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Goal-Directed Fluid Therapy Based on Stroke Volume Variations Improves Fluid Management and Gastrointestinal Perfusion in Patients Undergoing Major Orthopedic Surgery

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Key Words

Goal-directed fluid therapy · Stroke volume variation · FloTrac/Vigileo system · Orthopedic surgery

Abstract

Objective: To evaluate the influence of stroke volume variation (SVV)-based goal-directed therapy (GDT) on splanchnic organ functions and postoperative complications in orthopedic patients. Subjects and Methods: Eighty patients scheduled for major orthopedic surgery under general anesthesia were randomly allocated to one of two equal groups to receive either intraoperative volume therapy guided by SVV (GDT) or standard fluid management (control). In the SVV group, patients received colloid boluses of 4 ml/kg to maintain an SVV <10% when in the supine position or an SVV <14% if prone. In the control group, fluids were given to maintain a mean arterial pressure >65 mm Hg, a heart rate <100 bpm, a central venous pressure of 8–14 mm Hg, and a urine output >0.5 ml/kg/h. Intraoperative organ perfusion, hemodynamic data, hospitalization, postoperative complications, and mortality were recorded. Results: The heart rate at the end of surgery was significantly lower (p < 0.05), there were fewer hypotensive episodes (p < 0.05), the arterial and gastric intramucosal pH were higher (p < 0.05 for both), the gastric intramucosal PCO₂ was lower (p < 0.05), the intraop-

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E-Mail karger@karger.com www.karger.com/mpp This is an Open Access article licensed under the terms of the Creative Commons Attribution-NonCommercial 3.0 Unported license (CC BY-NC) (www.karger.com/OA-license), applicable to the online version of the article only. Distribution permitted for non-commercial purposes only. erative infused colloids and the total infused volume were lower (p < 0.05 for both), and the postoperative time to flatus was shorter (p < 0.05) in the GDT group than in the control group. No differences in the length of hospital stay, complications, or mortality were found between the groups. **Conclusion:** SVV-based GDT during major orthopedic surgery reduced the volume of the required intraoperative infused fluids, maintained intraoperative hemodynamic stability, and improved the perioperative gastrointestinal function.

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Introduction

Fluid balance is a major contributing factor to postoperative morbidity and mortality. Persistent hypovolemia is associated with organ hypoperfusion, systemic inflammatory response syndrome, sepsis, and multiple organ failure. Fluid overload, on the other hand, is associated with edema, ileus, postoperative nausea and vomiting, pulmonary complications, and increased cardiac demands [1]. Traditional methods to monitor the preload are based on measurements of pressure or volume, such

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Fu-hai Ji Department of Anesthesiology First Affiliated Hospital of Soochow University No. 188 Shizi Street, Suzhou, Jiangsu 215006 (China) E-Mail jifuhaisuda @163.com as the mean arterial pressure (MAP), the heart rate, or the central venous pressure (CVP). However, these are static parameters and do not accurately reflect fluid responsiveness [2]. The individualized 'goal-directed therapy' (GDT) concept [3–5] is a more rational strategy for perioperative fluid therapy, which achieves the maximal cardiac stroke volume via targeted administration of i.v. fluids, blood, and/or vasoactive substances. The FloTrac/ Vigileo system provides automatic and continuous monitoring of the cardiac output, stroke volume, and stroke volume variation (SVV) based on arterial pulse contour analysis. According to our previous study [6] and data from similar studies [7–9], SVV is a sensitive predictor of fluid responsiveness.

Fluid management is a crucial issue for patients undergoing major orthopedic surgery, in which large blood loss, transfusions, fluid shifts, and high incidences of postoperative complications are important concerns. Herein, we report the results of a prospective, randomized, controlled study that compared the intraoperative laboratory parameters of organ functions and postoperative complications between standard care management and SVV-based goaldirected fluid therapy. The primary study endpoint was gastrointestinal function, including the gastric intramucosal partial pressure of carbon dioxide (P_gCO₂), the gastric intramucosal pH (pH_i), and the time to passing the first flatus. We hypothesized that the use of goal-directed volume therapy would result in better gastrointestinal perfusion and fewer postoperative complications in patients undergoing major orthopedic surgery.

Subjects and Methods

Subjects

The Institutional Research Ethics Committee approved this trial and written informed consent was obtained from all patients. Patients scheduled for elective major orthopedic surgery, including total hip arthroplasty, spinal fusion surgery, femoral fracture surgery, and sacral tumor surgery under general anesthesia, with an anticipated blood loss >800 ml, were eligible for inclusion into this study. Patients were excluded if they were under 18 years old or had a BMI >40 or <15, coagulopathy, significant arrhythmia or cardiopulmonary dysfunction, or significant renal or liver diseases.

Eighty patients were randomized preoperatively into either a standard care management group (control, n = 40) or a goal-directed fluid therapy group (GDT, n = 40) using a random number generator in sealed envelopes. The anesthetist (K.P.) responsible for intraoperative management was aware of the group assignment, whereas all other members of the research team, other health care providers, and the patients were not.

Anesthesia and Monitoring

All patients fasted for 6 h before surgery and were premedicated with i.v. midazolam (0.01 mg/kg). A central venous catheter was inserted via the right internal jugular vein and an arterial line was inserted into the radial artery of the nondominant forearm. Standard monitoring included an ECG, MAP, CVP, pulse oximetry, temperature, end-tidal carbon dioxide, and the bispectral index.

In both groups, standard general anesthesia was induced with i.v. fentanyl ($3-4 \mu g/kg$), propofol (1.5-2 mg/kg), and vecuronium (0.15 mg/kg). After tracheal intubation, the lungs were ventilated at 8 ml/kg of tidal volume in a volume-controlled mode with 0-3 mm Hg positive end-expiratory pressure. The respiratory rate was set to maintain the end-tidal carbon dioxide at 35-40 mm Hg. The ventilator settings were unchanged during this study. Anesthesia was maintained with sevoflurane 2-3% in oxygen and fentanyl adjusted to maintain a bispectral index of 45-55. The body temperature was maintained at $>36^{\circ}$ C by a fluid warmer. A tonometry tube was inserted into the lumen of the stomach via the nasogastric route and was connected to a CO₂ monitor (Tonocap).

After induction of anesthesia, patients scheduled for spinal or sacral surgery were placed in the prone position on a prone pad with 4 small pads (2 shoulder and 2 pelvic supports) to allow the chest and abdomen to hang free. All of the patients received an i.v. prophylactic antibiotic and preemptive analgesia of 50 mg flurbiprofen axetil before skin incision. All operations were performed by the same surgical team.

Study Protocol

The intraoperative fluid management is shown in figure 1. In both groups, intraoperative basal fluid replacement was achieved by continuous infusion of 5 ml/kg/h crystalloid solution (Ringer's acetate). In the GDT group, an additional bolus of 4 ml/kg colloid solution (Voluven 130/0.4; 6%) was given when the SVV (measured by the FloTrac/Vigileo 3.0) increased >10% in the supine position or >14% in the prone position. Fluid boluses were repeated every 5 min if the criteria were met. In the control group, the anesthesiologist (K.P.) was free to give additional fluids, based on the subject's hemodynamic condition and responses, to maintain an MAP >65 mm Hg, a heart rate <100 bpm, a CVP of 8–14 mm Hg, and a urine output >0.5 ml/kg/h.

In both groups, anemia (hemoglobin level <80 g/l or hematocrit <28%) and an acute blood loss >20% of the calculated patient circulatory volume were corrected with transfusions of packed red blood cells and fresh frozen plasma in ratios approaching 2:1. Ephedrine boluses of 10 mg or phenylephrine boluses of 50 µg were given when fluid boluses failed to maintain a systolic arterial pressure >90 mm Hg or an MAP >65 mm Hg. These episodes were recorded as hypotensive events.

Arterial blood samples were taken at the time of skin incision and closure for blood counts, acid-base balance analysis, and other biochemical laboratory tests. At the same time, the P_gCO_2 was recorded. The mucosal-arterial PCO₂ gap was calculated as:

$$P_{g-a}CO_2 gap = P_gCO_2 - P_aCO_2,$$

where P_aCO_2 is the arterial carbon dioxide tension. In addition, the pH_i was calculated [10] as:

 $pH_i = arterial pH + log_{10}(P_aCO_2/P_gCO_2)$

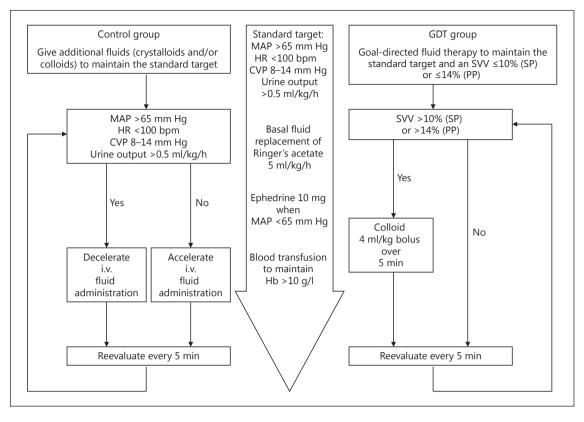


Fig. 1. Intraoperative fluid management. HR = Heart rate; Hb = hemoglobin; SP = supine position; PP = prone position.

Patients were transferred to either the intensive care unit or the postanesthesia care unit and were extubated when they fulfilled the standard clinical criteria (oxygenation, hemodynamics, and protective reflexes). Patient-controlled i.v. analgesia with fentanyl (background infusion of $0.4 \mu g/kg/h$, bolus dose of $0.4 \mu g/kg$, and lockout interval of 10 min) was used during the next 2 postoperative days. Regional anesthesia or analgesia (e.g. epidural catheters or nerve blocks) was not used in this study.

The same surgical team was in charge of the postoperative care, including fluid management (baseline crystalloid infusion of 40–50 ml/kg/day, colloids, and transfusions if required), daily antibiotics for 2–3 days, rescue analgesia (100 mg i.v. flurbiprofen axetil or 10 mg i.m. morphine), and antiemetics (10 mg i.v. metoclopramide or 3 mg i.v. granisetron). The discharge criteria were predefined by the Department of Orthopedics at our institution.

Postoperative complications were defined as follows: (a) cardiac complications: hypotension (MAP <65 mm Hg or SAP <90 mm Hg), arrhythmias (severe arrhythmias resulting in hemodynamic instability), or heart failure; (b) respiratory complications: ventilator support (need for mechanical ventilation in the intensive care unit), acute lung injury, or acute respiratory distress syndrome (PaO₂/FiO₂ <300 mm Hg); (c) abdominal complications: gastrointestinal hemorrhage (hematemesis and melena), hepatic dysfunction (transaminase >double the upper limit of normal), or hepatic failure (progressive jaundice rise and hepatic encephalopathy); (d) renal complications: renal dysfunction (creatinine >180 µmol/l) or renal failure (creatinine >450 µmol/l); (e) cerebral complications: postoperative cognitive dysfunction or coma; (f) infectious complications: wound infection or wound dehiscence, and (g) others: deep vein thrombosis, nausea, or vomiting. Postoperative complications, fluid management, drainage, length of hospital stay, and mortality were recorded.

Statistical Analysis

The statistical power analysis was based on a review of our hospital database, which showed an average time to passing the first flatus of 15.32 ± 5.16 h. Thirty-six patients per group were required for detection of a 20% difference in the postoperative first flatus time between the two groups with an α level of significance of 0.05 and a power of 80% (PASS 11.0.7). To compensate for dropped cases, 40 patients were studied in each group.

The statistical analysis was performed using SPSS 19.0 statistical software (IBM SPSS). Data were checked for normality using the Kolmogorov-Smirnov test. Continuous normally distributed data are presented as means \pm SD and were analyzed using paired or unpaired t tests. Nonnormally distributed data are presented as medians (IQR) and were tested using the Mann-Whitney U test and the Wilcoxon rank-sum test for unpaired and paired results, respectively. Categorical data are presented as numbers (%) and were compared using Fisher's exact test. p < 0.05 was considered statistically significant for all tests.

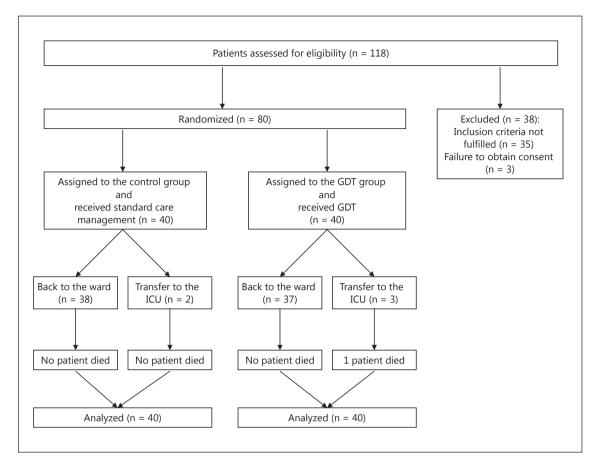


Fig. 2. Flowchart of the patients in this study. ICU = Intensive care unit.

Results

All 80 patients completed this study. No patient was excluded or dropped out of the study after randomization (fig. 2). There were no significant differences between the groups with regard to demographics and surgical characteristics (table 1).

In both groups, at the end of surgery, the MAP decreased and the CVP increased compared to the baseline values (table 2). Similarly, the mean hemoglobin and hematocrit decreased significantly in both groups (p < 0.05 for all). The mean heart rate in the GDT group (68 ± 13 bpm) was significantly lower than that in the control group (75 ± 13 bpm, p = 0.028), and there were fewer hypotensive episodes (n = 0, IQR 0–1, compared to n = 1, IQR 0–2, p = 0.021).

Also in the GDT group, at the end of surgery the mean SVV (7 ± 1) was significantly lower than the mean preoperative value $(9 \pm 2, p = 0.000)$, with a trend to-

ward higher cardiac outputs (4.79 \pm 1.24 l/min compared to 4.41 \pm 1.07 l/min, p = 0.057). The SVV and cardiac output were not evaluated in the control group.

The arterial and gastric pH_i decreased at the end of surgery, while the arterial and gastric intramucosal CO_2 tensions, $P_{g_{-a}}CO_2$, and the lactic acid concentration increased. At the end of surgery, compared to the control group, the mean gastric intramucosal CO_2 of the GDT group was significantly lower ($48.96 \pm 11.34 \text{ mm Hg compared to } 42.90 \pm 10.01 \text{ mm Hg}$, p = 0.013), while the arterial and gastric pH_i were higher (7.34 ± 0.05 compared to 7.36 ± 0.06 , p = 0.048, and 7.30 ± 0.11 compared to 7.37 ± 0.11 , p = 0.007, respectively).

The volume of infused intraoperative colloids was significantly lower in the GDT group than in the control group (500 ml, IQR 312–1,000, compared to 1,000 ml, IQR 500–1,000, p = 0.003). Similarly, the total infused volume in the GDT group was also significantly lower (1,850 ml, IQR 1,525–2,537, compared to 2,225 ml, IQR

Table 1. Demographic patient data and surgical characteristics

	GDT	Control	p value
Gender (male/female), n	17/23	18/22	0.822
Age, years	55±13	53 ± 10	0.505
Weight, kg	59±10	62±11	0.216
Height, cm	162±8	164±9	0.329
BMI	22.66 ± 3.22	23.25 ± 3.24	0.421
Position (supine/prone), n	25/15	22/18	0.496
ASA (I/II/III), n	17/22/1	17/21/2	0.837
Surgery			
Hip	21 (52.5)	20 (50.0)	0.823
Spine	16 (40.0)	18 (45.0)	0.651
Femur	2 (5.0)	1 (2.5)	1.000
Sacrum	1 (2.5)	1 (2.5)	1.000
Comorbidity			
Hypertension	11 (27.5)	15 (37.5)	0.340
Diabetes mellitus	2 (5.0)	7 (17.5)	0.157
Anemia	5 (12.5)	2 (5.0)	0.432
COPD	2 (5.0)	1 (2.5)	1.000
Multiple trauma	3 (7.5)	2 (5.0)	1.000
Tuberculosis	2 (5.0)	0 (0)	0.494
Coronary artery disease	0 (0)	1 (2.5)	1.000
Heart block	2 (5.0)	1 (2.5)	1.000
Cerebral infarction	1 (2.5)	2 (5.0)	1.000
Duration of surgery, min	175 ± 100	162 ± 80	0.520
Intraoperative BIS	52±2	53 ± 2	0.298

Values are presented as means \pm SD or numbers (%) unless otherwise stated. ASA = American Society of Anesthesiologists physical status classification; COPD = chronic obstructive pulmonary disease; BIS = bispectral index.

1,850–2,900, p = 0.036). The volume of blood loss, infused crystalloids, infused blood products, and urinary output did not differ between the groups.

The time to the first passage of flatus was significantly shorter in the GDT group (10.82 ± 5.83 h compared to 14.97 ± 11.17 h, p = 0.042; table 3). There was no significant difference between the two groups with regard to the postoperative amount of infused fluids, urine output and drainage volume at 0–24 or 24–48 h, or the volume of blood transfusion. There were also no differences in postoperative nausea and vomiting, patient-controlled analgesia requests, the number of patients who developed complications, the length of postoperative hospital stay, or mortality. One patient who underwent resection of a sacral tumor in the GDT group died from septic shock 17 days after surgery.

Fluid Management in Major Orthopedic Surgery

Discussion

The main findings of the present study are that SVVbased GDT during major orthopedic surgery reduced the required volume of intraoperative infused fluids, maintained intraoperative hemodynamic stability, and improved the perioperative gastrointestinal function.

Perioperative fluid management is challenging in high-risk surgical patients. The aim of volume therapy is not only to prevent hypovolemia but also to reduce the risk of fluid overload. Hypovolemia is recognized as a risk factor for adverse effects, ranging from minor organ dysfunction to multiple organ failure and even death. Conversely, fluid overload may impair pulmonary, cardiac, and gastrointestinal functions, contributing to postoperative complications and a prolonged recovery [11]. Therefore, appropriate hemodynamic monitoring is important for intraoperative fluid management. To our knowledge, this study is the first to directly compare conventional intraoperative management with SVV-based goal-directed fluid therapy with regard to organ functions and postoperative complications in patients undergoing major orthopedic surgery.

Ideally, a simple, affordable, and reliable method to improve intraoperative fluid management is desirable for routine use. Esophageal Doppler has been used to guide fluid management, with good results [12], but its use is partially limited by the need for experienced staff [13]. Also, the reliability of this method in major vascular procedures requiring cross-clamping of the descendent aorta has been questioned. On the other hand, arterial cannulation is routinely used in high-risk patients, and application of the FloTrac/Vigileo system is generally well tolerated by patients, without increased invasion and risk.

As reported in our previous study [6], the SVV obtained by the Vigileo/FloTrac system was a more sensitive predictor of fluid responsiveness than MAP, heart rate, or CVP measurements for intravascular volume assessment. Although the optimal cut-off value for SVV remains uncertain, the 10% threshold suggested by Manecke [14] for patients in the supine position and the 14% threshold for patients in the prone position [8] were considered to be the best available estimates for the Vigileo/FloTrac system. In our institution, patients scheduled for spinal or sacral surgery were placed in the prone position on a prone pad with 4 small pads (2 shoulder and 2 pelvic supports) to allow the chest and abdomen to hang free, which is similar to the method described by Biais et al. [8]. Therefore, we used an SVV of 10% for patients in the supine position and an SVV of 14% for patients in the prone

	GDT		Control		
	baseline	end of surgery	baseline	end of surgery	
Hemodynamic data					
Heart rate, bpm	68±11	68 ± 13^{a}	73±13	75±13	
MAP, mm Hg	88±9	79 ± 10^{b}	91 ± 10	81 ± 12^{b}	
CVP, mm Hg	8±3	10 ± 3^{b}	8±3	10 ± 3^{b}	
SVV, %	9±2	7 ± 1^{b}	NA	NA	
Cardiac output, l/min	4.41 ± 1.07	4.79 ± 1.24	NA	NA	
Hypotensive events	0 (0-1) ^a		1 (0-2)	1 (0-2)	
Laboratory parameters					
P _g CO ₂ , mm Hg	29.29 ± 5.57	42.90±10.01 ^{a, b}	30.81 ± 5.63	48.96±11.34 ^b	
P_aCO_2 , mm Hg	39.10 ± 6.83	42.11 ± 9.07^{b}	40.63 ± 6.11	44.26 ± 6.75^{b}	
P _{g-a} CO ₂ , mm Hg	-9.80 ± 9.44	0.78 ± 14.48^{b}	-9.82 ± 6.76	4.52 ± 11.48^{b}	
pHa	7.42 ± 0.04	$7.36 \pm 0.06^{a, b}$	7.42 ± 0.05	7.34 ± 0.05^{b}	
pH _i	7.55 ± 0.10	$7.37 \pm 0.11^{a, b}$	7.54 ± 0.08	7.30 ± 0.11^{b}	
Lactate, mmol/l	1.54 ± 0.32	2.12 ± 0.89^{b}	1.61 ± 0.57	2.35 ± 1.02^{b}	
Hemoglobin, g/l	12.03 ± 1.81	10.52 ± 1.54^{b}	11.94±1.69	10.28 ± 1.61^{b}	
Hematocrit	0.37 ± 0.05	0.33 ± 0.04^{b}	0.37 ± 0.04	0.32 ± 0.04^{b}	
Fluid management					
Blood loss, ml	800 (600-1,0	000)	800 (525-1,2		
Crystalloids infused, ml	1,000 (712-1,0	000)	1,000 (500-1,0	000)	
Colloids infused, ml	500 (312-1,000) ^a		1,000 (500-1,000)		
PRBC infused, ml	600 (400-600)		600 (400-800)		
FFP infused, ml	0 (0-200)		0 (0-200)		
Total volume infused, ml	1,850 (1,525-2,537) ^a		2,225 (1,850-2,900)		
Urinary output, ml	300 (200-400)		300 (200-475)		
Urinary output, ml/kg/h	1.98 (1.29-2.63)		2.20 (1.53-3.25)		

Table 2. Intraoperative hemodynamic data, laboratory parameters, and fluid management

Values are presented as means \pm SD or medians (IQR). NA = Not available; PRBC = packed red blood cells; FFP = fresh frozen plasma; pH_a = arterial pH; pH_i = intramucosal pH; P_gCO₂ = gastric intramucosal partial pressure of carbon dioxide; P_aCO₂ = arterial partial pressure of carbon dioxide; P_{g-a}CO₂ = mucosal-arterial PCO₂ gap. ^a p < 0.05 between the two groups. ^b p < 0.05 in comparison to baseline values.

position as the thresholds for hypovolemia to target volume optimization in our study.

In many studies [15–19], patients who had their fluid requirements managed with a goal-directed protocol received greater amounts of colloids than those who were treated with conventional or restrictive fluid management. Similarly, our results support that the use of intraoperative fluid therapy with accurate targeting of colloid fluid boluses may prevent excessive fluid administration. Greater amounts of infused fluids were less effective and jeopardized gastrointestinal function. In normovolemic healthy volunteers, 16% of colloids and more than 68% of the saline solution escaped into the extravascular fluid compartment 1 h after the infusion [20]. Edema of the intestines and other tissues may be responsible for poor tissue oxygenation and postoperative gut dysfunction [21]. Smaller volumes of infused fluids may help protect the gastrointestinal tract from dysfunction, which may help explain the shorter flatus time in the GDT group.

Regarding the lactate level, it is an indirect but sensitive measure of organ perfusion. Increased lactate correlates with an inadequate intravascular volume, tissue hypoxia, and energy failure due to blood flow redistribution [22]. In the present study, lactate-free fluids were used for volume substitution to exclude a potential bias. Our results showed that patients in both groups suffered a significant increase in lactate levels at the end of surgery relative to baseline values, indicating that these major surgical procedures had a great effect on organ perfusion. Tonometry, on the other hand, is a relatively noninvasive technique that measures the P_gCO_2 ; from this value, associated parameters such as the pH_i and the PCO₂ gap ($P_gCO_2 - P_aCO_2$) can be calculated. Low pH_i, high P_gCO_2 , and high

Table 3. Postoperative complications and fluid management

	GDT	Control	p value
Cardiovascular complications			
Hypotension	3 (7.5)	2 (5.0)	0.432
Arrhythmias	1 (2.5)	0 (0)	1.000
Heart failure	0 (0)	0 (0)	1.000
Respiratory complications			
Ventilator support	2 (5.0)	2 (5.0)	1.000
ALI/ARDS	1 (2.5)	0 (0)	1.000
Abdominal complications			
Flatus time, h	10±5	14 ± 11	0.042 ^a
Gastrointestinal hemorrhage	0 (0)	0 (0)	1.000
Hepatic dysfunction	5 (12.5)	6 (15.0)	0.745
Hepatic failure	1 (2.5)	0 (0)	1.000
Renal complications			
Urine output 0–24 h, ml	1,625 (1,175-2,412)	2,000 (1,150-2,700)	0.263
Urine output 24–48 h, ml	2,500 (1,800-3,100)	2,200 (1,700-3,525)	0.672
Renal dysfunction	1 (2.5)	3 (7.5)	0.615
Renal failure	1 (2.5)	0 (0)	1.000
Central nervous complications			
POCD	1 (2.5)	1 (2.5)	1.000
Coma	1 (2.5)	0 (0)	1.000
Infection-related complications			
Pneumonia	4 (10.0)	3 (7.5)	1.000
Wound infection	0 (0)	1 (2.5)	1.000
Wound dehiscence	0 (0)	0 (0)	1.000
Deep vein thrombosis	0 (0)	1 (2.5)	1.000
Nausea	5 (12.5)	8 (20.0)	0.363
Vomit	2 (5.0)	5 (12.5)	0.432
PCA requests	0 (0-2)	0 (0-2)	0.719
Fluid management, drainage			
Fluid infused 0–24 h, ml	$2,189\pm659$	$2,109\pm709$	0.606
Fluid infused 24–48 h, ml	$1,766 \pm 965$	$1,806 \pm 944$	0.852
Blood transfusion, ml	0 (0-200)	0 (0-200)	0.625
Drainage volume 0–24 h, ml	132 (100-263)	107 (38–187)	0.062
Drainage volume 24–48 h, ml	60 (25-120)	47 (16-135)	0.397
Drainage removal time, days	2 ± 0	2 ± 0	0.196
Postoperative stay, days	12±3	11±7	0.802
Mortality	1 (2.5)	0 (0)	1.000

Values are presented as numbers (%), medians (IQR), or means \pm SD. ALI/ARDS = Acute lung injury/acute respiratory distress syndrome; POCD = postoperative cognitive dysfunction; PCA requests = number of analgesic requirements with patient-control analgesia.^a p < 0.05 indicates a significant difference.

 PCO_2 gap values may indicate an inadequate oxygen supply to the bowel, leading to regional acidosis. In addition, intramucosal acidosis has been associated with a poor prognosis and multiple organ failure in critically ill patients, even in the absence of systemic acidosis or hypotension [23, 24]. Moreover, it has been shown that the correc-

tion of intramucosal acidosis may increase the survival rate of critically ill patients [25]. Recently, P_gCO_2 and pH_i were used as indicators of gastrointestinal function [26–28]. In this study, we found that the P_gCO_2 levels were lower and the pH_i levels were higher at the end of surgery in the GDT group. Similarly, in the GDT group, the post-

operative flatus time was shorter, indicating better gastrointestinal perfusion compared to the control group.

Our study has some limitations. First, the SVV and CO values were not available for the control group as the FloTrac/Vigileo device was not attached to these patients. Second, the inclusion of different surgical procedures might have influenced our results, because the pathophysiology and complications vary between joint and spinal surgery. Third, the absence of significant differences between the study groups in the incidence of nausea and vomiting, the length of hospital stay, and the incidence of mortality may have been due to our study not being powered enough to detect differences in these complications.

Conclusion

In patients undergoing major orthopedic surgery, SVV-based goal-directed intraoperative fluid therapy reduced the volume of intraoperative infused fluids, maintained intraoperative hemodynamic stability, and improved the perioperative gastrointestinal function relative to conventional treatment.

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