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# **Abstract**

**Background:** The optimal hemodynamic targets and management of patients with acute brain injury are not completely elucidated, but recent evidence points to important impact on clinical outcomes. We performed an international survey with the aim to investigate the practice in the hemodynamic targets, monitoring, and management of patients with acute ischemic stroke (AIS), intracranial hemorrhage (ICH) and subarachnoid hemorrhage (SAH).

**Methods:** This survey was endorsed by the European Society of Intensive Care (ESICM). An electronic questionnaire of 76 questions divided in 4 sections (general information, AIS, ICH, SAH specific questions) was available between January 2022 to March 2022 on the ESICM website.

**Results:** One hundred fifty-four healthcare professionals from 36 different countries and at least 98 different institutions answered the survey. Routine echocardiography is routinely performed in 37% of responders in AIS, 34% in ICH and 38% in SAH. Cardiac output monitoring is used in less than 20% of cases by most of the responders. Cardiovascular complications are the main reason for using advanced hemodynamic monitoring, and norepinephrine is the most common drug used to increase arterial blood pressure. Most responders target fluid balance to neutral (62% in AIS, 59% in ICH,44% in SAH), and normal saline is the most common fluid used. Large variability was observed regarding the blood pressure targets.

**Conclusions:** Hemodynamic management and treatment in patients with acute brain injury from cerebrovascular diseases vary largely in clinical practice. Further research is required to provide clear guidelines to physicians for the hemodynamic optimization of this group of patients.

**Keywords:** Subarachnoid hemorrhage, Acute ischemic stroke, Intracranial hemorrhage, Hemodynamic management



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# Take-home messages

- Heterogeneity exists in the approach of hemodynamic management of cerebrovascular disease.
- Pharmacologic strategies to achieve targets and monitoring targets are different across centers.
- Advanced, serial hemodynamic monitoring is not a standard of care in brain injured patients and it is reserved to most severe cases, although benefit might extend to hemodynamically stable patients.

#### Introduction

The hemodynamic management of cerebrovascular diseases is of fundamental importance in order to minimize secondary brain damage [1, 2]. Cerebral autoregulation may be impaired in acute brain injured (ABI) patients such as ischemic stroke (AIS), intracranial hemorrhage (ICH) and subarachnoid hemorrhage (SAH), and consequently brain perfusion and its oxygenation might be compromised. In these pathologies, cerebral blood flow is extremely sensitive to systemic blood variation, and therefore suboptimal hemodynamic management (including fluids and vasoactive administration policy and hemodynamic monitoring) of these patients may increase ischemic areas, promote hemorrhagic evolution and develop secondary brain injury [2–4].

Clinical strategies of hemodynamic management of these pathologies are still based on moderate/low quality of evidence retrieved from relatively small studies, and no strong recommendations are reported in the most updated consensuses on these topics [5–9]. Both the hemodynamic monitoring/targets and fluid management/balance should consider the physiological peculiarities of ABI patients, in particular given the disturbed cerebral autoregulation, tendency for increased intracranial pressure (ICP) and decreased cerebral perfusion pressure (CPP) and in those affected by an acute cardiac dysfunction associated with sympathetic stimulation and catecholamine release [9].

In the AIS the choice of the optimal pressure target is still debated. A recent multicentric randomized trial did not find difference in long-term clinical outcomes in AIS patients between intensive [target systolic arterial pressure (SAP) 130–140 mm Hg within 1 h] or "standard" (SAP < 180 mm Hg) blood pressure-lowering treatment over 72 h [10]. In ICH patients, despite contrasting results of trials investigating hemodynamic management, monitoring, as well as the targets of blood pressure to be achieved, the most recent guidelines recommend lowering SAP to a target range of 130 to 140 mm Hg in patients presenting with acute ICH of mild to moderate severity and SBP between 150 and 220 mm Hg [11–13]. In SAH patients, historical concepts of hemodynamic management, such as the "triple H" approach (hypertension, hypervolemia, and hemodilution) [14] have been challenged [15, 16]. The aim of this survey is to investigate the current hemodynamic practices worldwide for the management of AIS, ICH and SAH.

#### Methods

This is an international survey, proposed by the Neuro-intensive Care (NIC) and the Cardiovascular Dynamics (CD) sections of the European Society of Intensive Care Medicine (ESICM). No ethical approval was necessary for the development of this survey. The Steering committee, which included representatives of both ESICM sections, performed a non-systematic review of the literature (i.e., guidelines and consensus papers) on hemodynamic management of AIS, ICH and SAH patients and created a questionnaire of 76 questions divided in 4 sections (general information, acute ischemic stroke, intracranial hemorrhage, subarachnoid hemorrhage). The survey was reviewed and approved by the Research committee of the ESICM and was tested in a pilot cohort of potential respondents, including the Steering Committee. The survey was available on the ESICM website between January 2022 to March 2022. Three reminders were sent to potential responders. The questionnaire was created considering some issues around this topic, such as low levels of evidence, lack of good-quality studies and controversial results from observational trials. The survey was designed to identify (a) characteristics of the participants demographics, type of hospital/specialty and available neuromonitoring tools; (b) protocols for the hemodynamic management of this population; (c) hemodynamic targets used in this group of patients; (d) clinical management and use of hemodynamic monitoring in this population. Advanced hemodynamic monitoring was defined as the use of any further tool able to estimate cardiac output (i.e., calibrated/ uncalibrated hemodynamic monitoring based on arterial waveform analysis; bioreactance technology, etc.).

The target audience was ESICM members who had agreed to participate in ESICM surveys at the time of their membership registration and who treat patients with cerebrovascular diseases in their clinical practice. The investigators invited NIC or CD sections members, asking them to involve more respondents locally. Participants did not receive compensation for their participation in the survey, which was distributed via the ESICM office, thus protecting data confidentiality and anonymity. The survey was registered within the ESICM Survey portfolio.

# Statistical methods

Data from the questionnaire were exported as a comma-separated value report from the Surveymonkey<sup>®</sup> software package and subsequently stored as an Excel file (Microsoft Corp, Redmond, WA). Descriptive statistics were computed for all study variables. The results are presented as numbers and percentage. Continuous variables are reported as mean [standard deviation (SD)] or median [interquartile range (IQR)], whereas categorical variables were reported as frequency and proportion.

#### Results

# **General characteristics**

As shown in Table 1, 154 healthcare professionals from 36 different countries [87.1% from Europe] answered this survey. Answers came from 98 different institutions [mainly academic/teaching hospitals (83.1%) and neurocritical care centers (78.6%)] with

**Table 1** General characteristics of participants

General inform	ation					
World region o	f responders					
Europe	Australia	South America	India	Indonesia	Arabic coun- tries	North America
128 (87.1%)	1 (0,7%)	5 (3,4%)	7 (4,8%)	1 (0,7%)	2 (1,4%)	3 (2%)
Type of institu	ution:					
Academic/ teaching hospital	District Hos- pital	Non-teaching hospital	Private non-ad pital	cademic hos-		
128 (83,1%)	3 (1,9%)	19 (12,3%)	4 (2,6%)			
Number of be	ds in your institu	tion:				
< 250	250-500	500-750	750–1000	> 1000		
14 (9,1%)	27 (17,5%)	33 (21,4%)	37 (24%)	43 (28%)		
Catchment ar	ea population:					
< 100,000	100,000- 250,000	250,000- 500,000	500,000– 750,000	750,000- 1,000,000	> 1,000,000	
9 (5,8%)	24 (15,6%)	29 (18,8%)	30 (19,5%)	18 (11,7%)	44 (28,6%)	
Are you a cen	ter for neurocritic	cal care?				
Yes	No					
121 (78,6%)	33 (21,4%)					
Critically ill ne	euroscience patie	nts are generally	admitted to:			
Medical ICU	Mixed general- neurocritical care unit	Specialist neurocritical care unit	Surgical ICU	Other		
14 (9,1%)	82 (53,2%)	45 (29,2%)	8 (5,2%)	5 (3,2%)		
Number of ac	ute ischemic stro	ke patients admi	tted to ICU /yea	ır:		
Less than 5	5 to 10	10 to 20	20-30	>30		
3 (1,9%)	20 (13%)	39 (25,3%)	22 (14,3%)	70 (45,5%)		
Number of ac	ute hemorrhagic	stroke patients a	dmitted to ICU	/year:		
Less than 5	5 to 10	10 to 20	20-30	>30		
5 (3,2%)	17 (11%)	30 (19,5%)	31 (20,1%)	71 (46,1%)		
Number of su	barachnoid hem	orrhage patients	admitted to ICU	J /year:		
Less than 5	5 to 10	10 to 20	20-30	>30		
8 (5,2%)	20 (13%)	27 (17,5%)	14 (9,1%)	85 (55,2%)		
Medical staffi	ng of Neurocritic	al care unit/ICU a	dmitting neuro	critical care pati	ents:	
Anesthetist intensivist	General/ respiratory medicine Intensivist	Neurologist Intensivist	Neurosur- geon Inten- sivist	Other		
86 (55,8%)	35 (22,7%)	17 (11%)	2 (1,3%)	14 (9,1%)		
Medical staff	present 24/7:					
Qualified specialist	Fellow	Trainee	Specialist nurse	Telepresence	Other	None
134 (39%)	69 (20%)	68 (20%)	64 (19%)	5 (2%)	6 (2%)	0

ICU, intensive care unit

different capacity, [most of them with > 1,000 patients hospital capacity (28.0%)]. Most of healthcare professionals who responded generally manage > 30 AIS (45.5%), ICH (46.1%) and SAH (55.2%) patients per year.

According to the responders, most of the patients with neurological critical illness are admitted to mixed general-neurocritical care unit (53%), and less commonly to specialist

neurocritical care unit (29%) or other ICUs/wards. The majority of neurocritical patients are managed by anesthesiology intensivist (56%), followed by general/respiratory medicine intensivist (23%) and neurology intensivist (11%). Finally, in most of the centers (39%) a qualified specialist is 24/7 present.

# Hemodynamic management in AIS patients

A standardized protocol for arterial blood pressure management in patients with AIS is used by the 46% of the responders.

The use of echocardiography is quite heterogeneous: most of the participants (37%) declared to perform routine echocardiography during the first 24 h to rule out possible acute left ventricular dysfunction, while 9% only in selected cases. Cardiac output monitoring is used in less than 20% of cases by most (62%) of the participants; capillary refill time is used by less than half of the participants (40%) (Fig. 2).

An arterial pressure target of < 160/90 mmHg is reported in the 35% of patients eligible for thrombolysis, and in the 30% of patients after mechanical thrombectomy. In the population not eligible for intravenous thrombolysis, blood pressure targets of < 160/90 mmHg were reported in 29% of responders and < 185/110 mmHg in 27% of responders.

In AIS patients admitted to ICU, hemodynamic monitoring is mostly performed (60%) by electrocardiogram (ECG) and invasive blood pressure (IBP), while less frequently by non-invasive blood pressure, NIBP and ECG (34%). Advanced hemodynamic monitoring is rarely used (10%) (Table 2 and Fig. 1).

The most common indication for advanced hemodynamic monitoring was developing cardiovascular complications (34%), followed by sepsis/septic shock (25%), difficult blood pressure management (16%) and worsening of neurological symptoms (6%).

Initial management of low blood pressure is mostly based on MAP-based protocol (55%), no specific protocol is reported in the 14% of the answers, and norepinephrine is largely the most used first-line agent in the treatment of hypotension in AIS patients with preserved heart function (80%). The preferred first-line agent for lowering blood pressure is urapidil (24% of the answers), followed by beta-blockers (21%), calcium channel blockers (15%) and clonidine (14%).

Most of the participants monitor fluid balance every hour (31%) and a neutral fluid balance is the target according to 62% of the participants. A positive (13%) or negative (6%) balance are rarely defined as a hemodynamic target. When fluids administration is required, crystalloids [ringer lactate (35%) or normal saline (32%)] are usually preferred.

# Hemodynamic management in patients with hemorrhagic stroke

In ICH patients, a standardized protocol for arterial blood pressure management is used by the 43% of the responders. Echocardiography was reported to be performed by the 34% of the responders and CO monitoring is used in less than 20% of patients by the 53% of the responders. CRT is performed by 37% the participants. The commonest indication for advanced monitoring is the development of cardiovascular complications (35%), followed by the requirevent of vasoactive drugs (26%).

The most commonly reported arterial pressure target was < 140/90 mmHg (36%), followed by < 160/90 mmHg (29%). Hemodynamic monitoring is performed by NIBP and

 Table 2
 Questions and answers on acute ischemic stroke

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No answer <10%	<10%	10–30%	30-20%	> 20%
28 (18%) 23 (15%)	23 (15%)	37 (24%)	28 (18%)	38 (25%)
Which is th	e percentage of Al	S that undergo mech	anic thrombectomy/ye	Which is the percentage of AIS that undergo mechanic thrombectomy/year in your institution?
No answer	<10%	10–30%	30-20%	>20%
28 (18%) 48 (31%)	48 (31%)	32 (21%)	29 (19%)	17 (11%)
Do you hav	re a standardized p	protocol for arterial blo	ood pressure manager	Do you have a standardized protocol for arterial blood pressure management in patients with ischemic stroke?
No answer Yes	Yes	No		
28 (18%) 71 (46%)	71 (46%)	55 (36%)		
Do you rou	tinely perform an	echocardiography at t	the bedside in AIS pati	Do you routinely perform an echocardiography at the bedside in AIS patients during the first 24 h of admission to rule out possible acute LV dysfunction?

No a	No answer Yes	Yes	No	Selected cases
28 (1	28 (18%)	57 (37%)	55 (36%)	14 (9%)
How	/ many /	AIS patients receive sta	ndard cardiac output n	How many AIS patients receive standard cardiac output monitoring in your institution?
No a	ınswer	No answer <20%	30-50%	20%
28 (1	28 (18%)	95 (62%)	19 (12%)	12 (8%)
2		On you issued a see see and a see at the bedeine in AIS patients?	I test at the bedside in	AIS nationts?

No answer Yes	Yes	No		
28 (18%) 62 (40%)	62 (40%)	64 (42%)		
Which is th	e arterial blood	Which is the arterial blood pressure target that you use in patients eligible for iv thrombo	use in patients eligibl	e for iv thrombo
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Which is the	arterial blood pressur	Which is the arterial blood pressure target that you use in patients eligible for iv thrombolysis?	n patients eligible for iν	/ thrombolysis?	
No answer <140/90	<140/90	<160/90	185/110	Reduction of 15% of baseline ABP	Other
28 (18%)	16 (10%)	54 (35%)	33 (21%)	15 (10%)	8 (5%)
Which is the	arterial blood pressur	Which is the arterial blood pressure target that you use in patients not eligible for iv thrombolysis?	n patients not eligible f	or iv thrombolysis?	
No answer <140/90	<140/90	<160/90	185/110	Reduction of 15% of baseline ABP	Other
28 (18%)	13 (8%)	44 (29%)	42 (27%)	17 (11%)	10 (7%)
Which is the	arterial blood pressur	e target that you use ir	n patients eligible after	Which is the arterial blood pressure target that you use in patients eligible after mechanical thrombectomy?	
No answer <140/90	<140/90	<160/90	185/110	Reduction of 15% of baseline ABP	Other
28 (18%)	28 (18%)	46 (30%)	26 (17%)	11 (7%)	15 (10%)

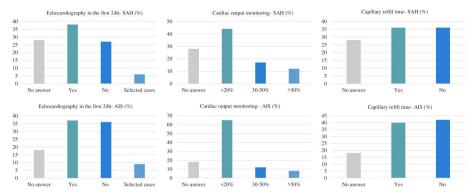
Table 2 (continued)

Which stand	dard hemodynamic m	Which standard hemodynamic monitoring do you use in	case of patient with isc	case of patient with ischemic stroke? (more than one anwer)	han one anwer)			
No answer	No answer Basic monitoring (NIBP, ECG)	Basic monitoring (invasive blood pressure, ECG)	ıre, ECG)	Advanced non-invasive hemodynamic monitoring	ve hemodynamic	Advanced invasive monitoring	Echocardiography	
28 (18%)	53 (34%)	92 (60%)		16 (10%)		(%9) 6	44 (29%)	
Which is the pressure)?	e commonest indicatic	Which is the commonest indication for starting an advanced hemodynamic monitoring in acute ischemic stroke patients in your unit? (i.e., the use of more than invasive arterial blood pressure)?	ced hemodynamic mo	nitoring in acute ische	mic stroke patients	in your unit? (i.e., the u	ise of more than invasi	ve arterial blood
No answer	Patients developing cardiovascular cor cations (i.e., pulmonary edema, cardio- genic shock, etc.)	Patients developing cardiovascular complications (i.e., pulmonary edema, cardiogenic shock, etc.)	Sepsis/septic shock	Sepsis/septic shock Difficult blood pressure management	re management	Patients developing neurological complications (i.e., neuroworsening)		
28 (18%)	52 (34%)		39 (25%)	25 (16%)		10 (6%)		
How do you	ı initially manage hyp	How do you initially manage hypotension in AIS patients						
No answer	No answer Mean arterial pressure-based protocol	No specific protocol	Fluids	Cardiac output-based protocol	protocol			
28 (18%)	85 (55%)	21 (14%)	14 (9%)	6 (4%)				
Which drug	is are used as first line	Which drugs are used as first line to increase blood pressure in your ICU in patients with preserved heart function?	ure in your ICU in patie	ents with preserved hea	art function?			
No answer	No answer Norepinephrine	Dobutamine	Vasopressin	Metaraminol				
28 (18%) 123 (80%)	123 (80%)	2 (1%)	1 (< 1%)	1 (< 1%)				
Which drug	is are used as second l	Which drugs are used as second line to increase blood pre	essure in your ICU in pa	essure in your ICU in patients with preserved heart function?	heart function?			
No answer	No answer Vasopressin	Epinephrine	Dobutamine	Dopamine	Norepinephrine	Terlipressin	Phenylephrine	None of the above
28 (18%)	58 (38%)	27 (18%)	19 (12%)	4 (3%)	5 (3%)	2 (1%)	1 (< 1%)	2 (1%)
Which drug	is are used as first line	Which drugs are used as first line to decrease blood pressure in your ICU?	ure in your ICU?					
No answer Urapidil	Urapidil	Beta blockers	Calcium channel blockers	Clonidine	Nitrates	ACE inhibitors	Angiotensin receptor tor antagonists	Azamethonium
28 (18%)	37 (24%)	33 (21%)	23 (15%)	21 (14%)	6 (4%)	3 (2%)	2 (1%)	1 (<1%)
Which drug	is are used as second l	Which drugs are used as second line to decrease blood pressure in your ICU?	ressure in your ICU?					
No answer	No answer Beta blockers	Calcium channel blockers	Clonidine	Urapidil	Nitrates	ACE inhibitors	Angiotensin receptor tor antagonists	Hydralazine

Table 2 (continued)

28 (18%) 34 (22%)	34 (22%)	32 (21%)	27 (18%)	11 (7%)	10 (6%)	7 (5%)	4 (3%)	1 (< 1%)
How freque	ently do you monitor	How frequently do you monitor fluid balance in patients	ts with ischemic stroke?	oke?				
No answer	No answer Every hour	Every two hours	Every 4 h	Every 8 h	Every 12 h	Every 24 h	We don't monitor fluid balance	
28 (18%) 48 (31%)	48 (31%)	22 (14%)	(%9) 6	1 (1%)	22 (14%)	22 (14%)	2 (1%)	
Which is yo	ur net daily fluid bal	Which is your net daily fluid balance target (i.e., all fluids INPUT minus all fluids OUTPUT)?	Is INPUT minus all f	luids OUTPUT)?				
No answer Negative	Negative	Neutral	Positive					
28 (18%) 10 (6%)	10 (6%)	96 (62%)	20 (13%)					
Which is th	Which is the first line fluid therapy you usually use?	py you usually use?						
No answer	No answer Ringer lactate	Normal saline	Plasmalyte	Albumin	Hypertonic saline Other	e Other		
28 (18%)	54 (35%)	49 (32%)	6 (4%)	3 (2%)	2 (1%)	30 (19%)		

ICU, intensive care unit; AIS, acute ischemic stroke; ABP, arterial blood pressure; NIBP, non-invasive blood pressure; ECG, electrocardiogram; ACE, angiotensin-converting enzyme



**Fig. 1** Hemodynamic monitoring in the considered subgroup of patients. Data are expressed as percentage of the overall answers obtained by the survey

ECG in 12%, and IBP and ECG in the 52% of the answers. Advanced hemodynamic monitoring is rarely used (6% of responders).

Initial management of low blood pressure is usually performed by a MAP-based protocol (53%) and the first-line agent in the treatment of hypotension is norepinephrine (74%). Drugs used in these patients as first line for lowering arterial blood pressure are usually beta-blockers (21%), calcium channel blockers (15%) and clonidine (10%). Fluid balance is mostly monitored hourly (31%) and the majority (59%) of participants aim at a neutral daily fluid balance, with the preferred fluids being ringer lactate (33%) and normal saline (28%) (Table 3 and Fig. 1).

#### Hemodynamic management in SAH patients

According to the responders, 44% use a standardized protocol for the blood pressure management of SAH patients.

Echocardiography during the first 24 h to rule out left ventricular dysfunction is performed by 38% of the survey responders. Cardiac output monitoring is usually used in a minority of SAH patients, and CRT is usually performed in these patients by the 36% of the responders. The commonest indication for advanced monitoring in SAH patients is the development of cardiovascular complications (47%), followed by the need for blood pressure drugs (28%), neurological complications (22%) and vasospasm (20%).

The most frequently reported goals for blood pressure before aneurysm treatment are <140/90 mmHg (31%) and <160/90 mmHg (22%). After aneurysm treatment, in patients without vasospasm the most common blood pressure targets are MAP 90 mmHg (24%) and MAP 80 mmHg (21%). Blood pressure in patients with vasospasm is mostly managed by targeting MAP on neurological status (32%) and targeting to a MAP of 100 mmHg (15%). Hemodynamic SAH monitoring is represented by IBP and ECG in the 44% of the answers, and NIBP and ECG in the 12%. Advanced hemodynamic monitoring is rarely used (9% of the answers).

Initial management of low blood pressure is usually performed by MAP-based protocol (49%), and the preferred first-line agent in the treatment of hypotension is norepinephrine (69%).

Fluid balance is mostly monitored every hour (29%) and a neutral daily fluid balance is targeted by most participants (44%), a positive balance by 38 answerers (25%), and a

**Table 3** Questions and answers on hemorrhagic stroke

Do you have a standardized protocol for arterial blood pressure management in patients with hemorrhagic stroke?

**No answer Yes No** 35 (23%) 66 (43%) 53 (34%)

Do you routinely perform an echocardiography at the bedside in patients with hemorrhagic stroke during the first 24 h of admission to rule out possible acute LV dysfunction?

 No answer
 Yes
 No
 Selected cases

 35 (23%)
 52 (34%)
 59 (38%)
 8 (5%)

How many patients with hemorrhagic stroke receive cardiac output monitoring in your institution?

**No answer < 20% 30–50% 50%** 35 (23%) 82 (53%) 23 (15%) 9 (8%)

Do you usually assess CRT (capillary refill time) at the bedside in patients with hemorrhagic stroke as part of routinely hemodynamic assessment?

**No answer Yes No** 35 (23%) 57 (37%) 62 (40%)

Which is the arterial blood pressure target that you use in patients after intracranial hemorrhage?

Which hemodynamic monitoring do you use in case of patient with hemorrhagic stroke?

No answer Basic Advanced Advanced Echocardimonitorinvasive monitornon-invaography ing (NIBP, ing (Invasive hemomonitor-ECG) sive blood dynamic ina pressure, monitor-ECG) ing 35 (23%) 19 (12%) 80 (52%) 2 (1%) 9 (6%) 9 (6%)

Which are the indications for advanced hemodynamic monitoring hemorrhagic stroke patients (i.e., the use of more than invasive arterial blood pressure)?

No answer In all In patients In patients In patients Never Other patients requiring with with neuadmitted drugs to cardiovasrological to the ICU cular (i.e., complicaincrease or reduce pulmonary tions (i.e., arterial edema, neurowblood cardioorsening) pressure genic shock. etc.) (neuroworsening) 35 (23%) 15 (10%) 40 (26%) 54 (35%) 2 (1%) 5 (3%) 3 (2%)

How do you manage hypotension in patients with hemorrhagic stroke?

No specific Cardiac No answer Mean Fluids arterial outputprotocol based pressurebased protocol protocol 35 (23%) 82 (53%) 25 (16%) 7 (5%) 5 (3%)

Which drugs are used as first line to increase blood pressure in your ICU?

No answer Norepi-Epineph-Metarami- Dobu-Vasopres-Dopamine nephrine sin rine nol tamine 35 (23%) 114 (74%) 2 (1%) 1 (< 1%) 1 (< 1%)1 (< 1%) 1 (< 1%)

Which drugs are used as second line to increase blood pressure in your ICU?

No answer Vasopres- Epineph- Dobu- Norepi- Dopamine Phenylesin rine tamine nephrine phrine

Table 3 (continued)

35 (23%)	61 (38%)	24 (16%)	22 (14%)	8 (5%)	4 (3%)	1 (< 1%)	
Which drug	s are used as	first line to	decrease bloo	od pressure ir	n your ICU?		
No answer	Beta blockers	Calcium channel blockers	Clonidine	Urapidil	Nitrates	ACE inhibitors	Angiotensin receptor antagonists
35 (23%)	33 (21%)	23 (15%)	16 (10%)	12 (8%)	7 (5%)	1 (< 1%)	1 (< 1%)
Which drug	s are used as	second line	to decrease l	blood pressui	re in your ICL	J?	
No answer	Beta blockers	Clonidine	Calcium channel blockers	ACE inhibitors	Urapidil	Nitrates	Angiotensin receptor antagonists
35 (23%)	35 (23%)	32 (21%)	22 (14%)	10 (6%)	12 (8%)	7 (5%)	1 (< 1%)
How freque	ently do you i	monitor fluid	l balance?				
No answer	<b>Every hour</b>	Every 2 h	Every 4 h	Every 12 h	Every 24 h		
35 (23%)	48 (31%)	18 (12%)	9 (6%)	23 (15%)	21 (14%)		
Which is yo	ur net daily f	luid balance	target?				
No answer	Negative	Neutral	Positive	Other			
35 (23%)	9 (6%)	91 (59%)	16 (10%)	3 (2%)			
Which is the	e first-line flu	id therapy y	ou usually us	e?			
No answer	Ringer lactate	Normal saline	Hyper- tonic saline	Plasma- lyte	Other		
35 (23%)	52 (33%)	43 (28%)	7 (5%)	6 (4%)	11 (7%)		

ICU, intensive care unit; ABP, arterial blood pressure; NIBP, non-invasive blood pressure; ECG, electrocardiogram; ACE, angiotensin-converting enzyme; CRT, capillary refill time.

negative balance by five (3%) survey responders. Fluids most commonly used in these patients include normal saline (30%) and Ringer lactate (28%) (Table 4 and Fig. 1) (Fig. 2).

# Discussion

Given the importance of hemodynamic monitoring and management in neurocritical care patients, we conducted this cross-sectional study to describe the current practice in hemodynamic management of patients with acute brain injury admitted to ICU. The results of the study may be summarized as follows: (1) the answers were mostly obtained from European centers with medium-to-high surge capacity, managing a high number of the considered neurological diseases; (2) at least one-third of the participants use echocardiography to investigate cardiac function of the patients during the first 24 h from admission, whereas advanced hemodynamic monitoring is rarely routinely adopted, being dedicated to those patients with neurological complications or cardiovascular impairment. CRT is used by about 40% of the participants; (3) only half of the participants use a standardized protocol for blood pressure management in AIS, ICH and SAH patients; (4) MAP or SAP-based protocols are adopted to titrate systemic pressure targets; however, these targets are not universally accepted and consistent across centers; (5) norepinephrine is widely the most commonly adopted drug as first-line drug for managing hypotensive events.

Although hemodynamic management in cerebrovascular diseases is known to be crucial to minimize secondary brain damage, hemodynamic goals that should be achieved are still debated [17]. The clinical pathways to reach these clinical targets (i.e., drugs, fluids) and the hemodynamic monitoring systems that should be used to

Table 4 Questions and answers on subarachnoid hemorrhage

Which is the percentage of patients who undergo aneurysm clipping/year in your institution?

**No answer** < **10%** 10–30% 30–50% 50% 43 (28%) 35 (23%) 29 (19%) 36 (23%) 11 (7%)

Which is the percentage of SAH that undergo aneurism coiling/year in your institution?

**No answer** < **10% 10–30% 30–50% 50%** 43 (28%) 21 (14%) 15 (10%) 27 (18%) 48 (31%)

Do you have a standardized protocol for arterial blood pressure management in patients with subarachnoid hemorrhage?

**No answer Yes No** 43 (28%) 67 (44%) 44 (29%)

Do you routinely perform an echocardiography at the bedside in SAH patients during the first 24 h of admission to rule out possible acute LV dysfunction?

 No answer
 Yes
 No
 Selected cases

 43 (28%)
 59 (38%)
 42 (27%)
 10 (6%)

How many SAH patients receive standard cardiac output monitoring in your institution?

**No answer** < **20**% **30–50**% **50**% 43 (28%) 67 (44%) 26 (17%) 18 (12%)

Do you usually assess CRT at the bedside in SAH patients?

**No answer Yes No** 43 (28%) 55 (36%) 56 (36%)

Which is the arterial blood pressure target that you use in patients before aneurism treatment?

MAP-based No answer < 140/90 < 160/90 185/110 Reduction Other of 15% of protocols the initial arterial blood pressure 43 (28%) 4 (3%) 6 (4%) 47 (31%) 34 (22%) 9 (6%) 11 (7%)

Which is the arterial blood pressure target that you use in patients after aneurism treatment without

vasospasm?

MAP MAP No answer MAP + 10 mmHg MAP Other compared to 80 mmHg 90 mmHg 100 mmHg baseline 43 (28%) 15 (10%) 33 (21%) 37 (24%) 8 (5%) 18 (12%)

Which is the arterial blood pressure target that you use in patients in patients with vasospasm?

No answer MAP + 10 mmHg MAP MAP MAP MAP target Other compared to 80 mmHg 90 mmHg 100 mmHg baseline rological status 43 (28%) 13 (8%) 10 (6%) 23 (15%) 50 (32%) 8 (5%)

Which hemodynamic monitoring do you use in case of patient with subarachnoid hemorrhage with no signs of vasospasm?

No answer Basic monitoring **Basic** Advanced Advanced Echocardi-(NIBP, ECG) monitoring non-invasive invasive ography (Invasive hemomonitoring blood presdynamic sure, ECG) monitoring 43 (28%) 18 (12%) 68 (44%) 5 (3%) 14 (9%) 6 (4%)

track them are not well defined. The role of the echocardiography in ICU has changed in the last decades, becoming patient-oriented, performed and interpreted by the intensivist to customize the therapy at the bedside by reassessing the effects of the

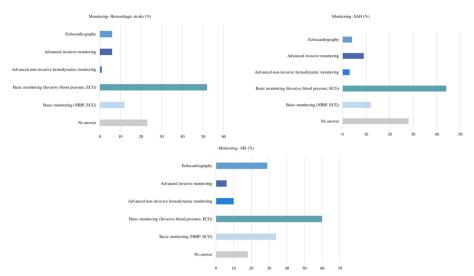
Table 4 (continued)

No answer	In all patients	In patients	In patients	In patients	Vasospasm	Other	Neve
NO answer	admitted to the	requiring drugs to increase or reduce arte- rial blood pressure	with cardio- vascular (i.e., pulmonary edema, cardiogenic shock, etc.) (neurowors- ening)	with neu-	vasuspasiii	Other	Nevel
43 (28%)	12 (8%)	43 (28%)	73 (47%)	34 (22%)	31 (20%)	2 (1%)	3 (2%)
How do you	u manage hypoten:	sion in SAH pati	ents?				
No answer	Mean arterial pressure-based protocol	No specific protocol	Cardiac out- put-based protocol	Fluids			
43 (28%)	76 (49%)	22 (14%)	7 (5%)	6 (4%)			
Which drug	s are used as first li	ine to increase l	olood pressure	in your ICU?			
No answer	Norepinephrine	Vasopressin	Dobutamine	Epinephrine	Metarami- nol		
43 (28%)	107 (69%)	3 (2%)	1 (< 1%)	1 (< 1%)	1 (< 1%)		
Which drug	s are used as secor	nd line to increa	se blood press	ure in your ICU	?		
No answer	Vasopressin	Epinephrine	Dobutamine	Norepi- nephrine	Dopamine		
43 (28%)	51 (33%)	21 (14%)	21 (14%)	11 (7%)	7 (5%)		
How freque	ently do you monito	or fluid balance	?				
No answer	Every hour	Every 2 h	Every 4 h	Every 12 h	Every 24 h		
43 (28%)	45 (29%)	21 (14%)	7 (5%)	20 (13%)	18 (12%)		
Which is yo	ur net daily fluid ba	alance target?					
No answer	Negative	Neutral	Positive				
43 (28%)	5 (3%)	68 (44%)	38 (25%)				
	e first-line fluid the	rapy you usuall	y use?				
No answer	Normal saline	Ringer lactate	Hypertonic saline	Plasmalyte	Albumin	Other	
43 (28%)	46 (30%)	43 (28%)	6 (4%)	5 (3%)	1 (< 1%)	10 (6%)	

ICU, intensive care unit; SAH, subarachnoid hemorrhage; CRT, capillary refill time; ABP, arterial blood pressure; NIBP, non-invasive blood pressure; ECG, electrocardiogram; ACE, angiotensin-converting enzyme.

strategies adopted [18]. The responses to this survey confirm the increasing use of this technique also in neurocritically ill patients.

In brain injured patients, systemic blood pressure is the most defined hemodynamic target, and an invasive monitoring is applied in selected cases of AIS, ICH and SAH (Tables 2, 3, 4). Both in AIS and ICH, blood pressure thresholds may vary accordingly to center protocols on initial management, to the time course of the disease and to the interventions performed (mechanical thrombectomy vs systemic thrombolysis). Although blood pressure threshold is the most studied hemodynamic target evaluated in literature, the survey participants declare a wide range of target MAP thresholds adopted (the most consistent answer was < 160/90 mmHg both before and after mechanical thrombectomy). Also, preferred first-line agents to lower blood pressure are different (more frequently urapidil and beta-blockers). By contrast, low blood pressure management is better standardized, with most participants using a MAP-driven protocol using norepinephrine.



**Fig. 2** Echocardiography, continuous hemodynamic monitoring and capillary refill time use in patients with subarachnoid hemorrhage (SAH) and acute ischemic injury (AIS). Data are expressed as percentage of the overall answers obtained by the survey

In AIS and ICH patients, standard hemodynamic management is the most oriented towards a strict control of blood pressure. In SAH patients, the findings of the survey are more heterogeneous. Besides invasive BP, echocardiography, CRT and advanced invasive hemodynamic monitoring are frequently used in SAH both in the initial phase of the disease and the vasospasm phase. One possible explanation is that in the early phase, echocardiography and advanced invasive monitoring are used to stabilize patients with cardiac stunning, frequently occurring in SAH patients. In the vasospasm phase, advanced monitoring and sequential echoes are frequently used, with the aim of increase MAP (and obtain MAP variations thresholds over time in SAH), while avoiding systemic complications. Moreover, the participants showed a more liberal attitude towards the fluid balance management in the vasospasm phase.

Overall, fluid therapy approach may be considered conservative in these patients, with most centers aiming at a neutral fluid balance. This finding may be explained by the intention of avoiding further swelling of the cerebral area surrounding lesion. A recent large prospective, multicentre, trial enrolling more than 2000 patients in two cohorts of traumatic brain injured patients, showed that a mean positive daily fluid balance was associated with higher ICU mortality per 0.1 L increase [19]. In SAH patients, few trials investigated the clinical effect of different hemodynamic strategies aimed at reducing cerebral vasospasm incidence, and a recent metanalysis showed low-quality evidence to support the use of advanced hemodynamic monitoring in selected SAH patients [20–22]. Regarding the type of fluid used, only for SAH patients the saline has been considered the first option (as compared to Ringer lactate for ICH and AIS patients). A recent large randomized-controlled trial showed a potential benefit of balanced solutions use for neurological patients, confirming the importance of the type of fluid adopted for fluid resuscitation/maintenance, irrespective of the hemodynamic targets [23].

Advanced invasive and non-invasive hemodynamic monitoring using echocar-diography is not frequently used; advanced monitoring is reserved to AIS and ICH patients developing cardiovascular or neurological complications, but not routinely used as standard management. This tool is widely adopted to obtain an initial assessment, but not as a part of the sequential dynamic monitoring system. However, the role of cardiac output in influencing cerebral blood flow is not widely acknowledged, especially regarding systemic blood pressure management [24]. As confirmation, a recent systematic review and meta-analysis including SAH patients identified some low-quality studies supporting advanced hemodynamic monitoring to guide clinical management [22].

#### Limitations and future directions

This survey presents several limitations which must be mentioned. The external validation of this survey is intrinsically limited by the bias regarding the selection of respondents and the accuracy of the questions in defining a specific issue. For this reason, it may hardly be considered as a complete snapshot of daily clinical practice, since most of the answers are obtained from very skilled centers, overrepresenting for a particular subgroup of physicians.

Respondents from specialized neuro ICUs were included together with those working in general ICUs, yielding more generalizable findings. This survey refers only to physicians' clinical practice in hemodynamic management of acute brain injuries without including patients' data and only considering ICU management. Finally, the response rate cannot be calculated considering the design of the study, since participants were encouraged to involve more respondents locally.

Highlighting the heterogeneity of hemodynamic monitoring and the lack of specific protocols of treatment on fluid management in neurocritical care patients, this survey may be also considered as a starting point to guide a research agenda in this field (Fig. 3). Starting from the evidence retrieved by several observational studies, large and specific randomized-controlled trials aiming at evaluating the clinical effect regarding the use of advanced hemodynamic monitoring and specific fluid/pressure targets should be designed in this field.

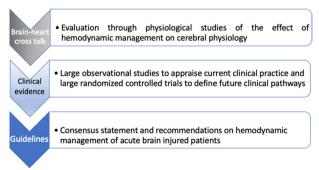


Fig. 3 Main steps for a research agenda for the hemodynamic management of acute brain injured patients

#### **Conclusions**

This ESICM survey shows important heterogeneity in the approach of hemodynamic monitoring and management of brain injured patients, including pharmacologic and non-pharmacological strategies. This survey may pave the way for the development of a research agenda, in the field of hemodynamic management of this population of patients, where evidence is importantly lacking.

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#### **Author contributions**

AM and CR drafted the manuscript and conceived the survey; GL, LG, FV, GM, MVJ, XM, PP and MC substantially contributed to manuscript preparation. All the authors approved the final version of the paper.

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# Availability of data and materials

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

#### **Declarations**

#### Ethics approval and consent to participate

Not applicable.

#### Consent for publication

Not applicable.

#### Competing interests

CR received fees for lecture from Masimo and GE; XM is a member of the Medical Advisory Board of Pulsion Medical Systems, Getinge. He received fees for consultancy and lectures; Dr. Messina received travel expenses and registration for meetings, congresses, and courses and lecture fees from Vygon, Edwards, Philips and Getinge. Prof. Cecconi is a consultant of Edwards Lifesciences (Directed Systems Consultancy); the other authors declare no competing interests for this paper.

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