

ORIGINAL

## 

# R. González-Cortés, J. López-Herce-Cid\*, A. García-Figueruelo, G. Tesorero-Carcedo, M. Botrán-Prieto, A. Carrillo-Álvarez

Servicio de Cuidados Intensivos Pediátricos, Hospital General Universitario Gregorio Marañón, Madrid, Spain

Received 17 November 2010; accepted 6 April 2011 Available online 21 November 2011

Abstract

### **KEYWORDS**

	Abstract
Critically ill children;	Objective: To analyze mortality and resource consumption in patients with long stays in pedi-
Pediatric intensive	atric intensive care units (Picos).
care unit;	Design: A retrospective, descriptive case series study.
Mortality:	Scope: Medical-surgical PICU in a third level hospital.
Length of stay;	Patients: Data were collected from patients with a stay of 28 days or more in PICU between
Tracheostomy:	2006 and 2010. Of the 2118 patients assisted in this period, 83 (3.9%) required prolonged stay.
Treatment	Study variables: Morbidity-mortality and resource consumption among patients with prolonged
withdrawal;	stay in the PICU.
Nosocomial infection;	Results: Mortality was higher in patients with a long stay (22.9%) than in the rest of the patients
Health resources	(2%) ( $p < 0.001$ ). In 52.6% of these patients, death occurred after withdrawal of treatment or
	after not starting resuscitation measures. Patients with prolonged stay showed a high inci-
	dence of nosocomial infection (96.3%) and an important consumption of healthcare resources
	(97.6% required conventional mechanical ventilation, 90.2% required transfusion of blood prod-
	ucts. 86.7% required intravenous vasoactive drugs and 22.9% required extracorporeal membrane
	oxygenation [ECMO]).
	Conclusions: Critical children with prolonged stay in the PICU show important morbidity and
	mortality and an important consumption of healthcare resources. The adoption of specific
	measures permitting early identification of patients at risk of prolonged stay is peeded in order
	the adapt the reporting early identification of patients at this of prioring distay is fielded in order
	to adapt the apeutic measures and available resources, and to improve treatment efficiency.
	© 2010 Elsevier Espana, S.L. and SEMICYUC. All rights reserved.

\* Corresponding author.

<sup>\*</sup> Please cite this article as: González-Cortés R, et al. Ingreso prolongado en la unidad de cuidados intensivos pediátricos: mortalidad y consumo de recursos asistenciales. Med Intensiva. 2011;35:417–23.

E-mail address: pielvi@hotmail.com (J. López-Herce-Cid).

<sup>2173-5727/</sup> $\ensuremath{\$}$  - see front matter  $\ensuremath{\$}$  2010 Elsevier España, S.L. and SEMICYUC. All rights reserved.

PALABRAS CLAVE

Niños en estado crítico; Unidad de cuidados intensivos pediátricos; Mortalidad; Duración de ingreso; Traqueostomía; Retirada de tratamiento; Infección nosocomial; Recursos sanitarios

#### Ingreso prolongado en la unidad de cuidados intensivos pediátricos: mortalidad y consumo de recursos asistenciales

#### Resumen

*Objetivo:* Analizar la mortalidad y el consumo de recursos de los niños con ingreso prolongado en unidades de cuidados intensivos pediátricos (UCIP).

Diseño: Estudio descriptivo retrospectivo de una serie de casos.

*Ámbito*: UCIP médico-quirúrgica de un hospital de tercer nivel.

*Pacientes:* Se recogieron los datos de los pacientes ingresados durante 28 o más días en la UCIP entre 2006 y 2010. De los 2.118 pacientes ingresados entre 2006 y 2010, 83 (3,9%) requirieron ingreso prolongado.

*Variables de interés*: Se analizaron la morbimortalidad y el consumo de recursos por los pacientes con ingreso prolongado.

*Resultados*: La mortalidad de los pacientes con ingreso prolongado fue mayor (22,9%) que la del resto de los pacientes (2%) (p < 0,001). En un 52,6% de estos pacientes el fallecimiento se produjo tras la limitación del esfuerzo terapéutico o por no iniciar medidas de reanimación. Los pacientes con ingreso prolongado presentaron una elevada incidencia de infección nosocomial (96,3%) y un elevado consumo de los recursos asistenciales (el 97,6% precisó ventilación mecánica; el 90,2%, transfusión de hemoderivados; el 86,7% fármacos vasoactivos intravenosos, y el 22,9%, oxigenación por membrana extracorpórea (ECMO)).

*Conclusiones:* Los niños en estado crítico con ingreso prolongado en la UCIP tienen una elevada morbimortalidad y requieren un elevado consumo de recursos asistenciales. Son necesarias medidas específicas que permitan identificar precozmente a los pacientes susceptibles de presentar ingreso prolongado para adecuar las medidas terapéuticas y los recursos disponibles y mejorar la eficiencia del tratamiento.

© 2010 Elsevier España, S.L. y SEMICYUC. Todos los derechos reservados.

#### Introduction

In the last few decades technological advances and the development of new treatments have led to a decrease in mortality in pediatric intensive care units (PICUs). However, these improvements in turn have led to the appearance of a new patient profile in these units, namely subjects who remain admitted to the PICU for very prolonged periods of time.

Such scientific-technological progress has given rise not only to a greater presence of patients in life-threatening situations for prolonged periods of time but also to an increasing number of patients in intensive care who are dependent upon a given type of technology or medical care which for different reasons cannot be provided outside these units.<sup>1</sup>

Patients with prolonged admission constitute a minority group, though as a result of their prolonged stay and their illnesses, they often consume an important amount of healthcare resources, with increased morbidity and mortality.<sup>2-4</sup>

The early identification of these patients could allow improved channeling of sociosanitary resources, thereby ensuring improved care and better utilization of the available resources.  $^{5-9}$ 

The present study analyzes the characteristics and evolution of the patients admitted for over 28 days to a PICU, and examines the mortality and the healthcare resource consumption generated by these patients.

#### Patients and methods

A retrospective descriptive study was made, based on a review of the clinical histories of all patients admitted to the PICU for 28 days or more during the period between January 2006 and March 2010.

Our PICU belongs to a third-level hospital with a recruitment population of approximately 700,000 in the Community of Madrid (Spain), and which moreover constitutes a national reference center in heart surgery. The PICU is an 11-bed medical-surgical unit with a heart surgery program and extracorporeal membrane oxygenation (ECMO).

The following data were collected on all patients: age, sex, diagnosis, presence of congenital anomalies and congenital heart disease, date and reason for admission, origin, existence of previous cardiorespiratory arrest, and type of surgery performed in the case of postsurgery patients. Severity scores were not recorded, since these are not systematically registered in our PICU.

As variables related to treatment, we registered the type of assisted ventilation (invasive mechanical ventilation, high-frequency ventilation, noninvasive ventilation, high-flow oxygen therapy) and its duration, the continuous perfusion of vasoactive drugs and its duration, extrarenal filtration techniques (specifying whether peritoneal dialysis or continuous venovenous extrarenal filtration was used) and their duration, treatment and duration of ECMO and/or ventricular assist procedures, other techniques (central venous catheterization, arterial catheterization, tracheostomy, gastrostomy and bladder catheterization), and the incidence and location of nosocomial infections. We likewise documented mortality, the causes of death, whether the patients were receiving vasoactive drugs, mechanical ventilation or ECMO at the time of death, and whether there had been limitation of therapeutic effort or a decision not to resuscitate. In the surviving patients we registered the destination at discharge from the PICU.

The data were analyzed using the SPSS version 15.0 statistical package for MS Windows. The  $\chi^2$  test was applied for the analysis of qualitative variables related to mortality, while the Student *t*-test was used to analyze the differences in the means or duration of the different treatments between deceased patients and survivors.

#### Results

During the study period we recorded a total of 2118 admissions to the PICU; of these, 83 (3.9%) involved a stay of 28 days or more. The prolonged stays accounted for 36.1% of the global stay in the PICU. The mean duration of stay in the patients with prolonged admission was  $55.6 \pm 30.1$  days (median 45 days, range 28–179 days)—the mean stay in the PICU for the total series being 6 days (range 0–179 days).

A total of 65.1% of the patients with prolonged admission were males, with a median age of 5 months (range 0-217 months); 65.1% were under 12 months of age. In turn, 81.9% presented congenital anomalies, and 73.5% of the total patients with prolonged admission were diagnosed with congenital heart disease.

Table 1 shows the diagnoses of the patients with prolonged admission. A total of 75.9% of the patients with prolonged admission suffered heart disease (the reason for admission being its surgical correction, heart transplantation or heart failure). In turn, 37.3% of the admissions were from the operating room (heart surgery in 92.7%) (Table 2). A percentage of the patients admitted after the correction of congenital heart disease were admitted not from the operating room but from other areas (neonatal ICU, hospitalization ward or hemodynamics room). The most frequent heart diseases in the studied group of patients were left ventricle hypoplasia (14 patients), Shone syndrome (10 patients), major vessel transpositioning (8 patients) and lung atresia (7 patients). Other pathologies were less common.

#### **Resources used**

Table 3 summarizes the healthcare resources used among the children with prolonged admission to the PICU. A full

Table 1Diagnoses of the patients with prolonged admission to the PICU.

Reason for admission	Patients, n (%)
Correction of congenital heart disease	37 (44.6)
Decompensated heart disease	16 (19.3)
Respiratory failure	7 (8.4)
Heart transplantation	5 (6)
Acquired myocardiopathy	5 (6)
Cardiorespiratory arrest	3 (3.6)
Airway surgery	3 (3.6)
Sepsis	3 (3.6)
Seizures	2 (2.4)
Head injuries	1 (1.2)
Spinal cord injuries	1 (1.2)
Total	83 (100)

Table 2Origin at admission of the patients with prolongedadmission to the PICU.

Origin at admission	Patients, n (%)
Operating room (elective surgery)	26 (31.3)
Operating room (emergency surgery)	5 (6)
Hospitalization ward	22 (26.5)
Neonatal Intensive Care	10 (12)
Emergencies	6 (7.2)
Hemodynamics room	4 (4.8)
Transfer from another hospital	10 (12)
Total	83 (100)

97.6% of the patients required invasive mechanical ventilation, 12% required high-frequency ventilation, and 65% required noninvasive ventilation. In turn, 33 patients (39.8%) required extrarenal filtration techniques: in all these cases venovenous hemodiafiltration was performed, with added peritoneal dialysis in 8 patients (9.6%).

A total of 86.8% of the patients required inotropic drugs in continuous perfusion (adrenalin in 75.9%), 22.9% were subjected to ECMO, and 4.8% required mechanical ventricular assistance. A total of 24.1% of the patients underwent heart transplantation (15 of them after admission to the PICU).

Bladder catheterization was carried out in 94.9% of the cases, with central venous accesses and arterial catheters in 92.8% and 92.6%, respectively. In turn, 90.2% of the patients required blood product transfusions during admission.

Table 3 Resources used by the patients with prolonged admission to the Pl	JCU
---	-----

Resource	Frequency, n (%)	Durat	Duration (days)	
		Mean $\pm$ SD	Median [range]	
Hospitalization	83 (100)	55.6 ± 30	45 [28-179]	
Conventional mechanical ventilation	81 (97.6)	$\textbf{42.6} \pm \textbf{30.6}$	34[0-166]	
Extrarenal filtration techniques	33 (39.8)	$21.7 \pm 16.2$	19[1-65]	
Vasoactive drugs	72 (86.7)	$36.5 \pm 24.3$	29 [1-137]	
ЕСМО	19 (22.9)	8.6 ± 7.1	7 [2-29]	
Ventricular assist	4 (4.8)	$\textbf{37.8} \pm \textbf{37.7}$	21.5 [14–94]	

 Table 4
 Comparison of deceased patients and survivors.

	Total	Deceased	Survivors	р
Total, n (%)	83 (100%)	19 (100%)	64 (100%)	-
Surgery, n (%)	41 (49.4%)	12 (63.2%)	29 (45.3%)	0.172
Congenital anomalies, n (%)	68 (81.9%)	17 (89.5%)	51 (79.7%)	0.33
Congenital heart disease, n (%)	61 (73.5%)	16 (84.2%)	45 (70.3%)	0.228
CRA prior to admission, n (%)	7 (8.4%)	3 (15.8%)	4 (6.3%)	0.189
Infection prior to admission, n (%)	26 (32.1%)	4 (22.2%)	22 (34.9%)	0.309
Nosocomial infection, n (%)	79 (96.3%)	19 (100%)	60 (95.2%)	0.333
Mechanical ventilation, n (%)	81 (97.6%)	19 (100%)	62 (96.9%)	0.435
Mean (days)	42.6	56.8	31.8	0.029
Median (days)	34	57	29	-
Range (days)	0-166	30-100	0-166	-
CPAP, n (%)	34 (41.5%)	4 (21.1%)	30 (47.6%)	0.039
BIPAP, n (%)	31 (37.8%)	6 (31.8%)	25 (39.7%)	0.523
High-flow oxygen therapy, n (%)	31 (39.2%)	1 (5.3%)	30 (50%)	0.001
Vasoactive drugs, n (%)	72 (86.7%)	16 (84.2%)	56 (87.5%)	0.71
Mean (days)	36.5	50.9	24.7	0.006
Median (days)	29	50	26	-
Range (days)	1-137	30-83	1-137	-
Adrenalin perfusion, n (%)	63 (75.9%)	15 (78.9%)	48 (75%)	0.724
ECMO, n (%)	19 (22.9%)	7 (36.8%)	12 (18.8%)	0.099
Mean (days)	8.6	13.3	5.9	0.023
Median (days)	7	11	4.5	-
Range (days)	2-29	2-29	2-15	-
Blood products, n (%)	74 (90.2%)	19 (100%)	55 (87.3%)	0.43
Extrarenal filtration, n (%)	33 (39.8%)	13 (68.4%)	20 (31.3%)	0.004
Mean (days)	21.7	23.3	20.8	0.675
Median (days)	19	19	18.5	-
Range (days)	1-65	4-50	1-65	-

BIPAP: noninvasive bi-level positive airway pressure ventilation; CPAP: continuous positive airway pressure; ECMO: extracorporeal membrane oxygenation; CRA: cardiorespiratory arrest.

The percentages in each column refer to the group of patients described in the column, not to the total.

Three patients (3.6%) presented a tracheostomy and four a (4.8%) gastrostomy at the time of admission to the PICU. During their stay, another 13 patients (15.7%) underwent tracheostomy and 9 (10.8%) gastrostomy.

#### Morbidity

Infection was suspected in 32.1% of the patients at the time of admission to the PICU; the percentage corresponding to infections of nosocomial origin could not be established. A total of 96.3% of the patients suffered at least one nosocomial infection during admission—the most frequent condition being pneumonia (72%), followed by urinary infections (34.6%) and catheter-related infections (28.4%).

#### Mortality

A total of 19 of the 83 patients with prolonged admission died. Mortality among the children with prolonged admission was significantly greater (22.9%) than in the rest of the patients (2%) (p < 0.001). The most frequent causes of death were withdrawal or limitation of therapeutic effort (42.1%) and the non-adoption of resuscitation measures (10.5%). A

total of 8.4% of the patients with prolonged admission had suffered cardiorespiratory arrest (CRA) before admission to the PICU, and 35.4% suffered CRA during their stay.

At the time of death, all the patients were subjected to mechanical ventilation (conventional mechanical ventilation in 89.5%, noninvasive ventilation in 5.3%, and high-frequency ventilation in 5.3%,); 89.5% were receiving vasoactive drugs in continuous perfusion; and 26.7% were subjected to ECMO.

There was no difference in mortality between patients under one year of age (23.3%) and the older children (22.6%). Table 4 depicts the relation of other variables to mortality. Although congenital heart disease, admission after surgery, ECMO, continuous extrarenal filtration techniques (CEFT), and blood product transfusions were more frequent in the patients who died, the differences only reached statistical significance in the case of CEFT. The patients who died presented a comparatively longer duration of mechanical ventilation, inotropic drug treatment and ECMO (Table 4).

Discharge of the survivors was to the pediatric ward (89%) or to other intensive care units (4.6%), while one patient (1.5%) was directly discharged home. Three patients (4.6%) were still in the PICU at the end of the study.

#### Discussion

The results of our study coincide with those published in other countries, and reveal that patients with prolonged admission to the PICU, while few in number, generate great concern and account for an important percentage of hospital stay.

The epidemiological characteristics of our patients are similar to those described by other authors, with a predominance of males and an important percentage of cases of congenital malformations—particularly heart diseases.<sup>1–3,10</sup> In effect, our series shows an important presence of patients with heart disease—this possibly being conditioned by the fact that our PICU is a reference center for disorders of this kind. This also explains why most admissions were from the operating room, other hospitalization areas, or from other centers.

#### Healthcare resources

We have observed important healthcare resource utilization among the patients with prolonged admission, in coincidence with the findings of earlier studies.<sup>1,5,11,12</sup>

Tracheostomy and gastrostomy can improve management, reduce the complications and improve the quality of life of patients with prolonged admission.<sup>13-16</sup> Although early tracheotomy is advocated in adults,<sup>17,18</sup> in children the best timing for this technique has not been established. The mean intubation time prior to elective tracheostomy in children subjected to prolonged mechanical ventilation varies from 31 to 51 days, depending on the data source.<sup>14,15</sup> In a survey of 63 pediatric intensivists in Canada regarding the use of programmed tracheostomy in children subjected to mechanical ventilation, 51% of the interviewed physicians considered programmed tracheostomy to be underused in children.<sup>14</sup> In our study, the procedure was only carried out in 15.8% of the patients.

#### Nosocomial infections

We recorded a high incidence of nosocomial infections,  $^{19-21}$  particularly respiratory infections probably related to the important use of invasive mechanical ventilation, in coincidence with the findings of earlier studies.  $^{22.23}$  Likewise, the prolonged use of central vascular accesses and bladder catheterization favors infection in these locations. Specific measures are therefore needed to reduce the duration of invasive techniques, and to prevent and quickly diagnose and treat nosocomial infectious processes in these patients.  $^{5,6,10,11,24-27}$ 

#### Discharge from the PICU

Most of our patients were discharged to other hospitalization areas. Patients with prolonged admission to the PICU often remain dependent upon technological systems fundamentally mechanical ventilation—and so require specific care measures not available in the hospitalization wards.<sup>28</sup> This fact often delays discharge from the PICU, despite the achievement of sufficient patient stability.

The creation of intermediate care units or units for technology-dependent patients, with sufficient personnel and specific material resources to safely treat subjects in clinical conditions that are unlikely to change (apart from the development of complications), could allow for more efficient healthcare resource utilization.<sup>29,30</sup>

Home mechanical ventilation facilitates the out-hospital care of some patients, increasing their quality of life and that of their relatives. However, this option requires the existence of care teams to conduct regular controls and to train the caregivers, and the family in turn must receive sufficient sociosanitary support. The scarcity of these aids often represents an obstacle for discharging patients from the PICU, contributing to further their chronification and institutionalization.<sup>28,31,32</sup>

#### Mortality

Mortality among patients with prolonged admission is much higher than in the rest of the patients admitted to the PICU. In effect, the mortality rate in our series was 10 times greater—in coincidence with the findings of other authors in pediatric patients.<sup>2,3,33</sup> It is important to mention that our PICU does not often admit patients for short-duration techniques or procedures—the low associated mortality of which would have contributed to lower our overall mortality figures.

The early identification of patients with a high mortality risk could allow more efficient use of the available resources, attempting to avoid the use of futile invasive procedures.<sup>7</sup> Some factors, such as heart diseases, or the need for extrarenal filtration techniques or ECMO, are associated with increased mortality, though in our study most of them showed no statistically significant differences. Nevertheless, it must be taken into account that most of these factors are only indicators or markers of patient severity.

In our study, and in the same way as in other series,  $^{3,34,35}$  the cause of death in almost one-half of the cases was the withdrawal or limitation of therapeutic effort. Despite this fact, in most cases mechanical ventilation or vasoactive drug treatment was maintained up until death. This may be due to the way in which the limitation of therapeutic effort is carried out in our PICU, though it could also imply the excessive prolongation of techniques that may be regarded as futile. It therefore seems necessary to optimize the approach to such patients when death draws near—addressing and planning the measures to be taken, with a view to avoiding inadequate treatment prolongation.<sup>3-5,8,34</sup>

Considering the retrospective design of our study, it is important to point out that its main limitation is the absence of comparisons between the group of patients with prolonged admission and those without prolonged admission—a situation that precluded a multivariate statistical analysis of the risk factors for prolonged admission.

On the other hand, our PICU has a series of specific characteristics due to the high percentage of patients admitted with congenital heart disease. Other studies involving PICUs with more general patients are therefore needed to confirm the results obtained.

In conclusion, although children subjected to prolonged admission represent a small percentage of the patients seen in the PICU, they suffer important morbidity-mortality, account for much of the occupation of the unit, and involve important healthcare resource utilization. Early identification is required of those patients with a high risk of requiring prolonged admission, in order to establish specific management strategies designed to allow more efficient care and use of the available resources. The creation of intermediate care units for the management of these patients may help reduce resource consumption and increase the quality of the provided healthcare.

#### References

- 1. Briassoulis G, Filippou O, Natsi L, Mavrikiou M, Hatzis T. Acute and chronic paediatric intensive care patients: current trends and perspectives on resource utilization. QJM. 2004;97:507–18.
- Martin CM, Hill AD, Burns K, Chen LM. Characteristics and outcomes for critically ill patients with prolonged intensive care unit stays. Crit Care Med. 2005;33:1922–7.
- Naghib S, Van der Starre C, Gischler SJ, Joosten KFM, Tibboel D. Mortality in very long-stay pediatric intensive care unit patients and incidence of withdrawal of treatment. Intensive Care Med. 2010;36:131–6.
- Sachdeva RC, Jefferson LS, Coss-Bu J, Brody BA. Resource consumption and the extent of futile care among patients in a pediatric intensive care unit setting. J Pediatr. 1996;128:742-7.
- 5. Marcin JP, Slonim AD, Pollack MM, Ruttimann UE. Long-stay patients in the pediatric intensive care unit. Crit Care Med. 2001;29:652–7.
- Parkman SE, Woods SL. Infants who have undergone cardiac surgery: what can we learn about lengths of stay in the hospital and presence of complications? J Pediatr Nurs. 2005;20:430–40.
- 7. Ruttimann UE, Pollack MM. Variability in duration of stay in pediatric intensive care units: a multiinstitutional study. J Pediatr. 1996;128:35–44.
- Goh AY, Mok Q. Identifying futility in a paediatric critical care setting: a prospective observational study. Arch Dis Child. 2001;84:265–8.
- 9. Gillespie M, Kuijpers M, Van Rossem M, Ravishankar C, Gaynor JW, Spray T, et al. Determinants of intensive care unit length of stay for infants undergoing cardiac surgery. Congenit Heart Dis. 2006;1:152–60.
- Van der Heide P, Hassing MBF, Gemke RJBJ. Characteristics and outcome of long-stay patients in a paediatric intensive care unit: a case-control study. Acta Paediatr. 2004;93: 1070-4.
- Auburtin B, Saizou C, Dauger S, Hartmann JF, Mercier JC, Beaufils F. Les séjours prolongés en réanimatión pédiatrique. Analyse rétrospective de 100 séjours. Arch Pediatr. 2001;8:158–65.
- López-Herce Cid J, Leyton Avilés P, Urbano Villaescusa J, Cidoncha Escobar E, Del Castillo Peral J, Carrillo Alvarez A, et al. Factores de riesgo de la ventilación mecánica prolongada de niños con cirugía cardiaca. Med Intensiva. 2008;32: 369–77.
- Parrilla C, Scarano E, Guidi ML, Galli J, Paludetti G. Current trends in paediatric tracheostomies. Int J Pediatr Otorhinolaryngol. 2007;71:1563–7.

- Principi T, Morrison GC, Matsui DM, Speechley KN, Seabrook JA, Singh RN, et al. Elective tracheostomy in mechanically ventilated children in Canada. Intensive Care Med. 2008;34:1498–502.
- Scales DC, Thiruchelvam D, Kiss A, Redelmeier DA. The effect of tracheostomy timing during critical illness on long-term survival. Crit Care Med. 2008;36:2547–57.
- Conlon NP, Breatnach C, O'Hare BP, Mannion DW, Lyons BJ. Health-related quality of life after prolonged pediatric intensive care unit stay. Pediatr Crit Care Med. 2009;10: 41–4.
- Griffiths J, Barber VS, Morgan L, Young JD. Systematic review and meta-analysis of studies of the timing of tracheostomy in adult patients undergoing artificial ventilation. BMJ. 2005;330:1243.
- Terragni PP, Antonelli M, Fumagalli R, Faggiano C, Berardino M, Pallavicini FB, et al. Early vs late tracheotomy for prevention of pneumonia in mechanically ventilated adult ICU patients: a randomized controlled trial. JAMA. 2010;303: 1483–9.
- Urrea M, Pons M, Serra M, Latorre C, Palomeque A. Prospective incidence study of nosocomial infections in a pediatric intensive care unit. Pediatr Infect Dis J. 2003;22: 490–4.
- Mireya UA, Martí PO, Xavier KV, Cristina LO, Miguel MM, Magda CM. Nosocomial infections in paediatric and neonatal intensive care units. J Infect. 2007;54:212–20.
- Stover BH, Shulman ST, Bratcher DF, Brady MT, Levine GL, Jarvis WR. Nosocomial infection rates in US children's hospitals' neonatal and pediatric intensive care units. Am J Infect Control. 2001;29:152–7.
- Elward AM, Warren DK, Fraser VJ. Ventilator-associated pneumonia in pediatric intensive care unit patients: risk factors and outcomes. Pediatrics. 2002;109:758–64.
- Bigham MT, Amato R, Bondurrant P, Fridriksson J, Krawczeski CD, Raake J, et al. Ventilator-associated pneumonia in the pediatric intensive care unit: characterizing the problem and implementing a sustainable solution. J Pediatr. 2009;154, 582–587.e2.
- 24. Brown KL, Ridout DA, Goldman AP, Hoskote A, Penny DJ. Risk factors for long intensive care unit stay after cardiopulmonary bypass in children. Crit Care Med. 2003;31:28–33.
- 25. García-Teresa MA, Casado-Flores J, Delgado Domínguez MA, Roqueta-Mas J, Cambra-Lasaosa F, Concha-Torre A, et al. Infectious complications of percutaneous central venous catheterization in pediatric patients: a Spanish multicenter study. Intensive Care Med. 2007;33:466–76.
- Guardia Camí MT, Jordan García I, Urrea Ayala M. Infección nosocomial en postoperados de cirugía cardiaca. An Pediatr (Barc). 2008;69:34-8.
- Principi N, Esposito S. Ventilator-associated pneumonia (VAP) in pediatric intensive care units. Pediatr Infect Dis J. 2007;26:841–3.
- Parker G, Bhakta P, Lovett C, Olsen R, Paisley S, Turner D. Paediatric home care: a systematic review of randomized trials on costs and effectiveness. J Health Serv Res Policy. 2006;11:110–9.
- 29. Kanter RK. Post-intensive care unit pediatric hospital stay and estimated costs. Crit Care Med. 2000;28:220-3.
- Tearl DK, Cox TJ, Hertzog JH. Hospital discharge of respiratorytechnology-dependent children: role of a dedicated respiratory care discharge coordinator. Respir Care. 2006;51: 744–9.
- Raymond JA. The integration of children dependent on medical technology into public schools. J Sch Nurs. 2009;25: 186-94.
- Lewis M, Noyes J. Discharge management for children with complex needs. Paediatr Nurs. 2007;19:26–30.

- Pollack MM, Wilkinson JD, Glass NL. Long-stay pediatric intensive care unit patients: outcome and resource utilization. Pediatrics. 1987;80:855–60.
- 34. Sprung CL, Cohen SL, Sjokvist P, Baras M, Bulow H, Hovilehto S, et al. End-of-life practices in European

intensive care units: the Ethicus Study. JAMA. 2003;290: 790-7.

35. Goh AY, Lum LC, Chan PW, Bakar F, Chong BO. Withdrawal and limitation of life support in paediatric intensive care. Arch Dis Child. 1999;80:424–8.