From awareness to action:

Can we do more to combat antimicrobial resistance in heavily immunocompromised patients?

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Introduction

- Antimicrobial resistance (AMR) is a global health threat, causing **4.71 million deaths** in 2021 (1.14 million directly from resistant infections).¹
- By 2050, AMR could lead to 8.22 million deaths annually and \$6.1 trillion in economic losses (3.8% of GDP).^{1,2}
- Transplant recipients and patients with cancer face a higher risk of AMR infections due to immunosuppression, frequent hospitalisations, invasive procedures and antibiotic exposure.³⁻⁵

Methods

- We conducted a keyword search using PubMed for publications (2018–2023) reporting burdens from AMR bacterial infections from Europe and North America.
- Fungal and viral infections were excluded to maintain a focused scope.

Objective: To assess AMR impact in heavily immunocompromised patients, focusing on treatment delays, patient experience and economic burden.

Results

- Search results (**Figure 1**) were reviewed in a virtual steering committee meeting, with additional publications included based on author expertise.
- Most studies identified in the literature reported retrospective cohort data (n=113), with the next most common being prospective cohorts (n=35).

Epidemiology of AMR and its mortality

- The impact of AMR on mortality varies significantly, with 30-day mortality rates ranging from 3% to 64.6%, depending on population and disease type.6-68
- Inappropriate empirical antibiotic therapy (IEAT) is a key factor contributing to AMR-associated mortality, with IEAT rates reported between **16.9** and **68.6%**. 11,13,36,42,43,62,69–79
- The frequency of AMR and associated mortality rates vary widely (Table 1).

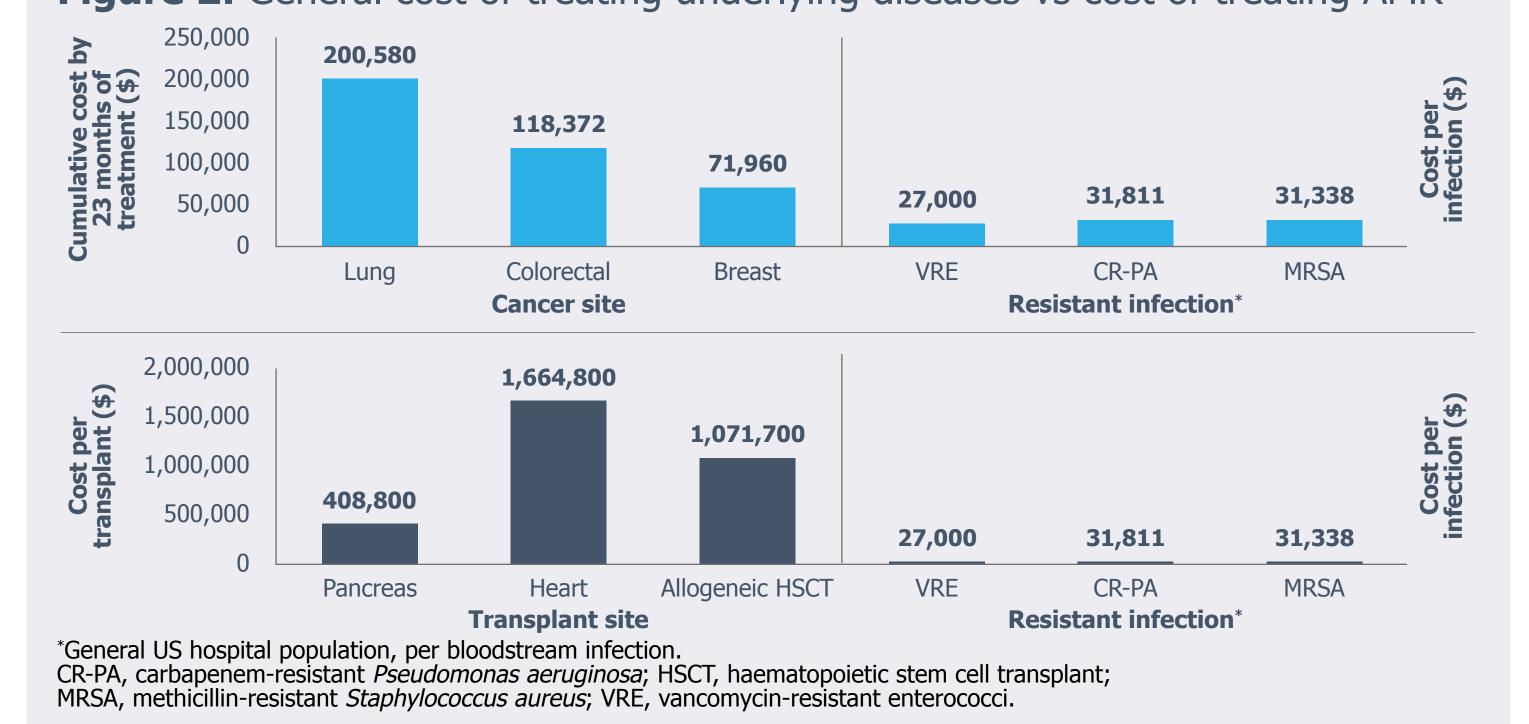
Table 1: Frequency of AMR and 30-day mortality across disease areas explored

	Solid cancer	Haematological malignancy	Solid organ transplant
Frequency of AMR	0.6-71.7%	1.7-52.3%	0-73.8%
30-day mortality	29.5–43%	6-50%	3.9-50%

Clinical management practices

- Many standard prophylactic regimens could be considered **inadequate** against AMR infections, particularly in oncology.^{24,89}
- The development of rapid^{90,91} diagnostic tests offers clinicians a valuable tool for timely microbiological assessment.
- **Appropriate stewardship** is pivotal to maintaining effectiveness of existing antibiotics. 92,93

Figure 2: General cost of treating underlying diseases vs cost of treating AMR

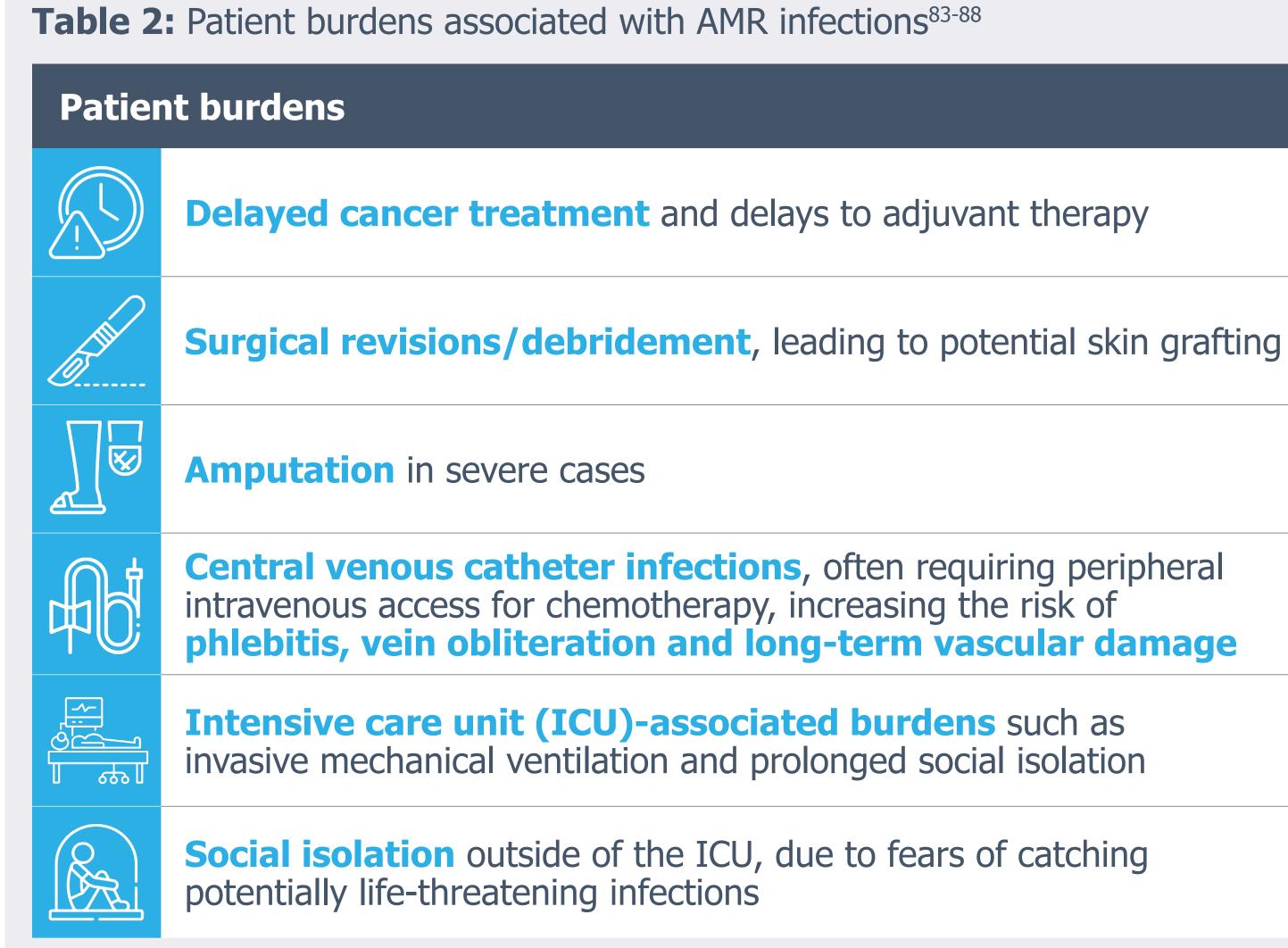


Consequences for patients

- There are no controlled studies exploring patient-reported experience measures, and real-world data on the impact of antibiotic stewardship programmes remain limited.
- Several case studies have highlighted the substantial burden of resistant infections on patients (Table 2).

Figure 1: Literature identification – flow chart of results identified and screened Publication on AMR n=410,088• 304,186 reports from 12 Nov 2018 1,517 non-English language 62,658 non-human studies • 14,869 irrelevant study types, or genome studies Identified studies of bacterial AMR • 2,971 reports of fungal, parasite, archaea or viral infections only within the selected date range 12 Nov 2018 to 11 Nov 2023 n=23,887 21,089 publications not in oncology, haematological malignancy or solid organ transplant • 1,192 not reported on a relevant patient burden keyword Publications reporting AMR Publications reporting AMR Publications reporting AMR burdens in **solid organ** burdens in **haematological** burdens in **oncology** transplant n=457 n = 348Excluded: 724 publications not including data from Europe or North America Publications reporting AMR Publications reporting AMR **Publications reporting AMR** burdens in **haematological** burdens in **solid organ** burdens in **oncology** in **transplant** in Europe **malignancy** in Europe Europe and North America and North America and North America Excluded: n = 200n = 295• 720 publications did not meet eligibility criteria for our search 34 publications of interest identified from report 882 publications screened for eligibility 12 publications of interest added after the data cut-off of 11 Dec 2023 (up to October 2024) 6 reports of interest from governmental or public health bodies 214 publications explored in this review

Table 2: Patient burdens associated with AMR infections⁸³⁻⁸⁸



Direct and indirect costs of AMR

- The **financial burden** of AMR infections is substantial, largely driven by prolonged hospitalisations, ICU admissions and ventilator dependence.¹⁵
- Managing an immunocompromised patient's underlying disease is often far more expensive than treating an AMR infection itself (Figure 2).
- Greater investment in infection control, rapid diagnostics and targeted therapies has the potential to reduce overall healthcare costs.
- Indirect costs, such as long-term economic burden and patient productivity loss, remain poorly characterised.



AMR continues to place a growing strain on healthcare resources, leading to treatment delays, increased patient burden, and social isolation. To address these challenges, we propose three key actions.



Increase **education** of clinicians across oncology and transplant to support the use of appropriate, tailored empirical antibiotic therapy.



Expand access to rapid diagnostic tools to guide treatment decisions.



Conduct costeffectiveness **studies** to assess AMR's broader economic impact.

