⁴⁰, Dreier Jens ⁶⁷, Dulière Guy-Loup ⁵⁷, Eapen George ⁶⁸, Engemann Heiko ⁵³, Ercole Ari ⁶⁶, Esser Patrick ⁵⁹, Ezer Erzsébet ⁶⁹, Fabricius Martin ⁷⁰, Feigin Valery L. ⁷¹, Feng Junfeng ⁶⁴, Foks Kelly ⁶⁵, Fossi Francesca ¹⁷, Francony Gilles ³¹, Freo Ulderico ⁷², Frisvold Shirin ⁷³, Furmanov Alex ⁷⁴, Gagliardo Pablo ⁷⁵, Galanaud Damien ²⁷, Gantner Dashiell ⁴⁰, Gao Guoyi ⁷⁶, Geleijns Karin ⁴², George Pradeep ¹, Ghuysen Alexandre ⁷⁷, Giga Lelde ⁷⁸, Giraud Benoit ²⁶, Glocker Ben ⁷⁹, Golubovic Jagos ⁹, Gomez Pedro A. ⁴³, Grossi Francesca ⁶¹, Gruen Russell L. ⁸⁰, Gupta Deepak ⁸¹, Haagsma Juanita A. ⁴⁷, Haitsma Iain ⁸², Hartings Jed A. ⁸³, Helbok Raimund ²³, Helseth Eirik ⁸⁴, Hertle Daniel ³⁰, Hoedemaekers Astrid 85, Hoefer Stefan 53, Horton Lindsay ⁸⁶, Huijben Jilske ⁴⁷, Hutchinson Peter J. ², Håberg Asta Kristine 87, Jacobs Bram 88, Jankowski Stefan 68, Jarrett Mike ³⁴, Jelaca Bojan⁹, Jiang Ji-yao ⁷⁶, Jones Kelly ⁸⁹, Kamnitsas Konstantinos ⁷⁹, Karan Mladen ⁶, Katila Ari ⁹⁰, Kaukonen Maija 91, Kerforne Thomas 26, Kivisaari Riku 91, Kolias Angelos G. ², Kolumbán Bálint ⁹², Kompanje Erwin 93, Kolundzija Ksenija 94, Kondziella Daniel 70, Koskinen Lars-Owe ³⁶, Kovács Noémi ⁹², Lagares Alfonso ⁴³, Lanyon Linda ¹, Laureys Steven ⁹⁵, Lecky Fiona ⁹⁶, Ledig Christian ⁷⁹, Lefering Rolf ⁹⁷, Legrand Valerie ⁹⁸, Lei Jin ⁶⁴, Levi Leon 99, Lightfoot Roger ¹⁰⁰, Lingsma Hester ⁴⁷, Loeckx Dirk ¹⁰¹, Lozano Angels ¹⁸, Maas Andrew I.R. ⁶⁰, MacDonald Stephen ¹⁰², Maegele Marc ¹⁰³, Majdan Marek ³³, Major Sebastian ¹⁰⁴, Manara Alex 105, Manley Geoffrey 106, Martin Didier 107, Martin Leon Francisco ¹⁰¹, Martino Costanza ³, Maruenda Armando 18, Maréchal Hugues 57, Masala Alessandro 3, Mattern Julia ²⁹, McFadyen Charles ⁶⁶, McMahon Catherine ¹⁰⁸, Melegh Béla ¹⁰⁹, Menon David ⁶⁶, Menovsky Tomas ⁶⁰, Morganti-Kossmann Cristina 110, Mulazzi Davide 39, Muraleedharan Visakh ¹, Murray Lynnette⁴⁰, Mühlan Holger 111, Nair Nandesh 60, Negru Ancuta 112, Nelson David 1, Newcombe Virginia 66, Nieboer Daan 47, Noirhomme Quentin ⁹⁵, Nyirádi József ⁵, Oddo Mauro ¹¹³, Oldenbeuving Annemarie 114, Oresic Matej 115, Ortolano Fabrizio 39, Palotie Aarno 116, 117, 118, Parizel Paul M. 119, Patruno Adriana 120,



Payen Jean-François ³¹, Perera Natascha ²², Perlbarg Vincent ²⁷, Persona Paolo ¹²¹, Peul Wilco ⁶³, Piippo-Karjalainen Anna ⁹¹, Pili Floury Sébastien ¹²², Pirinen Matti ¹¹⁶, Ples Horia ¹¹², Poca Maria Antonia ¹²³, Polinder Suzanne ⁴⁷, Pomposo Inigo ⁴¹, Posti Jussi ⁹⁰, Puybasset Louis ¹²⁴, Radoi Andreea ¹²³, Ragauskas Arminas 125, Raj Rahul 91, Rambadagalla Malinka 126, Real Ruben 53, Rehorčíková Veronika 33, Rhodes Jonathan 127, Ripatti Samuli 116, Rocka Saulius 125, Roe Cecilie ¹²⁸, Roise Olav ¹²⁹, Roks Gerwin ¹³⁰, Rosand Jonathan ¹³¹, Rosenfeld Jeffrey ¹¹⁰, Rosenlund Christina ¹³², Rosenthal Guy 74, Rossaint Rolf 48, Rossi Sandra 121, Rueckert Daniel 79, Rusnák Martin 133, Sacchi Marco 17, Sahakian Barbara ⁶⁶, Sahuquillo Juan ¹²³, Sakowitz Oliver 134, 135, Sala Francesca 120, Sanchez-Porras Renan 134, Sandor Janos ¹³⁶, Santos Edgar ²⁹, Sasu Luminita ⁶¹, Savo Davide ¹²⁰, Schäffer Nadine ¹⁰³, Schipper Inger ¹³⁷, Schlößer Barbara ²¹, Schmidt Silke ¹¹¹, Schoechl Herbert ¹³⁸, Schoonman Guus 130, Schou Rico Frederik 139, Schwendenwein Elisabeth ¹¹, Schöll Michael ²⁹, Sir Özcan ¹⁴⁰, Skandsen Toril ¹⁴¹, Smakman Lidwien ⁶³, Smeets Dirk 101, Smielewski Peter 56, Sorinola Abayomi 142, Stamatakis Emmanuel ⁶⁶, Stanworth Simon ⁵⁴, Steinbüchel Nicole ¹⁴³, Stevanovic Ana 48, Stevens Robert 144, Stewart William 145, Steyerberg Ewout W. 47, 146, Stocchetti Nino 147, Sundström Nina ³⁶, Synnot Anneliese ^{34, 148}, Taccone Fabio Silvio ¹⁸, Takala Riikka 90, Tamás Viktória 142, Tanskanen Päivi 91, Taylor Mark Steven ³³, Te Ao Braden ⁷¹, Tenovuo Olli ⁹⁰, Telgmann Ralph ⁵³, Teodorani Guido ¹⁴⁹, Theadom Alice ⁷¹, Thomas Matt ¹⁰⁵, Tibboel Dick ⁴², Tolias Christos ¹⁵⁰, Tshibanda Jean-Flory Luaba ¹⁵¹, Trapani Tony ⁴⁰, Tudora Cristina Maria 112, Vajkoczy Peter 67, Vallance Shirley 43, Valeinis Egils ⁷⁸, Van der Steen Gregory⁶⁰, van der Jagt Mathieu 152, van der Naalt Joukje 88, van Dijck Jeroen T.J.M. ⁶³, van Essen Thomas A. ⁶³, Van Hecke Wim ¹⁰¹, van Heugten Caroline ⁵⁹, Van Praag Dominique ⁶⁰, Vande Vyvere Thijs ¹⁰¹, Van Waesberghe Julia ⁴⁸, Vanhaudenhuyse Audrey ^{27, 95}, Vargiolu Alessia ¹²⁰, Vega Emmanuel ¹⁵³, Velt Kimberley ⁴⁷, Verheyden Jan ¹⁰¹, Vespa Paul M. ¹⁵⁴, Vik Anne ¹⁵⁵, Vilcinis Rimantas ¹⁵⁶, Vizzino Giacinta ¹⁷, Vleggeert-Lankamp Carmen 63, Volovici Victor 82, Voormolen Daphne ⁴⁷, Vulekovic Peter ⁹, Vámos Zoltán ⁶⁹, Wade Derick ⁵⁹, Wang Kevin K.W. ¹⁵⁷, Wang Lei ⁶⁴, Wessels Lars ¹⁵⁸, Wildschut Eno ⁴², Williams Guy ⁶⁶, Wilson Lindsay ⁸⁶, Winkler Maren K.L. ¹⁰⁴, Wolf Stefan ¹⁵⁸, Ylén Peter ¹⁵⁹, Younsi Alexander ²⁹, Zaaroor Menashe ⁹⁹, Zhihui Yang ¹⁶⁰, Ziverte Agate ⁷⁸, Zumbo Fabrizio ³.

- ¹ Karolinska Institutet, INCF International Neuroinformatics Coordinating Facility, Stockholm, Sweden.
- ² Division of Neurosurgery, Department of Clinical Neurosciences, Addenbrooke's Hospital & University of Cambridge, Cambridge, UK.
- ³ Department of Anesthesia & Intensive Care, M. Bufalini Hospital, Cesena, Italy.

- ⁴ Department of Clinical Neurosciences, Addenbrooke's Hospital & University of Cambridge, Cambridge, UK.
- ⁵ János Szentágothai Research Centre, University of Pécs, Pécs, Hungary.
 - ⁶ University of Cincinnati, Cincinnati, Ohio, United States.
- ⁷ Division of Surgery and Clinical Neuroscience, Department of Physical Medicine and Rehabilitation, Oslo University Hospital and University of Oslo, Oslo, Norway.
- ⁸ Department of Neurosurgery, University Hospital Northern Norway, Tromso, Norway.
- ⁹ Department of Neurosurgery, Clinical centre of Vojvodina, Faculty of Medicine, University of Novi Sad, Novi Sad, Serbia.
- Department of Physical Medicine and Rehabilitation, University hospital Northern Norway.
- ¹¹ Trauma Surgery, Medical University Vienna, Vienna, Austria.
- ¹² Department of Neurosurgery, Elisabeth-Tweesteden Ziekenhuis, Tilburg, the Netherlands.
- ¹³ Department of Anesthesiology & Intensive Care, University Hospital Nancy, Nancy, France.
 - ¹⁴ Riga Eastern Clinical University Hospital, Riga, Latvia.
- ¹⁵ Raymond Poincare hospital, Assistance Publique Hopitaux de Paris, Paris, France.
- ¹⁶ Department of Anesthesiology & Intensive Care, S Raffaele University Hospital, Milan, Italy.
 - ¹⁷ NeuroIntensive Care, Niguarda Hospital.
- ¹⁸ Department Anesthesiology and Surgical-Trauma Intensive Care, Hospital Clinic Universitari de Valencia, Spain.
- ¹⁹ Department of Neurosurgery, Radboud University Medical Center.
- ²⁰ Department of Neurosurgery, University of Szeged, Szeged, Hungary.
- ²¹ Institute for Transfusion Medicine (ITM), Witten/ Herdecke University, Cologne, Germany.
- ²² International Projects Management, ARTTIC, Munchen, Germany.
- ²³ Department of Neurology, Neurological Intensive Care Unit, Medical University of Innsbruck, Innsbruck, Austria.
- ²⁴ Department of Neurosurgery & Anesthesia & intensive care medicine, Karolinska University Hospital, Stockholm, Sweden.
- ²⁵ NIHR Surgical Reconstruction and Microbiology Research Centre, Birmingham, UK.
 - ²⁶ Intensive care Unit, CHU Poitiers, Poitiers, France.
- ²⁷ Anesthesie-Réanimation, Assistance Publique Hopitaux de Paris, Paris, France.
- ²⁸ Department of Anesthesia & ICU, AOU Città della Salute e della Scienza di Torino Orthopedic and Trauma Center, Torino, Italy.
- ²⁹ Department of Neurosurgery, University Hospital Heidelberg, Heidelberg, Germany.



- ³⁰ Department of Neurology, Odense University Hospital, Odense, Denmark.
- ³¹ Department of Anesthesiology & Intensive Care, University Hospital of Grenoble, Grenoble, France.
- ³² BehaviourWorks Australia, Monash Sustainability Institute, Monash University, Victoria, Australia.
- ³³ Department of Public Health, Faculty of Health Sciences and Social Work, Trnava University, Trnava, Slovakia.
 - ³⁴ Quesgen Systems Inc., Burlingame, California, USA.
- ³⁵ Australian & New Zealand Intensive Care Research Centre, Department of Epidemiology and Preventive Medicine, School of Public Health and Preventive Medicine, Monash University, Melbourne, Australia.
- ³⁶ Department of Neurosurgery, Umea University Hospital, Umea, Sweden.
- ³⁷ Department of Neurosurgery, University of Pecs and MTA-PTE Clinical Neuroscience MR Research Group and Janos Szentagothai Research Centre, University of Pecs, Hungarian Brain Research Program, Pecs, Hungary.
- ³⁸ Department of Medical Psychology, Universitätsklinikum Hamburg-Eppendorf, Hamburg, Germany.
- ³⁹ Neuro ICU, Fondazione IRCCS Cà Granda Ospedale Maggiore Policlinico, Milan, Italy.
- ⁴⁰ ANZIC Research Centre, Monash University, Department of Epidemiology and Preventive Medicine, Melbourne, Vitoria, Australia.
- ⁴¹ Department of Neurosurgery, Hospital of Cruces, Bilbao, Spain.
- ⁴² Intensive Care and Department of Pediatric Surgery, Erasmus Medical Center, Sophia Children's Hospital, Rotterdam, The Netherlands.
- ⁴³ Department of Neurosurgery, Hospital Universitario 12 de Octubre, Madrid, Spain.
- ⁴⁴ Department of Neuroscience, Azienda Ospedaliera Università di Padova, Padova, Italy.
 - ⁴⁵ NeuroIntensive Care, ASST di Monza, Monza, Italy.
- ⁴⁶ School of Medicine and Surgery, Università Milano Bicocca, Milano, Italy.
- ⁴⁷ Department of Public Health, Erasmus Medical Center-University Medical Center, Rotterdam, The Netherlands.
- ⁴⁸ Department of Anaesthesiology, University Hospital of Aachen, Aachen, Germany.
- ⁴⁹ Department of Anesthesia & Neurointensive Care, Cambridge University Hospital NHS Foundation Trust, Cambridge, UK.
- ⁵⁰ Movement Science Group, Oxford Institute of Nursing, Midwifery and Allied Health Research, Oxford Brookes University, Oxford, UK.
- ⁵¹ School of Public Health & PM, Monash University and The Alfred Hospital, Melbourne, Victoria, Australia.
- ⁵² Radiology/MRI department, MRC Cognition and Brain Sciences Unit, Cambridge, UK.

- ⁵³ Institute of Medical Psychology and Medical Sociology, Universitätsmedizin Göttingen, Göttingen, Germany.
 - ⁵⁴ Oxford University Hospitals NHS Trust, Oxford, UK.
- ⁵⁵ Department of Neurosurgery, University of Pecs and MTA-PTE Clinical Neuroscience MR Research Group and Janos Szentagothai Research Centre, University of Pecs, Hungarian Brain Research Program (Grant No. KTIA 13 NAP-A-II/8), Pecs, Hungary.
- ⁵⁶ Brain Physics Lab, Division of Neurosurgery, Dept of Clinical Neurosciences, University of Cambridge, Addenbrooke's Hospital, Cambridge, UK.
 - ⁵⁷ Intensive Care Unit, CHR Citadelle, Liège, Belgium.
 - ⁵⁸ Intensive Care Unit, CHU, Liège, Belgium.
- ⁵⁹ Movement Science Group, Faculty of Health and Life Sciences, Oxford Brookes University, Oxford, UK.
- ⁶⁰ Department of Neurosurgery, Antwerp University Hospital and University of Antwerp, Edegem, Belgium.
- ⁶¹ Department of Anesthesia & Intensive Care, Maggiore Della Carità Hospital, Novara, Italy.
- ⁶² Department of Neurosurgery, University Hospitals Leuven, Leuven, Belgium.
- ⁶³ Dept. of Neurosurgery, Leiden University Medical Center, Leiden, The Netherlands and Dept. of Neurosurgery, Medical Center Haaglanden, The Hague, The Netherlands.
- ⁶⁴ Department of Neurosurgery, Renji Hospital, Shanghai Jiaotong University School of Medicine, Shanghai, China.
- ⁶⁵ Department of Neurology, Erasmus MC, Rotterdam, the Netherlands.
- ⁶⁶ Division of Anaesthesia, University of Cambridge, Addenbrooke's Hospital, Cambridge, UK.
- ⁶⁷ Neurologie, Neurochirurgie und Psychiatrie, Charité Universitätsmedizin Berlin, Berlin, Germany.
- ⁶⁸ Neurointensive Care, Sheffield Teaching Hospitals NHS Foundation Trust, Sheffield, UK.
- ⁶⁹ Department of Anaesthesiology and Intensive Therapy, University of Pécs, Pécs, Hungary.
- Departments of Neurology, Clinical Neurophysiology and Neuroanesthesiology, Region Hovedstaden Rigshospitalet, Copenhagen, Denmark.
- National Institute for Stroke and Applied Neurosciences, Faculty of Health and Environmental Studies, Auckland University of Technology, Auckland, New Zealand.
- ⁷² Department of Medicine, Azienda Ospedaliera Università di Padova, Padova, Italy.
- ⁷³ Department of Anesthesiology and Intensive care, University Hospital Northern Norway, Tromso, Norway.
- ⁷⁴ Department of Neurosurgery, Hadassah-hebrew University Medical center, Jerusalem, Israel.
- ⁷⁵ Fundación Instituto Valenciano de Neurorrehabilitación (FIVAN), Valencia, Spain.
- ⁷⁶ Department of Neurosurgery, Shanghai Renji hospital, Shanghai Jiaotong University/school of medicine, Shanghai, China.

- ⁷⁷ Emergency Department, CHU, Liège, Belgium.
- ⁷⁸ Pauls Stradins Clinical University Hospital, Riga, Latvia.
- ⁷⁹ Department of Computing, Imperial College London, London, UK.
- ⁸⁰ Lee Kong Chian School of Medicine, Nanyang Technological University, Singapore; and Monash University, Australia.
- ⁸¹ Department of Neurosurgery, Neurosciences Centre & JPN Apex trauma centre, All India Institute of Medical Sciences, New Delhi-110,029, India.
- $^{\rm 82}$ Department of Neurosurgery, Erasmus MC, Rotterdam, the Netherlands.
- ⁸³ Department of Neurosurgery, University of Cincinnati, Cincinnati, Ohio, USA.
- ⁸⁴ Department of Neurosurgery, Oslo University Hospital, Oslo, Norway.
- 85 Department of Intensive Care Medicine, Radboud University Medical Center.
- ⁸⁶ Division of Psychology, University of Stirling, Stirling, UK.
- ⁸⁷ Department of Medical Imaging, St. Olavs Hospital and Department of Neuroscience, Norwegian University of Science and Technology, Trondheim, Norway.
- ⁸⁸ Department of Neurology, University Medical Center Groningen, Groningen, Netherlands.
- ⁸⁹ National Institute for Stroke & Applied Neurosciences of the AUT University, Auckland, New Zealand.
- ⁹⁰ Rehabilitation and Brain Trauma, Turku University Central Hospital and University of Turku, Turku, Finland.
 - ⁹¹ Helsinki University Central Hospital.
- ⁹² Hungarian Brain Research Program Grant No. KTIA13 NAP-A-II/8, University of Pécs, Pécs, Hungary.
- ⁹³ Department of Intensive Care and Department of Ethics and Philosophy of Medicine, Erasmus Medical Center, Rotterdam, The Netherlands.
- ⁹⁴ Department of Psichiatry, Clinical centre of Vojvodina, Faculty of Medicine, University of Novi Sad, Novi Sad, Serbia.
- ⁹⁵ Cyclotron Research Center, University of Liège, Liège, Belgium.
- ⁹⁶ Emergency Medicine Research in Sheffield, Health Services Research Section, School of Health and Related Research (ScHARR), University of Sheffield, Sheffield, UK.
- ⁹⁷ Institute of Research in Operative Medicine (IFOM), Witten/Herdecke University, Cologne, Germany.
- ⁹⁸ VP Global Project Management CNS, ICON, Paris, France
- ⁹⁹ Department of Neurosurgery, Rambam Medical Center, Haifa, Israel.
- ¹⁰⁰ Department of Anesthesiology & Intensive Care, University Hospitals Southhampton NHS Trust, Southhampton, UK.

- ¹⁰¹ icoMetrix NV, Leuven, Belgium.
- ¹⁰² Cambridge University Hospitals, Cambridge, UK.
- ¹⁰³ Cologne-Merheim Medical Center (CMMC), Department of Traumatology, Orthopedic Surgery and Sportmedicine, Witten/Herdecke University, Cologne, Germany.
- ¹⁰⁴ Centrum für Schlaganfallforschung, Charité Universitätsmedizin Berlin, Berlin, Germany.
- ¹⁰⁵ Intensive Care Unit, Southmead Hospital, Bristol, Bristol, UK.
- ¹⁰⁶ Department of Neurological Surgery, University of California, San Francisco, California, USA.
 - ¹⁰⁷ Department of Neurosurgery, CHU, Liège, Belgium.
- ¹⁰⁸ Department of Neurosurgery, The Walton centre NHS Foundation Trust, Liverpool, UK.
- ¹⁰⁹ Department of Medical Genetics, University of Pécs, Pécs, Hungary.
- ¹¹⁰ National Trauma Research Institute, The Alfred Hospital, Monash University, Melbourne, Victoria, Australia.
- ¹¹¹ Department Health and Prevention, University Greifswald, Greifswald, Germany.
- ¹¹² Department of Neurosurgery, Emergency County Hospital Timisoara, Timisoara, Romania.
 - ¹¹³ Centre Hospitalier Universitaire Vaudois.
- ¹¹⁴ Department of Intensive Care, Elisabeth-Tweesteden Ziekenhuis, Tilburg, the Netherlands.
- ¹¹⁵ Department of Systems Medicine, Steno Diabetes Center, Gentofte, Denmark.
- ¹¹⁶ Institute for Molecular Medicine Finland, University of Helsinki, Helsinki, Finland.
- Analytic and Translational Genetics Unit, Department of Medicine; Psychiatric & Neurodevelopmental Genetics Unit, Department of Psychiatry; Department of Neurology, Massachusetts General Hospital, Boston, MA, USA.
- ¹¹⁸ Program in Medical and Population Genetics; The Stanley Center for Psychiatric Research, The Broad Institute of MIT and Harvard, Cambridge, MA, USA.
- ¹¹⁹ Department of Radiology, Antwerp University Hospital and University of Antwerp, Edegem, Belgium.
- ¹²⁰ NeuroIntenisve Care Unit, Department of Anesthesia & Intensive Care, ASST di Monza, Monza, Italy.
- ¹²¹ Department of Anesthesia & Intensive Care, Azienda Ospedaliera Università di Padova, Padova, Italy.
- ¹²² Intensive Care Unit, CHRU de Besançon, Besançon, France.
- ¹²³ Department of Neurosurgery, Vall d'Hebron University Hospital, Barcelona, Spain.
- ¹²⁴ Department of Anesthesiology and Critical Care, Pitié -Salpêtrière Teaching Hospital, Assistance Publique, Hôpitaux de Paris and University Pierre et Marie Curie, Paris, France.
- ¹²⁵ Department of Neurosurgery, Kaunas University of technology and Vilnius University, Vilnius, Lithuania.
 - ¹²⁶ Rezekne Hospital, Latvia.



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- Department of Anesthesia, Critical Care & Pain Medicine NHS Lothian & University of Edinburg, Edinburgh, UK.
- Department of Physical Medicine and Rehabilitation, Oslo University Hospital/University of Oslo, Oslo, Norway.
- ¹²⁹ Division of Surgery and Clinical Neuroscience, Oslo University Hospital, Oslo, Norway.
- ¹³⁰ Department of Neurology, Elisabeth-TweeSteden Ziekenhuis, Tilburg, the Netherlands.
- ¹³¹ Broad Institute, Cambridge MA Harvard Medical School, Boston MA, Massachusetts General Hospital, Boston MA, USA.
- ¹³² Department of Neurosurgery, Odense University Hospital, Odense, Denmark.
- ¹³³ International Neurotrauma Research Organization, Vienna, Austria.
- ¹³⁴ Klinik für Neurochirurgie, Klinikum Ludwigsburg, Ludwigsburg, Germany.
 - ¹³⁵ University Hospital Heidelberg, Heidelberg, Germany.
- ¹³⁶ Division of Biostatistics and Epidemiology, Department of Preventive Medicine, University of Debrecen, Debrecen, Hungary.
- ¹³⁷ Department of Traumasurgery, Leiden University Medical Center, Leiden, The Netherlands.
- ¹³⁸ Department of Anaesthesiology and Intensive Care, AUVA Trauma Hospital, Salzburg, Austria.
- Department of Neuroanesthesia and Neurointensive Care, Odense University Hospital, Odense, Denmark.
- ¹⁴⁰ Department of Emergency Care Medicine, Radboud University Medical Center.
- ¹⁴¹ Department of Physical Medicine and Rehabilitation, St. Olavs Hospital and Department of Neuroscience, Norwegian University of Science and Technology, Trondheim, Norway.
- ¹⁴² Department of Neurosurgery, University of Pécs, Pécs, Hungary.
 - ¹⁴³ Universitätsmedizin Göttingen, Göttingen, Germany.
- ¹⁴⁴ Division of Neuroscience Critical Care, John Hopkins University School of Medicine, Baltimore, USA.
- ¹⁴⁵ Department of Neuropathology, Queen Elizabeth University Hospital and University of Glasgow, Glasgow, UK.
- ¹⁴⁶ Dept. of Department of Biomedical Data Sciences, Leiden University Medical Center, Leiden.
- ¹⁴⁷ Department of Pathophysiology and Transplantation, Milan University, and Neuroscience ICU, Fondazione IRCCS Cà Granda Ospedale Maggiore Policlinico, Milano, Italy
- Italy.

 148 Cochrane Consumers and Communication Review
 Group, Centre for Health Communication and Participation,
 School of Psychology and Public Health, La Trobe University,
 Melbourne, Australia.

- ¹⁴⁹ Department of Reahabilitation, M. Bufalini Hospital, Cesena, Italy.
- ¹⁵⁰ Department of Neurosurgery, Kings college London, London, UK.
 - ¹⁵¹ Radiology/MRI Department, CHU, Liège, Belgium.
- ¹⁵² Department of Intensive Care, Erasmus MC, Rotterdam, the Netherlands.
- ¹⁵³ Department of Anesthesiology-Intensive Care, Lille University Hospital, Lille, France.
- ¹⁵⁴ Director of Neurocritical Care, University of California, Los Angeles, USA.
- ¹⁵⁵ Department of Neurosurgery, St. Olavs Hospital and Department of Neuroscience, Norwegian University of Science and Technology, Trondheim, Norway.
- ¹⁵⁶ Department of Neurosurgery, Kaunas University of Health Sciences, Kaunas, Lithuania.
- ¹⁵⁷ Department of Psychiatry, University of Florida, Gainesville, Florida, USA.
- ¹⁵⁸ Interdisciplinary Neuro Intensive Care Unit, Charité Universitätsmedizin Berlin, Berlin, Germany.
 - ¹⁵⁹ VTT Technical Research Centre, Tampere, Finland.
 - ¹⁶⁰ University of Florida, Gainesville, Florida, USA

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