

University of Geneva Hospitals and Faculty of Medicine, Geneva, Switzerland



WHO Collaborating Centre Infection Prevention and Control and Antimicrobial Resistance



Le retrait systématique des cathéters veineux périphériques

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CONTENT

- Introduction
- Literature update: routine replacement *versus* clinically indicated replacement
- Large *«before-after-before»* study conducted in Geneva
- New data on dwell time
- Conclusions





INTRODUCTION

- A global audit across 13 countries:
 - almost 60% of inpatients had at least one peripheral venous catheter (PVC) in place
- In Switzerland, 48.6% of patients in acute care have a PVC in place every single day
- Approximately 330 million PVCs were sold annually in the US
- PVC-related complications:
 - Hematoma
 - Phlebitis
 - Extravasation
 - Bruising
- Little is known about the bloodstream infection (BSI) risk associated with PVCs





INTRODUCTION

PVCs and BSI risk

- Short-term PVCs accounted for:
 - a mean of 6.3% of nosocomial BSIs
 - 23% of nosocomial catheter-related BSIs
- Relative risk of CVC-BSI compared to PVC-BSI varied from 1.5 to 64
- Proportion of S. aureus PVC-BSI among CRBSI is high

Table 2. Risk of Staphylococcus aureus Bloodstream Infections due to Infected Peripheral Vascular Catheters CRBSI							
Study, First Author [Ref]	Staphylococcus aureus CR-BSIs due to PVCs	<i>Staphylococcus aureus</i> BSIs due to PVCs					
Mylotte [50]	50% of 28 CR-BSIs	18% of 79 BSIs					
Thomas ^a [51]	50% of 305 CR-BSIs						
Kok [52]	41% of 75 CR-BSIs	25% of 123 BSIs					
Bruno [55]		35% of 31 BSIs ^b					
Trinh [53]	12% of 196 CR-BSIs ^c						
Mestre [46]	64% of 14 CR-BSIs	28% of 32 BSIs					
Stuart [56]		24% of 583 BSIs					
Morris [54]	44% of 121 CR-BSIs	20% of 261 BSIs					
Rhodes [57]		24% of 151 BSIs ^d					
Austin ^a [49]		7.6% of 445 BSIs					

Prolonged dwell time and catheter insertion under emergent conditions increased risk of PVC-BSI...





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INTRODUCTION

PVCs and BSI risk – dwell time

- No correlation between the number of catheter days per site for patients with a peripheral IV device, and hospital-acquired bacteraemia
- The mean PVC dwell time in PVC-BSI cases was 3.9 days (±2.1 days)

- Horeover ... Several studies > reductions in risk of s. aureus in the studies in after instituting infection prevention initiatives involving limits in and mount prevenuou prevenuou muanves involving interventions and dwell time to 3 days or doing so along with other intervention The must be

Mermel L. Clir ous Diseases, 2017;65(10):1757–62 // Nystrom - J Hosp Infect 1983 //Pujol M et al. J Hosp Infect 2007; 67:22–9 //Collignon PJ et al. Med J Aust 2013; 199:750–1 //Freixas N et al. Clin Microbiol Infect 2013; 19:838–44. // Fry DE, Borzotta AP. Am J Surg 1994; 167:268–72. //Lolom I et al. Presse Med 2009; 38:34–42.





INTRODUCTION

Several prevention measures – one of them...

• Whether to replace PVCs routinely or when clinically indicated was categorized as an unresolved question by the US CDC:

Replacement of Peripheral and Midline Catheters Recommendations

- 1. There is no need to replace peripheral catheters more frequently than every 72–96 hours to reduce risk of infection and phlebitis in adults [36, 140, 141]. *Category IB*
- 2. No recommendation is made regarding replacement of peripheral catheters in adults only when clinically indicated [142–144]. *Unresolved issue*
- 3. Replace peripheral catheters in children only when clinically indicated [32, 33]. Category IB
- 4. Replace midline catheters only when there is a specific indication. Category II





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Recent systematic review and meta-analysis

- 7,412 catheters (from RCTs) included
- Routine replacement:
 - \downarrow infiltration of fluid into surrounding tissues
 - \downarrow rates of catheter failure due to **blockage**

Study or subgroup	Clinically indicated	Routine re- placement	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% CI
Rickard 2010	61/185	53/177		7.4%	1.1[0.81,1.49]
Rickard 2012	279/1593	235/1690		31.16%	1.26[1.07,1.48]
Vendramim 2018	172/672	134/647		18.66%	1.24[1.01,1.51]
Webster 2007	43/103	44/103		6.01%	0.98[0.71,1.35]
Webster 2008	135/379	120/376	- +	16.46%	1.12[0.91,1.36]
Xu 2017	144/553	161/645		20.31%	1.04[0.86,1.27]
Total (95% CI)	3485	3638	•	100%	1.16[1.06,1.26]
Total events: 834 (Clinically in	dicated), 747 (Routine repla	acement)			
Heterogeneity: Tau ² =0; Chi ² =3	.92, df=5(P=0.56); I ² =0%				
Test for overall effect: Z=3.36(F	P=0)				
	Fa	vours cl-indicated	0.5 0.7 1 1.5 2	Favours 3-day	

Study or subgroup	Clinically indicated	Routine re- placement	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% CI
Rickard 2010	4/185	5/177		1.01%	0.77[0.21,2.8]
Rickard 2012	344/1593	344/1690	H	66.22%	1.06[0.93,1.21]
Van Donk 2009	13/105	4/95		0.83%	2.94[0.99,8.71]
Vendramim 2018	80/672	61/647	+	12.33%	1.26[0.92,1.73]
Webster 2007	7/103	4/103		0.79%	1.75[0.53,5.8]
Webster 2008	30/379	20/376	++	3.98%	1.49[0.86,2.57]
Xu 2017	82/553	81/645	+	14.83%	1.18[0.89,1.57]
Total (95% CI)	3590	3733	•	100%	1.14[1.02,1.27]
Total events: 560 (Clinically ir	ndicated), 519 (Routine repla	acement)			
Heterogeneity: Tau ² =0; Chi ² =6	6.27, df=6(P=0.39); I ² =4.31%				
Test for overall effect: Z=2.34	(P=0.02)				





Recent systematic review and meta-analysis

- 7,412 catheters (from RCTs) included
- Clinically indicated removal:
 - \downarrow device-related costs

N		pta	acement	Mean Difference			Weight	Mean Difference	
	Mean(SD)	Ν	Mean(SD)		Fixe	d, 95% CI			Fixed, 95% CI
1593	61.7 (39.5)	1690	69.2 (43.5)		 			54.5%	-7.58[-10.42,-4.74
103	29.7 (16.4)	103	37.6 (20.2)	+				17.37%	-7.9[-12.92,-2.88]
379	41.1 (26.6)	376	46.2 (28.7)		•	-		28.13%	-5.17[-9.12,-1.22]
2075		2169						100%	-6.96[-9.05,-4.86]
1, df=2(P=0.5	7); I ² =0%								
0.0001)									
	1593 103 379 2075 1, df=2(P=0.5 0.0001)	1593 61.7 (39.5) 103 29.7 (16.4) 379 41.1 (26.6) 2075 1, df=2(P=0.57); l ² =0% 0.0001)	103 29.7 (16.4) 103 379 41.1 (26.6) 376 2075 2169 1, df=2(P=0.57); l ² =0%	1593 61.7 (39.5) 1690 69.2 (43.5) 103 29.7 (16.4) 103 37.6 (20.2) 379 41.1 (26.6) 376 46.2 (28.7) 2075 2169 1, df=2(P=0.57); l ² =0% 0.0001)	1593 61.7 (39.5) 1690 69.2 (43.5) 103 29.7 (16.4) 103 37.6 (20.2) 379 41.1 (26.6) 376 46.2 (28.7) 2075 2169 1, df=2(P=0.57); l ² =0%	1593 61.7 (39.5) 1690 69.2 (43.5) 103 29.7 (16.4) 103 37.6 (20.2) 379 41.1 (26.6) 376 46.2 (28.7) 2075 2169 1, df=2(P=0.57); l ² =0%	1593 61.7 (39.5) 1690 69.2 (43.5) 103 29.7 (16.4) 103 37.6 (20.2) 379 41.1 (26.6) 376 46.2 (28.7) 2075 2169 1, df=2(P=0.57); l ² =0%	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	1593 61.7 (39.5) 1690 69.2 (43.5) 54.5% 103 29.7 (16.4) 103 37.6 (20.2) 17.37% 379 41.1 (26.6) 376 46.2 (28.7) 28.13% 2075 2169 100% 1, df=2(P=0.57); l ² =0% 0.0001) 100%





Recent systematic review and meta-analysis

- 7,412 catheters (from RCTs) included
- No clear difference in the incidence of thrombophlebitis

	Clinically ind	icated	Routine replace	ement		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
1.2.1 Continuous infu	ision						
Rickard 2010	18	185	12	177	4.1%	1.44 [0.71, 2.89]	_ _
Rickard 2012	114	1593	114	1690	36.8%	1.06 [0.83, 1.36]	+
Vendramim 2018	55	672	64	647	21.7%	0.83 [0.59, 1.17]	
Webster 2007	1	103	2	103	0.7%	0.50 [0.05, 5.43]	
Webster 2008	16	379	12	376	4.0%	1.32 [0.63, 2.76]	
Xu 2017 Subtotal (95% Cl)	76	553 3485	77	645 3638	23.7% 90.9%	1.15 [0.86, 1.55] 1.05 [0.90, 1.23]	- <u>+</u>
Total events	280		281				ſ
Heterogeneity: Chi ^z = Test for overall effect:	: 3.74, df = 5 (P : Z = 0.64 (P = 0	= 0.59); l).52)	^z = 0%				
1.2.2 Intermittent info	usion						
Van Donk 2009 Subtotal (95% CI)	37	105 105	26	95 95	9.1% 9.1 %	1.29 [0.85, 1.96] 1.29 [0.85, 1.96]	-
Total events Heterogeneity: Not ap Test for overall effect:	37 oplicable : Z = 1.19 (P = 0).24)	26				
Total (95% CI)		3590		3733	100.0%	1.07 [0.93, 1.25]	L
Total events	317	1000	307	1100			ľ
rotar events	4.52 df - 6/P	- 0.61\.	Z= 0%				





Recent systematic review and meta-analysis (BSI):

• Similar incidences of **CRBSI**

Figure 4. Forest plot of comparison 1, clinically indicated versus routine change, outcome: 1.1 Catheter-related bloodstream infection

	Clinically ind	cated	Routine replace	ement		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
Rickard 2010	0	105	0	477		Not cotimable	
Rickard 2012	0	1593	1	1690	59.2%	0.35 [0.01, 8.67]	_
van Donk 2009	U	105	U	90		Notestimable	
Vendramim 2018	0	672	0	647		Not estimable	
Webster 2007	0	103	0	103		Not estimable	
Webster 2008	1	379	1	376	40.8%	0.99 [0.06, 15.80]	+
Xu 2017	0	553	0	645		Not estimable	
Total (95% CI)		3590		3733	100.0%	0.61 [0.08, 4.68]	
Total events	1		2				
Heterogeneity: Chi ² = Test for overall effect:	0.23, df = 1 (P Z = 0.47 (P = 0	= 0.63); .64)	²=0%				0.01 0.1 1 10 100 Favours cl-indicated Favours 3-day







Largest RCT 2012

- Exclusion criteria:
 - Bloodstream infection, planned removal of intravenous catheter within 24h, or intravenous catheter already in situ for more than 72 h.
 - PVCs inserted in an emergency were not eligible
- Other methodological problems:
 - Not blinded
 - Phlebitis assessed by research nurses







Largest RCT 2012

	Clinically indicated (n=1593)	Routine replacement (n=1690)	Risk (95%CI)	p value
Primary outcome, intention-to-treat analysis				
Phlebitis per patient, n (%)	114 (7%)	114 (7%)	RR 1·06 (0·83 to 1·36); ARD 0·41% (-1·33 to 2·15)	0.64
Phlebitis/1000 intravenous catheter days (95% CI)	13.08 (10.68–15.48)	13.11 (10.71–15.52)	HR 0.94 (0.73 to 1.23)	0.67
Primary outcome, per-protocol analysis*				
Phlebitis per patient	63/1351 (5%)	47/1186 (4%)	RR 1·18 (0·81 to 1·70); ARD 0·70% (-0·88 to 2·28)	0.39
Phlebitis/1000 intravenous catheter days (95% CI)	11.4 (8.6–14.2)	13.8 (9.9–17.8)	IRR 0.83 (0.56 to 1.23)	0.32
Secondary outcomes, n (n per 1000 intravenous o	atheter days)			
Any infusion failure†	670 (76.9)	636 (73·2)	HR 0.99 (0.89 to 1.11)	0.87
Infiltration	279 (32.0)	235 (27.0)	HR 1.06 (0.89 to1.27)	0.51
Occlusion	344 (39·5)	344 (39.6)	HR 0.92 (0.79 to 1.07)	0.92
Accidental removal	166 (19·0)	159 (1 8·3)	HR 0.98 (0.79 to 1.23)	0.88
CRBSI‡	0 (0)	1 (0.11)		
All BSI	4 (0.46)	9 (1.03)	HR 0.46 (0.14 to 1.48)	0.19
Venous (local) infection‡	0	0		
Mortality, n (%)§	4 (<1%)	4 (<1%)	RR 1.06 (0.27 to 4.23)	0.93

ARD=absolute risk difference. BSI=bloodstream infection. CRBSI=catheter-related bloodstream infection. HR=hazard ratio. IRR=incident rate ratio. RR=relative risk. *First catheter per patient only. †Combined endpoint of phlebitis, infiltration, occlusion, accidental removal, and CRBSI. ‡Risk and p value inestimable because of 0 incidence in one or both groups. §In all cases, mortality was unrelated to intravenous catheter treatment.

Table 3: Study outcomes by treatment group (per-patient analysis)

External validity:

- <1% mortality
- CR-BSI 1/3283

 (0.03%) patients
 (1/5907 PVCs)
- Data only from Australia...





New meta-analysis

• Two new studies from China







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Comparison of Routine Replacement With Clinically Indicated Replacement of Peripheral Intravenous Catheters

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Setting, patients and PVCs:

- Observational study: prospectively collected data at the University of Geneva Hospitals (ten sites)
- Included patients and PVCs: 1 January 2016 and 29 February 2020
- Hospital-wide prospective surveillance of all healthcare-associated bloodstream infections including PVC-BSI (IPC team)
- Individual-level data on PVC: electronic health record







Definitions:

- PVC-BSI:
- Catheter-related • BSI occurring from day of insertion until 48h after catheter removal and positive culture with the same microorganism of:
 - either a quantitative PVC tip culture ≥103 CFU/ml or
 - positive superficial culture with the same microorganism from pus from insertion site

OR

Catheter-associated • A BSI was associated with a catheter if occurring from day or insertion until 48h after catheter removal, the resolution of symptoms in 48h after catheter removal and the absence of any other infectious focus.





Intervention

1st January 2016-31st March 2018: **Routine replacement** of PVCs every 96h.

BASELINE







Statistics:

- Monthly aggregated data on PVCs and PVC-days were graphically summarized (2016 – 2020)
- Incidence rate ratios [IRR] were calculated for the intervention and reversion periods [baseline period as a reference] → segmented Poisson regression models on aggregated monthly data
 - Three sensitivity analyses:
 - Excluding catheters inserted during the year 2016
 - Excluding children
 - Excluding PVCs inserted in the ICU





Flow-chart:



Buetti N. et al. JAMA Intern Med. 2021 Nov 1;181(11):1471-1478.





RESULTS Description of the study population by study period Table. Characteristics of the Study Population by Study Period^a

Table. Characteristics of the Study Popula	tion by Study Perio	odª		
Characteristic	Baseline	Intervention	Reversion	P value
Sex ^b				
Female	47 114 (54.0)	31 259 (54.4)	10 555 (54.1)	.28
Male	40 207 (46.0)	26 225 (45.6)	8971 (45.9)	
Age, median (IQR) ^b	51 (33-71)	52 (33-72)	55 (35-74)	<.001
ICU admission	7120 (2.9)	2782 (2.1)	732 (1.8)	<.001
No. of catheters per patient, median (IQR) $^{\rm c}$	1 (1-2)	1 (1-2)	1 (1-2)	<.001
Dwell time, d				
>4	26 372 (10.9)	26 656 (20.4)	5170 (12.8)	<.001
>7	5745 (2.4)	10656 (8.1)	947 (2.3)	<.001
Insertion site				
Forearm	130 877 (54.2)	50 584 (38.7)	15 276 (37.8)	<.001
Arm	6930 (2.9)	2105 (1.6)	675 (1.7)	
Elbow	12 247 (5.1)	21 508 (16.4)	7530 (18.6)	
Hand	69 615 (28.8)	30 930 (23.7)	9141 (22.6)	
Other	6018 (2.5)	2636 (2.0)	771 (1.9)	
Wrist	15 745 (6.5)	23 016 (17.6)	7027 (17.4)	
Operator				
Out-of-hospital	18 909 (7.8)	10 573 (8.1)	2786 (6.9)	<.001
In-hospital	222 523 (92.2)	120 206 (91.9)	37 634 (93.1)	
PVC-BSI	11 (<0.1)	46 (<0.1)	4 (<0.1)	<.001





RESULTS <u>Number of PVCs</u> stratified by catheter duration during the three study periods.







Monthly incidence of PVC-BSIs during the three study periods.



The incidence rate of PVC-BSI during the intervention period was 0.9 per 10'000 catheter-days, compared to 0.13 per 10'000 catheterdays during the baseline period.





IRRs of PVC-BSI during intervention and reversion periods

Source	IRR (95% CI)				P value
Main analysis					
Intervention period (clinically indicated replacement)	7.20 (3.65-14.22)				<.001
Reversion period (routine replacement)	1.35 (0.30-6.17)				.69
Sensitivity analysis excluding 2016					
Intervention period (clinically indicated replacement)	5.94 (2.69-13.11)			_	<.001
Reversion period (routine replacement)	1.12 (0.23-5.37)		-		.89
Sensitivity analysis excluding children					
Intervention period (clinically indicated replacement)	7.18 (3.64-14.18)			_	<.001
Reversion period (routine replacement)	1.35 (0.30-6.15)				.70
	0.	.1	1 IRR (95%)	10 2 CI)	י 20

Reference: baseline period

Without ICU:

- Intervention: IRR 6.81, 95% CI 3.53-13.13, p<.001
- Reversion: IRR 1.26, 95% CI 0.28-5.68, p=0.76





Microbiological etiology of PVC-BSI, stratified by routine and clinically indicated replacement periods

	Routine	Clinically indicated	p-value*
	replacement°	replacement	
Achromobacter, n (%)	0 (0)	1 (2.2)	0.64
CoNS or other skin commensals, n	9 (60)	23 (50)	
(%)			
Enterobacter spp, n (%)	1 (6.7)	3 (6.5)	
Fungi, n (%)	1 (6.7)	1 (2.2)	
Klebsiella spp, n (%)	1 (6.7)	3 (6.5)	
MRSA <i>,</i> n (%)	0 (0)	3 (6.5)	
MSSA, n (%)	1 (6.7)	7 (15.2)	
Pseudomonas aeruginosa, n (%)	U (U)	3 (6.5)	
Serratia marcescens, n (%)	1 (6.7)	0 (0)	
Sphingomonas paucimobilis, n (%)	0 (0)	1 (2.2)	
Polymicrobial, n (%)	1 (6.7)	1 (2.2)	





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CONCLUSIONS

The debate is (still) open:

- Evidence from RCTs:
 - Two trials reported 2 suspected BSI and one CRBSI, respectively
 - RCTs usually did not specifically target PVC-BSI as primary outcome (rare occurrence)
 - Despite large numbers at first glance, all RCTs were underpowered for detecting differences in PVC-BSI
 - PVC inserted in non-critical situations
 - Randomization NOT at day 3 or 4
- Large observational studies:







CONCLUSIONS

The debate is (still) open:

- According to "insertion recommendations":
 - PVC should not be inserted for long infusions

Firming 0 Manager	Floure 2. Venous access douise recommandations for infusion of paripharally compatible infusion								
Figure 3. Venous ad	rigure 3. Venous access device recommendations for infusion of peripherally compatible infusate.								
	Proposed Duration of Infusion								
Device Type	≤5 d	6-14 d	15–30 d	≥31 d					
Peripheral IV catheter	No preference between peripheral IV and US-guided peripheral IV catheters for use ≤5 d								
US-guided peripheral IV catheter	US-guided peripheral IV cathe catheter if proposed	ter preferred to peripheral IV duration Is 6–14 d							
Nontunneled/acute central venous catheter	Central venous catheter pr or If hemodynamic moni	eferred in critically III patients toring is needed for 6–14 d							
Midline catheter	Midline catheter preferred to P	ICC If proposed duration is ≤14 d							



Chopra V et al. Annals of Internal Medicine, Vol. 163 No. 6 (Supplement), 15 September 2015 // Timsit J et al. Intensive Care Med (2022) 48:1422–1425



CONCLUSIONS

The debate is (still) open:

- 1 PVC-BSI per 10'000 catheter-days \rightarrow justification for routine replacement of PVCs?
 - PVC-BSI is the rarest among many complications around vascular access.
- Routine replacement
 high number of used catheters per hospital stay
- Clinically indicated removal maybe reduced device-related costs
- Repeated insertions
 - Patient discomfort and decreased venous capital for patients
 - HCWs \rightarrow increased risks of needle-stick injuries and is time-consuming for vascular access teams

PR/

- Phlebitis: unclear new data suggest a reduction with routine replacement
- Routine replacement reduces extravasation
- Rates of catheter failure due to **blockage** were probably lower in the routine replacement IIIBANALIZATION!!! group
- CRBSI
 - CRBSI \rightarrow morbidity & mortality.
 - S. aureus infections?

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