Katie Harron Angie Wade **Berit Muller-Pebody Harvey Goldstein** Roger Parslow Jim Grav John C. Hartley **Quen Mok Ruth Gilbert**

Risk-adjusted monitoring of blood-stream infection in paediatric intensive care: a data linkage study

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K. Harron (⋈) · A. Wade · H. Goldstein · R. Gilbert MRC Centre for Epidemiology of Child Health, Institute of Child Health, University College London, 30 Guilford Street, London WC1 N 1EH, UK e-mail: k.harron@ich.ucl.ac.uk;

k.harron@ucl.ac.uk Tel.: +44-207-9052764 Fax: +44-207-9052793

B. Muller-Pebody Healthcare Associated Infection and Antimicrobial Resistance Department, Health Protection Agency, London, UK

R. Parslow Paediatric Epidemiology Group, University of Leeds, Leeds, UK

J. Grav Department of Microbiology, Birmingham Children's Hospital, Birmingham, UK

J. C. Hartley Department of Microbiology, Virology and Infection Prevention and Control, Great Ormond Street Hospital, London, UK

O Mok Paediatric Intensive Care Unit, Great Ormond Street Hospital, London, UK

Abstract *Purpose:* National monitoring of variation in the quality of infection control in paediatric intensive care units (PICUs) requires comparisons of risk-adjusted rates. To inform the development of a national monitoring system, we evaluated the effects of risk-adjustment and outcome definition on comparisons of blood-stream infection (BSI) rates in PICU, using linkage of risk-factor data captured by national audit (PIC-ANet) with laboratory records of BSI. Admission data for two Methods: children's hospitals 2003-2010 were extracted from PICANet and linked using multiple identifiers with laboratory BSI records. We calculated trends of PICU-acquired BSI, defined as BSI occurring between at least 2 days after admission until up to 2 days following discharge. In one PICU, we compared rates of all PICU-acquired BSI with clinically significant PICU-acquired BSI submitted to the national surveillance system. Results: Of 20,924

admissions, 1,428 (6.8 %) were linked to 1,761 PICU-acquired BSI episodes. The crude incidence rateratio for PICU-acquired BSI between PICUs was 1.15 [95 % confidence interval (CI) 1.05-1.26] but increased to 1.26 (1.14-1.39) after risk-adjustment. Rates of PICU-acquired BSI were 13.44 (95 % CI 12.60–14.28) per 1,000 bed-days at PICU 1 and 18.05 (95 % CI 16.80–19.32) at PICU 2. Of PICU-acquired BSI at PICU 2, 41 % was classified as clinically significant. Rates of PICU-acquired BSI decreased by 10 % per year between 2003 and 2010 for skin organisms and 8 % for non-skin organisms. Conclusions: Risk-adjustment and standardisation of outcome measures are essential for fair comparisons of BSI rates between PICUs. Linkage of risk-factor data and BSI surveillance is feasible and could allow national risk-adjusted monitoring.

Keywords Blood-stream infection · Surveillance · Data linkage · Paediatric intensive care · Risk-adjustment · Bacteraemia

Introduction

Healthcare-associated infection (HAI) is an important of nosocomial blood-stream infection (BSI) of any clini-

National Health Service (NHS), and paediatric intensive care units (PICUs) have one of the highest reported rates cause of adverse clinical outcome and cost to the UK cal specialty [1-4]. Interventions aimed at reducing HAI infection in the US and have led the way for similar interventions in the UK, including the introduction of the Department of Health Saving Lives CVC care bundle in culture with: 2005 [5-8].

Monitoring of infection outcomes is vital for determining the impact of these interventions and for sustaining improvements in practice [9, 10]. Comparison of healthcare associated BSI rates and assessment of variation of quality of care between PICUs would allow improved practices to be targeted at those most likely to benefit.

Reliable risk-adjustment is required for meaningful comparisons between PICUs [11]. At a national level, no single dataset accurately captures both clinical and microbiological BSI data [12]. LabBase2 is a national voluntary microbiological surveillance database collated by the Health Protection Agency (HPA), based on clinically significant isolates identified at laboratories serving NHS hospitals [13]. The Paediatric Intensive Care Audit Network (PICANet) collects data on all children admitted to PICUs in Britain, including clinical details such as diagnosis and morbidity score on admission [14]. Linkage of these existing datasets offers an efficient method for providing risk-adjusted national surveillance.

To demonstrate the effect of risk-adjustment using routinely collected data and the importance of standardised outcome measures, we linked PICANet data with microbiology data from two large UK PICUs, to assess reductions in BSI acquired within PICU.

Methods

Data and case definition

Admission data for all children admitted to Birmingham Children's Hospital (BCH) PICU (General and Cardiac) and Great Ormond Street Hospital (GOSH) PICU (General and Cardiac) between March 2003 and December 2010 were extracted from the PICANet database. Variables extracted for analysis were admission and discharge dates, Paediatric Index of Mortality (PIM2) score at admission, age, sex, admission type (planned or unplanned), admission source (same hospital or elsewhere), primary diagnosis at admission [based on primary diagnosis coded with the International Classification of Disease (ICD10)], and variables indicating renal support and mechanical ventilation [15].

BCH and GOSH laboratories routinely record microbiology data for all bacterial isolates from blood culture. From these laboratory systems, we extracted data on all positive isolates for children admitted to hospital between

have succeeded in dramatically decreasing rates of March 2003–2010, regardless of whether they were considered contaminants or clinically significant.

We defined an episode of BSI as any positive blood

- 1. One or more organisms isolated from any blood sample taken on the same day
- Repeated samples with positive cultures of the same organism within 14 days

A child could have more than one BSI episode during an admission if different organisms were isolated on different days, or if the same organism was isolated on more than 1 day (>14 days apart).

We defined PICU-acquired BSI as any BSI from samples taken up to 2 days after discharge from PICU, but excluding samples taken on the day of admission or the day after admission to PICU. As laboratory records included the date but not the time the specimen was taken, some included samples may have been taken between 24 and 48 h after admission to PICU, thereby possibly resulting in a slight overestimation of rates. Laboratory data after discharge from PICU was available for the majority of children (81 %) as they were discharged to another ward within the same hospital.

We separately analysed PICU-acquired BSI due to skin and non-skin organisms, as we hypothesised that infection control practices might have more impact on BSI due to skin organisms, which can be acquired during invasive procedures or sample contamination [16, 17]. Skin organisms were defined as coagulase negative staphylococcus, Staphylococcus epidermidis, Corynebacterium spp. or Proprionibacterium spp. All other organisms were classified as non-skin organisms. PICUacquired BSI episodes could include either skin organisms, non-skin organisms, or both, and so the number of episodes of PICU-acquired BSI due to skin organisms and due to non-skin organisms does not add up to the total number of episodes. We separately analysed PICUacquired BSI in children staying at least 48 h to account for any differences in the number of very short-term admissions (with a lower risk of infection) between PICUs.

We assessed the effect of different case definitions for BSI reporting. The national voluntary surveillance system LabBase2, coordinated by the HPA, requires reporting of clinically significant BSI. At one of the two laboratories, an additional variable indicated positive BSI that had been considered to be clinically significant and were reported to LabBase2, based on prospectively recorded consultant microbiologist opinion following regular meetings with the clinical team and with reference to the CDC/NHSN definition of laboratory-confirmed bloodstream infection [18]. At the other laboratory, no classification of clinically significant BSI was available.

Data linkage

We generated our dataset by deterministic linkage of PICANet and microbiology records based on unique identifiers (NHS number, hospital number, name, date of birth and sex). Linkage was manually verified to ensure there were no false matches or missed matches.

Statistical analysis

PICU-acquired BSI rates were calculated as the number of PICU-acquired BSI per 1,000 bed-days. Because of the low event rate, we assumed a log-linear trend and fitted Poisson regression models¹ to the data, with length of stay (in hours) as the exposure variable. Model fits were compared using Akaike's Information Criterion, goodness of fit tests and likelihood-ratio tests for nested models. Models with best-fit were used to identify significant riskfactors for PICU-acquired BSI from the patient characteristics extracted from PICANet (Table 1). For these models, the quarter-year of admission, age in months, and PIM2 score were treated as continuous variables. Quarter of year (3-month calendar period), sex, admission type, admission source, renal support, mechanical ventilation and diagnosis group (cardiovascular, respiratory, infection or other) were categorical variables. We aimed to evaluate how relatively simple risk-adjustment affects comparative measures rather than to inform direct comparisons between PICUs, as BSI risk-factors were limited within our data. A PICU-year interaction term was added to the model to test for differences in trends between PICUs. The two PICUs were randomly coded as PICU 1 and PICU 2 so that individual units were not identified (same coding used throughout analysis) and to deter erroneous conclusions about relative performance.

Incidence rate ratios (IRR) with 95 % confidence intervals (CI) were derived from models of best fit and used to compare rates between PICUs. Risk-adjusted rates were derived for each PICU by adjusting for all risk-factors found to be significant in the Poisson regression. Kernel density estimates of rates were estimated for graphical display.

The time to infection was calculated as the number of days between admission and the date of PICU-acquired BSI. Cox proportional hazards models were fitted to the data to estimate the survival function for each PICU and to identify significant risk-factors for time to infection.

Analysis was performed using Stata 11 [19].

Table 1 PICANet data: characteristics of all children admitted to two PICUs in England, 2003–2010

	n admissions	%	
Length of stay			
<48 h	9,059	43.29	
>=48 h	11,865	56.71	
Year of admission			
2003	2,361	11.28	
2004	2,762	13.20	
2005	2,423	11.58	
2006	2,647	12.65	
2007	2,638	12.61	
2008	2,664	12.73	
2009	2,936	14.03	
2010	2,493	11.91	
Quarter of year	,		
Jan–Mar	4,833	23.10	
Apr–Jun	5,425	25.93	
Jul-Sep	5,564	26.59	
Oct–Dec	5,102	24.38	
PICU	-,		
PICU 1	12,310	58.83	
PICU 2	8,614	41.17	
Admission type	-,		
Planned	8,583	41.02	
Unplanned	12,341	58.98	
Admission source	12,5 .1	20.50	
Same hospital	12,772	61.04	
Elsewhere	8,152	38.96	
Diagnosis group	-,		
Other	7,163	34.23	
Cardiovascular	8,493	40.59	
Respiratory	4,600	21.98	
Infection	668	3.19	
Renal support		0.17	
No	20,081	95.97	
Yes	843	4.03	
Mechanical ventilation			
No	16,195	77.4	
Yes	4,729	22.6	
Sex	.,.=>	22.0	
Female	11,813	56.46	
Male	9,110	43.54	
Unknown	1	0.00	
Age group	-	0.00	
<1 year	11,347	54.23	
1–4 years	4,839	23.13	
5–10 years	2,611	12.48	
11–15 years	2,127	10.17	
Total	20,924	10.17	

Results

There were 13,376 records of bacterial isolates from blood culture dating between March 2003 and December 2010, inclusive, extracted from the microbiology systems at the two hospitals (n = 5,589 PICU 1, n = 7,787 PICU 2; hospital-wide). These corresponded to 6,842 episodes of BSI as defined in the case definition (hospital-wide).

Records for 20,924 admissions were extracted from the PICANet database for the two PICUs over the same time period, comprising a total of 117,800 bed-days (approximately 15,000 bed-days or 2,600 admissions per

¹Full model: $log(infections) = \beta_1 \times quarter$ -year of admission + $\beta_2 \times PICU + \beta_3 \times admission$ type + $\beta_4 \times diagnosis$ group + $\beta_5 \times renal$ support + $\beta_6 \times sex + \beta_7 \times age + \beta_8 \times PIM2 + \beta_9 \times admission$ source + $\beta_{10} \times mechanical$ ventilation + $\beta_{11} \times quarter$ of year + log(length of stay).

characteristics of the children admitted to the two PICUs between 2003 and 2010.

PICU-acquired BSI

A total of 1.761 PICU-acquired BSI episodes were linked to 1,428 admissions. Of these, 144 (8.2 %) BSI episodes included more than one type of organism isolated on the same day. Of these, 220 (15 %) admissions had multiple PICU-acquired BSI episodes during their stay in PICU.

The rate of PICU-acquired BSI per 1,000 bed-days was 15.17 (14.45–15.86) per 1,000 bed-days (Table 2). Quarter-year of admission, quarter of calendar year, age, admission type, admission source, renal support, PICU and diagnosis group were found to be significant at the 5 % level, but there was no evidence of an effect of sex. mechanical ventilation or PIM2 score on the rate of PICU-acquired BSI.

The crude IRR underestimated the difference between PICUs (1.15; 95 % CI 1.05-1.26) and the IRR increased to 1.26 (95 % CI 1.14-1.39) after adjusting for differences in significant risk-factors between PICUs. Although the median length of stay differed by PICU (70 and 48 h in PICUs 1 and 2 respectively), the exclusion of shortterm admissions (<48 h, n episodes = 119) did not account for the difference between PICUs. The adjusted PICU IRR for admissions at least 48 h was 1.20 (95 % CI 1.08–1.33). The median time to PICU-acquired infection was 7 days from admission (95 % binomial CI 6–7). For any given time point, a greater proportion of children in PICU 2 had PICU-acquired BSI than in PICU 1, after adjusting for significant risk-factors [mechanical ventilation, age, PIM2, renal status, admission type, admission source, diagnosis group and quarter-year at admission (Fig. 1)]. Rates were lower in summer months.

The rate of PICU-acquired BSI decreased by 9 % (95 % CI 7–11 %) each year during the study period. This corresponded to a 44 % reduction (a reduction of 7.54 per 1,000 bed-days) in the rate of PICU-acquired BSI between 2003 and 2010. The percentage of admissions each year with PICU-acquired BSI decreased by 45 % over the study period (from 8.3 to 4.5 % of admissions). There was no evidence of a difference in trends between PICUs (p value = 0.43 for trend-PICU interaction).

Skin organisms and non-skin organisms

The rate of PICU-acquired BSI due to skin organisms was higher than the rate of PICU-acquired BSI due to non-skin organisms, and risk-factors for skin organisms differed to those for non-skin organisms (Table 3).

Rates of PICU-acquired BSI due to both skin organisms and due to non-skin organisms decreased between

year at the two PICUs combined). Table 1 shows the Table 2 Rate and risk-adjusted incidence rate ratios (IRRs) for PICUacquired BSI

	PICU-acquired BSI				
Rate (95 % CI) per 1,000 bed-days		15.17 (14.45–15.86)			
% (95 % CI) of admissions with BSI		6.82 % (6.48–7.17 %)			
	n episodes	Incidence rate ratio (95 % CI)	p value		
Year of admission		0.91 (0.89-0.93)	< 0.001		
Quarter of year					
Jan–Mar	472	1			
Apr–June	429	0.83 (0.73-0.95)			
Jul-Sep	410	0.83 (0.73-0.95)			
Oct-Dec	450	0.96 (0.85-1.10)	0.005		
Age (months)		0.99 (0.99-0.99)	< 0.001		
PICU					
PICU 1	976	1			
PICU 2	785	1.26 (1.14–1.39)	< 0.001		
Admission type					
Planned	448	1			
Unplanned	1313	1.45 (1.28–1.64)	0.001		
Admission source					
Same hospital	1,007	1			
Elsewhere	754	0.72 (0.65–0.80)	< 0.001		
Diagnosis group					
Other	632	1			
Cardiovascular	601	0.81 (0.72–0.92)			
Respiratory	441	0.77 (0.68–0.87)			
Infection	87	1.26 (1.01–1.59)	< 0.001		
Renal support					
No	1,482	1			
Yes	279	1.47 (1.29–1.68)	< 0.001		
Total	1,761				

IRRs were adjusted for all significant risk-factors listed in the table and the IRR for year of admission was converted from the IRR for quarter-year of admission

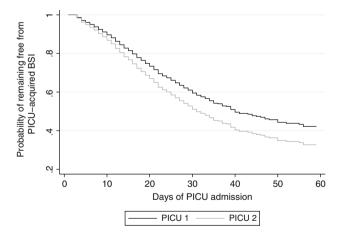


Fig. 1 Probability of remaining free from PICU-acquired BSI in first 60 days of admission by PICU, adjusted for mechanical ventilation, age, PIM2, renal status, admission type, admission source, diagnosis group and quarter-year at admission

2003 and 2010. As expected, there was a larger reduction seen in PICU-acquired BSI due to skin organisms (a 47 % reduction of 5.00 per 1,000 bed-days) compared with the decline due to non-skin organisms (a 34 % reduction of 2.64 per 1,000 bed-days). The percentage of admissions per year with PICU-acquired BSI due to skin organisms decreased by 52 % (from 5.8 to 2.81) compared with 41 % (from 3.6 to 2.1) due to non-skin organisms.

The difference between PICUs was due to differences in rates of non-skin organisms—there was no evidence of a difference in rates of skin organisms between PICUs (IRR = 1.10; 95 % CI 0.97-1.25). The crude IRR for non-skin organisms underestimated the difference between PICUs (1.38; 95 % CI 1.21-1.57), rising to 1.62 (95 % CI 1.41-1.85) after adjusting for differences in population at risk between PICUs. Figure 2 shows the risk-adjusted rate of PICU-acquired BSI due to skin overall PICU-acquired BSI at PICU 2. organisms and non-skin organisms in each PICU.

Clinically significant BSI

321 (41 %) were classified as clinically significant. This was largely due the number of coagulase negative staphylococcus classified as not clinically significant—of the 445 PICU-acquired BSI due to skin organisms, only 155 (34.5 %) were clinically significant. However, a reduction in the number of non-skin organisms was also seen—of the 419 PICU-acquired BSI due to non-skin organisms, 284 (67.8 %) were clinically significant.

The rate of clinically significant BSI was 7.39 (95 % CI 6.58-8.18) per 1,000 bed-days, compared with an overall rate of 18.05 (95 % CI 16.80-19.32) for all PICU-acquired BSI at PICU 2 and 13.44 (95 % CI 12.60–14.28) at PICU 1.

Figure 3 shows the rate of clinically significant and

Discussion

Classification of BSI as clinically significant or not, based Our study demonstrates the importance of risk-adjustment on prospectively recorded clinical opinion, was available and standardised outcome measures for monitoring of for PICU 2. Of the 785 PICU-acquired BSI in this PICU, changes in BSI rates within and between PICUs. We show

Table 3 Rates and risk-adjusted incidence rate ratios (IRRs) of PICU-acquired BSI due to skin organisms and non-skin organisms

Rate (95 % CI) per 1,000 bed-days % (95 % CI) of admissions with >=1 specimen		sms		Non-skin organisms		
		9.29 (8.74–9.84) 4.62 % (4.34–4.92 %)		7.37 (6.86–7.85) 3.06 % (2.83–3.30 %)		
						n
	0.90 (0.87-0.93)	< 0.001		0.92 (0.89-0.94)	< 0.001	
290	1		227			
266	0.83 (0.71–0.99)		212	Not significant	0.230	
250	0.82 (0.69-0.97)		200			
272	0.96 (0.81–1.14)	0.0457	216			
	0.99 (0.99-0.99)	< 0.001		0.99 (0.99-0.99)	< 0.001	
595	1		513	1		
483	0.75 (0.66–0.86)	< 0.001	342	0.70 (0.60-0.81)	< 0.001	
				· · ·		
633	1		436	1		
445	1.10 (0.97–1.25)	0.134	419	1.51 (1.31–1.74)	< 0.001	
	(,			,		
272	1		229	1		
806	1.39 (1.20–1.63)	< 0.001	626	1.25 (1.04–1.51)	0.015	
	,			,		
406	1		289	1		
	0.70 (0.60–0.81)			0.87 (0.73–1.04)		
	` /	< 0.001		,	< 0.001	
	0.75 (0.62 1.07)	10.001	01	2.00 (1.02 2.00)	10.001	
921	1		709	1		
	•	< 0.001		•	0.001	
137	1.12 (1.17 1.00)	<0.001	110	1.25 (1.20 1.01)	0.001	
192	Not significant		180	1		
	110t Significant			-	0.002	
000	Not significant		013		0.002	
1.078	110t Significant		855	1.07 (1.00-2.57)	0.021	
	290 266 250 272 595 483 633 445 272	## A.62 % (4.34–4.92 %) ## Incidence rate ratio (95 % CI) ## CID ## O.90 (0.87–0.93) ## CID ## O.90 (0.87–0.93) ## CID ## O.90 (0.87–0.99) ## CID ## O.83 (0.71–0.99) ## CID ## O.83 (0.71–0.99) ## CID ## O.84 (0.69–0.97) ## CID ## O.99 (0.99–0.99) ## CID ## CID ## O.75 (0.66–0.86) ## CID ## CID ## O.75 (0.60–0.81) ## O.70 (0.60–0.81) ## O.75 (0.52–1.07) ## O.75	9.29 (8.74–9.84) 4.62 % (4.34–4.92 %) Incidence rate ratio p value (95 % CI) 0.90 (0.87–0.93) <0.001 290 1	9.29 (8.74–9.84) 4.62 % (4.34–4.92 %) Incidence rate ratio (95 % CI) 0.90 (0.87–0.93) 200 1 227 266 0.83 (0.71–0.99) 212 250 0.82 (0.69–0.97) 272 0.96 (0.81–1.14) 0.0457 216 0.99 (0.99–0.99) 301 595 1 483 0.75 (0.66–0.86) 310 436 445 1.10 (0.97–1.25) 272 1 806 1.39 (1.20–1.63) 289 338 0.70 (0.60–0.81) 301 0.81 (0.70–0.94) 33 0.75 (0.52–1.07) 310 709 157 1.42 (1.19–1.68) Not significant 180 886 Not significant 180 180	9.29 (8.74–9.84) 7.37 (6.86–7.85) 4.62 % (4.34–4.92 %) 3.06 % (2.83–3.30 %) n Incidence rate ratio (95 % CI) 0.90 (0.87–0.93) 200 1 227 266 0.83 (0.71–0.99) 212 Not significant 250 0.82 (0.69–0.97) 200 272 0.96 (0.81–1.14) 0.0457 216 0.99 (0.99–0.99) 255 1 513 1 0.99 (0.99–0.99) 595 1 483 0.75 (0.66–0.86) <0.001 342 0.70 (0.60–0.81) 633 1 436 1 445 1.10 (0.97–1.25) 0.134 419 1.51 (1.31–1.74) 272 1 806 1.39 (1.20–1.63) <0.001 626 1.25 (1.04–1.51) 406 1 289 1 338 0.70 (0.60–0.81) 310 0.87 (0.73–1.04) 301 0.81 (0.70–0.94) 195 0.75 (0.62–0.90) 33 0.75 (0.52–1.07) <0.001 61 2.00 (1.52–2.66) 921 1 709 1 157 1.42 (1.19–1.68) <0.001 146 1.53 (1.28–1.84) Not significant 180 1 Not significant 180 1.67 (1.08–2.57)	

IRRs were adjusted for all significant risk-factors listed in the table

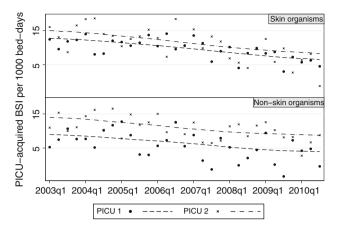


Fig. 2 Rates of PICU-acquired BSI due to skin organisms and non-skin organisms by PICU and quarter-year at admission, adjusted for quarter of year, admission type, admission source, age, renal support and diagnosis group. *Symbols* quarterly crude rates, *Lines* Kernel density estimate of risk-adjusted trends

that simple risk-adjustment can be achieved through linkage of routinely collected data. As variation between units is a key measure for monitoring the quality of care in PICU, these potential sources of bias need to be taken into account to ensure fair and accurate assessment of variation in outcomes [20].

This study spans a period when national and local guidance led to major changes in infection control and a near halving in rates of BSI. The dramatic reduction in PICU-acquired BSI rates seen from 2007, particularly in BSI due to skin organisms, corresponds to the introduction of the Saving Lives CVC care bundle [8]. These guidelines for insertion and maintenance of CVCs to prevent BSI led to the introduction of 2 % chlorhexidine swabs at GOSH and infection control audits of hand

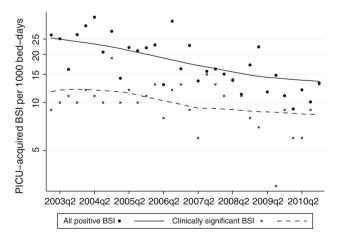


Fig. 3 Rates of clinically significant and all PICU-acquired BSI at PICU 2 by quarter-year at admission, adjusted for quarter of year, admission type, admission source, age, renal support and diagnosis group. *Symbols* quarterly crude rates *Lines* Kernel density estimate of risk-adjusted rates

hygiene and high impact interventions consistently scoring >90 % in BCH. This is the first study to link microbiology data with hospital admissions and to quantify this reduction in PICUs, though such changes are well recognised by clinicians [21].

Although there is broad recognition of the importance of risk-adjustment, current PICU risk-adjustment scores for mortality (PIM2) or indicators of severity of illness (renal support, mechanical ventilation) do not accurately capture all risk-factors for BSI. The overall difference in rates between PICUs did not appear to be due to skinorganisms, which are likely to correlate most closely with infection control practices. The variation in rates of PICU-acquired BSI due to pathogens may be due to differences in PICU casemix that are not routinely captured. More detailed risk-factor information may further explain variation between PICUs and improvements in routine data quality and methodology will allow a wider range of risk-factors to be explored. Use of information on trajectories of care preceding PICU admission using longitudinal hospital administrative data may enhance risk-adjustment [22–25].

We identified large differences between rates of total BSI and clinically significant BSI for both skin and nonskin organisms. Although definitions for classification of healthcare-associated BSI exist, there are no clear criteria to guide clinical judgement on classification of clinically significant positive isolates that are reported to the national surveillance system LabBase2 [26-28]. Clinicians need to make daily judgements for the care and treatment of patients, but national monitoring requires objective outcome measures to achieve fair comparisons. Automated downloads of laboratory data to the voluntary surveillance system, introduced in recent years, offer the opportunity to capture all positive BSI data. Analyses based on these data would overestimate the total burden of clinically significant BSI acquired in PICU, as in our study. However, it is also important to monitor contamination, as this may lead to overtreatment and possible increased length of stay in PICU. Measuring all positive BSI would ensure valid comparisons, undistorted by variations in clinical opinion on clinical significance.

The success of risk-adjusted monitoring on a national scale will depend on a linkage infrastructure that ensures data quality and minimises error due to linkage of imperfect or incomplete identifiers, while allowing for standardised, risk-adjusted analyses of infection rates. Our analyses quantify substantial reductions in BSI rates in two PICUS, but a national system is required to identify units where further improvements are needed and to determine which practice innovations have most impact.

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Service Division NHS Scotland, the Royal Belfast Hospital for Sick Children, Our Lady's Children's Hospital, Crumlin, Children's University Hospital, Temple Street and The Harley Street Clinic, London.

Ethical approval Collection of personally identifiable data has been approved by the National Information Governance Board (Formerly the Patient Information Advisory Group) http://www.nigb.nhs.uk/s251/registerapp and ethical approval granted by the Trent Medical Research Ethics Committee, ref. 05/MRE04/17. Consent for the use of the data in this study was obtained by the PICANet unit leads.

References

- Lakshmi KS, Jayashree M, Singhi S, Ray P (2007) Study of nosocomial primary bloodstream infections in a pediatric intensive care unit. J Trop Pediatr 53(2):87–92. doi: 10.1093/tropej/fml073
- Abou Elella Ř, Najm H, Balkhy H, Bullard L, Kabbani M (2010) Impact of bloodstream infection on the outcome of children undergoing cardiac surgery. Pediatr Cardiol 31(4):483–489. doi: 10.1007/s00246-009-9624-x
- 3. Elward A, Hollenbeak C, Warren D, Fraser V (2005) Attributable cost of nosocomial primary bloodstream infection in pediatric intensive care unit patients. Pediatrics 115(4):868
- 4. Yogaraj J, Elward A, Fraser V (2002) Rate, risk factors, and outcomes of nosocomial primary bloodstream infection in pediatric intensive care unit patients. Pediatrics 110(3):481
- Pronovost P, Goeschel C, Colantuoni E, Watson S, Lubomski L, Berenholtz S, Thompson D, Sinopoli D, Cosgrove S, Sexton J (2010) Sustaining reductions in catheter related bloodstream infections in Michigan intensive care units: observational study. BMJ 340:c309
- Bhutta A, Gilliam C, Honeycutt M, Schexnayder S, Green J, Moss M, Anand K (2007) Reduction of bloodstream infections associated with catheters in paediatric intensive care unit: stepwise approach. BMJ 334(7589):362–365
- 7. Miller MR, Griswold M, Harris JM II, Yenokyan G, Huskins WC, Moss M, Rice TB, Ridling D, Campbell D, Margolis P, Muething S, Brilli RJ (2010) Decreasing PICU catheter-associated bloodstream infections: nACHRI's quality transformation efforts. Pediatrics 125(2):206–213. doi: 10.1542/peds.2009-1382

- 8. Department of Health (2005) Saving Lives: a delivery programme to reduce healthcare associated infections including MRSA. Department of Health, London
- Pronovost PJ, Berenholtz SM, Needham DM (2008) Translating evidence into practice: a model for large scale knowledge translation. BMJ 337:a1714. doi:10.1136/bmj.a1714
- Lilford R, Mohammed MA, Spiegelhalter D, Thomson R (2004) Use and misuse of process and outcome data in managing performance of acute medical care: avoiding institutional stigma. Lancet 363(9415):1147–1154
- Davenport RJ, Dennis MS, Warlow CP (1996) Effect of correcting outcome data for case mix: an example from stroke medicine. BMJ 312(7045):1503–1505
- García Álvarez L, Aylin P, Tian J, King C, Catchpole M, Hassall S, Whittaker-Axon K, Holmes A (2011) Data linkage between existing healthcare databases to support hospital epidemiology. J Hosp Infect 79(3):231–235
- Lamagni T, Potz N, Powell D, Pebody R, Wilson J, Duckworth G (2011) Mortality in patients with meticillinresistant Staphylococcus aureus bacteraemia, England 2004–2005. J Hosp Infect 77(1):16–20
- 14. Universities of Leeds and Leicester (2012) Paediatric Intensive Care Audit Network National Report 2009–2011. ISBN 9780853163121 Available: http://wwwpicanetorguk/Documents/ General/Annual_Report_2012/ Ninth_PICANet_Annual_Report_ 2009_2011_Summary_Reportpdf Accessed 19 September 2012
- 15. Slater A, Shann F, Pearson G (2003) PIM2: a revised version of the Paediatric Index of Mortality. Intens Care Med 29(2):278–285

- Safdar N, Maki D (2004) The pathogenesis of catheter-related bloodstream infection with noncuffed short-term central venous catheters. Intens Care Med 30(1):62–67. doi: 10.1007/s00134-003-2045-z
- O'Grady N, Alexander M, Dellinger E, Gerberding J, Heard S, Maki D, Masur H, McCormick R, Mermel L, Pearson M (2002) Guidelines for the prevention of intravascular catheter-related infections. Am Acad Pediatr Policy 110(5):e51–e74
- Horan TC, Andrus M, Dudeck MA (2008) CDC/NHSN surveillance definition of health care—associated infection and criteria for specific types of infections in the acute care setting. Am J Infect Control 36(5):309–332
- StataCorp (2009) Stata statistical software: release 11. StataCorp LP, College Station
- Shann F, Pearson G, Slater A, Wilkinson K (1997) Paediatric index of mortality (PIM): a mortality prediction model for children in intensive care. Intens Care Med 23(2):201–207
- Harron K, Ramachandra G, Mok Q, Gilbert R (2011) Consistency between guidelines and reported practice for reducing the risk of catheter-related infection in British paediatric intensive care units. Intens Care Med 37(10):1641–1647
- 22. Holman CDAJ, Preen DB, Baynham NJ, Finn JC, Semmens JB (2005) A multipurpose comorbidity scoring system performed better than the Charlson index. J Clin Epidemiol 58(10):1006–1014
- 23. Armitage J, van der Meulen J (2010) Identifying co-morbidity in surgical patients using administrative data with the Royal College of Surgeons Charlson Score. Brit J Surg 97(5):772–781

- Quan H, Sundararajan V, Halfon P, Fong A, Burnand B, Luthi JC, Saunders LD, Beck CA, Feasby TE, Ghali WA (2005) Coding algorithms for defining comorbidities in ICD-9-CM and ICD-10 administrative data. Med Care Nov 43(11):1130–1139
- Preen DB, Holman CDAJ, Spilsbury K, Semmens JB, Brameld KJ (2006)
 Length of comorbidity lookback period affected regression model performance of administrative health data. J Clin Epidemiol 59(9):940–946
- 26. Centers for Disease Control and Prevention (2012) Central Line-Associated Bloodstream Infection (CLABSI) event: guidelines and procedures for monitoring CLABSI. National Healthcare Safety Network (NHSN) manual Atlanta, GA: NHSN (http://www.cdc.gov/nhsn/PDFs/pscManual/4PSC_CLABScurrent.pdf)
- 27. Laupland KB, Gregson DB, Vanderkooi OG, Ross T, Kellner JD (2009) The changing burden of pediatric bloodstream infections in Calgary, Canada, 2000–2006. Pediatr Infect Dis J 28(2):114
- 28. Kollef MH, Zilberberg MD, Shorr AF, Vo L, Schein J, Micek ST, Kim M (2011) Epidemiology, microbiology and outcomes of healthcare-associated and community-acquired bacteremia: a multicenter cohort study. J Infection 62(2):130–135