

Reducing the Risk of Iatrogenic Anemia and Catheter-Related Bloodstream Infections Using Closed Blood Sampling

INTRODUCTION

In the Intensive Care Unit (ICU), critically ill patients are more numerous and severely ill than ever before.¹ To effectively care for these patients, clinicians rely on physiologic monitoring of blood-flow, oxygen transport, coagulation, metabolism, and organ function. This type of monitoring has made the collection of blood for testing an essential part of daily management of the critically ill patient, yet it is widely recognized that excessive phlebotomy has a deleterious effect on patient health. The result is a clinical paradox in which diligent care may contribute to iatrogenic anemia.

RISKS ASSOCIATED WITH CONVENTIONAL DIAGNOSTIC BLOOD SAMPLING

Iatrogenic Anemia

The process of obtaining a blood sample from an indwelling central venous or arterial catheter requires a volume of diluted blood (2–10 mL) to be discarded or “cleared” from the catheter before a sample can be taken.^{2,3} Studies have shown that patients with central venous or arterial catheters have more blood sampling than ICU patients who don’t have these catheters and the total blood volume drawn from patients with arterial catheters is 44% higher than patients without arterial catheters (See Table 1).^{4,5}

Use of blood sampling techniques that rely on discarding a volume of blood for each sample may contribute to iatrogenic anemia, which remains a prevalent issue affecting the vast majority of patients in the ICU.

It has also been reported that mean blood loss per cardiothoracic ICU patient stay is approximately 377 mL, 240 mL per patient stay in general surgical ICUs and 41.5 mL per patient stay in medical-surgical ICUs.^{4,6} Another study found that the total average volume of blood drawn over a 7-day medical intensive care unit (MICU) stay was 257.4 mL (See Figure 1).⁷ More recently, an ICU-based study found an average blood draw volume in 24 hours was 41.1 mL per patient.⁸ Because the most critically ill patients may have up to 24 diagnostic blood samples drawn in a day, this frequent sampling can contribute to 17% of the total blood loss while in the ICU.^{9,10,11}

Loss of blood volume causes anemia, a condition in which lowered hematocrit (HCT) and hemoglobin (Hgb) in the blood limits the ability of red blood cells (RBC) to transport oxygen to the body’s tissues. Use of blood sampling techniques that rely on discarding a volume of blood for each sample may contribute to iatrogenic anemia, which remains a prevalent issue affecting the vast majority of patients in the ICU, especially those with prolonged stays.

Almost 95% of patients admitted to an intensive care unit have an Hgb concentration that is below normal by day 3 of admission, often requiring blood transfusion.¹² It has also been shown that phlebotomy accounts for 49% of the variation in the amount of RBCs transfused.²

TABLE 1. COMPARISON OF NUMBER OF PROCEDURES AND VOLUME OF BLOOD DRAWN IN ARTERIAL LINE AND NONARTERIAL LINE PATIENT GROUPS* ⁵

	ARTERIAL LINE	NONARTERIAL LINE	PERCENT DIFFERENCE	P VALUE
No. of procedures				
1	8.1 ± 4.7	5.8 ± 2.8	28	0.048
2	5.0 ± 2.8	3.4 ± 1.5	32	0.012
T	13.1 ± 6.8	9.2 ± 3.4	30	0.014
Volume of blood, mL				
1	70.9 ± 37.2	42.4 ± 22.1	40	0.002
2	43.5 ± 24.4	22.0 ± 11.6	49	<0.001
T	114.7 ± 53.9	63.6 ± 28.4	44	<0.001

*1=first 24-h period; 2=second 24-h period; T=total over 48 h

Despite evidence to support a restrictive transfusion practice, the transfusion of packed red blood cells (PRBC) remains a primary intervention for the treatment of ICU patients with anemia.¹³ In two large, multi-facility cohort studies, 44% of patients in ICUs in the United States and 37% of those in ICUs in Western Europe received blood transfusions.^{8,14} Blood transfusions are associated with negative effects on patient outcomes, including increased risk for infection, which may explain the positive correlation between organ dysfunction and the number of blood draws.^{8,10,11,15} Additional risks of transfusions include allergic, anaphylactic, and hemolytic transfusion reactions and acute respiratory distress syndrome, all contributing to significant morbidity and mortality.^{16,17,18,19}

Catheter-related bloodstream infections (CRBSI)

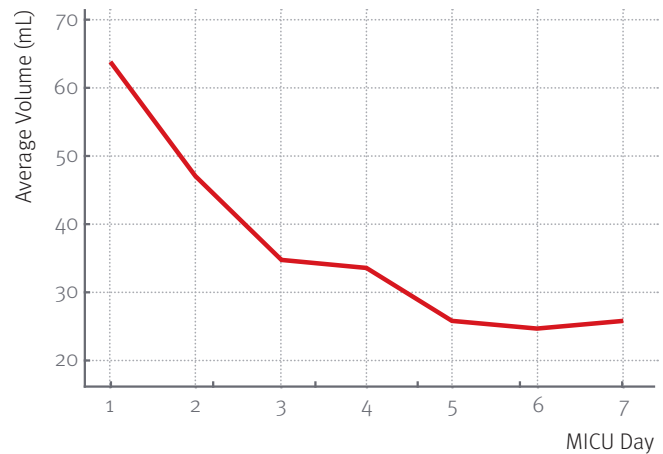
CRBSI is the most common nosocomial bacteremia in critically ill patients.²⁰ It affects nearly 50,000 patients each year in the US, with an attributable mortality of up to 35% and a financial cost of up to \$30,000 per case.²¹ Studies also show that CRBSIs stemming specifically from arterial catheters occur at a rate less than infection rates of short-term central venous catheters.^{22,23,24,25,26}

High levels of catheter handling can facilitate hub colonization with micro-organisms derived from the patient's skin or from contact by healthcare workers.²⁷ The risk for bacterial ingress and arterial line contamination is also increased by catheter manipulation such as opening the system for blood sampling.^{28,29} In particular, using a 3-way stopcock without a self-sealing port for blood sampling may increase contamination due to access frequency, insufficient aseptic technique, or residual blood within the ports.³⁰

One study found that patients using a blood conservation system had a 48% reduction in PRBC transfusion requirements.³¹

the sterile reservoir back to the patient after a sample has been drawn, reducing blood loss as well as the potential for bacterial ingress to the closed system. The closed, in-line BCS also reduces clinician exposure to potential bloodborne pathogens during the sampling process.

FIGURE 1. LABORATORY BLOOD VOLUME⁷



The average laboratory blood volume drawn over a 7-day MICU stay.
Total average volume of blood drawn for 7 days was 257.4 mL

CLOSED, IN-LINE BLOOD CONSERVATION SYSTEMS (BCS)
BCS such as SafeSet® (ICU Medical Inc., San Clemente, CA) eliminate the need to discard the clearing volume associated with sampling through indwelling arterial catheters.³¹ When performing a sample collection using a BCS, blood and flush solution are drawn into a reservoir distal to the sampling port. Then, while maintaining aseptic sampling technique, a clinician is able to return the blood clearing volume held in

Avoiding transfusions by conserving blood

In-line BCSs have been associated with a 50% reduction in daily diagnostic blood loss, and reducing blood loss helps reduce cases of anemia in the ICU, as well as risks associated with blood transfusions.^{7,32,33,34}

In a survey of members of the Society of Critical Care Medicine, most agreed that in-line BCSs could be very useful in preventing anemia.⁷ Another study found that patients using a BCS had a 48% reduction in PRBC transfusion requirements as well as a smaller decrease in Hgb levels between ICU admission and discharge.³¹ This finding is significant given the current worldwide shortage of PRBCs, the cost of transfusions (estimated between \$500 and \$1,200),³⁵ and the desire to avoid the significant risk of morbidity and mortality associated with transfusions.

Reducing catheter contamination and the risk of CRBSI

By eliminating open systems and minimizing points for bacterial ingress, closed, in-line BCSs may significantly reduce arterial and central line contamination.³⁶ One study found the use of a BCS correlated with lower rates of intraluminal fluid contamination compared to a traditional 3-way open-port stopcock system. Cultures of the intraluminal fluid from the open-port stopcock system yielded growth of various species of micro-organisms, compared to any positive cultures from the BCS, which yielded growth of a single species. Another study reported that use of a closed BCS (n=60) resulted in fewer instances of intraluminal fluid contamination compared to use of a conventional 3-way stopcock system (n=70), 7% vs 61%, respectively.³⁷

The ability of a BCS to prevent microbial contamination is further enhanced by incorporating needlefree connectors into the sampling port stopcock. An in vitro study found that closed systems, combined with needlefree, self-sealing valve sampling ports, maintained a barrier that minimized bacterial ingress into the catheter and reduced colonization of the sampling hub.³⁸ Another study comparing conventional open sampling systems to self-sealing valve connectors within a post-surgical cardiothoracic ICU reported a 4.3% hub colonization rate with valve connectors and a 14.2% colonization rate with open sampling systems.³⁹ The study also reported 10.9% catheter tip colonization with the self-sealing valve connectors and 17.2% colonization rate with open sampling systems (See Figure 2).

CONCLUSION

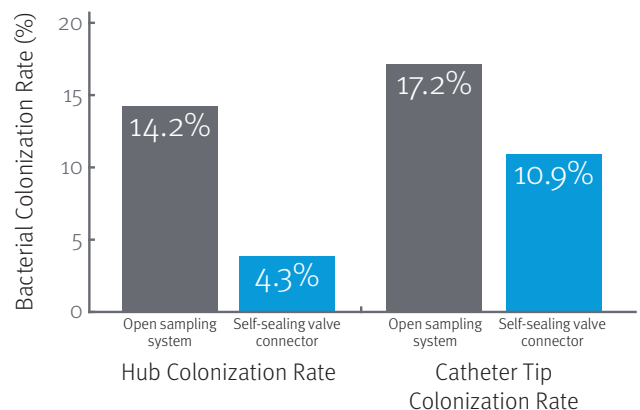
The pervasive anemia experienced by the majority of patients in the ICU is minimized by the utilization of closed, in-line blood sampling and conservation systems.

By reducing blood loss and the potential for iatrogenic anemia, closed blood sampling and conservation systems help reduce the need for and inherent risk of transfusions in the ICU. Finally, the application of closed blood sampling and conservation systems prevents the transfer of bacteria into the catheter and helps clinicians in their efforts to minimize catheter-related bloodstream infections.



The SafeSet Closed Blood Conservation System (ICU Medical Inc., San Clemente, CA) allows clinicians to conserve blood by reinfusing the clearing volume drawn during blood sampling.

FIGURE 2. COMPARISON OF BACTERIAL COLONIZATION RATES USING OPEN SAMPLING SYSTEMS AND SELF-SEALING VALVE CONNECTORS³⁹



References

- Barie, P. Phlebotomy in the intensive care unit: strategies for blood conservation. *Critical Care* 2004, 8(Suppl 2):S34-S36
- Corwin HL, Parsonnet KC, Gettinger A: RBC transfusion in the ICU. Is there a reason? *Chest* 1995, 108:767-771.
- Zimmerman JE, Seneff MG, Sun X, Wagner DP, Knaus WA: Evaluating laboratory usage in the intensive care unit: patient and institutional characteristics that influence frequency of blood sampling. *Crit Care Med* 1997, 25:737-748.
- Smoller BR, Kruskall MS. Phlebotomy for diagnostic laboratory tests in adults. Pattern of use and effect on transfusion requirements. *N Engl J Med* 1986;314:1233-5.
- Low LL, Harrington GR, Stoltzfus DP. The Effect of Arterial Lines on Blood-Drawing Practices and Costs in Intensive Care Units. *CHEST* 1995; 108:216-19
- Henry ML, Garner WL, Fabri PJ. Iatrogenic anemia. *Am J Surg* 1986;151:362-3.
- Silver MJ, Li YH, Gragg LA, Jubran F, Stoller JK: Reduction of blood loss from diagnostic sampling in critically ill patients using a blood-conserving arterial line system. *Chest* 1993, 104:1711-1715.
- Vincent JL, Baron JF, Reinhart K, et al. Anemia and blood transfusion in critically ill patients. *JAMA* 2002;288:1499-507.
- Fowler RA, Berenson M. Blood conservation in the intensive care unit. *Crit Care Med* 2003, 31:5715-5720.
- Tinmouth AT, McIntyre LA, Fowler RA: Blood conservation strategies to reduce the need for red blood cell transfusion in critically ill patients. *CMAJ* 2008, 178:49-57.
- von Ahsen N, Muller C, Serke S, Frei U, Eckardt KU: Important role of nondiagnostic blood loss and blunted erythropoietic response in the anemia of medical intensive care patients. *Crit Care Med* 1999, 27:2630-2639.
- Corwin HL. Erythropoietin use in critically ill patients: forest and trees. *CMAJ* 2007; 177(7):747-749.
- Hebert PC, Wells G, Blajchman MA, Marshall J, Martin C, Pagliarello G, Tweeddale M, Schweitzer I, Yetisir E, The Transfusion Requirements in Critical Care Investigators for the Canadian Critical Care Trials Group: A Multicenter, Randomized, Controlled Clinical Trial of Transfusion Requirements in Critical Care. *N Engl J Med* 1999, 340:409-417.
- Corwin HL, Gettinger A, Pearl RG, Fink MP, Levy MM, Abraham E, MacIntyre NR, Shabot MM, Duh MS, Shapiro MJ: The CRIT Study: Anemia and blood transfusion in the critically ill--current clinical practice in the United States. *Crit Care Med* 2004, 32:39-52.
- Ottino G, De Paulis R, Pansini S, Rocca G, Tallone MV, Comoglio C, Costa P, Orzan F, Morea M: Major sternal wound infection after open-heart surgery: a multivariate analysis of risk factors in 2,579 consecutive operative procedures. *Ann Thorac Surg* 1987, 44:173-179.
- Kleinman S, Chan P, Robillard P: Risks associated with transfusion of cellular blood components in Canada. *Transfus Med Rev* 2003, 17:120-162.
- Shorr AF, Jackson WL, Kelly KM, Fu M, Kollef MH: Transfusion practice and blood stream infections in critically ill patients. *Chest* 2005, 127:1722-1728.
- Gong MN, Thompson BT, Williams P, Pothier L, Boyce PD, Christiani DC: Clinical predictors of and mortality in acute respiratory distress syndrome: potential role of red cell transfusion. *Crit Care Med* 2005, 33:1191-1198.
- Zilberberg MD, Carter C, Lefebvre P, Raut M, Vekeman F, Duh MS, Shorr AF: Red blood cell transfusions and the risk of acute respiratory distress syndrome among the critically ill: a cohort study. *Crit Care* 2007, 11:R63.
- Valles J, Leon C, Alvarez-Lerma F. Nosocomial bacteremia in critically ill patients: a multicenter study evaluating epidemiology and prognosis. *Clin Infect Dis* 1997;24:387e395.
- Wenzel RP, Edmond MB. Team-based prevention of catheter-related infections. *N Engl J Med* 2006;355:2781e2783.
- Traoré O, Liotier J, Souweine B. Prospective study of arterial and central venous catheter colonization and of arterial- and central venous-related bacteremia in intensive care units. *Crit Care Med* 2005;33:1276-80.
- O'Grady NP, Alexander M, Burns LA, Dellinger EP, Garland J, Heard SO, et al. Guidelines for the prevention of intravascular catheter-related infections. *Am J Infect Control* 2011;39:S1-34.
- Khalifa R, Dahyot-Fizelier C, Laksiri L, Ragot S, Petipas F, Nanadoumgar H, et al. Indwelling time and risk of colonization of peripheral arterial catheters in critically ill patients. *Intensive Care Med* 2008;34:1820-6.
- Maki DG, Kluger DM, Crnich CJ. The risk of bloodstream infection in adults with different intravascular devices: a systematic review of 200 published prospective studies. *Mayo Clin Proc* 2006;81:1159e1171.
- Koh DB, Gowardman JR, Rickard CM, Robertson IK, Brown A. Prospective study of peripheral arterial catheter infection and comparison with concurrently sited central venous catheters. *Crit Care Med* 2008;36:397e402.
- Sitges-Serra A, Linˆares J, Garau J. Catheter sepsis: the clue is the hub. *Surgery* 1985;97:355e357.
- Adal KA, Farr BM: Central venous catheter-related infections: a review. *Nutrition* 1996, 12:208-213.
- Crow S, Conrad SA, Chaney-Rowell C, King JW. Microbial contamination of arterial infusions using for hemodynamic monitoring: a randomized trial of contamination with sampling through conventional stopcock versus a novel closed system. *Infect Control Hosp Epidemiol* 1989;10:557-61.
- Maki DG. Pathogenesis, prevention, and management of infectious due to intravascular devices used for inpatient therapy. In: Bisano AL, Waldvogel FA, editors. *Infections associated with in-dwelling medical devices*. Washington [DC]: American Society for Microbiology; 1989. p. 161-77.
- Mukhopadhyay A, Yip HS, Prabhuswamy D, Chan YH, Phua J, Lim TK, et al. The use of blood conservation device to reduce red blood cell requirements: a before and after study. *Crit Care* 2010;14:R7.
- MacIsaac CM, Presneill JJ, Boyce CA, Byron KL, Cade JF: The influence of a blood conserving device on anaemia in intensive care patients. *Anaesth Intensive Care* 2003, 31:653-657.
- Peruzzi WT, Parker MA, Lichtenthal PR, et al. A clinical evaluation of a blood conservation device in medical intensive care unit patients. *Crit Care Med* 1993;21:501-6.
- Gleason E, Grossman S, Campbell C: Minimizing diagnostic blood loss in critically ill patients. *Am J Crit Care* 1992, 1:85-90.
- Shander A, Hofmann A, Ozawa S, et al. Activity-based costs of blood transfusions in surgical patients at four hospitals. *Transfusion*. 2010 Apr; 50(4):753-65.
- Peruzzi WT, Noskin GA, Moen SG, Yungbluth M, Lichtenthal P, Shapiro BA: Microbial contamination of blood conservation devices during routine use in the critical care setting: results of a prospective, randomized trial. *Crit Care Med* 1996, 24:1157-1162.
- Crow S, Conrad SA, Chaney-Rowell C, King JW. Microbial contamination of arterial infusions using for hemodynamic monitoring: a randomized trial of contamination with sampling through conventional stopcock versus a novel closed system. *Infect Control Hosp Epidemiol* 1989;10:557-61.
- Yébenes JC, Sauca G, Solsona M, Martinez R, Serra-Prat M, Gil P, et al. Safety of positive-pressure valve connectors in arterial catheters inserted into critically ill patients. *J Hosp Infect* 2008;70:341-5.
- Bouza E, Munoz P, Lopez-Rodriguez J, et al. A needleless closed system device (CLAVE) protects from intravascular catheter tip and hub colonization: a prospective randomized study. *J Hosp Infect* 2003;54:279e287