Characteristics and outcome of long-stay patients in a paediatric intensive care unit in Cape Town, South Africa

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Background. Paediatric intensive care is a costly, specialised and limited resource in low- and middle-income countries. The implications of extended paediatric intensive care unit (PICU) stay in South Africa (SA) are not known.

Objectives. To describe the characteristics, outcomes and resource consumption of long-stay patients (LSPs) and to identify predictive factors for long PICU stay.

Methods. A retrospective review of routinely collected data on all children admitted to an SA PICU over one calendar year. Long PICU stay was defined statistically as >19 days. Long- and short-stay patient (SSP) groups were compared, and variables significantly associated with long stay on univariate analysis were entered into a stepwise multiple regression model.

Results. Over the study period, 1 126 children (median age 8 months, 60.9% male) were admitted to the PICU, occupying 5 936 bed-days; 54 LSPs (4.8%) utilised 1 807 (30.4%) bed-days. Mortality and the standardised mortality ratio (actual/mean predicted mortality) in LSPs and SSPs were 29.6% v. 12% (p=0.002) and 2.4 v. 0.7 (p=0.002), respectively. Median duration of stay for LSPs and SSPs was 29.5 days and 2 days, respectively (p<0.0001). LSPs were younger than SSPs (median 4 months (interquartile range 2 - 17) v. 9 months (2 - 34); p=0.03), and fewer were male (48% v. 61.6%, p=0.049). On multivariate analysis, only female gender was independently associated with long PICU stay. Conclusions. LSPs represent a small proportion of PICU admissions, yet have a higher mortality rate than SSPs and consume disproportionate PICU resources. No predictive model could be established for early recognition of potential LSPs to plan PICU bed allocation effectively.

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Paediatric intensive care is a costly, specialised and limited resource that should be used as effectively and efficiently as possible. Paediatric intensive care unit (PICU) costs can range from about USD70 to over >USD1 000 per patient day, depending on the sophistication and complexity of services offered.[1] In the context of low- and middle-income countries (LMICs), where there is often limited access to PICU care, it is particularly important to ensure that resources are used rationally, while upholding the ethical principle of distributive justice. Rational PICU use includes careful monitoring of patient outcomes relative to resource utilisation and applying appropriate admission criteria to contain financial, staffing and social costs.[2]

Intensive care is now offered to children with complex and chronic conditions who may not have been admitted to PICUs in the past. [3] Advances in PICU care have resulted in increased survival of critically ill patients, some of whom require long-term PICU stay. Compared with short-stay patients (SSPs), long-stay patients (LSPs) require a different spectrum of resource allocation, with greater involvement of the extended multidisciplinary team, to improve their PICU experience and optimise their functional outcomes. This constitutes a challenge in terms of staffing needs, costs, and the PICU system, structure and function.[4]

It has been suggested that patients who have a long duration of PICU stay use a disproportionate amount of resources compared with SSPs. Together with the perceived vulnerability of LSPs to increased mortality and morbidity, there is a concern about appropriate resource allocation to this group. There is a paucity of outcome data for LSPs, especially from Africa. Existing outcome data from other settings describe a wide spectrum of LSPs, with outcomes ranging

from excellent to poor. [3,5] On an individual level, the expected outcome for a child with Guillaine-Barré syndrome, for example, will differ from that of a child with end-stage Duchenne's muscular dystrophy, although both may be LSPs in the PICU.

While it is accepted that length of stay (LOS) may reflect severity of illness and PICU quality and performance, there is no uniform definition of what constitutes long PICU stay. Previous studies have defined LSPs as having a PICU LOS of anywhere beyond 7 days to >30 days, [6-8] with various methods used to identify these thresholds.

It is useful to describe the characteristics, impact and outcomes of LSPs in order to plan their care more economically, ensure optimal involvement of the multidisciplinary team, optimise patient outcome, enable appropriate counselling of family members, and develop guidelines for limitation or withdrawal of medical care.[2] Most existing literature on LSPs in PICUs report findings from developed countries, which may have different population and burden of disease profiles from developing countries.

Objectives

To determine an appropriate threshold for defining long PICU stay in a South African (SA) PICU, to describe the characteristics and outcomes of these LSPs, and to determine any predictive factors associated with long PICU stay.

Methods

Study design and setting

The setting was the 22-bed multidisciplinary PICU at Red Cross War Memorial Children's Hospital, Cape Town, SA. This PICU admits ~1 400 children per annum, of whom ~500 are emergency

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admissions, mostly for the management of infectious diseases, and the majority require invasive mechanical ventilation. Extracorporeal membrane circulation was not available at the time of the study. PICU beds are in great demand and there is pressure to maintain rapid patient turnover. Long-term invasive and non-invasive ventilation is available for stable patients not requiring PICU-level care in other wards in the hospital.

This was a retrospective descriptive study of all children admitted to the PICU over one calendar year. Clinical data were extracted from a pre-existing PICU database.

Permission to conduct the study was obtained from the institutional human research ethics committee (ref. no. 105/2011), and the need for informed consent was waived owing to the retrospective nature of the study.

Statistical analysis

The definition of long PICU LOS in our setting was modelled using the following techniques, described by Weissman:^[7] (i) for normally distributed data, two standard deviations (SDs) above the mean LOS; (ii) five times the median LOS; (iii) beyond the 75th and 95% percentile of the median LOS; and (iv) visual examination of the frequency distribution graph of LOS v. number of admissions. A specific LOS was identified from the start of the 'tail' of the distribution curve (if not normally distributed). The bestfit model for our data was then chosen to distinguish between LSPs and SSPs.

Data were tested for normality using the Shapiro-Wilks test. Data were not normally distributed and are therefore presented throughout this article as medians (interquartile ranges, IQRs) for continuous data and proportions for categorical data. The characteristics and outcomes of the long-stay group were compared with those of the short-stay group using the Mann-Whitney *U*-test and the χ^2 or Yates corrected χ^2 test as appropriate.

Variables significantly associated with the binary outcome of interest (LSPs) on univariate analysis were entered into a forward, stepwise, logistic regression model to determine independent predictive factors for long stay, with the intention of developing a clinical predictive algorithm.

The proportion of ICU days consumed by LSPs was calculated to determine resource allocation.

Statistical analyses were conducted using Statistica version 11 (StatSoft, USA) and a significance level of *p*<0.05 was chosen.

Results

Over the study period, 1 126 children (median age 8 months (IQR 2 -32), 60.9% male), occupying 5 936 bed days, were admitted to the PICU. Baseline patient data are presented in Table 1.

Defining the LSP

Considering that the data were not normally distributed, it was not considered appropriate to identify long stay as two SDs above the mean LOS.[7]

The model using the >75th percentile of the median LOS corresponded to PICU LOS of 6 days. Two hundred and fifty-three patients (22.5%) fulfilled this criterion, with a median LOS of 11 days (IQR 8 - 17) and a 14.0% mortality rate.

When using five times the median PICU LOS (>15 days) for identification of LSPs, 76 (6.7%) of patients fulfilled the definition, with a median LOS of 25.5 days (IQR 18.5 - 35.5) and 30.0%

Modelling long stay using the >95th percentile of the median LOS was found to be equivalent to the visual examination of the start of the 'tail' of the distribution curve (Fig. 1), with 54 patients (4.8%) spending ≥20 days in the PICU, with a median LOS of 29.5 days (IQR 25 - 40) and a 29.6% mortality rate.

We therefore chose to define LSPs as those having a duration of stay of >19 days according to the two best-fit models.

LSP characteristics and outcomes

In comparison with SSPs, LSPs were significantly younger, and greater proportions were female and admitted for emergency care. The main reason for admission of LSPs was management of paediatric illness, including sepsis, pneumonia and congenital heart disease, whereas SSPs were more commonly admitted for postoperative care, traumarelated injury and poisoning (Tables 1 and 2).

LSPs had significantly higher mortality and a significantly higher standardised mortality ratio (actual/mean predicted mortality using the Paediatric Index of Mortality Score version 2 (PIM2) compared with SSPs (Table 1). There were no significant differences between LSPs who died and those who survived (Table 3).

In the final multiple regression model, only female gender was independently associated with the outcome of long stay, making it impossible to develop a predictive model for LSPs (Table 4).

Resource utilisation

LSPs comprised 4.8% of the PICU population yet utilised 30.4% (1 807) of the total PICU bed days during the study period (Fig. 2).

Discussion

This study presents data from an SA setting that support international reports that LSPs are a small but consistent proportion of the PICU population who utilise substantial PICU resources yet have worse outcomes than other PICU patients.[2,3,6]

Although LSPs are increasingly acknowledged as an integral part of PICU planning and care, they have been a difficult group to prospectively identify and investigate, [6] mainly owing to the lack of a uniform definition of what constitutes an LSP. A particular LOS may be designated as a threshold to identify the LSP, but this threshold varies widely in the literature. [3,6-8] We established a rational basis for defining long PICU stay in our context, and identified significant differences between our short- and long-stay populations.

Similar to previous studies,^[6,9] we used the >95th percentile of the median PICU LOS to define LSPs, which also corresponded to the start of the tail of the frequency distribution curve. This duration of >19 days was also reported by Jeena et al.[11] in another SA PICU over

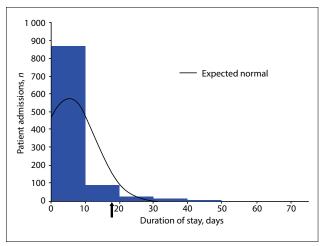


Fig. 1. LOS distribution curve. The arrow reflects the start of the visual tail and the 95th percentile.

a decade ago, but is lower than that reported by other authors using $different\ methods.^{\tiny [3,5,8,9,11]}$

It is important to standardise institutional definitions of the LSP in order to engage in ongoing audits and practise improvement initiatives; to plan the care of LSPs more economically, including counselling of families about additional risks and complications of an extended PICU stay; to motivate for involvement of the extended multidisciplinary team to provide holistic care to the child while he or she is in the PICU; and to ensure prompt recognition of medical futility. It would also be useful to have a uniform, global definition of an LSP for appropriate data comparison among PICUs. The definition delineated in this study could provide a rational comparison for further studies conducted in similar contexts around the world.

On univariate analysis, LSPs were found to be younger than SSPs, and a greater proportion were female and admitted for emergency reasons, similar to previous reports. [6] Although lower admission weight was associated with long PICU stay, this was not adjusted for patient age or height and cannot therefore be interpreted as malnutrition, wasting or stunting. Capturing nutritional characteristics in future studies would be helpful, as these data reflect the health of the population and are potentially modifiable factors.

Previous studies from SA^[10] and India^[12] described most admission diagnoses in LSPs as being of infective origin, while studies from developed countries list congenital heart disease, neurological disease and metabolic disorders as the more common diagnoses in LSPs. [12] Our LSP PICU admission diagnoses and profile were diverse owing to the varied case mix in our multidisciplinary setting. It was notable, however, that LSPs were diagnosed more frequently with pneumonia, congenital heart disease without surgery, and tracheo-oesophageal fistula or oesophageal atresia than the short-stay group. The high incidence of pneumonia is expected in the SA context of significant infectious disease burden. We did not record comorbid conditions other than HIV, as this is the most prevalent comorbidity in our population group. In future studies it would be useful to capture

	All	LSPs	SSPs	
	(N=1 126)	(n=54)	(n=1 072)	p-value
Gender (males/females)	686:440	26:28	660:412	0.049
	(60.9% male)	(48.1% male)	(61.6% male)	
Age (mo.), median (IQR)	8 (2 - 32)	4 (2 - 17)	9 (2 - 34)	0.03
Weight (kg), median (IQR)	7 (3.6 - 13.2)	4.75 (3.1 - 10.2)	7.15 (3.6 - 13.6)	0.04
Risk of mortality using PIM2 score	0.0723 (0.0259 - 0.1923)	0.096 (0.0487 - 0.1490)	0.0712 (0.0255 - 0.1938)	0.4
Emergency admissions, n (%)	832 (73.9)	46 (85.2)	786 (73.3)	0.05
Reason for admission, n (%)				
Paediatric illness	676 (60.0)	41 (75.9)	635 (58.3)	0.02
Poisoning	15 (1.3)	0 (0)	15 (1.4)	0.8
Non-accidental injury	4 (0.4)	1 (1.9)	3 (0.3)	0.5
Post cardiac surgery	188 (16.7)	9 (16.7)	179 (16.7)	0.9
Post thoracic surgery	16 (1.4)	0 (0)	16 (1.5)	0.8
Post abdominal surgery	74 (6.6)	1 (1.9)	73 (6.7)	0.2
Post cranial surgery	50 (4.4)	1 (1.9)	49 (4.6)	0.5
Post spinal surgery	15 (1.3)	0 (0)	15 (1.4)	0.8
Post airway surgery	16 (1.4)	0 (0)	16 (1.5)	0.8
Post surgery, other	15 (1.3)	0 (0)	15 (1.4)	0.8
MVA	26 (2.3)	0 (0)	26 (2.4)	0.5
Other accident	31 (2.8)	1 (1.9)	30 (2.8)	0.99
HIV status				
HIV test not done	544 (48.3)	22 (40.7)	522 (48.6)	0.3
HIV-positive, not symptomatic of AIDS	78 (6.9)	1 (1.9)	77 (7.2)	0.2
HIV-negative	409 (36.3)	22 (40.7)	387 (36.1)	0.5
HIV-positive and mildly symptomatic	95 (8.4)	7 (13.0)	88 (8.2)	0.3
Mortality, n (%)	145 (12.9)	16 (29.6)	129 (12.0)	0.0002
Risk-adjusted mortality (actual/mean predicted)	0.77	2.4	0.7	0.002
Duration of PICU stay (days), median (IQR)	3.0 (1 - 6)	29.5 (25 - 40)	2 (1 - 5)	<0.0001
MVA = motor vehicle accident.				

	LSPs	SSPs	
Specific primary diagnoses, n (%)	(n=41)	(n=1 072)	<i>p</i> -value
Sepsis (including meningococcal)	2 (4.9)	54 (5.0)	0.75
Pneumonia/pneumonitis	13 (31.7)	195 (18.2)	0.03
Cardiomyopathy/endo-/myo-/pericarditis	3 (7.3)	24 (2.2)	0.1
Congenital heart disease without surgery	13 (31.7)	58 (5.4)	< 0.0001
Necrotising enterocolitis	1 (2.4)	19 (1.8)	0.8
Tracheo-oesophageal fistula or oesophageal atresia	3 (7.3)	2 (0.2)	< 0.0001
Burns related	1 (2.4)	7 (0.7)	0.7
Non-HIV immunodeficiency	1 (2.4)	4 (0.4)	0.5
Chronic lung disease	1 (2.4)	1 (0.1)	0.1
Other shock	1 (2.4)	2 (0.2)	0.2
Congenital diaphragmatic hernia	1 (2.4)	5 (0.5)	0.5
Rheumatic heart disease	0 (0)	5 (0.5)	0.5
Gastroschisis	0 (0)	4 (0.4)	0.3
Central nervous system	0 (0)	62 (5.8)	0.2
Guillain-Barré syndrome	0 (0)	2 (0.2)	0.1
Gastroenteritis	0 (0)	61 (5.7)	0.2
Upper airway obstruction	0 (0)	17 (1.6)	0.9
Bronchiolitis	0 (0)	11 (1.0)	0.9
Apnoea	0 (0)	11 (1.0)	0.9
Asthma	0 (0)	5 (0.45)	0.5
Gastrointestinal tract	0 (0)	21 (2.0)	0.7
Tuberculosis	0 (0)	3 (0.3)	0.2
Renal failure	0 (0)	5 (0.5)	0.5
Kwashiorkor	0 (0)	2 (0.2)	0.1
Inhaled foreign body	0 (0)	2 (0.2)	0.1

all comorbid diseases and chronic care device use, as these may predispose to long PICU stay. $^{[6]}$

Diagnosis-specific mortality rates may guide critical care pathways^[6] and admission policies. There is the perception that certain conditions, such as Guillain-Barré syndrome, require prolonged LOS but have an excellent outcome, while other conditions such as extensive burns or cardiomyopathy may also result in prolonged LOS but have a poorer prognosis. This requires further research to inform admission policies in individual units.

We used the PIM2 model for the prediction of mortality during the study period, which has been validated in this PICU. [13] We found that although the admission PIM2 score was an excellent predictor of overall mortality (p<0.0001), it was not a predictive factor for long PICU stay.

PICU LOS is a marker of severity of illness, resource utilisation and performance of a PICU. International studies have consistently described small PICU LSP groups (1 - 7.5% of the total PICU population) consuming a disproportionately large amount of available resources (18 - 50%). [3,6,8,11] Our data support these findings, with LSPs constituting <5% of the PICU population, yet calculated to have utilised 30.4% of the total PICU bed days throughout the study period, and having an almost three-fold risk of death compared with the SSP group. LSP mortality rates have previously been reported to range from two-fold to almost ten-fold higher than those of SSPs. [3,6,8,11] The fact that the standardised mortality ratio was

also significantly higher than that of SSPs suggests that the actual outcomes of LSPs were worse than admission predictions of outcome.

We were unable to identify any clinically relevant predictive factors associated with long PICU stay from admission data. It is suggested that in future prospective studies, patients be reassessed after ≥2 weeks of PICU stay to attempt to identify factors predictive of outcome at that point. Once a patient has been recognised as being an LSP, it would be appropriate to discuss the ongoing care plan as a multidisciplinary team, including family counselling as appropriate. [11,14] Ongoing assessment of the LSP would ensure timeous review of medical futility criteria in order to avoid prolonging suffering and ensure rational resource allocation.

An Australian study concluded that more than two-thirds of children who spent >28 days in a PICU had an unfavourable outcome (moderate disability, severe disability or death), with almost half the LSPs having died at 6 months' follow-up after PICU discharge. [3] A Canadian prospective observational study found longer PICU stays to be independently associated with poor quality of life and functional outcome at follow-up after 1 month. [15] PICU LOS is a treatment-related exposure that has been highlighted as a predictor of post-critical illness psychiatric morbidity, including post-traumatic stress disorder and depression. [16] In our study, outcome was measured using PICU survival and non-survival only. Owing to the retrospective nature of this study, we were unable to measure quality of life or morbidity of survivors of long PICU stay,

	Alive (<i>n</i> =38)	Died (n=16)	<i>p</i> -value
fale gender, n (%)	18 (47.4)	8 (50.0)	0.9
ge (mo.), median (IQR)	4.5 (2.0 - 17.0)	3.5 (2.0 - 15.0)	0.9
Veight (kg), median (IQR)	4.8 (3.1 - 10.2)	4.9 (3.3 - 10.0)	0.9
isk of mortality using PIM2 score	0.099 (0.056 - 0.141)	0.090 (0.023 - 0.254)	0.8
mergency admissions, n (%)	34 (89.5)	12 (75.0)	0.3
eason for admission, n (%)			
Paediatric illness	30 (78.9)	11 (68.8)	0.7
Poisoning	0	0	-
Non-accidental injury	1 (2.6)	0	0.7
Post cardiac surgery	5 (13.2)	4 (25.0)	0.5
Post thoracic surgery	0	0	-
Post abdominal surgery	0	1 (6.3)	0.7
Post cranial surgery	1 (2.6)	0	0.7
Post spinal surgery	0	0	-
Post airway surgery	0	0	-
Post surgery, other	0	0	-
MVA	0	0	-
Other accident	1 (2.6)	0	0.7
IIV status, n (%)			
Test not done	17 (44.7)	5 (31.3)	0.5
HIV-positive, not symptomatic of AIDS	1 (2.6)	0	0.7
HIV-negative	14 (36.8)	8 (50.0)	0.6
HIV-positive and mildly symptomatic	4 (10.5)	3 (18.8)	0.8
Duration of ICU stay before discharge or death (d), nedian (IQR)	27 (25 - 36)	38.5 (26.5 - 45.0)	0.08

	Adjusted odds ratio	95% confidence interval	<i>p</i> -value
Female gender	1.7	1.001 - 3.03	0.047
Age	0.99	0.98 - 1.01	0.2
Weight	1.01	0.96 - 1.05	0.8
Paediatric illness as reason for admission	1.5	0.71 - 3.33	0.3
Emergency admission	1.46	0.60 - 3.55	0.4

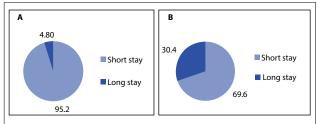


Fig. 2. (A) Proportion of LSPs in the PICU population, and (B) their resource consumption (in terms of bed utilisation) relative to SSPs.

and this constitutes a study limitation. We did not collect data related to previous PICU admissions and how this influenced subsequent outcome, and this is recommended for future studies.

This study did not address the nature of PICU care required by LSPs. In many cases the nursing, therapeutic input and psychosocial needs of these children and their families may differ profoundly from those of acutely ill SSPs. Parents too are at higher risk of adverse consequences of their child's admission. A prospective study in the USA highlighted frequent conflicts in the families of LSPs. [14]

Long-term functional outcome and quality of life measures may be more useful in the context of improved PICU survival, and it is recommended that these outcomes be used in future prospective studies of LSPs.^[3] There is much scope for future research on the quality of life of PICU survivors, particularly in the SA context where there is a low ratio of health professionals to patients, the social welfare disability grant offered to children with a moderate to severe disability is currently USD120 per month, and there

is poor infrastructure and inaccessibility to transport in much of the country. $^{[17]}$ Identifying whether high PICU mortality has been traded for a higher prevalence of morbidity has potential ramifications for provision and delivery of healthcare to this vulnerable population.

The poor outcome of LSPs reported here and in previous studies raises concerns around inappropriate resource consumption in the PICU. We recognise that in the SA setting, equitable distribution of healthcare services is an important ethical consideration when budgeting and planning for healthcare resource allocation.

Conclusions

It is important to standardise institutional definitions of LSPs. Our study defined LSPs as patients with a PICU LOS of >19 days based on the two best-fit models: >95th centile of duration of stay and the visual 'tail' of the LOS distribution curve. The definition delineated in this study could provide a rational basis for comparison between similar PICUs across the world.

LSPs represent a small percentage of PICU admissions yet have significantly increased mortality (greater than expected at the time of admission) and consume a disproportionate amount of resources compared with SSPs. This pattern from an SA setting is in keeping with international trends.

We were unable to establish a predictive model for the early recognition of potential LSPs in order to plan PICU bed allocation effectively.

Further investigations are needed to assess the quality of life of survivors of long PICU stay, particularly in LMICs.

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