## Are management decisions in critical patients changed with use of hemodynamic parameters from transpulmonary thermodilution technique?

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**Background:** The assessment of hemodynamic variables is a mainstay in the management of critically ill patients. Hemodynamic variables may help physicians to choose among use of a vasopressor, an inotropic agent, or discontinuation of drugs. In this study, we aimed to investigate the usefulness of advanced hemodynamic variables in clinical decision-making.

**Methods:** Surveys regarding the case were administered to 25 surgeons working in nationally designated trauma centers or on trauma teams, using a voting system at a medical conference. The patient was a 67-year-old male with a crush injury of the left leg after a pedestrian traffic accident, who had aggravated pulmonary edema after leg amputation. Three clinical situations were given and the decision choices were: immediately after amputation, in 8 hours, and on the second day after amputation. Three kinds of variables from hemodynamic monitoring systems were provided for each clinical situation: conventional hemodynamic variables, including central venous pressure; variables from pulse contour analysis (PCA) [cardiac output (CO), stroke volume index, stroke volume variation (SVV), and systemic vascular resistance index); and variables from transpulmonary thermodilution (TPTD) technique (global ejection fraction and extravascular lung water index). The changes in decisions according to each provided hemodynamic variable were investigated and analyzed.

**Results:** The advanced hemodynamic parameters were considered to have a decisive effect on choosing vasopressors and inotropic agents. The decision was changed in 88% (22/25) of physicians using variables from the advanced monitoring systems. Among them, 82% (18/22) of physicians chose hemodynamic variables from the TPTD technique as their reason for change regarding management of a patient with severe pulmonary edema.

**Conclusions:** Advanced monitoring systems might be helpful in decision-making for critically ill patients. Multiple parameters and trends in change could be more important than a single value. Clinicians should select the system most appropriate according to its advantages and limitations, and interpret the variables obtained correctly.

**Keywords:** Hemodynamic monitoring; intensive care; transpulmonary thermodilution (TPTD); pulmonary edema; trauma

Submitted Feb 22, 2019. Accepted for publication Jun 24, 2019. doi: 10.21037/atm.2019.06.78 View this article at: http://dx.doi.org/10.21037/atm.2019.06.78

#### Introduction

Hemodynamic monitoring plays an important role in the management of today's acutely ill patient, particularly in the early stages of resuscitation. Monitoring modalities may help to identify underlying pathophysiological processes so that appropriate forms of therapy can be selected (1). Various monitoring systems are being developed and are available, in addition to conventional hemodynamic variables. The pulse contour analysis (PCA) and indicator dilution techniques [e.g., transpulmonary thermodilution (TPTD)] were introduced in the early 2000s (2). These advanced monitoring systems provide variables such as cardiac output (CO), stroke volume variation (SVV), pulmonary vascular permeability index (PVPI), and global end-diastolic volume (GEDV) using techniques that are less invasive than pulmonary artery catheterization (PAC). Despite development of hemodynamic monitoring, there are no optimal and universally applicable hemodynamic variables, which makes it difficult for physicians to choose and apply advanced hemodynamic monitoring.

In this survey on the utilization of advanced monitoring, survey respondents were provided variables of conventional monitoring, PCA, and TPTD for a patient with respiratory distress after a pedestrian traffic accident. We investigated the changes in decisions regarding patient management according to each variable, to evaluate whether advanced hemodynamic monitoring is helpful in decision-making for treatment.

#### Methods

Using a voting system at a medical conference, surveys were administered to 25 surgeons working in a nationally designated trauma center or on a trauma team. The survey involved the case of 67-year-old male who presented to the emergency room after a pedestrian traffic accident. On arrival, he was drowsy, with a Glasgow Coma Scale score of 10 and sluggish pupils reflex of 3 mm. Vital signs were as follows: blood pressure, 86/50 mmHg; pulse rate, 81 beats/min; respiratory rate, 20 breaths/min; and body temperature, 36.0 °C. The patient's left leg below the knee joint was severely damaged, with active bleeding and extensive soft-tissue loss. Brain computed tomography (CT) scan showed tiny subdural hemorrhage, subarachnoid hemorrhage, and temporal bone fracture without mass effect. There was no chest or abdominal injury. Enhanced CT of the lower extremity showed an open, comminuted, displaced fracture in the neck of the fibula and dislocation

of the proximal tibiofibular joint. During exploration of the injured leg, total destruction of the peroneal nerve, popliteal artery and vein were detected. Saphenous vein interposition was performed for the ruptured popliteal artery, and then open reduction and external fixation was done. A total 15 units of packed red blood cells and 16 units of fresh frozen plasma were transfused during the first 24 hours. Thirty hours later, color changes in the lower leg were observed due to graft failure; subsequently, leg amputation above the knee was performed. The VolumeView<sup>®</sup> system (Edwards Lifesciences Corp., Irvine, CA, USA) was used for patient monitoring. A central venous catheter with a thermistor was placed in the right subclavian vein, and a 4 French arterial catheter to assess volumetric parameters was placed in the right common femoral artery.

In the surveys, three clinical situations were given, and the decision choices given for each situation were as follow: immediately, 8 hours, and on the second day after amputation. Three kinds of variables from hemodynamic monitoring systems were provided for each clinical situation: conventional hemodynamic variables, including central venous pressure; variables from PCA; and variables from the TPTD technique. The variables from the Swan-Ganz catheter were not provided, given that the catheter was not used in this patient. Changes according to each provided hemodynamic variable were investigated and analyzed. Surveys were conducted blindly using the voting system.

#### **Results**

#### Physician demographics

Survey respondents comprised 12 general surgeons and 10 thoracic and cardiovascular surgeons. A total of 16 physicians had experience in managing critically ill injured patients of more than 5 years after completing their specialty training. Among them, 80% (20/25) of physicians knew how to interpret hemodynamic variables from PCA, and 68% (17/25) had used the FloTrac<sup>®</sup> system (Edwards Lifesciences Corp.) in an intensive care unit (ICU). Seventeen (68%) physicians were aware of the TPTD technique but only 12 (48%) had experience with it, including use of the VolumeView<sup>®</sup> system (*Table 1*).

# Situation #1: immediate postoperative status after amputation

Chest radiograph and conventional monitoring data are

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Table 1	Demograp	ohics of	physicians
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Parameters	N=25	
Specialty, n (%)		
General surgery	12 (48.0)	
Thoracic and cardiovascular surgery	10 (40.0)	
Neurosurgery	2 (8.0)	
Urology	1 (4.0)	
Years working as a specialist treating critically injured patients, n (%)		
1–2 years	5 (20.0)	
3–4 years	4 (16.0)	
5–9 years	8 (32.0)	
>10 years	8 (32.0)	
Pulse contour analysis, n (%)		
Physicians who know about the pulse contour analysis	20 (80.0)	
Physicians who have experienced the pulse contour analysis	17 (68.0)	
TPTD technique, n (%)		
Physicians who know about the TPTD technique	17 (68.0)	
Physicians who have experienced the TPTD technique	12 (48.0)	

TPTD, transpulmonary thermodilution.

shown in *Figure 1A*. The total amount of fluids infused was 120 mL/h and urine output were 250–300 mL/h. Body weight was reduced from 89.0 to 84.5 kg after amputation. Administered drugs were as follows: norepinephrine, 25 µg/min; remifentanil, 0.05 µg/kg/min; and dexmedetomidine, 0.6 µg/kg/h. Arterial blood gas analysis (ABGA) showed pH 7.46, PaCO<sub>2</sub> 38 mmHg, PaO<sub>2</sub> 67 mmHg, HCO<sub>3</sub> 27 mmol/L, and SaO<sub>2</sub> 94%.

*Figure 1B* shows the hemodynamic variables from PCA: CO 7.9 L/min (normal range, 4.0–8.0 L/min); cardiac index (CI) 3.9 L/min/m<sup>2</sup> (normal range, 2.5–4.0 L/min/m<sup>2</sup>); stroke volume index (SVI) 36 mL/beat/m<sup>2</sup> (normal range, 33–47 mL/beat/m<sup>2</sup>); SVV 12% (normal range, <10–15%); systemic vascular resistance index (SVRI) 2,553 dyn·s/cm<sup>-5</sup>/m<sup>2</sup> (normal range, 1,970–2,390 dyn·s/cm<sup>-5</sup>/m<sup>2</sup>).

Variables from the TPTD technique were as follows: global ejection fraction (GEF) 19% (normal range, 25–35%); extravascular lung water index (EVLWI) 14.1 mL/kg (normal range, 3–7 mL/kg); PVPI 2.7 (>3, lung injury; <3,

hydrostatic or cardiogenic edema) (Figure 1C).

Surveys queried the choice of patient management with the variables given from each hemodynamic monitoring system. When only conventional variables were given, observation (n=8) and decrease of norepinephrine (n=8) were the most common responses of survey participants. With variables from PCA, 13 respondents chose to decrease norepinephrine. When variables from the TPTD technique were presented, 10 respondents chose to increase dobutamine (*Figure 2*).

In this case, dobutamine (10 µg/kg/min) was initiated. Dexmedetomidine was stopped, and infusions of cisatracurium (5 mg/h) and propofol (30 µg/kg/min) were started for treatment of acute respiratory distress syndrome (ARDS), as a deep sedation strategy. The mode and setting of mechanical ventilation were changed as follows: pressure support/pressure-controlled ventilation mode; fraction of inspired oxygen (FiO<sub>2</sub>) 0.6; peak inspiratory pressure (PIP) 29 cmH<sub>2</sub>O; positive end-expiratory pressure (PEEP) 13 cmH<sub>2</sub>O; pressure support of 16 cmH<sub>2</sub>O; and respiratory rate 24/min. Two hours later, urine output was decreased to 100 mL/h; subsequently, continuous furosemide (8 mg/h) was started.

#### Situation #2: 8 hours after amputation

Chest radiograph and conventional monitoring data are shown in *Figure 3A*. The total amount of fluids infused was 100 mL/h and urine output were 200–300 mL/h. Body weight of the patient was decreased to 83.2 kg, 1.3 kg less than the previous day. Administered drugs were as follows: norepinephrine, 20 µg/min; dobutamine, 10 µg/kg/min; remifentanil, 0.05 µg/kg/min; propofol, 30 µg/kg/min; cisatracurium, 5 mg/h; and furosemide, 8 mg/h. ABGA showed pH 7.45, PaCO<sub>2</sub> 44 mmHg, PaO<sub>2</sub> 70 mmHg, HCO<sub>3</sub> 30.6 mmol/L, and SaO<sub>2</sub> 95%. Hemodynamic variables from PCA were as follows: CO 8.0 L/min; CI 3.9 L/min/m<sup>2</sup>; SVI 37 mL/beat/m<sup>2</sup>; SVV 11%; SVRI 1,981 dyn·s/cm<sup>-5</sup>/m<sup>2</sup> (*Figure 3B*). Variables from the TPTD technique were GEF of 20%, EVLWI of 13.0 mL/kg, and PVPI of 2.5 (*Figure 3C*)

Surveys were conducted again, using the same method. Observation was the most common response with conventional variables (n=16), variables from PCA (n=16), and variables from TPTD techniques (n=11). Other responses included volume restriction and increasing the amount of furosemide infusion (*Figure 4*).

In clinical practice situation #2, primarily observation was

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**Figure 1** Immediate postoperative status after amputation. (A) Chest radiograph and conventional monitoring data; (B) variables from pulse contour analysis; (C) variables from TPTD. TPTD, transpulmonary thermodilution; CI, cardiac index; CO, cardiac output; CVP, central venous pressure; FiO<sub>2</sub>, fraction of inspired oxygen; H/U, hourly urine output; MV, mechanical ventilation; PEEP, positive end-expiratory pressure; PIP, peak inspiratory pressure; PS, pressure support; RASS, Richmond Agitation Sedation Scale; RR, respiratory rate; SVI, stroke volume index; SIMV, synchronized intermittent mandatory ventilation; SVRI, systemic vascular resistance index; SVV, stroke volume variation; V/S, vital sign.

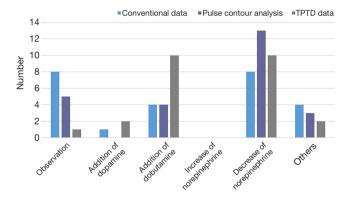


Figure 2 Results of a survey on the management of immediate postoperative status after amputation. TPTD, transpulmonary thermodilution.

chosen. Twelve hours later, norepinephrine (15 µg/min) and furosemide (5 mg/h) were decreased because vital signs were stable and hourly urine output was increased (>300 mL/h for 2 hours).

#### Situation #3: second postoperative day after amputation

Chest radiograph and conventional monitoring data are shown

in *Figure 5A*. The total amount of fluids infused was 100 mL/h and urine output were 150–300 mL/h. Body weight decreased a total 2.2 kg, compared with the previous day. Drugs administered were as follows: norepinephrine 15 µg/min; dobutamine 10 µg/kg/min; remifentanil 0.05 µg/kg/min; propofol 30 µg/kg/min; cisatracurium 5 mg/h; and furosemide 5 mg/h. ABGA showed pH 7.55, PaCO<sub>2</sub> 33 mmHg, PaO<sub>2</sub> 108 mmHg, HCO<sub>3</sub> 30.9 mmol/L, and SaO<sub>2</sub> 99%.

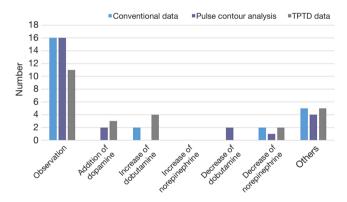
Variables from PCA were as follows: CO 5.2 L/min; CI 2.6 L/min/m<sup>2</sup>; SVI 29 mL/beat/m<sup>2</sup>; SVV, 11%; and SVRI, 3,115 dyn-s/cm<sup>-5</sup>/m<sup>2</sup> (*Figure 5B*). Variables from the TPTD technique were GEF of 12%, EVLWI of 9.5 mL/kg, and PVPI of 1.8 (*Figure 5C*).

In the survey, decrease of norepinephrine was the most common response with conventional variables (n=16), variables from PCA (n=24), and variables from TPTD techniques (n=8). Seven and five participants changed to addition of dopamine or dobutamine, respectively, when variables from TPTD were given (*Figure 6*).

For patient management in situation #3, primarily dopamine (10 µg/kg/min) was initiated; norepinephrine was then slowly tapered off. Because point-of-care ultrasound showed bilateral fluid accumulation in the thorax, closed thoracostomy with a small-bore catheter was performed. Annals of Translational Medicine, Vol 7, No 16 August 2019



**Figure 3** Eight hours after amputation. (A) Chest radiograph and conventional monitoring data; (B) variables from pulse contour analysis; (C) variables from TPTD. TPTD, transpulmonary thermodilution; CI, cardiac index; CO, cardiac output; CVP, central venous pressure; FiO2, fraction of inspired oxygen; H/U, hourly urine output; MV, mechanical ventilation; PEEP, positive end-expiratory pressure; PIP, peak inspiratory pressure; PS, pressure support; RASS, Richmond Agitation Sedation Scale; RR, respiratory rate; SVI, stroke volume index; SIMV, synchronized intermittent mandatory ventilation; SVRI, systemic vascular resistance index; SVV, stroke volume variation; V/S, vital sign.



**Figure 4** Survey results regarding patient management 8 hours after amputation. Observation was the most common response for all three methods. Other responses included volume restriction and increasing the amount of furosemide infusion.

#### Clinical course on the third day after amputation

Clinical and hemodynamic variables are shown in *Figure* 7. Infused drugs and their amounts were as follows: norepinephrine 8 µg/min; dobutamine 10 µg/kg/min; dopamine 10 µg/kg/min; remifentanil 0.05 µg/kg/min; dexmedetomidine 0.5 µg/kg/h; and furosemide 2 mg/h. ABGA showed pH 7.46, PaCO<sub>2</sub> 45 mmHg, PaO<sub>2</sub> 69 mmHg, HCO<sub>3</sub> 32.8 mmol/L, and SaO<sub>2</sub> 98%. The patient's respiratory and hemodynamic condition was stable throughout the treatment course and he was discharged without any complication.

#### Final survey results

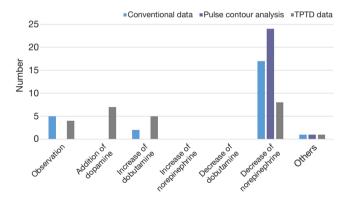
After completion of the patient management decision surveys, a questionnaire was administered querying whether the decisions made were influenced by advanced hemodynamic monitoring methods. A total 88% (22/25) of physicians stated that their decision changed when variables from advanced monitoring were presented. Among them, 82% (18/22) of physicians chose hemodynamic variables from the TPTD technique as their reason for the change (*Figure 8*).

#### Discussion

The key principles of hemodynamic monitoring include providing accurate and reproducible measurements of relevant variables, which can guide therapy. The method chosen must also be easy to use, readily available, operator independent, and cost effective (1). Physicians should try to choose devices that have these features and that are appropriate for each clinical situation. During the past several decades, technologies of hemodynamic monitoring



**Figure 5** The second postoperative day after amputation. (A) Chest radiograph and conventional monitoring data; (B) variables from pulse contour analysis; (C) variables from TPTD. TPTD, transpulmonary thermodilution; CI, cardiac index; CO, cardiac output; CVP, central venous pressure; FiO<sub>2</sub>, fraction of inspired oxygen; H/U, hourly urine output; MV, mechanical ventilation; PEEP, positive end-expiratory pressure; PIP, peak inspiratory pressure; PS, pressure support; RASS, Richmond Agitation Sedation Scale; RR, respiratory rate; SVI, stroke volume index; SIMV, synchronized intermittent mandatory ventilation; SVRI, systemic vascular resistance index; SVV, stroke volume variation; V/S, vital sign.



**Figure 6** Survey results regarding patient management on the second postoperative day after amputation. Decrease of norepinephrine was most common response with all three methods. Seven and five participants changed to addition of dopamine or dobutamine, respectively, when variables from TPTD were given. TPTD, transpulmonary thermodilution.

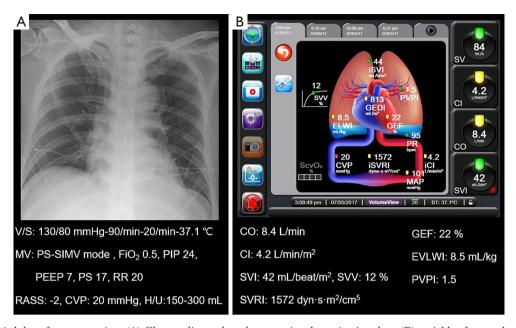
have improved, from being invasive to noninvasive and from a static to a functional approach (3). With the help of advanced monitoring, the management of critically ill patients has also evolved.

However, as yet, there are no optimal hemodynamic variables that are universally applicable to all patients.

Even though thermodilution with PAC, which was introduced in the early 1970s, has been considered optimal for CO monitoring, this approach is extremely invasive and has failed to show improvement in mortality when applied in critically ill patients (4-6). The use of PAC has decreased with the advent of newer technologies. According to a Swiss web-based survey regarding the kind of monitoring used in the adult ICU, echocardiography (95%), PAC (85%), and TPTD (82%) were the most commonly available. It seems that the PAC method is being progressively replaced by new monitoring techniques, such as TPTD, which was the preferred technique in our survey (7). Another survey conducted among ICU clinicians in 16 European countries also showed that TPTD was the preferred technique for measuring CO (8). However, a survey performed in Belgium and Germany on knowledge of the TPTD technique revealed that knowledge, interpretation, and measurement techniques were all suboptimal among ICU personnel, although these methods were used regularly (9).

As for the choice of monitoring system, clinicians must be aware of the most appropriate modality for each clinical situation, according to the system's advantages and limitations. With regard to quantifying pulmonary edema, the measurement of EVLWI can provide early

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**Figure 7** The third day after amputation. (A) Chest radiograph and conventional monitoring data; (B) variables from pulse contour analysis and TPTD. TPTD, transpulmonary thermodilution; CI, cardiac index; CO, cardiac output; CVP, central venous pressure; FiO<sub>2</sub>, fraction of inspired oxygen; H/U, hourly urine output; MV, mechanical ventilation; PEEP, positive end-expiratory pressure; PIP, peak inspiratory pressure; PS, pressure support; RASS, Richmond Agitation Sedation Scale; RR, respiratory rate; SVI, stroke volume index; SIMV, synchronized intermittent mandatory ventilation; SVRI, systemic vascular resistance index; SVV, stroke volume variation; V/S, vital sign.

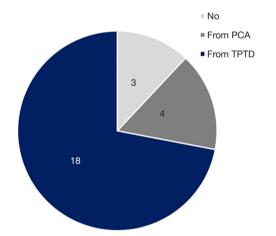


Figure 8 Results of questionnaire regarding whether decisions were changed with use of advanced hemodynamic monitoring methods. TPTD, transpulmonary thermodilution; PCA, pulse contour analysis.

detection of lung water accumulation, to more optimally guide fluid therapy by differentiating hydrostatic versus increased pulmonary capillary permeability (2), as in this case. Measured EVLWI that is utilized in managing fluid balance in ARDS (10,11) has shown lower cumulative fluid balance, decreased ICU mortality (12), reduced duration of mechanical ventilation, and shorter length of ICU stay (13). A previous study that aimed to evaluate the relationship between measured EVLWI, pulmonary vascular permeability using the TPTD technique, and severity categories as defined by the Berlin definition (14) showed an association between increased EVLWI and pulmonary vascular permeability (15). Moreover, most studies consider that TPTD is as reliable as PCA (16-19) and outperforms uncalibrated devices (20-22). As SVV measured by PCA has been shown to be a good predictor of fluid responsiveness in various clinical settings (23), TPTD can also be utilized to guide fluid therapy, such as during the early resuscitation phase, to achieve a minimum level of preload to allow for sufficient vital organ perfusion (24). TPTD may show less respiratory phase-dependent variation (25).

Despite the theoretical advantages of advanced monitoring, its clinical effectiveness has been inconsistent as monitoring itself does not guarantee improvement in patient survival unless it is paired with a proper therapeutic protocol. In a recent study, TPTD-based fluid management failed to show improved outcome and caused a more

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positive fluid balance in non-septic shock patients (26). With respect to septic shock, TPTD is considered superior to PAC by intensivists (8); however, TPTD has not been associated with improved survival (26,27). TPTD, as well as PCA, becomes unreliable when CO is very low. Frequent recalibration is required when the vasomotor tone changes with both PCA and TPTD (5,28-30). TPTD also remains considerably invasive because it requires a specific femoral artery catheter (3).

This study has several limitations. First, only surgeons participated, and the number of participants was relatively small. Intensivists, who could have provided additional valuable information, were not available to participate in this survey. Second, half of the respondents had never used hemodynamic monitoring with the TPTD technique before the survey. Third, the patient was not fully sedated and was mechanically ventilated using lung-protective settings; therefore, there might be technical errors because the recommendations for ventilation, such as full sedation, high tidal volume, and low PEEP, were not followed. High PEEP and low tidal volume may have interfered with heart-lung interaction and resulted in reduced accuracy of the variables. Fourth, the clinical efficacy of TPTD was unevaluable because this study was based on a single case. Fifth, the variables from Swan-Ganz catheter were not provided, given that the catheter was not used in this patient. Such information could have aided further comparisons between invasive and non-invasive techniques.

However, in this case, with the help of PAC and TPTD monitoring, respiratory distress in the patient was managed for cardiogenic pulmonary edema, as EVLWI and PVPI were within the normal ranges and GEF was only 12–19%. In addition, decisions were not made using a single value but rather using trends in the variables. On the first postoperative day, two-dimensional echocardiography was done to check heart function, which showed stress-induced cardiomyopathy with lowered ejection fraction (below 40%); both results suggest that the main cause of edema was cardiogenic. Therefore, TPTD provided valuable information for establishing an optimal treatment strategy in a patient with ARDS.

#### Conclusions

In summary, decisions regarding patient management may be changed using various parameters from different hemodynamic monitoring systems. Although there is no

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optimal method, use of an advanced monitoring system might be helpful in decision-making for critically ill patients. Information of multiple parameters and trends in change from advanced hemodynamic monitoring may be more valuable than a single value in making such decisions. If clinicians select the system that is most appropriate in each case, according to the system's advantages and limitations, and interpretation is based on correctly obtained variables, use of parameters from different monitoring systems could boost the effectiveness of patient management. The accumulation of experiences, research, and technical developments in hemodynamic monitoring will lead to improved management of critically ill patients.

#### **Acknowledgments**

None.

#### Footnote

*Conflicts of Interest*: This abstract was presented by oral presentation to the English session in the 45th annual meeting of the Japanese society of intensive care medicine.

*Ethical Statement*: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The need for ethics approval and consent of the present study was waived by Institutional Review Board of the Dankook University Hospital. Written informed consent was obtained from all patients.

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**Cite this article as:** Chang YR, Choi SH, Chang SW. Are management decisions in critical patients changed with use of hemodynamic parameters from transpulmonary thermodilution technique? Ann Transl Med 2019;7(16):370. doi: 10.21037/atm.2019.06.78

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