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Clinical review: Update on hemodynamic monitoring - a consensus of 16

Jean-Louis Vincent¹*, Andrew Rhodes³, Azriel Perel¹, Greg S Martin⁴, Giorgio Della Rocca⁵, Benoit Vallet⁶, Michael R Pinsky⁷, Christoph K Hofer⁸, Jean-Louis Teboul⁹, Willem-Pieter de Boode¹⁰, Sabino Scioletta¹¹, Antoine Vieillard-Baron¹², Daniel De Backer¹, Keith R Walley¹³, Marco Maggiorini¹⁴ and Mervyn Singer¹⁵

Abstract
Hemodynamic monitoring plays a fundamental role in the management of acutely ill patients. With increased concerns about the use of invasive techniques, notably the pulmonary artery catheter, to measure cardiac output, recent years have seen an influx of new, less-invasive means of measuring hemodynamic variables, leaving the clinician somewhat bewildered as to which technique, if any, is best and which he/she should use. In this consensus paper, we try to provide some clarification, offering an objective review of the available monitoring systems, including their specific advantages and limitations, and highlighting some key principles underlying hemodynamic monitoring in critically ill patients.

Introduction
Hemodynamic monitoring plays an important role in the management of today’s acutely ill patient. Essentially, hemodynamic monitoring can be helpful in two key settings. The first is when a problem has been recognized; here, monitoring can help to identify underlying pathophysiological processes so that appropriate forms of therapy can be selected. A typical scenario is the patient in shock for whom options are to give more fluids or to give a vasopressor or an inotropic agent, depending on the hemodynamic evaluation. The second setting is more preventative, with monitoring allowing preemptive actions to be performed before a significant problem arises. A typical scenario here is the perioperative patient in whom monitoring can be used to detect hypovolemia or low oxygen delivery (DO₂) early, enabling timely corrective therapy to be initiated.

Although microcirculatory changes are believed to play a major role in the development of organ dysfunction and multiple organ failure and there is increasing interest in new techniques to monitor the microcirculation, these are not yet available for clinical practice, and hemodynamic monitoring, therefore, still focuses on the macrocirculation. Current hemodynamic monitoring therefore includes measurement of heart rate, arterial pressure, cardiac filling pressures or volumes, cardiac output, and mixed venous oxygen saturation (SvO₂). Although not perfect, the pulmonary artery catheter (PAC) has long been considered the optimal form of hemodynamic monitoring, allowing for the almost continuous, simultaneous recording of pulmonary artery and cardiac filling pressures, cardiac output and SvO₂. However, although the incidence of complications with the PAC is relatively low, the technique is still quite invasive and there is no clear evidence for improved outcomes associated with its insertion and use to guide therapy [1]. As a result, interest in alternative monitoring systems has surged in recent years.

There are now many different monitoring systems available, and physicians may feel somewhat confused by the multiple possibilities. These systems can be easily listed in order of degree of invasiveness, from the highly invasive PAC to the completely non-invasive bioimpedance/bioreactance technique and transthoracic echo-Doppler. Classifying them according to how accurate (closeness of measured values to the ‘true’ value, expressed as the bias) or precise (variability of values due to random errors of measurement) [2] they are is more difficult, in part because of the lack of a perfect ‘gold’ standard for comparison. Most devices have been evaluated by comparing their results with those obtained by intermittent thermodilution from the PAC as the reference, although this technique has its own limitations and may not represent the best choice of comparator [2].

Our purpose in this consensus article is not to review the technology or modus operandi of the various systems
in any detail, not to provide readers with a shopping list, nor to identify one system that would be suitable in all patients; rather, we will briefly review the advantages and limitations of each system, and propose ten key principles to guide choice of monitoring system(s) in today’s acutely ill patients.

**Available systems for monitoring cardiac output**

Examples of the main systems that are available for estimating cardiac output are listed in Table 1.

**Thermodilution (pulmonary artery catheter)**

The intermittent thermodilution technique, in which boluses of ice-cold fluid are injected into the right atrium via a PAC and the change in temperature detected in the blood of the pulmonary artery used to calculate cardiac output, is still widely considered as the standard method of reference. Adaptation of the PAC to incorporate a thermal filament (Vigilance™, Edwards Life Sciences, Irvine, CA, USA) or thermal coil (OptiQ™, ICU Medical, San Clemente, CA, USA) that warms blood in the superior vena cava and measures changes in blood temperature at the PAC tip using a thermistor, provides a continuous measure of the trend in cardiac output, with the displayed values representing an average of the values over the previous 10 minutes. The averaged values have the advantage of eliminating variability in the presence of arrhythmias, but the disadvantage of not being real-time values, thus limiting the usefulness of this approach for assessing rapid hemodynamic changes in unstable patients.

The PAC has a key advantage over many other systems in that it provides simultaneous measurements of other hemodynamic parameters in addition to cardiac output, including pulmonary artery pressures, right-sided and left-sided filling pressures, and \( \text{SvO}_2 \).

**Transpulmonary or ultrasound indicator dilution**

The PiCCO® (Pulsion Medical Systems, Munich, Germany), LiDCO™ (Edwards Life Sciences), and COstatus™ (Transonic Systems Inc., Ithaca, NY, USA) systems allow cardiac output to be investigated less invasively, using a central venous (to allow calibration) and an arterial catheter, rather than needing to introduce a catheter into the pulmonary artery. The PiCCO® and recently launched VolumeView™ systems require a femoral artery catheter. These devices use the same basic principles of dilution to estimate the cardiac output as with PAC thermodilution. PiCCO® and VolumeView™ use injections of ice cold intravenous fluid...
as the indicator, measuring change in temperature down-stream to calculate cardiac output, whereas LiDCO™ uses minute amounts of lithium chloride as the indicator and measures levels using a lithium-selective electrode. COstatus® calculates cardiac output by using ultrasound technology to measure changes in blood ultrasound velocity and blood flow following an injection of warm saline solution.

Cardiac output values measured using transpulmonary or ultrasound indicator dilution techniques correlate well with those measured using PAC thermodilution [3-6] and may show less respiratory phase-dependent variation [4]. The PiCCO® and the VolumeView™ systems provide variables in addition to cardiac output, such as global end-diastolic volume and measurements of extravascular lung water. The COstatus® system also provides some derived variables, including total end diastolic volume index.

Arterial pressure trace-derived estimation of cardiac output
In addition to the intermittent indicator dilution cardiac output measurements discussed above, the PiCCO® and LiDCO™ systems can also estimate cardiac output on a continuous basis from the arterial pressure waveform with (PiCCO2® and LiDCOPlus®) or without (LiDCORapid®) the need for recalibration when changes in vascular compliance may have occurred. The PiCCO® system uses a pulse contour analysis and the LiDCO™ system a pulse power analysis. In addition to these, Vigileo™ (Edwards Life Sciences) and MostCare™ (Vytech, Padova, Italy, using the Pressure Recording Analytical Method (PRAM)) systems have been developed for arterial waveform analysis without external calibration. Each of these systems contains a proprietary algorithm for converting a pressure-based signal into a flow measurement. The specific algorithms have individual characteristics and make different assumptions - for example, related to arterial compliance (Vigileo™) or pressure (MostCare™) - which can make them more or less accurate depending on the clinical circumstances. The level of accuracy and precision of each device needs to be understood as the data cannot be superimposed from one system to another. The advantages of these arterial pressure-based cardiac output monitoring systems over PAC-derived measurements is primarily their less invasive nature.

The major weakness of all these devices is the drift in values whenever there is a major change in vascular compliance, as, for example, in vascular leak syndrome with increased vessel wall edema leading to decreased arterial compliance. Aortic valve regurgitation may further decrease the accuracy of these techniques. Over- or under-damped arterial pressure waveforms will also decrease the precision of these monitors.

Echocardiography and echo-Doppler
Echocardiography allows measurement of cardiac output using standard two-dimensional imaging or, more commonly, Doppler-based methods. The main interest in echocardiography in general is that it can be used not only for measurement of cardiac output but also for the additional assessment of cardiac function. Echocardiography is particularly useful as a diagnostic tool because it allows the visualization of cardiac chambers, valves and pericardium. Small ventricles (‘kissing ventricles’) may incite fluid administration whereas a poorly contractile myocardium may suggest that a dobutamine infusion is a better choice. Right ventricular dilatation may orient towards the diagnosis of massive pulmonary embolism or myocardial infarction whereas the presence of pericardial fluid may suggest a diagnosis of pericardial tamponade. Severe valvulopathy can also be recognized promptly. However, echocardiography instruments and expertise may not be readily available everywhere; in some institutions, this is still the domain of the cardiologists and they need to be called to do the procedure.

If an ultrasound beam is directed along the aorta using a probe, part of the ultrasound signal will be reflected back by the moving red blood cells at a different frequency. The resultant Doppler shift in the frequency can be used to calculate the flow velocity and volume and hence cardiac output. Echo-Doppler evaluation can provide reasonable estimates of cardiac output, but again is operator-dependent and continuous measurement of cardiac output using this technique is not possible. Echo-Doppler evaluation may be applied either transthoracically or transeosophageally. However, transthoracic techniques do not always yield good images and transeosophageal techniques are more invasive such that some sedation, and often endotracheal intubation, is required in order to obtain reliable measurements. Moreover, the esophageal probe is uncomfortable in non-intubated patients, although may be better tolerated if inserted nasally, and should be used cautiously in patients with esophageal lesions. The signal produces different waveforms that can be used to distinguish to some extent changes in preload, afterload and left ventricular contractility. Doppler flow studies focusing on the descending thoracic aorta may not provide a reliable measurement of the total cardiac output (for example, with epidural use), and are invalid in the presence of intra-aortic balloon pumping. Echo-Doppler cardiac output estimates vary considerably for several reasons, including difficulty in assessment of the velocity time integral, calculation error due to the angle of insonation, and problems with correct measurement of the cross-sectional area. Some training is required when using these techniques. Esophageal-Doppler techniques have been shown to be useful for optimizing fluid administration in high risk surgical patients [7,8].
Simplified transesophageal Doppler techniques can be convenient as the probe is smaller than for standard esophageal echocardiography techniques. Simplified trans-thoracic Doppler systems allow estimation of aortic blood flow and may be even less invasive; however, although these techniques can be simple to perform in healthy volunteers, access to good images may be more difficult in critically ill patients. Moreover, there is a fairly prolonged learning curve for correct use of this system [9]. These methods need further validation in critically ill patients.

CO₂ rebreathing
CO₂ rebreathing systems, based on the Fick principle, use a CO₂ sensor, a disposable airflow sensor and a disposable rebreathing loop. CO₂ production is calculated from minute ventilation and its CO₂ content, and the arterial CO₂ content is estimated from end-tidal CO₂. Partial rebreathing reduces CO₂ elimination and increases the end-tidal CO₂. By combining measurements taken during and without rebreathing, venous CO₂ content can be calculated. Bioreactance has developed out of bioimpedance and measures changes in the frequency of the electrical currents traversing the chest, rather than changes in impedance, potentially making it less sensitive to noise. These techniques are non-invasive and can be applied quickly. They have been used for physiological studies in healthy individuals and may be useful in perioperative applications [10], but are less reliable in critically ill patients [11]. Electrical interference may also occur in the ICU environment.

Bioimpedance and bioreactance
Bioimpedance is based on the fact that the conductivity of a high-frequency, low-magnitude alternating current passed across the thorax changes as blood flow varies with each cardiac cycle. These changes can be measured using electrodes placed on a patient’s chest and used to generate a waveform from which cardiac output can be calculated. Bioreactance has developed out of bioimpedance and measures changes in the frequency of the electrical currents traversing the chest, rather than changes in impedance, potentially making it less sensitive to noise. These techniques are non-invasive and can be applied quickly. They have been used for physiological studies in healthy individuals and may be useful in perioperative applications [10], but are less reliable in critically ill patients [11]. Electrical interference may also occur in the ICU environment.

Key principles of hemodynamic monitoring
Having briefly discussed some of the advantages and limitations of the available systems, we now consider some key principles than can help when considering which hemodynamic monitoring system to use.

Principle 1: no hemodynamic monitoring technique can improve outcome by itself
Hemodynamic monitoring can only improve outcomes if three conditions are met: the data obtained from the monitoring device must be sufficiently accurate to be able to influence therapeutic decision making; the data obtained from the monitoring system must be relevant to the patient being monitored; and changes in management made as a result of the data obtained need to be able to improve outcomes. If the data are interpreted or applied incorrectly, or the therapies themselves are ineffective or harmful, the resultant change in management will not improve patient status and may be deleterious. If these three conditions are not met, monitoring is unlikely to be associated with improved outcomes, and this may account for the lack of evidence of improved outcomes in acutely ill patients with use of any monitoring device, not only the PAC [12].

Principle 2: monitoring requirements may vary over time and can depend on local equipment availability and training
The optimal monitoring system will depend on the individual patient, the problem already present or potentially arising for which the monitoring is required, and the devices and expertise available at the institution in question.

For initial evaluation of the critically ill patient, an invasive approach is still often needed, which includes insertion of an arterial catheter and a central venous catheter; this is because of the need for secure intravenous and arterial access in such patients and the presumed increased accuracy of measurements based on direct pressure monitoring. The data provided can already guide initial treatment. Analysis of the arterial pressure trace can identify fluid responsiveness in mechanically ventilated patients, although there are some limitations to this technique, including adaptation to the respirator (often with high doses of sedative agents and even paralysis), need for absence of arrhythmias, and use of relatively large tidal volumes. Response to passive leg raising can be used if beat-by-beat measurements of stroke volume are monitored. Once stabilized, less invasive monitoring techniques should be employed. Importantly, monitoring systems are not necessarily mutually exclusive and can sometimes be used to complement each other. For example, echocardiography can provide additional information in the early assessment of critically ill patients (Figure 1).

There is still a place for the PAC (Swan-Ganz), which has the advantage of allowing measurement of cardiac filling pressures and pulmonary artery pressures, cardiac output and SvO₂ (and now also extravascular lung water). However, although in the past a PAC was inserted early in all critically ill patients, today its insertion is no longer necessary during initial resuscitation, but should rather be reserved for complex cases, for example, patients with right ventricular dysfunction, difficult assessment of
optimal fluid management, or specific cases of cardiac failure.

**Principle 3: there are no optimal hemodynamic values that are applicable to all patients**

Although it may be appealing to have some simple targets, such as keeping the mean arterial pressure above 65 mmHg, the central venous pressure (CVP) above 8 mmHg, or DO₂ above 600 mL/minute/M², such targets are overly simplistic and may even be potentially dangerous. For example, the acceptable minimal arterial pressure may be very different in a young individual without co-morbidity compared to an elderly atherosclerotic, previously hypertensive patient. Likewise, the CVP may remain low in adequately resuscitated patients or may be high at baseline in patients with pulmonary hypertension due to underlying chronic lung disease. Similarly, it is difficult to define an optimal level of cardiac output as cardiac output is an adaptive parameter for which there is no single ‘normal’ value, but only normal ranges. Since the purpose of the cardiovascular system is to match blood flow to metabolic demand, and this demand may vary widely even over short time intervals, targeting a specific cardiac output or even sustaining a threshold value may be inappropriate. For example, keeping a cardiac output above 5 or 6 liters per minute in an adult would be like driving constantly at 80 km/h, whether in a small town or on the freeway. In the critically ill, cardiac output increases in sepsis, as in anemia, but may be reduced with sedation or anesthesia. Multiple factors therefore need to be considered when determining whether cardiac output is optimal for a particular patient, including the degree of tissue perfusion as estimated from a careful clinical examination and blood lactate levels (Figure 2). Alarms should thus be individualized for each patient and reevaluated regularly.

**Principle 4: we need to combine and integrate variables**

Any variable on its own provides relatively little information - it is just one piece of a large puzzle. We need rather to integrate all the available data from multiple sources. For example, a hypotensive patient with a low cardiac output will present different diagnoses (hypovolemia, decreased contractility or obstruction) and hence require different treatments to a hypotensive patient with a high cardiac output (decreased vascular tone). Likewise, as discussed earlier, correct interpretation of a low cardiac output involves consideration of many factors (Figure 2), including some assessment of cardiac filling (pressures or dimensions) to assess ventricular preload.

**Principle 5: measurements of SvO₂ can be helpful**

SvO₂ reflects the balance between oxygen consumption (VO₂) and DO₂ and thus provides an indication of the adequacy of tissue oxygenation. If there is no PAC in situ, the oxygen saturation in the superior vena cava (ScvO₂) can be measured using a central venous catheter and has been proposed as a surrogate for SvO₂. Importantly, ScvO₂ represents just an approximation of the SvO₂ [13] and the absolute values of ScvO₂ and SvO₂ are not interchangeable. The difference between these two parameters is influenced by the sampling site of central venous blood, the presence of left-to-right shunts, incomplete mixing of venous blood, oxygen extraction in the renal and the splanchnic beds, redistribution of blood flow through the upper and lower body, level of consciousness (anesthesia) and myocardial VO₂. The reliability of ScvO₂ is also dependent on the position of the tip of the catheter, with right atrial measurements closely
approximating SvO₂ and high vena cava measurements often deviating substantially from SvO₂. In general, SvO₂ is more useful when the value is below normal (see below), even though in these conditions it may not reflect a hemodynamic problem. Simultaneous measurements of blood lactate levels can be helpful. A diagnostic algorithm based on SvO₂ and cardiac output is shown in Figure 3.

**Principle 6: a high cardiac output and a high SvO₂ are not always best**

Although ICU physicians may like to increase cardiac output and SvO₂ by giving more fluid and inotropic agents, is this always good? Excessive fluid administration to increase cardiac output may result in fluid overload with massive edema formation and this may be associated with worse outcomes [14]; some systems measure extravascular lung water, which can help document this. Similarly, excessive doses of dobutamine can be detrimental, compromising myocardial function, especially in patients with coronary artery disease; giving inotropic agents in the presence of coronary artery disease is like trying to stimulate a tired horse. Using vasoactive agents and fluids to increase DO₂ to supranormal levels in all patients may result in excessive mortality rates and this strategy has been abandoned [15]. A high ScvO₂ has been suggested as a target for some high risk patients or in shock resuscitation, with Rivers and colleagues [16] reporting that septic patients assigned to an early goal-directed therapy algorithm had higher ScvO₂ values and reduced mortality rates. However, this was a strategy for early resuscitation of patients with severe sepsis in a single institution, and needs further validation in multicenter studies, several of which are currently ongoing. Importantly, applying the same strategy in general ICU patients may not improve outcomes [17]. Indeed, in patients with sepsis, a high SvO₂ may be the result of maldistribution of peripheral blood flow and altered oxygen extraction, rather than adequate perfusion, such that patients may still deteriorate even with a high SvO₂. In sepsis, a high cardiac output, like a high SvO₂, can be associated with worse outcomes.

**Principle 7: cardiac output is estimated, not measured**

No bedside method is available to directly assess cardiac output, so all values obtained are estimates. As such, comparison of measurements obtained with different techniques results in relatively poor agreement and significant bias. The intermittent thermodilution technique is generally considered as the ‘reference’ standard, but has its own limitations. A measurement obtained by a less invasive technique may be preferable if it can be obtained more rapidly and easily, even if it is slightly less accurate. Importantly, the accuracy of absolute values may be less important if one is following trends, for example, to track the short-term effects of therapies, such as fluid loading.

**Principle 8: monitoring hemodynamic changes over short periods of time is important**

Monitoring of acute changes in cardiac output can be important, for example, in patients at risk of acute bleeding or in assessing the response to fluid administration to separate fluid responders from non-responders. Evaluating the response to a dobutamine or to a nitrates infusion is another example of this functional monitoring that may also have sound clinical applications. This assessment of hemodynamic variations observed during the challenge of the cardiovascular system has been termed ‘functional hemodynamic monitoring’ [18]. The study of slow changes in cardiac output over several days may be less relevant in most patients, although can be useful to follow the clinical course of the cardiac patient. Combining measures of multiple variables and their dynamic interactions in response to time and specific treatments often increases the sensitivity and specificity of these monitoring modalities to identify specific disease processes and quantify whether therapy is effective or not.

**Principle 9: continuous measurement of all hemodynamic variables is preferable**

Although there are no data to demonstrate the superiority of continuous cardiac output measurements over intermittent monitoring, there has been a global evolution towards more continuous measurement of variables. We can now routinely measure various hemodynamic variables continuously, including heart rate, arterial pressure and CVP. Using the thermodilution technique, one may not wish to go back to intermittent measurements of cardiac output by repeated injections of cold water boluses. It may even be preferable to have real-time (beat-by-beat) continuous cardiac output measurement rather than a built-in delay like the semi-continuous
cardiac output provided with the Vigilance™ system. Systems that are not continuous (for example, echocardiography) or that require calibration (for example, transpulmonary indicator dilution) may not provide the real-time data needed for optimal acute management of unstable critically ill patients, whereas systems that provide continuous non-calibrated cardiac output measurements suffer from reduced accuracy.

Principle 10: non-invasiveness is not the only issue
Non-invasiveness is not the only goal. Although it is always preferable to be less invasive, being non-invasive is not always possible and may even be counter effective. For example, continuous monitoring of arterial pressure is more invasive than intermittent monitoring but is helpful in hypotensive (or severe hypertensive) states. Likewise, a central venous catheter can be helpful to monitor the CVP and the ScvO₂ (and also facilitates the rapid administration of fluids). Whenever possible, we should of course try to be as non-invasive as possible, but arterial pressure monitoring and CVP monitoring are still invasive. Echocardiography must be promoted more for its ability to offer a direct evaluation of cardiac function than for its non-invasiveness. Even though it is the most invasive method, the PAC is still of value in very sick patients with complex problems, for example, respiratory failure with shock and oliguria. At the other extreme, completely non-invasive bioimpedance has a place in healthy individuals, but little place in critically ill patients. Other monitoring systems are of use in patients with conditions somewhere between these two extremes. The optimal device depends on the type of patient, the question being asked, and the condition being managed or anticipated.

Conclusion
The ideal hemodynamic monitoring system should comprise all the key factors listed in Table 2; however, such a system does not currently exist so we must try and choose devices that have a maximum of these attributes, bearing in mind that there is no ‘one size fits all’ type of system and one should, therefore, select the system most appropriate for each patient and, perhaps even more importantly, for each type of problem. It is important to be familiar with the technology being used, profiting from its advantages but recognizing its limitations. Most systems now offer (almost) continuous measurements, with the possible exception of echocardiography techniques because it is difficult to leave the probe in place for prolonged periods. Hemodynamic monitoring can be particularly helpful in the early stages of resuscitation, but is less useful when organ failure is established. Most importantly, one must never forget that it is not the monitoring itself that can improve outcomes but the changes in therapy guided by the data obtained.

Abbreviations
CVP, central venous pressure; DO₂, oxygen delivery; PAC, pulmonary artery catheter; ScvO₂, central venous oxygen saturation; SvO₂, mixed venous oxygen saturation; VO₂, oxygen consumption.

Competing interests
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Author details
1Department of Intensive Care, Erasme Hospital, Université Libre de Bruxelles, 808 route de Lenon, 1070-Brussels, Belgium. 2Department of Intensive Care Medicine, St George’s Healthcare NHS Trust, Blackshaw Road, London, SW1 7 QX, UK. 3Department of Anesthesiology and Intensive Care, Sheba Medical Center, Tel Aviv University, Tel Aviv, 52621 Israel. 4Division of Pulmonary, Allergy and Critical Care, Emory University School of Medicine, Grady Memorial Hospital, 615 Michael Street, Suite 205, Atlanta, GA 30322, USA. 5Department of Anesthesia and Intensive Care Medicine, University Hospital, Medical School, University of Udine, P.le S. Maria della Misericordia, 15331 Udine, Italy. 6Department of Anesthesiology and Critical Care Medicine, Pôle d’Anesthésie Réanimation, Hôpital Claude Huriez, rue Michel Polonoski, CHU Univ Nord de France, 59000 Lille, France. 7Department of Critical Care Medicine, 606 Scaife Hall, 3550 Terrace Street, Pittsburgh, PA 15261, USA. 8Institute of Anaesthesiology and Intensive Care Medicine, Trenkwalder City Hospital, Zürich, Birmensdorferstr. 497, 8063 Zürich, Switzerland. 9Service de réanimation médicale, Centre Hospitalo-Universitaire de Bicêtre, Assistance Publique-Hôpitaux de Paris, EA 4046, Université Paris Sud, 78 rue du Général Leclerc, 94 270 Le Kremlin-Bicêtre, France. 10Department of Neonatology, Radboud University Nijmegen Medical center, PO Box 9101, 6500 NJN Nijmegen, The Netherlands. 11Department of Anaesthesia and Intensive Care, University of Siena, Viale Bracci, 1, 53100 Siena, Italy. 12Réanimation Medical and Covidien; Deltex Medical provide unrestricted research grants.
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