



cdc.gov.au/cdi • Electronic publication date: 18.11.2025 • doi.org/10.33321/cdi.2025.49.052

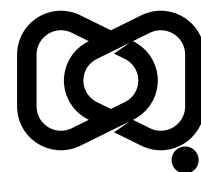
Australian Group on Antimicrobial Resistance (AGAR) surveillance outcome programs – bloodstream infections and antimicrobial resistance patterns in Australian children and adolescents, January 2022 – December 2023

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Australian Government

Department of Health,
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Communicable Diseases Intelligence (CDI) is a peer-reviewed scientific journal published by the interim Australian Centre for Disease Control within the Department of Health, Disability and Ageing.

The journal aims to disseminate information on the epidemiology, surveillance, prevention and control of communicable diseases of relevance to Australia and the near region.

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ISSN: 2209-6051 Online

This journal is indexed by Index Medicus and Medline.

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Abstract

Between January 2022 and December 2023, there were 1,827 bloodstream infection (BSI) isolates in 1,745 children and adolescents reported to the Australian Group on Antimicrobial Resistance (AGAR) surveillance outcome programs, with 40% of episodes in children aged < 12 months. Two-thirds of BSIs were community-onset.

Of 1,034 gram-negative isolates, 932 (90%) were Enterobacterales. Gram-negative BSI episodes were more commonly community-onset and in children < 12 months of age. Of Enterobacterales isolates, 17.9% were ciprofloxacin resistant; 14.0% were ceftriaxone and/or ceftazidime resistant; 9.5% were gentamicin and/or tobramycin resistant; and 8.9% were piperacillin-tazobactam resistant. Increasing ciprofloxacin resistance was noted, primarily due to the increase in *Salmonella* Typhi BSI. Overall, 13% of Enterobacterales were extended spectrum β -lactamase producers, and 18.5% were multi-drug resistant (MDR).

Of 601 *Staphylococcus aureus* isolates, 13.6% were methicillin-resistant (MRSA), and 5.5% were MDR. Overall, 14.4% of *S. aureus* isolates were erythromycin resistant; 10.3% were clindamycin resistant; and 5.0% were ciprofloxacin resistant. Erythromycin, clindamycin, and ciprofloxacin resistance in MRSA were significantly higher than in methicillin-sensitive isolates. No co-trimoxazole resistant *S. aureus* was isolated.

There were 192 enterococcal isolates reported; 70.8% were *E. faecalis* and 17.2% were *E. faecium*. All ampicillin-resistant, vancomycin-resistant, and MDR enterococci were *E. faecium*.

The 2022–2023 AGAR Kids Biennial Report shows relative stability in the antimicrobial resistance landscape within the Australian paediatric population, with few significant differences detected when compared to the 2020–2021 report. Small increases in the proportion of resistant Enterobacterales and *Enterococcus* spp. isolates highlight the importance of ongoing surveillance to inform stewardship and infection prevention interventions.

Keywords: Australian Group on Antimicrobial Resistance (AGAR); antimicrobial resistance surveillance; paediatrics; bacteraemia; Enterobacterales; *Staphylococcus aureus*; *Enterococcus*

Introduction

The Australian Group on Antimicrobial Resistance (AGAR) performs targeted whole-of-population antimicrobial resistance (AMR) surveillance in *Staphylococcus aureus*, *Enterococcus* spp., and key gram-negative bloodstream isolates. In 2023, a total of 51 hospitals reported bloodstream infection (BSI) data to AGAR, with all Australian states and mainland territories represented and all tertiary paediatric hospitals participating.

The AGAR reports feed into the key pillar objective #5 'Integrated Surveillance and Response to Resistance and Usage' of the National Antimicrobial Resistance Strategy of Australia.¹ The reports allow healthcare professionals and policy makers to 'use evidence-based surveillance and monitoring data to inform actions and responses to contain antimicrobial resistance'.¹

The AGAR Kids committee, formed in 2022, was established to investigate and present neonatal- and paediatric-specific AMR data from the AGAR programs. As called for by the World Society for Paediatric Infectious Diseases (WSPID),^{2,3} accessible neonatal and paediatric data is required to optimise prescribing and antimicrobial stewardship efforts. The first AGAR Kids Biennial report described the epidemiology of AMR in bacterial isolates, reported to the AGAR programs from children and adolescents < 18 years of age, in 2020 and 2021.^{4,5} A subsequent report from the AGAR Kids committee described the trends and shifts in AMR across nine years (2013–2021).^{6–8}

This second biennial AGAR Kids report describes the epidemiology, antimicrobial susceptibility patterns and antimicrobial resistance trends of bacterial isolates, and the patient outcomes of children and adolescents < 18 years old with bacterial BSIs as reported to AGAR in 2022–2023.

Methods

From 1 January 2022 to 31 December 2023, participating laboratories submitted unique BSI episodes of *S. aureus*, *Enterococcus* spp., Enterobacterales, *Pseudomonas aeruginosa*, and *Acinetobacter* spp.

Participants

Forty-one laboratories processing blood cultures from 53 hospitals submitted data from patients < 18 years old to the AGAR surveillance outcome programs: 52 hospitals in 2022 and 51 hospitals in 2023. All tertiary paediatric hospitals (n = 8), and 25 of the 29 principal referral centres in Australia, reported data during the two years, as well as five public acute group A hospitals, three private group A hospitals, two private group B hospitals and all public group C hospitals in the north-west of Western Australia.

Data collection

A BSI episode was defined as a clinical event associated with a positive blood culture, irrespective of the number of bacterial species identified. A new episode of bacteraemia in the same patient was recorded if the blood culture was collected more than 14 days after the initial positive culture. An episode was defined as community-onset if the first positive blood culture of the episode was collected \leq 48 hours after hospital admission, and as hospital-onset if collected > 48 hours after admission.

The data submitted by each laboratory to AGAR for each episode included a laboratory accession number; the date of collection; the organism isolated (genus and species); and the antimicrobial susceptibility test results (minimum inhibitory concentrations [MICs]). The patient's age, sex and jurisdiction of residence were also provided. If the patient was admitted to hospital, the dates of admission and discharge were reported. Depending on the laboratory's level of participation in AGAR, limited clinical and outcome data were also provided, including the principal clinical manifestation, device related infection (yes or no), and the outcome (died, all-cause or survived) at seven and 30 days. For the AGAR Kids Biennial Report, data from AGAR were restricted to patients aged < 18 years (0–17 years inclusive).

Laboratory methods

Where possible, isolates were identified to species level by each laboratory, using the laboratory's routine method. Species identification methods included the Vitek[®] and BD Phoenix[™] automated microbiology systems, and matrix assisted laser desorption ionization—time of flight (MALDI-TOF) mass spectrometry (Bruker MALDI biotyper[®] or Vitek[®] MS). Antimicrobial susceptibility testing was performed using one of two commercial semi-automated systems: the Vitek[®] 2 (bioMérieux) or BD Phoenix[™] (BD), both of which are calibrated to the ISO (International Organization for Standardization) reference standard method of broth microdilution.

All methicillin-resistant *S. aureus* (MRSA) isolates, all *E. faecium* isolates, and select gram-negative isolates with specific antimicrobial resistant profiles underwent whole genome sequencing (WGS) to determine the presence of specific antimicrobial resistant determinants, including cefotaximases (e.g. CTX-15); plasmid mediated AmpC β -lactamases; and carbapenemases such as New Delhi metallo- β -lactamases (NDMs). A detailed description of the molecular laboratory methods employed by AGAR is available in previously published reports.^{6,7,9}

Data analysis

The AMR for R package (v2.0) was used to transform MIC data as per European Committee on Antimicrobial Susceptibility Testing (EUCAST) 2024 (v14) breakpoints.^{10,11} Multi-drug resistance was defined as resistance to one or more agents in three or more antimicrobial classes. Intrinsic resistance mechanisms were not included in determining if an organism was multi-drug resistant (MDR). Apart from *Enterobacter* spp., Enterobacterales were phenotypically classified as an extended spectrum β -lactamase (ESBL) producer if the organism had a ceftriaxone and/or ceftazidime MIC \geq 2 mg/L. For *Enterobacter* spp., isolates with a cefepime MIC \geq 1 mg/L were classified as ESBL-producing.

Descriptive statistics for the overall population and per year were stratified by age, sex, and state/territory. Proportions were only calculated when the total number for a category was > 10. Categorical data were compared using the chi-square or Fisher's exact test. Continuous data were compared using the Mann-Whitney U-test. Associations between exposure and outcome were estimated using odds ratios (OR). Proportions for the organisms classified in the World Health Organisation (WHO) Bacterial Priority Pathogen List (BPPL) were calculated with 95% confidence intervals (95% CI).¹²

Ethics

Approval to conduct the prospective data collection was given by the research ethics committee associated with each participating healthcare facility.

Results

Patient characteristics

Between 1 January 2022 and 31 December 2023, there were 1,827 isolates, from 1,745 children and adolescents < 18 years old, reported to the AGAR surveillance outcome programs. Across all Australian states and mainland territories, Enterobacterales were the most frequently reported isolates (n = 932; 51.0%). Over the two years, *Staphylococcus aureus* (n: 601) and *Escherichia coli* (n: 457) comprised more than half of all isolates reported to AGAR (57.9%; 1,058/1,827). The commonest isolates remained unchanged between 2020–2021 and 2022–2023, with the exception of *Salmonella* Typhi for which significantly more isolates were reported in 2022–2023 than in 2020–2021 (50 vs 14 isolates; χ^2 : 17; $p < 0.001$; 95% CI: 1.0–2.8) (see Appendix A, Tables A.1.1 and A.1.2).⁴ Approximately half of the *S. Typhi* isolates were from children aged 5–11 years (n = 23; 46.0%); a similar proportion were from those living in Victoria (24/50; 48.0%).

Over the two years, 40.5% of episodes were in children aged < 12 months (n = 706/1,745), with a quarter of all episodes occurring in children \leq 90 days old. Two-thirds of all episodes were community-onset (n = 1,184; 67.8%). However, as observed in 2020–2021, *Enterococcus* spp. BSIs were more frequently hospital-onset than were BSIs involving *S. aureus* or Enterobacterales (67.6% vs 27.9%; χ^2 : 117, $p < 0.001$). As a proportion of total BSIs, hospital-onset infections were highest in Victoria (41.7%) and New South Wales (36.4%), in children < 12 months old (44.1%), in device-related infections (60.1%), and in BSIs involving at least one MDR isolate (87.5%) (Table 1).

The most frequently reported primary clinical manifestations were febrile neutropenia (n = 218; 12.5%), osteomyelitis/septic arthritis (n = 213; 12.2%), and urinary tract infections (UTI; n = 204; 11.7%). Episodes with at least one MDR isolate were most frequently reported in children and adolescents with febrile neutropenia (53/213; 24.3%) or UTIs (26/204; 12.7%) reported as the principal clinical manifestation.

Overall, 19.3% of episodes were device-related; a decrease from the 23.5% of episodes in the previous report χ^2 : 9; $p = 0.003$). This decrease was primarily observed in Victoria, which decreased from 32.4% in 2020–2021 to 20.7% in 2022–2023 (χ^2 : 16; $p < 0.001$). Thirteen percent of BSI episodes were polymicrobial (227/1,745); such episodes were more commonly observed in hospital-onset infections ($p = 0.001$) (Table 1).

The median length of stay (LoS) following blood culture collection was 11 days (interquartile range, IQR: 6–22 days) (Table 1). Children and adolescents with an episode involving at least one MDR isolate remained in hospital longer than did patients who did not have a MDR isolate (14 days vs 9 days; χ^2 : 16; $p < 0.001$).

One in five patients were still admitted to hospital at thirty days post-blood culture collection (380/1,745; 21.8%) (Table 1). Remaining in hospital thirty days post-collection was more frequent in children and adolescents with either an enterococcal BSI (OR: 2.7; 95% CI: 1.9–3.8), aged < 12 months (OR: 2.6; 95% CI: 2.0–3.2), a hospital-onset infection (OR: 11.2; 95% CI: 8.6–14.6), or a device-related infection (OR: 2.3; 95% CI: 1.7–2.9).

Over the two years, 46 children and adolescents died with a BSI (2.6%). *E. coli* was reported in a quarter of patients who died (n = 13/46; 28.3%). Seven of the patients who died had an MDR isolate (15.2%) (Table 2).

Table 1: Characteristics of patients aged < 18 years with a bacteraemic event reported to the Australian Group on Antimicrobial Resistance, per survey, January 2022 – December 2023

| Category | Characteristic | <i>Enterococcus</i> spp. (N = 185) | | <i>S. aureus</i> (N = 593) | | Gram-negative bacteria (N = 967) | | Total patients (N = 1,745) ^a | |
|---------------------------------|------------------|---------------------------------------|------|-------------------------------|------|-------------------------------------|------|--|------|
| | | n | % | n | % | n | % | n | % |
| Age (years) | Median | < 1 | — | 6 | — | 1 | — | 2 | — |
| | IQR ^b | 0–5 | — | 0–12 | — | 0–9 | — | 0–10 | — |
| Age group | ≤ 28 days | 46 | 24.9 | 50 | 8.4 | 194 | 20.1 | 290 | 16.6 |
| | 29–90 days | 29 | 15.7 | 47 | 7.9 | 105 | 10.9 | 181 | 10.4 |
| | 91–364 days | 32 | 17.3 | 59 | 9.9 | 144 | 14.9 | 235 | 13.5 |
| | 1– 4 years | 30 | 16.2 | 110 | 18.5 | 186 | 19.2 | 326 | 18.7 |
| | 5–11 years | 16 | 8.6 | 174 | 29.3 | 168 | 17.4 | 358 | 20.5 |
| | 12–17 years | 32 | 17.3 | 153 | 25.8 | 170 | 17.6 | 355 | 20.3 |
| Sex | Female | 75 | 40.5 | 237 | 40.0 | 411 | 42.5 | 723 | 41.4 |
| | Male | 110 | 59.5 | 356 | 60.0 | 556 | 57.5 | 1,022 | 58.6 |
| Jurisdiction ^c | ACT | 5 | 2.7 | 19 | 3.2 | 30 | 3.1 | 54 | 3.1 |
| | NSW | 70 | 37.8 | 209 | 35.2 | 336 | 34.7 | 615 | 35.2 |
| | NT | 4 | 2.2 | 36 | 6.1 | 32 | 3.3 | 72 | 4.1 |
| | Qld | 10 | 5.4 | 70 | 11.8 | 102 | 10.5 | 182 | 10.4 |
| | SA | 17 | 9.2 | 36 | 6.1 | 68 | 7.0 | 121 | 6.9 |
| | Tas. | 5 | 2.7 | 16 | 2.7 | 29 | 3.0 | 50 | 2.9 |
| | Vic. | 53 | 28.6 | 135 | 22.8 | 260 | 26.9 | 448 | 25.7 |
| | WA | 21 | 11.4 | 72 | 12.1 | 110 | 11.4 | 203 | 11.6 |
| Onset location | Community | 60 | 32.4 | 466 | 78.6 | 658 | 68.0 | 1,184 | 67.8 |
| | Hospital | 125 | 67.6 | 127 | 21.4 | 309 | 32.0 | 561 | 32.2 |
| Device-related BSI ^d | Yes | 72 | 39.6 | 95 | 16.5 | 170 | 19.4 | 337 | 19.3 |
| | No | 111 | 59.4 | 460 | 77.2 | 733 | 74.0 | 1,304 | 74.7 |
| | Unknown | 2 | 1.0 | 38 | 6.3 | 64 | 6.6 | 104 | 6.0 |

| Category | Characteristic | <i>Enterococcus</i> spp. (N = 185) | | <i>S. aureus</i> (N = 593) | | Gram-negative bacteria (N = 967) | | Total patients (N = 1,745) ^a | |
|--------------------------------|----------------|---------------------------------------|------|-------------------------------|------|-------------------------------------|------|--|------|
| | | n | % | n | % | n | % | n | % |
| Polymicrobial BSI ^d | — | 72 | 38.9 | 54 | 9.1 | 101 | 10.4 | 227 | 13.0 |
| Length of stay (days) | Median | 16.5 | — | 12 | — | 10 | — | 11 | — |
| | IQR | 7–35.2 | — | 7–23 | — | 5–20 | — | 6–22 | — |
| All-cause mortality at 30 days | Died | 7 | 3.8 | 7 | 1.2 | 32 | 3.3 | 46 | 2.6 |
| | Survived | 151 | 81.6 | 435 | 73.4 | 681 | 70.4 | 1,267 | 72.6 |
| | Unknown | 27 | 14.6 | 151 | 25.5 | 254 | 26.3 | 432 | 24.8 |

- a The total column may not equate to the sum of each organism group, as some patients have polymicrobial bacteraemic episodes and are reported in each respective column to their episode but only counted once in the total number of patients.
- b IQR: interquartile range.
- c ACT: Australian Capital Territory; NSW: New South Wales; NT: Northern Territory; Qld: Queensland; SA: South Australia; Tas: Tasmania; Vic: Victoria; WA: Western Australia.
- d BSI: bloodstream infection.

Table 2: Characteristics of patients aged < 18 years with a bacteraemic event reported to the Australian Group on Antimicrobial Resistance for which one or more isolate was multidrug resistant, *Enterobacteriales* isolates that produced extended-spectrum β -lactamases, or *Staphylococcus aureus* isolates that were methicillin-resistant, January 2022 – December 2023

| Category | Characteristic | <i>Enterobacteriales</i> ^{a,b} | | | | | | | | <i>Enterococcus</i> ^a | | | | <i>Staphylococcus aureus</i> ^{a,c} | | | | | | | |
|---------------------------|------------------|---|------|---------|------|------|------|----------|------|----------------------------------|------|---------|------|---|------|------|------|------|------|---------|------|
| | | MDR | | Not MDR | | ESBL | | Not ESBL | | MDR | | Not MDR | | MRSA | | MSSA | | MDR | | Not MDR | |
| | | n | % | n | % | n | % | n | % | n | % | n | % | n | % | n | % | n | % | n | % |
| Overall | — | 172 | 18.5 | 760 | 81.5 | 125 | 13.4 | 807 | 86.6 | 33 | 5.5 | 568 | 94.5 | 82 | 13.6 | 519 | 86.4 | 15 | 7.8 | 177 | 92.2 |
| Age (years) | Median | 2 | — | 1 | — | 1 | — | 1 | — | 1 | — | 6 | — | 5 | — | 6 | — | 13 | — | < 1 | — |
| | IQR ^d | 0–11 | — | 0–8 | — | 0–11 | — | 0–8 | — | 0–4 | — | 0–12 | — | 1–10 | — | 0–12 | — | 1–15 | — | 0–4 | — |
| Age group | ≤ 28 days | 40 | 20.7 | 153 | 79.3 | 32 | 16.6 | 161 | 83.4 | 9 | 17.3 | 43 | 82.7 | 5 | 9.6 | 47 | 90.4 | 1 | 2.1 | 46 | 97.9 |
| | 29–90 days | 12 | 11.1 | 96 | 88.9 | 10 | 9.3 | 98 | 90.7 | 2 | 4 | 48 | 96 | 5 | 10 | 45 | 90 | 1 | 3.4 | 28 | 96.6 |
| | 91–364 days | 20 | 14 | 123 | 86 | 18 | 12.6 | 125 | 87.4 | 5 | 8.3 | 55 | 91.7 | 8 | 13.3 | 52 | 86.7 | 1 | 2.9 | 33 | 97.1 |
| | 1–4 years | 35 | 21 | 132 | 79 | 23 | 13.8 | 144 | 86.2 | 9 | 8.1 | 102 | 91.9 | 23 | 20.7 | 88 | 79.3 | 1 | 3.1 | 31 | 96.9 |
| | 5–11 years | 26 | 17.1 | 126 | 82.9 | 15 | 9.9 | 137 | 90.1 | 7 | 4 | 167 | 96 | 26 | 14.9 | 148 | 85.1 | 1 | 6.3 | 15 | 93.7 |
| | 12–17 years | 39 | 23.1 | 130 | 76.9 | 27 | 16 | 142 | 84 | 1 | 0.6 | 153 | 99.4 | 15 | 9.7 | 139 | 90.3 | 10 | 29.4 | 24 | 70.6 |
| Sex | Female | 62 | 15.7 | 334 | 84.3 | 47 | 11.9 | 349 | 88.1 | 16 | 6.6 | 226 | 93.4 | 37 | 15.3 | 205 | 84.7 | 5 | 6.4 | 73 | 93.6 |
| | Male | 110 | 20.5 | 426 | 79.5 | 78 | 14.6 | 458 | 85.4 | 17 | 4.7 | 342 | 95.3 | 45 | 12.5 | 314 | 87.5 | 10 | 8.8 | 104 | 91.2 |
| Jurisdiction ^e | ACT | 6 | 18.8 | 26 | 81.3 | 6 | 18.8 | 26 | 81.2 | 0 | — | 19 | 100 | 1 | 5.3 | 18 | 94.7 | 0 | — | 5 | — |
| | NSW | 69 | 21.3 | 255 | 78.7 | 60 | 18.5 | 264 | 81.5 | 12 | 5.7 | 200 | 94.3 | 22 | 10.4 | 190 | 89.6 | 6 | 8.1 | 68 | 91.9 |
| | NT | 5 | 16.1 | 26 | 83.9 | 3 | 9.7 | 28 | 90.3 | 4 | 11.1 | 32 | 88.9 | 19 | 52.8 | 17 | 47.2 | 1 | — | 3 | — |
| | Qld | 6 | 6.1 | 92 | 93.9 | 4 | 4.1 | 94 | 95.9 | 3 | 4.3 | 67 | 95.7 | 4 | 5.7 | 66 | 94.3 | 1 | 10 | 18 | 90 |
| | SA | 14 | 23 | 47 | 77 | 10 | 16.4 | 51 | 83.6 | 1 | 2.6 | 38 | 97.4 | 4 | 10.3 | 35 | 89.7 | 0 | — | 5 | — |
| | Tas. | 4 | 13.3 | 26 | 86.7 | 2 | 6.7 | 28 | 93.3 | 0 | — | 16 | 100 | 0 | — | 16 | 100 | 0 | — | 46 | 100 |
| | Vic. | 53 | 21 | 199 | 79 | 33 | 13.1 | 219 | 86.9 | 13 | 9.6 | 123 | 90.4 | 18 | 13.2 | 118 | 86.8 | 7 | 13.2 | 23 | 86.8 |
| | WA | 15 | 14.4 | 89 | 85.6 | 7 | 6.7 | 97 | 93.3 | 0 | — | 73 | 100 | 14 | 19.2 | 59 | 80.8 | 0 | — | 23 | 100 |
| Onset location | Community | 105 | 16.3 | 539 | 83.7 | 70 | 10.9 | 574 | 89.1 | 19 | 4.1 | 450 | 95.9 | 68 | 14.5 | 401 | 85.5 | 1 | 1.6 | 61 | 98.4 |
| | Hospital | 67 | 23.3 | 221 | 76.7 | 55 | 19.1 | 233 | 80.9 | 14 | 10.6 | 118 | 89.4 | 14 | 10.6 | 118 | 89.4 | 14 | 10.8 | 116 | 89.2 |

| Category | Characteristic | <i>Enterobacterales</i> ^{a,b} | | | | | | | | <i>Enterococcus</i> ^a | | | | <i>Staphylococcus aureus</i> ^{a,c} | | | | | | | |
|---------------------------------|------------------|--|------|---------|------|------|------|----------|------|----------------------------------|-----|---------|------|---|------|------|------|------|------|---------|------|
| | | MDR | | Not MDR | | ESBL | | Not ESBL | | MDR | | Not MDR | | MRSA | | MSSA | | MDR | | Not MDR | |
| | | n | % | n | % | n | % | n | % | n | % | n | % | n | % | n | % | n | % | n | % |
| Device-related BSI ^f | Yes | 34 | 20 | 136 | 80 | 29 | 17.1 | 141 | 82.9 | 7 | 7.1 | 92 | 92.9 | 8 | 8.1 | 91 | 91.9 | 5 | 6.6 | 71 | 93.4 |
| | No | 126 | 18.1 | 569 | 81.9 | 88 | 12.7 | 607 | 87.3 | 22 | 4.7 | 442 | 95.3 | 65 | 14 | 399 | 86 | 10 | 9.6 | 104 | 90.4 |
| | Unknown | 12 | 17.9 | 55 | 82.1 | 8 | 11.9 | 59 | 88.1 | 3 | 8.1 | 34 | 91.9 | 9 | 23.7 | 29 | 76.3 | 0 | — | 2 | 100 |
| Polymicrobial BSI ^f | — | 26 | 18.8 | 112 | 81.2 | 16 | 11.6 | 122 | 88.4 | 2 | 3.2 | 60 | 96.8 | 5 | 8.1 | 57 | 91.9 | 4 | 5.1 | 75 | 94.9 |
| Length of stay (days) | Median | 14 | — | 9 | — | 15 | — | 9 | — | 17 | — | 11 | — | 16 | — | 11 | — | 17 | — | 17 | — |
| | IQR ^d | 8–24 | — | 4–17 | — | 8–28 | — | 4–18 | — | 8–39 | — | 7–22 | — | 8–29 | — | 7–22 | — | 9–24 | — | 7–36 | — |
| All-cause mortality at 30 days | Died | 5 | 17.9 | 23 | 82.1 | 5 | 17.9 | 23 | 82.1 | 0 | — | 7 | 100 | 3 | — | 4 | — | 1 | 14.3 | 6 | 85.7 |
| | Survived | 134 | 20.7 | 512 | 79.3 | 98 | 15.2 | 548 | 84.8 | 26 | 5.9 | 414 | 94.1 | 64 | 14.5 | 376 | 85.5 | 14 | 8.9 | 144 | 91.1 |
| | Unknown | 33 | 12.8 | 225 | 87.2 | 22 | 8.5 | 236 | 91.5 | 7 | 4.5 | 147 | 95.5 | 15 | 9.7 | 139 | 90.3 | 0 | — | 27 | 100 |

a MDR: multidrug resistant.

b ESBL: extended-spectrum β-lactamase.

c. MRSA: methicillin-resistant *Staphylococcus aureus*. MSSA: methicillin-susceptible *Staphylococcus aureus*.

d IQR: interquartile range.

e ACT: Australian Capital Territory; NSW: New South Wales; NT: Northern Territory; Qld: Queensland; SA: South Australia; Tas: Tasmania; Vic: Victoria; WA: Western Australia.

f BSI: bloodstream infection.

Gram negatives

There were 1,034 gram-negative bacterial isolates from 967 episodes reported in children and adolescents to AGAR in 2022–2023; this was more than the 902 isolates from 867 episodes reported in 2020–2021. Ninety percent of gram-negative isolates were identified as Enterobacterales ($n = 932/1,034$, 90.1%); 6.6% were *Pseudomonas aeruginosa* ($n = 68/1,034$); and 3.3% were *Acinetobacter* spp. ($n = 34/1,034$). A majority of BSI episodes with gram-negative bacteria were community-onset (68.0%) and in males (57.5%); infants < 12 months of age accounted for the highest proportion of gram-negative BSI episodes (48.4%) (Table 1).

Enterobacterales

Overall, 17.9% of Enterobacterales were ciprofloxacin resistant; 14.0% were ceftriaxone and/or ceftazidime resistant; 9.5% were gentamicin and/or tobramycin resistant; and 8.9% were piperacillin-tazobactam resistant. Only seven Enterobacterales isolates were meropenem resistant (0.8%). No meropenem-resistant isolates were reported in infants < 12 months old (Table 3).

The proportion of ciprofloxacin-resistant Enterobacterales was significantly higher than reported in 2020–2021 (17.9% vs 13.2%; $\chi^2: 7$; $p = 0.009$). However, if *Salmonella* spp. were excluded from the analysis, the ciprofloxacin-resistant proportion of isolates was similar in 2022–2023 to the proportions reported in 2020–2021 (12.7% vs 12.5%). Similarly, if *Salmonella* spp. were removed from the analysis, the proportions of ciprofloxacin-resistant Enterobacterales in Victoria (27.1% including *Salmonella* spp. vs 19.3% excluding *Salmonella* spp.; $\chi^2: 4$; $p = 0.05$), in neonates (17.2% vs 8.3%; $\chi^2: 5$; $p = 0.02$), and in community-onset BSIs (19.3% vs 8.1%, $\chi^2: 29$, $p < 0.001$) were all similar to the 2021–2022 reporting period.

For *Salmonella* spp., almost half of the isolates reported were ciprofloxacin resistant (67/143; 46.9%; 95% CI: 38.5–55.4); a significant increase when compared to 2020–2021 (46.9% vs 20.6%; $\chi^2: 12$; $p = 0.007$). The increase was primarily attributed to the increase in *S. Typhi* reported as a proportion of all *Salmonella* isolates during the reporting period (34.0% vs 15.6% of all *Salmonella* isolates were *S. Typhi*), given that 70.1% of ciprofloxacin-resistant *Salmonella* isolates in 2022–2023 were *S. Typhi* (47/67). Of the *S. Typhi* isolates, 94% were ciprofloxacin resistant (47/50), compared with only 9.9% of non-typhoidal *Salmonella* (8/81; $\chi^2: 86.4$; $p < 0.001$). There were only three ceftriaxone-resistant *Salmonella* spp. reported in 2022–2023, including one *S. Typhi*.

One hundred and twenty-five (13.4%) of the 932 Enterobacterales had an ESBL phenotype. In the neonatal population there was a significant increase in the proportion of Enterobacterales with an ESBL phenotype, from 8.3% in 2020–2021 to 16.6% in 2022–2023 ($\chi^2: 5$, $p = 0.03$). Additionally, an ESBL phenotype was identified more frequently in patients with a hospital-onset BSI than in patients with a community-onset BSI (OR: 1.9; 95% CI: 1.3–2.8; $p < 0.001$) (Table 2).

The proportion of MDR Enterobacterales increased from 14.5% in 2020–2021 to 18.5% in 2022–2023 ($n = 172/932$; $\chi^2: 5$; $p = 0.03$). Over two thirds of MDR Enterobacterales were identified as *E. coli* (69.8%; 120/172), and one in four *E. coli* were MDR (26.3%; $n = 120/457$). The likelihood of an Enterobacterales isolate being MDR was greater for hospital-onset BSIs (OR: 1.6; 95% CI: 1.1–2.2; $p = 0.01$). The median length of stay for patients with an MDR Enterobacterales BSI was significantly longer than for those with a non-MDR Enterobacterales BSI (14 days vs 9 days; $p < 0.001$) (Table 2).

Table 3: Number of *Enterobacteriales* isolates tested and proportion of isolates resistant to gentamicin/tobramycin, ceftriaxone/ceftazidime, piperacillin–tazobactam, and ciprofloxacin from patients aged < 18 years as reported to the Australian Group on Antimicrobial Resistance, January 2022 – December 2023

| Category | Characteristic | Gentamicin/tobramycin ^a | | | | Ceftriaxone/ceftazidime ^a | | | | Piperacillin–tazobactam ^a | | | | Ciprofloxacin ^a | | | |
|---------------------------|----------------|------------------------------------|----|------|-----------|--------------------------------------|-----|------|-----------|--------------------------------------|----|------|-----------|----------------------------|-----|------|-----------|
| | | N | n | R% | 95% CI | N | n | R% | 95% CI | N | n | R% | 95% CI | N | n | R% | 95% CI |
| Overall | — | 927 | 88 | 9.5 | 7.7–11.6 | 927 | 130 | 14.0 | 11.9–16.4 | 917 | 82 | 8.9 | 7.2–11.0 | 927 | 166 | 17.9 | 15.5–20.6 |
| Jurisdiction ^b | ACT | 32 | 3 | 9.4 | 2.0–25.0 | 32 | 6 | 18.8 | 7.2–36.4 | 32 | 3 | 9.4 | 2.0–25 | 32 | 7 | 21.9 | 9.3–40.0 |
| | NSW | 322 | 34 | 10.6 | 7.4–14.4 | 322 | 62 | 19.3 | 15.1–24 | 321 | 42 | 13.1 | 9.6–17.3 | 321 | 52 | 16.2 | 12.3–20.7 |
| | NT | 31 | 2 | 6.5 | 0.8–21.4 | 31 | 3 | 9.7 | 2.0–25.8 | 31 | 0 | 0 | — | 30 | 2 | 6.7 | 0.8–22.1 |
| | Qld | 97 | 5 | 5.2 | 1.7–11.6 | 97 | 4 | 4.1 | 1.1–10.2 | 97 | 5 | 5.2 | 1.7–11.6 | 98 | 6 | 6.1 | 2.3–12.9 |
| | SA | 60 | 7 | 11.7 | 4.8–22.6 | 60 | 11 | 18.3 | 9.5–30.4 | 60 | 5 | 8.3 | 2.8–18.4 | 60 | 10 | 16.7 | 8.3–28.5 |
| | Tas. | 29 | 1 | 3.4 | 0.1–17.8 | 29 | 2 | 6.9 | 0.8–22.8 | 29 | 0 | 0 | — | 29 | 3 | 10.3 | 2.2–27.4 |
| | Vic. | 252 | 32 | 12.7 | 8.9–17.5 | 252 | 35 | 13.9 | 9.9–18.8 | 245 | 23 | 9.4 | 6.0–13.8 | 251 | 68 | 27.1 | 21.7–33.0 |
| | WA | 104 | 4 | 3.8 | 1.1–9.6 | 104 | 7 | 6.7 | 2.7–13.4 | 102 | 4 | 3.9 | 1.1–9.7 | 104 | 18 | 17.3 | 10.6–26.0 |
| Age group | ≤ 28 days | 192 | 18 | 9.4 | 5.7–14.4 | 192 | 30 | 15.6 | 10.8–21.5 | 192 | 10 | 5.2 | 2.5–9.4 | 192 | 33 | 17.2 | 12.1–23.3 |
| | 29–90 days | 108 | 9 | 8.3 | 3.9–15.2 | 108 | 11 | 10.2 | 5.2–17.5 | 105 | 8 | 7.6 | 3.3–14.5 | 107 | 4 | 3.7 | 1.0–9.3 |
| | 91–364 days | 141 | 13 | 9.2 | 5.0–15.3 | 141 | 21 | 14.9 | 9.5–21.9 | 136 | 12 | 8.8 | 4.6–14.9 | 142 | 11 | 7.7 | 3.9–13.4 |
| | 1–4 years | 166 | 15 | 9 | 5.1–14.5 | 166 | 25 | 15.1 | 10.0–21.4 | 165 | 16 | 9.7 | 5.6–15.3 | 165 | 37 | 22.4 | 16.3–29.6 |
| | 5–11 years | 152 | 10 | 6.6 | 3.2–11.8 | 152 | 16 | 10.5 | 6.1–16.5 | 151 | 14 | 9.3 | 5.2–15.1 | 151 | 52 | 34.4 | 26.9–42.6 |
| | 12–17 years | 168 | 23 | 13.7 | 8.9–19.8 | 168 | 27 | 16.1 | 10.9–22.5 | 168 | 22 | 13.1 | 8.4–19.2 | 168 | 29 | 17.3 | 11.9–23.8 |
| Onset location | Community | 641 | 47 | 7.3 | 5.4–9.6 | 641 | 71 | 11.1 | 8.8–13.8 | 636 | 29 | 4.6 | 3.1–6.5 | 639 | 123 | 19.3 | 16.3–22.5 |
| | Hospital | 286 | 41 | 14.3 | 10.5–18.9 | 286 | 59 | 20.6 | 16.1–25.8 | 281 | 53 | 18.9 | 14.5–23.9 | 286 | 43 | 15 | 11.1–19.7 |

a N: number of isolates tested; n: number of resistant isolates; R%: proportion of resistant isolates; 95% CI: 95% confidence interval.

b ACT: Australian Capital Territory; NSW: New South Wales; NT: Northern Territory; Qld: Queensland; SA: South Australia; Tas: Tasmania; Vic: Victoria; WA: Western Australia.

Pseudomonas aeruginosa

Over the two years, 64 *P. aeruginosa* isolates were reported: 36 in 2022 and 28 in 2023. No *P. aeruginosa* isolates were reported from the Northern Territory (as in 2020–2021) or from Tasmania. Thirteen *P. aeruginosa* isolates were piperacillin-tazobactam resistant (13/68; 19.1%; 95% CI: 10.6–30.5); nine were cefepime and/or ceftazidime resistant (13.2%; 95% CI: 6.2–23.6); seven were ciprofloxacin resistant (10.3%; 95% CI: 4.2–20.1). One isolate was meropenem resistant (1.5%; 95% CI: 0.0–7.9) and no *P. aeruginosa* isolates were tobramycin resistant. The two MDR *P. aeruginosa* isolates reported (3.1%) reported were from children aged 5–11 years old with hospital-onset BSIs.

Acinetobacter spp.

In total, 34 *Acinetobacter* spp. isolates from 30 patients were reported over the two years; however, 71% of such isolates were reported in 2023 (24/34; 70.6%). One *Acinetobacter* isolate, identified as *A. lwoffii*, was ciprofloxacin and co-trimoxazole resistant. No other *Acinetobacter* isolates were resistant to the antimicrobials tested, including meropenem or amikacin.

Molecular analysis

One in four gram-negative isolates were referred for molecular analysis of resistance genes in 2022–2023 (266/1,034; 25.7%), with results available for 264 at the time of analysis. In approximately 40% of the isolates referred for WGS, a β -lactamase resistance gene was detected (38.3%; 101/264), including the majority of *E. coli* (73/81) and *Klebsiella pneumoniae* complex (22/26) isolates. More detailed genomic analysis is available in the 2022 and 2023 AGAR amalgam reports.^{9,13} The proportion of ESBL gene detection in referred isolates was not significantly different to that seen in previous years (37.5% in 2018–2019 and 38.7% in 2020–2021).

Genes conferring resistance to the carbapenems were detected in seven gram-negative isolates: $bla_{\text{OXA-23}}$ was detected in an *A. baumannii* isolate, bla_{NDM} in two *E. coli* isolates, and $bla_{\text{IMP-4}}$ in four isolates: an *E. coli*, an *Enterobacter cloacae* complex isolate, a *K. pneumoniae* complex and a *Serratia marcescens*.

Gram positives

Staphylococcus aureus

There were 601 *S. aureus* isolates reported from 593 children and adolescents in 2022–2023; 13.6% were MRSA and 5.5% were MDR. The median age of patients with *S. aureus* bacteraemia (SAB) was 6 years, and 60.0% of patients were male (356/593). Most SABs were community-onset (78.6%; 466/593) and monomicrobial (90.1%; 539/593). Only 16.0% of SABs were device-related. One in five SAB patients remained in hospital after 30 days (120/593; 20.2%) (Table 1).

Overall, 14.4% of *S. aureus* were erythromycin resistant; 10.3% were clindamycin resistant; and 5.0% were ciprofloxacin resistant. Proportions of resistance in MRSA were significantly higher for erythromycin ($p = 0.02$), clindamycin ($p = 0.04$) and ciprofloxacin ($p < 0.001$). No *S. aureus* was resistant to co-trimoxazole (Table 4). Five percent of *S. aureus* were MDR; two-thirds of all MDR *S. aureus* were MRSA (21/33; 63.6%). More than half of MDR SABs were community-onset (19/33; 57.5%) (Table 2).

Although the proportion of *S. aureus* that were methicillin resistant in the Northern Territory was higher in 2022–2023 than in 2020–2021, the difference was not statistically significant (52.8% vs 48.3%; $p = 0.9$). Of the total 82 MRSA isolates, WGS results were available for 72 isolates at the time of analysis. The most frequently identified MRSA clone was ST93-IV (29/72; 40.3%). The Panton-Valentine Leucocidin (PVL) toxin associated-genes were detected in 71.0% of MRSA (54/72), with the Northern Territory having the highest proportion of MRSA that were PVL positive (17/18; 94.4%).

Table 4: Number of *Staphylococcus aureus* isolates tested and proportion of isolates resistant to clindamycin, ciprofloxacin, and erythromycin from patients aged < 18 years as reported to the Australian Group on Antimicrobial Resistance, January 2022 – December 2023

| Category | Characteristic | Clindamycin ^a | | | | Ciprofloxacin ^a | | | | Erythromycin ^a | | | |
|-------------------------------------|----------------|--------------------------|----|------|-----------|----------------------------|----|-----|-----------|---------------------------|----|------|-----------|
| | | N | n | R% | 95% CI | N | n | R% | 95% CI | N | n | R% | 95% CI |
| Overall | — | 600 | 62 | 10.3 | 8.0–13.1 | 600 | 30 | 5.0 | 3.4–7.1 | 584 | 84 | 14.4 | 11.6–17.5 |
| Jurisdiction ^b | ACT | 19 | 3 | 15.8 | 3.4–39.6 | 19 | 0 | 0 | — | 19 | 3 | 15.8 | 3.4–39.6 |
| | NSW | 211 | 22 | 10.4 | 6.7–15.4 | 211 | 13 | 6.2 | 3.3–10.3 | 195 | 25 | 12.8 | 8.5–18.3 |
| | NT | 36 | 5 | 13.9 | 4.7–29.5 | 36 | 3 | 8.3 | 1.8–22.5 | 36 | 5 | 13.9 | 4.7–29.5 |
| | Qld | 70 | 9 | 12.9 | 6.1–23 | 70 | 1 | 1.4 | 0.0–7.7 | 70 | 9 | 12.9 | 6.1–23.0 |
| | SA | 39 | 1 | 2.6 | 0.1–13.5 | 39 | 1 | 2.6 | 0.1–13.5 | 39 | 12 | 30.8 | 17.0–47.6 |
| | Tas. | 16 | 2 | 12.5 | 1.6–38.3 | 16 | 0 | 0 | — | 16 | 2 | 12.5 | 1.6–38.3 |
| | Vic. | 136 | 14 | 10.3 | 5.7–16.7 | 136 | 12 | 8.8 | 4.6–14.9 | 136 | 22 | 16.2 | 10.4–23.5 |
| | WA | 73 | 6 | 8.2 | 3.1–17 | 73 | 0 | 0 | — | 73 | 6 | 8.2 | 3.1–17.0 |
| Age group | ≤ 28 days | 52 | 12 | 23.1 | 12.5–36.8 | 52 | 3 | 5.8 | 1.2–15.9 | 52 | 17 | 32.7 | 20.3–47.1 |
| | 29–90 days | 50 | 3 | 6.0 | 1.3–16.5 | 50 | 3 | 6 | 1.3–16.5 | 48 | 6 | 12.5 | 4.7–25.2 |
| | 91–364 days | 60 | 13 | 21.7 | 12.1–34.2 | 60 | 4 | 6.7 | 1.8–16.2 | 58 | 13 | 22.4 | 12.5–35.3 |
| | 1–4 years | 110 | 12 | 10.9 | 5.8–18.3 | 110 | 9 | 8.2 | 3.8–15 | 106 | 19 | 17.9 | 11.2–26.6 |
| | 5–11 years | 174 | 12 | 6.9 | 3.6–11.7 | 174 | 6 | 3.4 | 1.3–7.4 | 169 | 17 | 10.1 | 6.0–15.6 |
| | 12–17 years | 154 | 10 | 6.5 | 3.2–11.6 | 154 | 5 | 3.2 | 1.1–7.4 | 151 | 12 | 7.9 | 4.2–13.5 |
| Onset location | Community | 468 | 43 | 9.2 | 6.7–12.2 | 468 | 20 | 4.3 | 2.6–6.5 | 458 | 60 | 13.1 | 10.1–16.5 |
| | Hospital | 132 | 19 | 14.4 | 8.9–21.6 | 132 | 10 | 7.6 | 3.7–13.5 | 126 | 24 | 19.0 | 12.6–27.0 |
| Methicillin resistance ^c | MRSA | 82 | 13 | 15.9 | 8.7–25.6 | 82 | 18 | 22 | 13.6–32.5 | 82 | 18 | 22.0 | 13.6–32.5 |
| | MSSA | 518 | 49 | 9.5 | 7.1–12.3 | 518 | 12 | 2.3 | 1.2–4.0 | 502 | 66 | 13.1 | 10.3–16.4 |

a N: number of isolates tested; n: number of resistant isolates; R%: proportion of resistant isolates; 95% CI: 95% confidence interval.

b ACT: Australian Capital Territory; NSW: New South Wales; NT: Northern Territory; Qld: Queensland; SA: South Australia; Tas: Tasmania; Vic: Victoria; WA: Western Australia.

c MRSA: methicillin-resistant *Staphylococcus aureus*; MSSA: methicillin-susceptible *Staphylococcus aureus*.

Enterococcus spp.

Over the two years there were 192 *Enterococcus* spp. isolates reported from 182 patients. The majority of isolates were *E. faecalis* (70.8%; 136/192), and 17.2% of enterococci were *E. faecium* (33/192). More than half of all *E. faecium* isolates were isolated in New South Wales (19/33; 57.6%). *E. faecium* BSI episodes were more frequently hospital-onset (χ^2 : 5.3; p = 0.02), and most neonatal enterococcal BSIs were caused by *E. faecalis* (41/46, 89.1%).

Although only 14.2% of *Enterococcus* isolates were ampicillin resistant (26/183), all ampicillin-resistant isolates were identified as *E. faecium*. Overall, 81.3% of *E. faecium* were ampicillin resistant (26/32). Ampicillin-resistant enterococcal BSIs were primarily hospital-onset or in adolescents 12–17 years of age (Table 5).

Vancomycin resistance was only detected in *E. faecium* isolates; 40.6% of *E. faecium* isolates were vancomycin resistant (13/32). While there was an increase in the proportion of vancomycin-resistant *E. faecium* in 2022–2023 compared to 2020–2021, this increase did not attain statistical significance because of the small numbers of isolates reported (40.6% vs 19.5%; χ^2 : 3; p = 0.09). Vancomycin-resistant enterococci (VRE) were 21.5% of all hospital-onset enterococcal infections; most VRE occurred in adolescents aged 12–17 years (61.5%; 8/13). Significant differences were also noted between different jurisdictions (Table 5). Of the 13 *E. faecium* isolates phenotypically resistant to vancomycin, 12 were *vanB* positive, and one was *vanA* positive. Only one *E. faecium* isolate was teicoplanin resistant (Table 5).

Fifteen *E. faecium* isolates (45.5%) were MDR, of which 14 were hospital-onset (14/15; 93.3%) (Table 3). Of the 44 *E. faecium* isolates were referred for WGS, there were 23 different sequence types (ST) identified, with ST78 the most frequent (11/44; 25.0%).

WHO Priority Pathogens

From the WHO Bacterial Priority Pathogens List (BPPL), two of the three ‘critical priority’ group pathogens were detected in children and adolescents over the two years: 14.6% of Enterobacterales were third-generation cephalosporin resistant and 1.2% of Enterobacterales were carbapenem resistant. No carbapenem-resistant *A. baumannii* were reported. Five of the six ‘high priority’ group resistant isolates surveyed by AGAR were detected (Table 6).

Table 5: Number of *Enterococcus* spp. isolates tested and proportion of isolates resistant to ampicillin and vancomycin from patients aged < 18 years as reported to the Australian Group on Antimicrobial Resistance, January 2022 – December 2023

| Category | Characteristic | Ampicillin ^a | | | | Vancomycin ^a | | | |
|---------------------------|---------------------------|-------------------------|----|------|-----------|-------------------------|----|------|-----------|
| | | N | n | R% | 95% CI | N | n | R% | 95% CI |
| Overall | — | 183 | 26 | 14.2 | 9.5–20.1 | 187 | 13 | 7.0 | 3.8–11.6 |
| Species | <i>E. faecalis</i> | 131 | 0 | 0 | — | 133 | 0 | 0 | — |
| | <i>E. faecium</i> | 32 | 26 | 81.3 | 63.6–92.8 | 32 | 13 | 40.6 | 23.7–59.4 |
| | Other <i>Enterococcus</i> | 20 | 0 | 0 | — | 22 | 0 | 0 | — |
| Jurisdiction ^b | ACT | 5 | 0 | 0 | — | 5 | 0 | 0 | — |
| | NSW | 65 | 17 | 26.2 | 16–38.5 | 69 | 4 | 5.8 | 1.6–14.2 |
| | NT | 4 | 1 | 25 | — | 4 | 1 | 25 | — |
| | Qld | 10 | 1 | 10 | 0.3–44.5 | 10 | 1 | 10 | 0.3–44.5 |
| | SA | 18 | 0 | 0 | — | 18 | 0 | 0 | — |
| | Tas | 5 | 0 | 0 | — | 5 | 0 | 0 | — |
| | Vic | 53 | 7 | 13.2 | 5.5–25.3 | 53 | 7 | 13.2 | 5.5–25.3 |
| | WA | 23 | 0 | 0 | — | 23 | 0 | 0 | — |
| Age group | ≤ 28 days | 46 | 2 | 4.3 | 0.5–14.8 | 46 | 1 | 2.2 | 0.1–11.5 |
| | 29–90 days | 26 | 1 | 3.8 | 0.1–19.6 | 28 | 1 | 3.6 | 0.1–18.3 |
| | 91–364 days | 34 | 3 | 8.8 | 1.9–23.7 | 34 | 1 | 2.9 | 0.1–15.3 |
| | 1–4 years | 28 | 5 | 17.9 | 6.1–36.9 | 30 | 1 | 3.3 | 0.1–17.2 |
| | 5–11 years | 16 | 4 | 25 | 7.3–52.4 | 16 | 1 | 6.3 | 0.2–30.2 |
| | 12–17 years | 33 | 11 | 33.3 | 18–51.8 | 33 | 8 | 24.2 | 11.1–42.3 |
| Onset location | Community | 58 | 2 | 3.4 | 0.4–11.9 | 59 | 0 | 0 | — |
| | Hospital | 125 | 24 | 19.2 | 12.7–27.2 | 128 | 13 | 10.2 | 5.5–16.7 |

a N: number of isolates tested; n: number of resistant isolates; R%: proportion of resistant isolates; 95% CI: 95% confidence interval.

b ACT: Australian Capital Territory; NSW: New South Wales; NT: Northern Territory; Qld: Queensland; SA: South Australia; Tas: Tasmania; Vic: Victoria; WA: Western Australia.

Table 6: Number and proportion of World Health Organization (WHO) priority pathogens and resistance profiles detected in patients aged < 18 years within the Australian Group on Antimicrobial Resistance (AGAR) surveillance outcomes programs, January 2022 – December 2023

| Priority | Organism | Resistance to ^b | Resistance characteristics ^a | | | |
|----------|--|----------------------------|---|-----|------|-----------|
| | | | N | n | R% | 95% CI |
| Critical | Enterobacterales | Carbapenems | 932 | 11 | 1.2 | 0.5–1.9 |
| | Enterobacterales | 3GC | 932 | 130 | 13.9 | 11.7–16.2 |
| | <i>A. baumannii</i> complex | Carbapenems | 15 | 0 | 0 | — |
| High | <i>Salmonella</i> Typhi | Fluroquinolones | 50 | 47 | 94.0 | 87.4–100 |
| | <i>Shigella</i> spp. | Fluroquinolones | 1 | 0 | 0 | — |
| | <i>E. faecium</i> | Vancomycin | 32 | 13 | 40.6 | 23.7–59.4 |
| | <i>P. aeruginosa</i> | Carbapenems | 68 | 1 | 1.5 | 0–4.3 |
| | Non-typhoidal <i>Salmonella</i> | Fluroquinolones | 85 | 8 | 9.4 | 3.2–15.6 |
| | <i>S. aureus</i> | Methicillin | 601 | 82 | 13.6 | 10.9–16.4 |
| | <i>Neisseria gonorrhoeae</i> ^c | 3GC and/or fluroquinolones | — | — | — | — |
| Medium | Group A <i>Streptococcus</i> ^c | Macrolides | — | — | — | — |
| | <i>Streptococcus pneumoniae</i> ^c | Macrolides | — | — | — | — |
| | <i>Haemophilus influenzae</i> ^c | Ampicillin | — | — | — | — |
| | Group B <i>Streptococcus</i> ^c | Penicillin | — | — | — | — |

a N: number of isolates tested; n: number of resistant isolates; R%: proportion of resistant isolates; 95% CI: 95% confidence interval.

b Carbapenems: meropenem, ertapenem, imipenem; 3GC (third-generation cephalosporins): ceftazidime, ceftriaxone; fluroquinolones: ciprofloxacin, levofloxacin, moxifloxacin, norfloxacin.

c Organisms not collected within AGAR surveillance outcome programs, but listed here to provide context of the full WHO pathogen list.

Discussion

The 2022–2023 AGAR Kids Biennial Report shows stability in the AMR landscape within the Australian paediatric population, with few significant differences detected when compared to the 2020–2021 report. Minor variations observed between the two reporting periods were possibly due to the easing of coronavirus disease 2019 (COVID-19) public health measures across Australia in 2022.

Although the WHO BPPL was updated in 2024 and thus is different to the list reported against in the 2020–2021 AGAR Kids report, when compared to global estimates, the proportion of resistant isolates detected in the Australian paediatric population is limited. Whilst the global median resistance of *A. baumannii* to carbapenems was 69.0–73.4% in 2022, no resistant isolates were reported in AGAR Kids. Conversely, whilst the global median resistance to ciprofloxacin for *Salmonella* spp. was 16.7% (5.4–30.4%), the proportion in Australian children and adolescents was 46.9%. Though this result is higher than the global median estimates, the majority of ciprofloxacin-resistant *Salmonella* observed in this report were *S. Typhi*, cases of which in Australia are almost always acquired whilst travelling to endemic regions.¹⁴ This is reflected by the significant reduction of salmonellosis notifications across the population to the National Notifiable Disease Surveillance System (NNDSS) during the Australian border lockdown period, and by the absence of any cases of *S. Typhi* or *S. Paratyphi* reported in the 2020–2021 AGAR Kids report.^{4,15} Considering children and adolescents have increased risk to Typhi acquisition when travelling overseas, increased messaging around *S. Typhi* vaccine uptake before overseas travel may be helpful in reducing the number of typhoid infections.^{14,16} The Australian Immunisation Handbook recommends typhoid vaccination for all travellers aged ≥ 2 years to endemic regions.¹⁷

Whilst most ciprofloxacin-resistant *Salmonella* were identified as *S. Typhi* or *S. Paratyphi*, ~10% of non-typhoidal *Salmonella* were also ciprofloxacin resistant, which is similar to other Australian AMR surveillance reports.¹⁸ Although ciprofloxacin-resistant *Salmonella* BSIs are associated with overseas travel, several papers have described endogenous fluoroquinolone resistance in Australian *Salmonella* isolates, as well as concordant resistance to third-generation cephalosporins or azithromycin.^{19–21} Monitoring fluoroquinolone resistance in animal *Salmonella* isolates, particularly by surveillance of plasmid-mediated quinolone resistance (PMQR) genes, may provide oversight of endogenous ciprofloxacin resistance transmission in the circumstances of low numbers in humans.²²

Although not statistically significant, the proportion of third-generation cephalosporin resistant Enterobacterales has continued to increase, with 15.9% of Enterobacterales resistant to third-generation cephalosporins in 2023; this is the highest proportion observed in the paediatric population across 11 years of AGAR surveillance.⁶ Research conducted from five Australian children's hospitals during 2019–2021 indicates a slightly higher proportion of third-generation cephalosporin resistant Enterobacterales BSIs at 21.9%.²³ As was observed in this report, children with third-generation cephalosporin resistant Enterobacterales BSIs, as well as those with MDR Enterobacterales BSIs, had longer hospital stays and higher mortality.²³ In recent years, penicillins with extended-spectrum (J01CA), first-generation cephalosporins (J01DB), and β -lactam/ β -lactamase inhibitor combinations (J01CR) were some of the more frequently dispensed systemic antibacterial classes under the Pharmaceutical Benefits Scheme (PBS) and Repatriation PBS (RPBS).¹⁸ As third-generation cephalosporins are often used as first-line empirical treatment for paediatric sepsis and meningitis, ongoing strategies to combat increasing resistance must include antimicrobial stewardship and ongoing attention to the appropriateness of third-generation cephalosporin use.

Over the past 10 years there has been a small but clinically important increase in the number of carbapenem-resistant Enterobacterales (CPE) in the AGAR paediatric population; from one isolate in 2014–2015 to 11 isolates in 2022–2023 (χ^2 : 5; $p = 0.02$). A similar increase of CPE has been observed in the AGAR adult population. This trend has occurred in parallel with an increase in meropenem use since 2017.¹⁸ However, there has been no similar increase of carbapenem resistance observed in the *Acinetobacter* spp. or *P. aeruginosa* in either the paediatric or adult population. A more detailed analysis of CPE BSIs in Australia indicates that most isolates were locally acquired (endemic) rather than overseas.^{18,24}

There was no change in the proportion of antimicrobial resistance reported in *P. aeruginosa* isolates during 2022–2023 in the paediatric AGAR populations, despite a national increase in piperacillin-tazobactam use from 2018–2022.^{18,24,25} In a recent analysis of paediatric *P. aeruginosa* BSIs in Queensland, Slack et al. observed increases in piperacillin-tazobactam and ciprofloxacin resistances between 2015–2019.²⁶ The piperacillin-tazobactam resistant proportions observed by Slack et al. are similar to those in the paediatric AGAR population, both during the same time frame (2015–2019) and in 2022–2023. Conversely, the proportion of ciprofloxacin resistance in the paediatric AGAR population is greater than that observed in the Queensland report.⁶ Resistance in *P. aeruginosa* should be monitored carefully, as *P. aeruginosa* BSIs are more frequent in patients with serious underlying comorbidities or frequent healthcare contact.

There were no statistically significant changes in the proportions of *S. aureus* resistance when compared to the previous AGAR Kids Biennial report. As previously observed, the Northern Territory had the highest proportion of *S. aureus* that were methicillin resistant, as was observed in the adult population.^{4,7,27} In a recent analysis of MRSA data across the north of Australia (2016–2021), an increase in MRSA infection was associated with remoteness index; i.e., the more remote, the higher the odds of an MRSA infection. However, no significant association between socio-economic indexes and MRSA infection was observed.²⁸ Using Bayesian modelling, Wozniak et. al., found the remote regions of east and west Pilbara (Western Australia), Daly-West Arnhem and Alice Springs (Northern Territory), and the Outback North (Queensland) had the highest predicted prevalence of MRSA across all age groups and sexes, whilst urban settings had the lowest predicted prevalence.²⁸ Future geographic analysis of AGAR Kids data using similar methods across a national dataset are planned, with results expected to mirror these findings.

Although there was an increase in the proportion of *E. faecium* resistant to vancomycin in the paediatric population between the two biennial reports, the total number of vancomycin-resistant *E. faecium* (VREfm) was small and should be interpreted with caution. It should be noted, however, that during the same time period the proportion of VREfm reported in the adult population increased significantly, from 37.2% in 2020–2021 to 48.9% in 2022–2023 (χ^2 : 29; $p < 0.001$). As observed in children and adolescents, all VRE reported in adults in 2022–2023 were *E. faecium*.⁹ When compared internationally, Australia continues to have higher proportions of *E. faecium* that are vancomycin resistant: over the past five years, the United Kingdom observed increases in VREfm from 20.6% in 2019 to 22.4% in 2023 ($p = 0.07$), and the EU/EEA population-weighted mean percentage of VREfm was 19.8% in 2023; however, national percentages of VRE across the EU/EEA ranged from 0.0% to 60.9%.^{29,30} As well as increasing resistance to vancomycin, there was an increase in the proportion of MDR enterococci in both adults and children in Australia. As *Enterococcus* spp. have intrinsic resistance to several antimicrobial classes, increasing resistance to the few antimicrobial agents available to treat enterococcal BSIs poses a critical threat.

Conclusions

The 2022–2023 AGAR Kids Biennial Report shows relative stability in the AMR landscape within the Australian paediatric population, with few significant differences detected when compared to the 2020–2021 report. From BSIs in children and adolescents during 2022–2023, there have been small increases in the proportions of resistant Enterobacterales and *Enterococcus* spp. isolates reported, whilst resistance in *S. aureus* has remained stable. The lifting of public health restrictions and the increase in international travel may have impacted the epidemiology of BSIs and antimicrobial resistance patterns in the paediatric AGAR population during 2022–2023. The second edition of the AGAR Kids Biennial report reiterates the importance of reporting age-delimited data in AMR surveillance reports, with distinct differences observed not only between adults and children, but amid age groups.

Acknowledgments

This report was the work of the AGAR Kids committee and was funded by the Australian Government Department of Health, Disability and Ageing. The AGAR Kids committee would like to thank the participants to the AGAR surveillance program in 2022/2023:

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Children's Hospital Westmead
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Gold Coast University Hospital
Jennifer Robson and Marianne Allen – Greenslopes
Private Hospital and Mater Private Hospital
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Kelly Papanoum and Xiao Ming Chen –
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Marcel Leroi and Elizabeth Grabsch – Austin Health
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Dandenong Hospital
Tony Korman and Despina Kotsanas – Monash
Medical Centre and Monash Children's Hospital
Katherine Bond and Rose Cotronei –
Royal Melbourne Hospital
Andrew Daley and Gena Gonis –
Royal Women's Hospital
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St Vincent's Hospital

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Denise Daley and Shakeel Mowlaboccus –
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Appendix A

Table A.1.1: Number and proportion of the ten most frequently reported isolates to the Australian Group on Antimicrobial Resistance in patients aged < 18 years, January 2020 – December 2021

| Organism | 2020 N: 823 | | 2021 N: 856 | | 2020–2021 N: 1,679 | |
|--|----------------|------|----------------|------|-----------------------|------|
| | n | % | n | % | n | % |
| <i>Staphylococcus aureus</i> | 303 | 36.8 | 304 | 35.5 | 607 | 36.2 |
| <i>Escherichia coli</i> | 163 | 19.8 | 215 | 25.1 | 378 | 22.5 |
| <i>Enterococcus faecalis</i> | 60 | 7.3 | 62 | 7.2 | 122 | 7.3 |
| <i>Klebsiella pneumoniae</i> complex | 61 | 7.4 | 61 | 7.1 | 122 | 7.3 |
| <i>Enterobacter cloacae</i> complex | 60 | 7.3 | 46 | 5.4 | 106 | 6.3 |
| <i>Salmonella</i> (non-Typhi) | 22 | 2.7 | 52 | 6.1 | 74 | 4.4 |
| <i>Pseudomonas aeruginosa</i> | 31 | 3.8 | 30 | 3.5 | 61 | 3.6 |
| <i>Enterococcus faecium</i> | 32 | 3.9 | 9 | 1.1 | 41 | 2.4 |
| <i>Klebsiella oxytoca</i> | 13 | 1.6 | 17 | 2.0 | 30 | 1.8 |
| <i>Acinetobacter baumannii</i> complex | 10 | 1.2 | 8 | 0.9 | 18 | 1.1 |

Table A.1.2: Number and proportion of the ten most frequently reported isolates to the Australian Group on Antimicrobial Resistance in patients aged < 18 years, January 2022 – December 2023

| Organism | 2022 N: 791 | | 2023 N: 1,036 | | 2022–2023 N: 1,827 | |
|--------------------------------------|----------------|------|------------------|------|-----------------------|------|
| | n | % | n | % | n | % |
| <i>Staphylococcus aureus</i> | 278 | 35.1 | 323 | 31.2 | 601 | 32.9 |
| <i>Escherichia coli</i> | 200 | 25.3 | 257 | 24.8 | 457 | 25.0 |
| <i>Enterococcus faecalis</i> | 67 | 8.5 | 69 | 6.7 | 136 | 7.4 |
| <i>Klebsiella pneumoniae</i> complex | 45 | 5.7 | 62 | 6.0 | 107 | 5.9 |
| <i>Enterobacter cloacae</i> complex | 37 | 4.7 | 54 | 5.2 | 91 | 5.0 |
| <i>Salmonella</i> (non-Typhi) | 34 | 4.3 | 51 | 4.9 | 85 | 4.7 |
| <i>Pseudomonas aeruginosa</i> | 38 | 4.8 | 30 | 2.9 | 68 | 3.7 |
| <i>Salmonella</i> Typhi | 15 | 1.9 | 35 | 3.4 | 50 | 2.7 |
| <i>Serratia marcescens</i> | 12 | 1.5 | 23 | 2.2 | 35 | 1.9 |
| <i>Enterococcus faecium</i> | 15 | 1.9 | 18 | 1.7 | 33 | 1.8 |