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# Antimicrobial stewardship programs: Progress and challenges in Brazilian pediatric intensive care units

Programa de gerenciamento de antimicrobianos: progressos e desafios nas unidades de terapia intensiva pediátricas brasileiras

Programa de gestión del uso de antimicrobianos: avances y desafíos en las unidades de cuidados intensivos pediátricos brasileñas

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#### **ABSTRACT**

**Objective:** to assess Leadership Support and Definition of Responsibilities for antimicrobial stewardship programs in Brazilian pediatric intensive care units. **Methods:** a multicenter, cross-sectional study conducted in 219 hospitals using a Leadership Support and Definition of Responsibilities assessment form. The data were analyzed based on location, service profile and institutional size. **Results:** hospitals in the Southeast region, with a private institutional profile, over 200 beds and up to 20 intensive care beds stood out. It was observed that 73.30% of the hospitals had formal documentation of their antimicrobial stewardship program, 90.50% had dedicated teams, 66.21% had sufficient financial resources, and 67.58% had designated a responsible manager. **Conclusion:** inadequate funding and the need to standardize activities are challenges. It is essential to improve multidisciplinary team engagement and to integrate strategies into antimicrobial stewardship programs for antimicrobial safety in children.

Descriptors: Health Resources; Drug Resistance, Microbial; Antimicrobial Stewardship; Leadership; Efficacy.

#### **RESUMO**

**Objetivo:** avaliar o Apoio das Lideranças e a Definição de Responsabilidades dos programas de gerenciamento de antimicrobianos em unidades de terapia intensiva pediátrica no Brasil. **Métodos:** estudo multicêntrico, transversal, realizado em 219 hospitais por meio de formulário de avaliação do Apoio das Lideranças e Definição de Responsabilidades. Os dados foram analisados de acordo com localização, perfil de atendimento e tamanho porte da instituição. **Resultados:** destacaram-se os hospitais localizados na região sudeste, perfil institucional privado, mais de 200 leitos e até 20 leitos de terapia intensiva. Observou-se que 73,30% dos hospitais possuíam documentação formal do programa de gerenciamento de antimicrobiano, 90,50% tinham equipes dedicadas, 66,21% dispunham de recursos financeiros suficientes e 67,58% definiram um gestor responsável. **Conclusão:** a falta de financiamento adequado e a necessidade de padronização das atividades são desafios. Melhorar a adesão da equipe multiprofissional e integrar estratégias dos programas de gerenciamento de antimicrobianos são essenciais para a segurança antimicrobiana em crianças.

Descritores: Recursos em Saúde; Resistência Microbiana a Medicamentos; Gestão de Antimicrobianos; Liderança; Eficácia.

### **RESUMEN**

**Objetivo:** evaluar el Apoyo de los Directivos y la Definición de Responsabilidades de los programas de gestión de antimicrobianos en unidades de cuidados intensivos pediátricos en Brasil. **Métodos:** estudio multicéntrico, transversal realizado en 219 hospitales mediante un formulario de evaluación de Apoyo de los Directivos y Definición de Responsabilidades. Los datos se analizaron según ubicación, perfil del servicio y tamaño de la institución. **Resultados:** se destacaron los hospitales ubicados en la región Sudeste, el perfil institucional privado y aquellos con más de 200 camas y hasta 20 camas de cuidados intensivos. Se observó que el 73,30% de los hospitales contaba con documentación formal del programa de gestión de antimicrobianos, el 90,50% contaba con equipos dedicados, el 66,21% contaba con recursos financieros suficientes y el 67,58% contaba con un gerente designado. **Conclusión:** los desafíos son la falta de financiamiento adecuado y la necesidad de estandarización de las actividades. Es fundamental mejorar la adherencia del equipo multidisciplinario e integrar estrategias de los programas de gestión de antimicrobianos para la seguridad antimicrobiana en niños.

**Descriptores:** Recursos en Salud; Farmacorresistencia Microbiana; Programas de Optimización del Uso de los Antimicrobianos; Liderazgo; Eficacia.

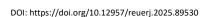
## INTRODUCTION

Antimicrobial resistance (AMR) poses a growing threat to global public health, compromising the effectiveness of treatments and increasing morbidity and mortality rates. This problem is particularly concerning in Latin America, where several factors, such as inappropriate use of antimicrobials and a lack of control programs, contribute to the rise of

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bacterial resistance<sup>1,2</sup>. In response to this crisis, the Antimicrobial Stewardship Program (ASP) has emerged as a crucial strategy to promote the rational use of antimicrobials and control the spread of resistant pathogens<sup>3</sup>.

The World Health Organization (WHO) recognized AMR as a global priority, launching the Global Action Plan to combat antimicrobial resistance in 2015<sup>4</sup>. This plan aims to improve awareness and understanding of AMR, strengthen surveillance and research, reduce the incidence of infections, optimize the use of antimicrobial drugs, and ensure sustainable investments in the AMR response<sup>5</sup>.

In Latin America, several countries have implemented policies and programs to address AMR<sup>5</sup>. In Brazil, the Brazilian Health Regulatory Agency (*Agência Nacional de Vigilância Sanitária*, ANVISA) plays a central role in coordinating actions to promote rational antimicrobial use. In 2017, the country launched its National Plan for Antimicrobial Resistance Prevention and Control, aligned with WHO guidelines<sup>6</sup>. This plan includes ASP implementation in hospitals, aiming to reduce inappropriate antimicrobial use and monitor bacterial resistance<sup>7</sup>.

In 2019 and 2022, national investigations into the Antimicrobial Stewardship Program (ASP) scenario in Brazil were conducted, coordinated by ANVISA. These studies provided a detailed overview of the implementation and effectiveness of ASPs in healthcare facilities, highlighting progress and identifying areas for improvement<sup>8</sup>. National actions to reduce infections include promoting healthcare practices, active surveillance of hospital-acquired infections, and training healthcare professionals on this scenario and the appropriate use of antimicrobials<sup>9</sup>.

The success of an ASP depends heavily on compliance with elements 1 and 2 of the program's guidelines. The first element emphasizes Leadership Support, which is crucial to ensuring the allocation of human, financial, and technological resources necessary for effective program implementation. Without leadership commitment, the ASP's sustainability and effectiveness are compromised <sup>10</sup>. The second element highlights the Definition of Responsibilities, ensuring that all team members have clear roles and objectives. Appointing specific leaders and defining their responsibilities is essential to coordinating ASP activities and ensuring that program goals are achieved in an organized and efficient manner <sup>11</sup>.

Furthermore, the quality of relationships between the stakeholders involved in the ASP is crucial. A good relationship, based on positive behaviors, between the hospital's clinical leadership (senior management) and all professionals committed to AMR control is essential to the program's effectiveness. This relationship fosters the integration of all involved with a common goal: providing patient-centered care. Cooperation and effective communication between senior management and the clinical team foster a collaborative environment where ASP initiatives can be implemented more efficiently and with greater adherence by healthcare professional <sup>12,13</sup>.

The Brazilian scenario is particularly challenging in pediatric intensive care units (PICU), where antimicrobial resistance is a critical concern. Studies show that the rate of infections by multidrug-resistant pathogens is high, exacerbated by the extensive and often inappropriate use of antimicrobials<sup>14</sup>. Implementing ASP in Brazil is crucial, but it faces significant obstacles, including a lack of human and financial resources, technological limitations, and the need for specialized training for healthcare professionals<sup>15</sup>.

Measuring and monitoring antimicrobial use in pediatrics presents specific and complex challenges, especially when compared to the adult setting. Children are not just "small adults"; they have unique physiological and immunological characteristics, in addition to a distinct nosological profile, often marked by acute respiratory infections, gastroenteritis, urinary tract infections, exanthematous diseases, and febrile conditions of viral etiology, which are frequent causes of hospitalization in this population<sup>16</sup>.

This context demands individualized and judicious therapeutic approaches, since inappropriate use of antimicrobials, whether in terms of dose, duration, spectrum, or route of administration, can have significant impacts on the development of bacterial resistance and pediatric patient safety. Furthermore, the scarcity of robust data and the limited availability of effective monitoring systems hinder the accurate assessment of antimicrobial use and the true prevalence of microbial resistance in hospitalized pediatric patients<sup>14,16</sup>.

Therefore, strengthening ASP programs in PICUs is crucial to mitigating AMR in Brazil. These programs not only improve medication use but also promote patient safety and quality of care, contributing to a more resilient and sustainable healthcare system<sup>17,18</sup>. The objective of this study was to assess Leadership Support and Definition of Responsibilities for antimicrobial stewardship programs in pediatric intensive care units in Brazil.

## **M**ETHOD

This is a cross-sectional, multicenter study carried out in Brazilian hospitals.



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As the largest country in South America and the fifth largest in the world, Brazil has over 210 million inhabitants and a territorial extension of approximately 8.5 million km². Composed of 5,570 municipalities and 27 federative units (26 states and the Federal District), it is divided into five macro-regions (North, Northeast, Midwest, Southeast and South), based on natural, social, cultural, political, and economic criteria<sup>19</sup>. According to the Brazilian Health Regulatory Agency (ANVISA), in 2022, there were 662 hospitals with Pediatric ICUs (PICUs) in Brazil, with the Southeast region having 346 (52.26%), followed by the Northeast with 110 (16.62%) and South with 83 (12.54%).

The sample consisted of data from a segment of the 2nd national ASP survey in Brazil, coordinated by ANVISA/Ministry of Health. The study focused on the PICU, with an invitation sent by email to all hospitals in the three healthcare spheres (federal, state and regional). Recruitment was conducted through ANVISA's communication channels, and participants voluntarily filled out an electronic form (Google Forms®).

Data collection was conducted between October 2022 and January 2023 by filling out an electronic form sent to the PICUs in Brazil. The questionnaire made it possible to analyze the level of adherence to six essential elements recommended for structuring an effective ASP: (Component 1) Leadership Support; (Component 2) Definition of Responsibilities; (Component 3) Education; (Component 4) Monitoring of antimicrobial use; (Component 5) Evidence-based interventions for the rational use of antimicrobials; and (Component 6) Monitoring of process and outcome indicators<sup>20</sup>.

The data collection instrument assessed program implementation, classifying each component as inadequate, basic, intermediate, or advanced, based on the score obtained. Responses, which varied in points, included some questions with multiple answers, where all options selected within the same component were counted<sup>20</sup>. The survey, organized into sections, addressed the identification of the health service profile and the existence or absence of the ASP. The questionnaire contained 46 dichotomous questions (YES or NO) and 15 questions with multiple answer options, all mandatory depending on the ASP implementation.

A specific analysis was conducted only for hospitals with implemented ASP, focusing on components 1 (Leadership Support) and 2 (Definition of Responsibilities). Component 1 assessed the availability of human, financial, and technological resources, while component 2 examined the appointment of ASP team members and the definition of their leaders and roles (11). A sub-analysis was conducted regarding the existence of a team responsible for implementing the ASP in the hospital. Hospital characteristics such as location, service profile, hospital size, and the size of the PICU were also considered. Hospitals were classified as private (managed by private, charitable, or philanthropic entities) or public (managed by municipal, state, or federal government entities)<sup>21</sup>. Categorization was performed according to the number of hospital beds (up to 100, 101/199, and ≥200 beds), Brazilian regions, and the number of PICU beds (up to 20 and >20 beds)<sup>21</sup>.

All analyses were conducted using the Statistical Package for the Social Sciences (SPSS®), version 23 (IBM, Armonk, NY). The comparison of continuous scores for components 1 (Leadership Support) and 2 (Definition of Responsibilities) between the service profile and hospital size and PICU size was performed using two approaches: the first with parametric tests using the Bootstrapping procedure with 5,000 resamplings and the bias-corrected and accelerated method (BCa); and the second with non-parametric tests due to non-normal distribution of variables.

The comparison between the type of institution used Student's t-test for independent samples and Mann-Whitney's U test, while the comparisons between the number of available beds used one-way analysis of variance (ANOVA) and the Kruskal-Wallis test<sup>22</sup>.

Finally, comparisons of the relative frequencies of 'yes' responses between the type of institution and the number of available beds were performed using the a posteriori z-test in the chi-square analysis, adjusting p-values using the Bonferroni method, according to the number of comparisons between groups. p-values≤0.05 were considered statistically significant<sup>22</sup>.

The research protocol respected the ethical aspects related to research with human beings, according to National Health Council Resolution No. 466 dated December 12<sup>th</sup>, 2012, and was approved by the Research Ethics Committee.

## **RESULTS**

Of the 393 (59.36%) participating PICUs in Brazil, 219 that reported having implemented ASP were included in the final study sample and had their data analyzed. Of these, the overall level of adherence to components 1 (Leadership Support) and 2 (Definition of Responsibilities) differed between states, but the overall average for both was classified as basic (Figure 1).





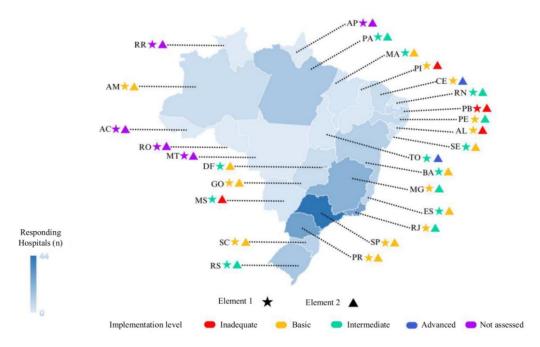


Figure 1: Number of hospitals that responded to the survey on the ASP in PICU implemented by state and adherence level to the Leadership Support and Definition of Responsibilities components (n=219). Brazil, 2023.

Component 1 scored 117.85 (±39.65) and level 2 scored 84.79 (±47.02). In level 1, the classifications "basic" with 78 institutions (35.62%) and "advanced" with 64 (29.22%) stood out, followed by "intermediate" with 61 (27.85%) and "inadequate" with 16 (7.31%). Level 2 also obtained the same classifications: "basic" with 88 (40.18%) and "advanced" with 49 (22.38%) of the institutions, followed by "intermediate" with 43 (19.63%) and "inadequate" with 39 (17.81%).

Figure 2 describes the findings related to the existence of a formal document approved by the institution's board of directors for the ASP.

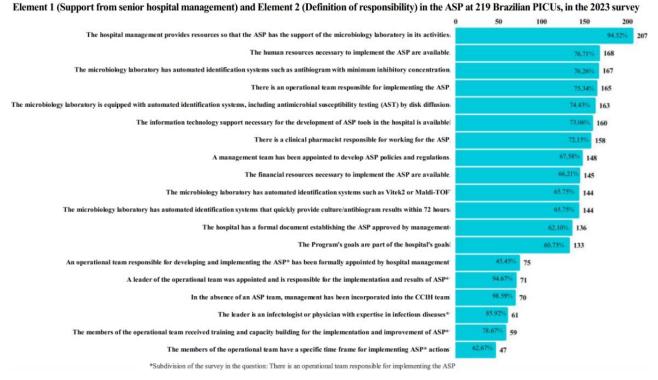


Figure 2: Number of hospitals that responded to the survey on the ASP in PICU implementation and the level of adherence to the Leadership Support and Definition of Responsibilities components (n=219). Brazil, 2023.





Regarding Component 1 (Leadership Support), 62.1% of hospitals have a formal document approved by the board of directors for the ASP, 60.73% have integrated the program's goals with the hospital's strategic goals, and 76.71% have the necessary human resources. Furthermore, 94.52% of hospitals have laboratory support, and 73.06% have technological support.

Regarding Component 2 (Definition of Responsibilities), 67.58% defined a management team for the ASP, 75.34% have an operational team, and 94.67% have a leader responsible for the program, with 85.92% of these leaders being infectologists. In addition, 72.15% of hospitals have clinical pharmacists in the ASP. Table 1 presents data related to the nature of the institutions.

**Table 1:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), according to the nature of the hospitals with the team responsible for implementing the ASP, according to the nature of the institution (n=165). Brazil, 2023.

	Pro	file	
	Private	Public	p-
Characterization of the operational team responsible for implementing the ASP	n(%)	n(%)	value*
The hospital has a formal document establishing the ASP approved by the management	85 (73.3)	31 (26.7)	0.02
The ASP goals are part of the hospital's goals	81 (77.1)	24 (22.9)	0.01
The financial resources necessary to implement the ASP are available	86 (72.9)	32 (27.1)	
The human resources necessary to implement the ASP are available	99 (74.4)	34 (25.6)	0.088
The hospital management provides resources so that the ASP has the support of the microbiology	119		
laboratory in its activities	(75.8)	38 (24.2)	
The information technology support necessary for the development of ASP tools in the hospital			
is available	94 (76.4)	29 (23.6)	
A management team has been appointed to develop ASP policies and regulations	97 (73.5)	35 (26.5)	
In the absence of an ASP team, management has been incorporated into the CCIH team	25 (75.8)	8 (24.2)	0.071
An operational team responsible for developing and implementing the ASP* has been formally			
appointed by hospital management	33 (70.2)	14 (29.8)	
The members of the operational team have a specific time frame for implementing ASP* actions	33 (70.2)	14 (29.8)	
The members of the operational team received training and capacity building for the implementation			
and improvement of ASP*	44 (74.6)	15 (25.4)	
A leader of the operational team was appointed and is responsible for the implementation and results			
of ASP*	55 (77.5)	16 (22.5)	
The leader is an infectologist or physician with expertise in infectious diseases*	47 (77.0)	14 (23.0)	
There is a clinical pharmacist responsible for working for the ASP	98 (77.2)	29 (22.8)	

It is observed that 70.05% of the responding institutions are private (n=160), while 59 are public (26.94%). Data distribution according to the Federation states is described in Tables 2, 3, 4, 5 and 6.

**Table 2:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), of the hospitals that had a team responsible for implementing the ASP in the states of the Southern Region (n=165). Brazil, 2023.

	PR	RS	SC
Characterization of the operational team responsible for implementing the ASP	n (%)	n (%)	n (%)
The hospital has a formal document establishing the ASP approved by the management	6 (5.2)	9 (7.8)	5 (4.3)
The ASP goals are part of the hospital's goals	10 (9.5)	7 (6.7)	3 (2.9)
The financial resources necessary to implement the ASP are available	10 (8.5)	8 (6.8)	6 (5.1)
The human resources necessary to implement the ASP are available	17 (12.8)	11 (8.3)	4 (3.0)
The hospital management provides resources so that the ASP has the support of the microbiology laboratory in its activities	19 (11.5)	12 (7.3)	7 (4.2)
The information technology support necessary for the development of ASP tools in the hospital is available	17 (13.8)	8 (6.5)	7 (5.7)
A management team has been appointed to develop ASP policies and regulations	14 (10.6)	12 (9.1)	6 (4.5)
In the absence of an ASP team, management has been incorporated into the CCIH team	5 (15.2)	0 (0.0)	1 (3.0)
An operational team responsible for developing and implementing the ASP* has been formally appointed by hospital management	6 (8.0)	4 (5.3)	1 (1.3)
The members of the operational team have a specific time frame for implementing ASP* actions	3 (6.5)	3 (6.4)	1 (2.1)
The members of the operational team received training and capacity building for the implementation and improvement of ASP*	6 (10.2)	2 (3.4)	1 (1.7)
A leader of the operational team was appointed and is responsible for the implementation and results of ASP*	6 (8.5)	4 (5.6)	1 (1.4)
The leader is an infectologist or physician with expertise in infectious diseases*	5 (8.2)	3 (4.9)	1 (1.6)
There is a clinical pharmacist responsible for working for the ASP	16 (12.6)	10 (7.9)	6 (4.7)

**Notes:** PR – Paraná; SC – Santa Catarina; RS – Rio Grande do Sul; \*Subdivision of the survey question: There is an operational team responsible for implementing ASP.





**Table 3:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), of the hospitals that had a team responsible for implementing the ASP in the states of the Southeast Region. Brazil, 2023.

	ES	MG	RJ	SP
Characterization of the operational team responsible for implementing the ASP	n (%)	n (%)	n (%)	n(%)
The hospital has a formal document establishing the ASP approved by the management	4 (3.4)	15 (12.9)	15 (12.9)	27 (23.3)
The ASP goals are part of the hospital's goals	6 (5.7)	11 (10.5)	18 (17.1)	16 (15.2)
The financial resources necessary to implement the ASP are available	7 (5.9)	13 (11.0)	16 (16.6)	25 (21.2)
The human resources necessary to implement the ASP are available	6 (4.5)	12 (9.0)	20 (15.0)	25 (18.8)
The hospital management provides resources so that the ASP has the support of the				
microbiology laboratory in its activities	7 (4.2)	17 (10.3)	20 (13.3)	32 (20.6)
The information technology support necessary for the development of ASP tools in the				
hospital is available	6 (4.9)	11 (8.9)	12 (9.8)	24 (19.5)
A management team has been appointed to develop ASP policies and regulations	5 (3.8)	15 (11.4)	18 (13.6)	23 (17.4)
In the absence of an ASP team, management has been incorporated into the CCIH team	2 (6.1)	2 (6.1)	4 (12.1)	11 (33.3)
An operational team responsible for developing and implementing the ASP* has been				
formally appointed by hospital management	3 (4.0)	10 (13.3)	10 (13.3)	16 (21.3)
The members of the operational team have a specific time frame for implementing				
ASP* actions	2 (4.3)	4 (8.5)	4 (8.5)	11 (23.4)
The members of the operational team received training and capacity building for the				
implementation and improvement of ASP*	2 (3.4)	9 (15.3)	6 (10.2)	11 (18.6)
A leader of the operational team was appointed and is responsible for the implementation				
and results of ASP*	2 (2.8)	9 (12.7)	9 (12.7)	15 (21.1)
The leader is an infectologist or physician with expertise in infectious diseases*	2 (3.3)	7 (11.5)	8 (13.1)	15 (24.6)
There is a clinical pharmacist responsible for working for the ASP	6 (4.7)	12 (9.4)	20 (15.7)	22 (17.3)

**Notes:** ES – Espírito Santo; MG – Minas Gerais; RJ – Rio de Janeiro; SP – São Paulo; \*Subdivision of the survey in the question: There is an operational team responsible for implementing the ASP.

**Table 4:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), of the hospitals that had a team responsible for implementing the ASP in the states of the Midwest Region. Brazil, 2023.

	DF	GO	MS	MT#
Characterization of the operational team responsible for implementing the ASP	n(%)	n(%)	n(%)	n(%)
The hospital has a formal document establishing the ASP approved by the management	6(5.2)	0(0.0)	1(0.9)	0(0.0)
The ASP goals are part of the hospital's goals	5(4.8)	1(1.0)	0(0.0)	0(0.0)
The financial resources necessary to implement the ASP are available	5(4.2)	1(0.8)	2(1.7)	0(0.0)
The human resources necessary to implement the ASP are available	5(3.8)	1(0.8)	1(0.8)	0(0.0)
The hospital management provides resources so that the ASP has the support of the microbiology laboratory in its activities	5(3.6)	2(1.2)	2(1.2)	0(0.0)
The information technology support necessary for the development of ASP tools in the hospital is available	6(4.9)	2(1.6)	2(1.6)	0(0.0)
A management team has been appointed to develop ASP policies and regulations	6(4.5)	0(0.0)	1(0.8)	0(0.0)
In the absence of an ASP team, management has been incorporated into the CCIH team	0(0.0)	2(6.1)	1(3.0)	0(0.0)
An operational team responsible for developing and implementing the ASP* has been formally appointed by hospital management	3(4.0)	1(1.3)	0(0.0)	0(0.0)
The members of the operational team have a specific time frame for implementing ASP* actions	3(6.4)	0(0.0)	0(0.0)	0(0.0)
The members of the operational team received training and capacity building for the implementation and improvement of ASP*	2(3.4)	1(1.7)	0(0.0)	0(0.0)
A leader of the operational team was appointed and is responsible for the implementation and results of ASP*	3(4.2)	1(1.4)	0(0.0)	0(0.0)
The leader is an infectologist or physician with expertise in infectious diseases*	3(4.3)	1(1.6)	0(0.0)	0(0.0)
There is a clinical pharmacist responsible for working for the ASP	6(4.7)	1(0.8)	0(0.0)	0(0.0)

**Notes:** DF – *Distrito Federal*; GO – Goiás; MS – Mato Grosso do Sul; MT – Mato Grosso; "States that did not participate in the survey; \*Subdivision of the survey in the question: There is an operational team responsible for implementing the ASP.





Research Article Artigo de Pesquisa Artículo de Investigación

**Table 5:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), of the hospitals that had a team responsible for implementing the ASP in the states of the Northeast Region. Brazil, 2023.

Characterization of the operational team responsible for	AL	ВА	CE	MA	PB	PE	PI	RN	SE
implementing the ASP	n(%)	n(%)	n(%)	n(%)	n(%)	n(%)	n(%)	n(%)	n(%)
The hospital has a formal document establishing the ASP									
approved by the management	0(0.0)	7(6.0)	4(3.4)	7(6.0)	0(0.0)	4(3.4)	0(0.0)	1(0.9)	2(01.7)
The ASP goals are part of the hospital's goals	0(0.0)	7(6.7)	5(4.8)	7(6.7)	0(0.0)	2(1.9)	0(0.0)	1(1.0)	2(1.9)
The financial resources necessary to implement the ASP are available	0(0.0)	8(6.8)	2(1.7)	7(5.9)	0(0.0)	4(3.4)	0(0.0)	0(0.0)	1(0.8)
The human resources necessary to implement the ASP are available	1(0.8)	8(6.0)	4(3.0)	8(6.0)	0(0.0)	3(2.3)	0(0.0)	1(0.8)	2(1.5)
The hospital management provides resources so that the ASP has									
the support of the microbiology laboratory in its activities	1(0.6)	7(4.8)	5(3.0)	9(5.5)	1(0.6)	5(3.6)	0(0.0)	1(0.6)	2(1.2)
The information technology support necessary for the									
development of ASP tools in the hospital is available	1(0.8)	6(4.9)	3(2.4)	9(7.3)	0(0.0)	5(4.1)	0(0.0)	1(0.8)	0(0.0)
A management team has been appointed to develop ASP policies									
and regulations	1(0.8)	6(4.5)	5(3.8)	8(6.1)	1(0.8)	4(3.0)	0(0.0)	1(0.8)	2(1.5)
In the absence of an ASP team, management has been									
incorporated into the CCIH team	0(0.0)	2(6.1)	0(0.0)	1(3.0)	0(0.0)	2(6.1)	0(0.0)	0(0.0)	0(0.0)
An operational team responsible for developing and									
implementing the ASP* has been formally appointed by									
hospital management	0(0.0)	4(5.3)	4(5.3)	7(9.3)	0(0.0)	3(4.0)	0(0.0)	0(0.0)	1(1.3)
The members of the operational team have a specific time frame									
for implementing ASP* actions	0(0.0)	2(4.3)	2(4.3)	7(14.9)	0(0.0)	2(4.3)	0(0.0)	0(0.0)	1(2.1)
The members of the operational team received training and capacity									
building for the implementation and improvement of ASP*	0(0.0)	4(6.8)	4(6.8)	6(10.2)	0(0.0)	2(3.4)	0(0.0)	0(0.0)	1(1.7)
A leader of the operational team was appointed and is responsible									
for the implementation and results of ASP*	0(0.0)	4(5.6)	4(5.6)	7(9.9)	0(0.0)	3(4.2)	0(0.0)	0(0.0)	1(1.4)
The leader is an infectologist or physician with expertise in									
infectious diseases*	0(0.0)	4(6.6)	1(1.6)	6(9.8)	0(0.0)	3(4.9)	0(0.0)	0(0.0)	1(1.6)
There is a clinical pharmacist responsible for working for the ASP	0(0.0)	6(4.7)	5(3.9)	8(6.3)	0(0.0)	3(2.4)	0(0.0)	1(0.8)	1(0.8)

**Notes:** AL – Alagoas; BA – Bahia; CE – Ceará; MA – Maranhão; PB – Paraíba; PE – Pernambuco; PI – Piauí; RN – Rio Grande do Norte; SE – Sergipe. \*Subdivision of the survey in the question: There is an operational team responsible for implementing the ASP.

**Table 6:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), of the hospitals that had a team responsible for implementing the ASP in the states of the Northern Region. Brazil, 2023.

	AC#	AM	AP#	RO#	RR#	то
Characterization of the operational team responsible for implementing the ASP	n(%)	n(%)	n(%)	n(%)	n(%)	n(%)
The hospital has a formal document establishing the ASP approved by the management	0(0.0)	1(0.9)	0(0.0)	0(0.0)	0(0.0)	1(0.9)
The ASP goals are part of the hospital's goals	0(0.0)	2(1.9)	0(0.0)	0(0.0)	0(0.0)	1(1.0)
The financial resources necessary to implement the ASP are available	0(0.0)	1(0.8)	0(0.0)	0(0.0)	0(0.0)	1(0.8)
The human resources necessary to implement the ASP are available	0(0.0)	2(1.5)	0(0.0)	0(0.0)	0(0.0)	1(0.8)
The hospital management provides resources so that the ASP has the support of the						
microbiology laboratory in its activities	0(0.0)	2(1.2)	0(0.0)	0(0.0)	0(0.0)	1(0.6)
The information technology support necessary for the development of ASP tools in the						
hospital is available	0(0.0)	2(1.6)	0(0.0)	0(0.0)	0(0.0)	1(0.8)
A management team has been appointed to develop ASP policies and regulations	0(0.0)	2(1.5)	0(0.0)	0(0.0)	0(0.0)	1(0.8)
In the absence of an ASP team, management has been incorporated into the CCIH team	0(0.0)	0(0.0)	0(0.0)	0(0.0)	0(0.0)	0(0.0)
An operational team responsible for developing and implementing the ASP* has been formally						
appointed by hospital management	0(0.0)	1(1.3)	0(0.0)	0(0.0)	0(0.0)	1(1.3)
The members of the operational team have a specific time frame for implementing						
ASP* actions	0(0.0)	1(2.1)	0(0.0)	0(0.0)	0(0.0)	1(2.1)
The members of the operational team received training and capacity building for the						
implementation and improvement of ASP*	0(0.0)	1(1.7)	0(0.0)	0(0.0)	0(0.0)	1(1.7)
A leader of the operational team was appointed and is responsible for the implementation						
and results of ASP*	0(0.0)	1(1.4)	0(0.0)	0(0.0)	0(0.0)	1(1.4)
The leader is an infectologist or physician with expertise in infectious diseases*	0(0.0)	0(0.0)	0(0.0)	0(0.0)	0(0.0)	1(1.6)
There is a clinical pharmacist responsible for working for the ASP	0(0.0)	2(1.6)	0(0.0)	0(0.0)	0(0.0)	1(0.8)

**Notes:** AC – Acre; AM – Amazonas; AP – Amapá; RO – Rondônia; RR – Roraima; TO – Tocantins; \*States that did not participate in the survey; \*Subdivision of the survey in the question: There is an operational team responsible for implementing the ASP.

The Southeast region has the largest number of hospitals with 98 units (44.75%), followed by the South with 49 (22.37%) and the Northeast with 36 (16.44%). In terms of capacity, most hospitals have 200 or more beds (n=112; 51.14%), with the Southeast region standing out, comprising 49 hospitals in this category (50%). Furthermore,





the Midwest region has the highest proportion of private hospitals (82.40%) and the South has the highest proportion of hospitals with more than 200 beds (55.10%). Regarding ICU sizes, 198 have up to 20 PICU beds (90.41%).

Tables 7 and 8 show the data related to the characteristics of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), according to the total number of beds and PICU of the hospitals studied.

**Table 7:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), according to the total number of beds in hospitals with a team responsible for implementing the ASP (n=165). Brazil, 2023.

	Hospital Beds (n - %)			
Characterization of the operational team responsible for implementing the ASP	Up to 100	101-199	>200	
The hospital has a formal document establishing the ASP approved by the management	16 (13.8)	34 (29.3)	66 (56.9)	
The ASP goals are part of the hospital's goals	7 (25.0)	7 (25.0)	14 (50.0)	
The financial resources necessary to implement the ASP are available	18 (15.3)	35 (29.7)	65 (55.1)	
The human resources necessary to implement the ASP are available	21 (15.8)	37 (27.8)	75 (56.4)	
The hospital management provides resources so that the ASP has the support of the microbiology				
laboratory in its activities	26 (16.6)	47 (29.9)	84 (53.5)	
The information technology support necessary for the development of ASP tools in the hospital				
is available	18 (14.6)	41 (33.3)	64 (52.0)	
A management team has been appointed to develop ASP policies and regulations	19 (14.4)	36 (27.3)	77 (58.3)	
In the absence of an ASP team. management has been incorporated into the CCIH team	10 (30.3)	12 (36.4)	11 (33.3)	
An operational team responsible for developing and implementing the ASP* has been formally				
appointed by hospital management	10 (13.3)	25 (33.3)	40 (53.3)	
The members of the operational team have a specific time frame for implementing ASP* actions	5 (10.6)	15 (31.9)	27 (57.4)	
The members of the operational team received training and capacity building for the				
implementation and improvement of ASP*	8 (13.6)	20 (33.9)	31 (52.5)	
A leader of the operational team was appointed and is responsible for the implementation and				
results of ASP*	9 (12.7)	23 (32.4)	39 (54.9)	
The leader is an infectologist or physician with expertise in infectious diseases*	9 (14.8)	18 (29.5)	34 (55.7)	
There is a clinical pharmacist responsible for working for the ASP	19 (15.0)	37 (29.1)	71 (51.9)	

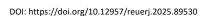
**Table 8:** Characterization of the operational team in the responses of components 1 (Leadership Support) and 2 (Definition of Responsibilities), according to the number of PICU beds of hospitals with a team responsible for implementing the ASP (n=165). Brazil, 2023.

PICU Beds (n - %)				
Characterization of the operational team responsible for implementing the ASP	Up to 20	Over 20	p-value	
The hospital has a formal document establishing the ASP approved by the management	105 (90.5)	11 (9.5)	0.02	
The ASP goals are part of the hospital's goals	95 (90.5)	10 (9.5)	0.01	
The financial resources necessary to implement the ASP are available	104 (88.1)	14 (11.9)		
The human resources necessary to implement the ASP are available	118 (88.7)	15 (11.3)	0.056	
The hospital management provides resources so that the ASP has the support of the microbiology				
laboratory in its activities	143 (91.1)	14 (8.9)		
The information technology support necessary for the development of ASP tools in the hospital				
is available	108 (87.8)	15 (12.2)		
A management team has been appointed to develop ASP policies and regulations	119 (90.2)	13 (9.8)	0.017	
In the absence of an ASP team, management has been incorporated into the CCIH team	30 (90.9)	3 (9.1)	0.017	
An operational team responsible for developing and implementing the ASP* has been formally				
appointed by hospital management	67 (89.3)	8 (10.7)		
The members of the operational team have a specific time frame for implementing ASP* actions	39 (83.0)	8 (17.0)		
The members of the operational team received training and capacity building for the				
implementation and improvement of ASP*	51 (86.4)	8 (13.6)		
A leader of the operational team was appointed and is responsible for the implementation and				
results of ASP*	63 (88.7)	8 (11.3)		
The leader is an infectologist or physician with expertise in infectious diseases*	53 (86.9)	8 (13.1)		
There is a clinical pharmacist responsible for working for the ASP	114 (89.8)	13 (10.2)		

Notes: \*Subdivision of the survey in the question: Is there an operational team responsible for implementing the ASP.

A total of 165 (75.34%) services were identified as having an operational team working in primary care. In terms of the hospitals' profile, most of those with such teams are private. The distribution of hospitals varies in







terms of capacity, ranging from small (up to 100 beds) to large (over 200 beds), with the majority having over 200 beds.

Regarding PICU beds, most hospitals have up to 20 pediatric ICU beds, with a smaller proportion having more than 20 beds. However, when analyzed individually, the group with more than 20 PICU beds stands out with a higher proportion of YES responses (66.06%) in most scenarios analyzed. Table 1 also presents the comparisons between hospitals with PICUs up to 20 beds and more than 20 beds for components 1 and 2, with p<0.05. Furthermore, variables were identified concerning nature with p values not less than 0.05, but which draw attention, such as: the existence of human resources to implement the ASP in the hospital (p=0.088) and the management of the ASP incorporated into the Hospital Infection Control Committee (p=0.071), indicating relevant aspects that deserve consideration in the health services analyzed.

#### **DISCUSSION**

An analysis of the ASP implementation levels in Brazilian states reveals significant variation, which can be correlated with the regions' GDP and population density. International studies highlight that regions with higher GDP and population density tend to have better infrastructure and resources for implementing AMR programs <sup>23,24</sup>. States such as São Paulo, Rio de Janeiro, and Minas Gerais have higher GDP and higher population density, providing better resources and support for ASP implementation. However, São Paulo is only at the basic level for both elements, indicating that, despite the available resources, there is still room for improvement in implementation, despite having a larger number of participants. On the other hand, states such as Acre, Amapá, Mato Grosso, Rondônia, and Roraima face additional challenges due to the lower availability of financial and technological resources, reflected in the inadequate level of implementation due to non-participation in the survey or non-adherence to the ASP.

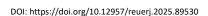
The findings on ASP implementation in PICU in Brazilian hospitals strongly correlate with recent international literature, highlighting both the progress and the challenges faced<sup>25,26</sup>. The support of senior hospital management, highlighted in Figure 2 with 94.52% of hospitals providing resources for the ASP, is essential for the success of the implementation<sup>27,28</sup>. This finding is corroborated by studies that emphasize the importance of institutional support and leadership for the effectiveness of ASP<sup>23,24,27</sup>. The availability of adequate human and financial resources for ASP implementation is another crucial factor (76.71% and 66.21%, respectively). These results are consistent with findings from other research indicating that resource availability is a significant barrier in Latin American and Caribbean countries<sup>24</sup>. Lack of funding and qualified personnel limits the effective implementation of these programs, as also observed in low- and middle-income countries<sup>28</sup>.

Furthermore, Information Technology (IT) support, present in 73.06% of Brazilian hospitals, reflects a global trend toward integrating technological tools into the ASP. A study demonstrated that the use of methodologies such as Lean Six Sigma, which rely heavily on data and technology, can optimize processes and reduce antimicrobial use without compromising clinical outcomes<sup>25</sup>. This alignment highlights the importance of IT in the continuous monitoring and assessment of programs. The presence of teams responsible for implementing ASP (75.34%) stands out, a strategy widely recognized in the literature as fundamental to the success of AMR programs. Other studies reinforce that the formation of multidisciplinary teams is vital to addressing AMR in a coordinated and effective manner<sup>29,30</sup>. The integration of clinical pharmacists, mentioned in 72.15% of Brazilian hospitals, is a recommended practice and has been successfully implemented in several regions of the world, including the Middle East<sup>25</sup>.

Despite notable progress in implementing ASP in Brazilian PICUs, some weaknesses have been revealed that deserve attention. A critical aspect is that only 66.21% of hospitals have the necessary financial resources to implement ASP. This financial limitation is a common challenge, especially in low- and middle-income countries. A review highlights that lack of adequate funding is a significant barrier in Latin America and the Caribbean, directly affecting the ability to maintain sustainable and effective programs<sup>24</sup>. Furthermore, several studies indicate that resource scarcity is a critical obstacle preventing the advancement of robust AMR programs in many parts of the world, especially in resource-limited settings<sup>31-34</sup>.

Another vulnerability identified is that only 67.58% of hospitals designated a manager responsible for developing ASP policies and regulations. Appointing dedicated leaders is essential to ensure the effective implementation and oversight of AMR programs. According to studies, institutional leadership is crucial to the success of programs, as it facilitates the coordination of multidisciplinary efforts and the implementation of evidence-based strategies. The absence of clear leadership can result in fragmented and less effective efforts in the fight against antimicrobial resistance <sup>23,26-28,34</sup>. This study also indicates that only 60.73% of the hospitals have







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ASP goals aligned with their strategic goals. This strategic alignment is crucial to ensuring that AMR programs receive the necessary support from all levels of hospital administration. Studies show that integrating ASP goals with the hospital's overall strategy improves guideline adherence and intervention success. A lack of alignment can lead to inappropriate prioritization of ASP activities, resulting in reduced effectiveness and impact on reducing inappropriate antimicrobial use<sup>23-27</sup>.

An analysis of the operational team responsible for implementing the ASP in 165 Brazilian hospitals reveals several strengths and weaknesses, as corroborated by international literature. It is important to note the differences and similarities in the factors assessed, both in terms of the nature of the hospitals (private vs. public) and their bed capacity. Private hospitals are more likely to have a formal document establishing the ASP and are more likely to integrate the ASP goals into the hospital's strategic goals, suggesting greater availability of formal structures and dedicated resources. Hospitals with larger bed capacities (>200) are significantly more likely to have a formal document and integrate ASP goals, reflecting greater administrative and resource capacity. Most hospitals with up to 20 PICU beds have a formal document and integrate ASP goals, indicating an operational need to standardize processes and ensure quality of care<sup>24,26,28</sup>.

Although some differences observed between private and public hospitals are not statistically significant (p>0.05), some aspects suggest that institutional nature may influence policy implementation and the integration of ASP goals. Therefore, although there was no robust statistical significance across all variables analyzed, the trend noticed indicates a need to consider the hospitals' profile when formulating and implementing these policies. Observed in 85 (73.30%) of the hospitals with p-value=0.02, the existence of a formal document establishing the ASP suggests an association between program formalization and its effective implementation. A study emphasizes that formal documentation is essential for the success of ASPs, as it eases resource mobilization and ensures health professionals' commitment<sup>23</sup>. This institutional support is corroborated by results from other countries that demonstrate the importance of formalizing policies for program effectiveness<sup>28,31,32</sup>.

The presence of teams responsible for implementing ASP in 105 (90.50%) of the hospitals with up to 20 beds is a crucial factor for the effectiveness of the programs, with p-value=0.020. This result is consistent with the literature, which highlights the importance of dedicated multidisciplinary teams in managing antimicrobial resistance. Studies indicate that well-structured teams with clear responsibilities are more effective in implementing ASP, especially in low- and middle-income settings<sup>23,28</sup>. The alignment of ASP goals with the hospital's strategic goals, observed in 81 (77.10%) of the private hospitals, presents p-value=0.01, indicating significant statistical significance.

The combined analysis of these results provides a comprehensive understanding of the factors that affected ASP implementation in Brazil, from internal hospital conditions to regional disparities. While the operational teams' characterization reveals the strengths and weaknesses within hospitals, the implementation in the states highlights the need for specific policies to overcome regional challenges and ensure effective implementation nationwide.

This survey offers valuable contributions to antimicrobial stewardship by providing a comprehensive assessment of ASP in PICUs in Brazil. Despite the existence of pediatric ASP in the country, its activities vary significantly, necessitating initiatives toward standardization. Understanding the impact of antimicrobial resistance (AMR) and identifying key weaknesses and strengths in prevention are essential to inform policy and institutional decisions tailored to each location and care setting. This is particularly important for reducing the use of broad-spectrum antibiotics. Incorporating ASP strategies is believed to promote the judicious use of antimicrobials, reducing infant morbidity and mortality and associated costs. This approach is expected to increase multidisciplinary team adherence to the program in clinical practice and improve safety regarding AMR in PICUs.

#### Study limitations

The study had some limitations due to the hospitals' voluntary participation. Most participating hospitals had ASPs, which may not adequately represent all hospitals with PICUs in Brazil. Responses to the survey varied across regions, which may compromise data representativeness when compared to each other. The specific characteristics of hospitals in terms of type of care and number of beds in different regions were not identified. The multiple-choice questions may not have covered all possible options, and it was not possible to add free-text responses. In addition, the lack of previous research in this context limited the discussion and comparison of the results obtained.



#### **CONCLUSION**

This study shows that the effectiveness of implementing ASPs in PICU in Brazil depends directly on effective support from institutional leadership and a clear Definition of Operational Responsibilities. The national analysis shows that, although relevant initiatives are underway, adherence to Components 1 (Leadership Support) and 2 (Definition of Responsibilities) remains uneven across states, being more consistent in regions with greater infrastructure and hospital management capacity.

Component 1, which reflects leadership commitment, is a critical pillar: where there is active and engaged leadership, there is more significant progress in ASP implementation. Component 2, which addresses the definition of roles and responsibilities, remains a weak point in many institutions, especially in states with less socioeconomic development, where inadequate levels of program structure persist.

The lack of committed technical leadership and formal accountability structures limits an ASP transformative potential in PICUs. Therefore, the program's existence on paper is not enough: genuine leadership engagement and institutionalized responsibilities are essential to ensure its effectiveness. Institutions that already have organized teams and clear leadership demonstrate that change is possible, provided there is strategic direction.

Therefore, to advance in tackling antimicrobial resistance in PICUs, it is essential that Brazil adopts structural measures, prioritizes training and retention of committed leaders and implements mechanisms that ensure definition and accountability of the teams involved. Only with this commitment will it be possible to consolidate ASPs as a robust and effective institutional policy, contributing to patient safety and strengthening public health in all territories.

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## Use of artificial intelligence tools

The authors declare that no artificial intelligence tools were used in the composition of the manuscript "Antimicrobial stewardship programs: Progress and challenges in Brazilian pediatric intensive care units".

