



ORIGINAL ARTICLE

Selective decontamination of the digestive tract in a burns unit reduces the incidence of hospital-acquired infections: A retrospective before-and-after cohort study



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KEYWORDS

Selective decontamination of

Abstract

Objective: To evaluate the effect of selective decontamination of the digestive tract (SDD) on hospital-acquired infections (HAIs) in patients with acute burn injury requiring admission to a Burns Unit (BU).

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Abbreviations: ABLs, Advanced Burn Life Support; ABSI, Abbreviated Burn Severity Index; AIC, Akaike's Information Criterion; APACHE, Acute Physiology And Chronic Health Evaluation; ASP, Antimicrobial Stewardship Program; ATLS, Advanced Trauma Life Support; BET, Biological Engineering Technology; BU, Burns Unit; CFU, Colony-Forming Units; CI, Confidence Interval; EMR, Electronic Medical Record; ENVIN-HELICS, Estudio Nacional de Vigilancia de Infección Nosocomial (National Surveillance Study of Nosocomial Infection) – Hospitals in Europe Link for Infection Control through Surveillance; HAI, Hospital-Acquired Infection; ICU, Intensive Care Unit; IQR, Interquartile Range; MDRB, Multidrug-Resistant Bacteria; MV, Mechanical Ventilation; No., Number; OR, Odds Ratio; RCT, Randomized, Controlled Trial; SDD, Selective Decontamination of the Digestive tract; SEMICYUC, Sociedad Española de Medicina Intensiva, Crítica y Unidades Coronarias; STROBE, STrengthening the Reporting of OBservational studies in Epidemiology; TBSA, Total Body Surface Area; VAP, Ventilator-Associated Pneumonia.

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the digestive tract;
 Selective digestive
 decontamination;
 Burns unit;
 Burn patients;
 Prevention;
 Infection;
 Hospital-acquired
 infections;
 Ventilator-associated
 pneumonia;
 Bloodstream
 infections;
 Bacteremia

Design: Retrospective before-and-after cohort study, between January 2017 and June 2023. SDD was implemented in March 2019, dividing patients into two groups.

Setting: Four-bed BU, in a referral University Hospital in Spain.

Patients: All the patients admitted during the study period were eligible for analysis. Patients who died or were discharged within 48 hours of admission, and patients with an estimated survival less than 10% not considered for full escalation of therapy were excluded.

Intervention: SDD comprised the administration of a 4-day course of an intravenous antibiotic, and an oral suspension and oral topical paste of non-absorbable antibiotics during the stay in the BU.

Main variable of interest: Incidence of HAIs during the stay in the BU. Secondary outcomes: incidence of specific types of infections by site (bacteremia, pneumonia, skin and soft tissue infection) and microorganism (Gram-positive, Gram-negative, fungi), and safety endpoints.

Results: We analyzed 72 patients: 27 did not receive SDD, and 45 received SDD. The number of patients who developed HAIs were 21 (77.8%) and 21 (46.7%) in the non-SDD and the SDD groups, respectively ($p=0.009$). The number of hospital-acquired infectious episodes were 2.52 (1.21–3.82) and 1.13 (0.54–1.73), respectively ($p=0.029$).

Conclusions: SDD was associated with a reduced incidence of bacterial HAIs and a decrease in the number of infectious episodes per patient.

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PALABRAS CLAVE

Descontaminación
 selectiva del tracto
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 Descontaminación
 digestiva selectiva;
 Unidad de quemados;
 Paciente quemado;
 Prevención;
 Infección;
 Infección de
 adquisición
 hospitalaria;
 Neumonía asociada a
 ventilación mecánica;
 Infección del torrente
 sanguíneo;
 Bacteriemia

La descontaminación digestiva selectiva en una Unidad de Quemados disminuye la incidencia de infecciones de adquisición hospitalaria: estudio de cohortes retrospectivo antes-después

Resumen

Objetivo: Evaluar el efecto de la descontaminación digestiva selectiva (DDS) en las infecciones de adquisición hospitalaria (IAH) en pacientes con quemadura en una Unidad de Quemados (UQ).

Diseño: Cohorte retrospectiva antes-después, entre enero de 2017 y junio de 2023. La DDS se implementó en marzo de 2019, dividiendo los pacientes en dos grupos.

Ámbito: UQ de cuatro camas, en hospital universitario de referencia en España.

Pacientes: Todos los pacientes ingresados durante el periodo de estudio fueron elegibles para el análisis. Excluimos a aquellos que fallecieron o recibieron el alta en las primeras 48 horas de ingreso, y a aquellos con una supervivencia predicha menor del 10% que no fueran candidatos a tratamiento completo.

Intervención: La DDS incluyó la administración de un ciclo de antibiótico intravenoso durante 4 días, y una suspensión y pasta tópica oral de antibióticos no absorbibles durante la estancia en UQ.

Variable de interés principal: Incidencia de IAH durante la estancia en UQ. Desenlaces secundarios: incidencia de infecciones por localización (bacteriemia, neumonía, piel/partes blandas) y microorganismo (Gram-positivos, Gram-negativos, hongos), y desenlaces de seguridad.

Resultados: Analizamos 72 pacientes: 27 no recibieron DDS, y 45 recibieron DDS. El número de pacientes que desarrolló IAH fue 21 (77.8%) y 21 (46.7%) en los grupos sin y con DDS, respectivamente ($p=0.009$). El número de episodios de IAH fue 2.52 (1.21–3.82) y 1.13 (0.54–1.73), respectivamente ($p=0.029$).

Conclusiones: La DDS se asocia a una reducción en la incidencia de IAH, y de episodios de infección por paciente.

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Background

Acute burn injury leads to significant disturbances in homeostasis, resulting in immunodepression. Changes in both

cellular and humoral immunity render patients more susceptible to microbial invasion. Consequently, burned patients experience a compromised primary defense against environmental microorganisms. The concomitant presence of

necrotic and avascular tissue further fosters the colonization and proliferation of microorganisms, resulting in poor penetration of systemic antibiotics. Moreover, the release of toxic substances from such tissue may exacerbate the local immune response.^{1–4}

Infection is the leading cause of death for patients who survive more than 72 hours following a burn injury.⁵ Infection precedes multiorgan dysfunction in over 80% of patients with acute burn injuries and is considered the direct cause of death in more than one-third of the patients.⁶ The incidence of sepsis ranges from 3% to 30% in individuals with burns that account for more than 20% of their total body surface area (TBSA).^{7,8}

Preventing infections is associated with increased survival, a reduced length of hospital stay, and lower healthcare-related costs.⁹ Selective decontamination of the digestive tract (SDD) is a strategy to prevent hospital-acquired infections (HAIs). It consists of a short course of parenteral broad-spectrum antibiotics, along with non-absorbable topical antimicrobials in the oropharynx and the gut.¹⁰ The implementation of this strategy, together with a bundle of measures aiming to reduce the burden of ventilator-associated pneumonia (VAP) in intensive care units (ICUs), was suggested in current clinical practice guidelines.^{11,12} A recent meta-analysis of general critical care patients concluded that SDD is associated with a lower risk of bacteremia and VAP, which translates to increased survival.¹³ A contemporary systematic review of burn patients suggests that SDD may reduce VAP and bacteremia episodes from Enterobacteriaceae. This may potentially improve survival and reduce the incidence of organ dysfunction by modulating the inflammatory response.¹⁴ Moreover, a small size, retrospective study found a downtrend in the incidence of burn wound infection with the use of SDD.¹⁵ However, the use of SDD in burn units is scarce. This may be due to the limited high-quality evidence in this population and the difficulties of conducting multicenter studies with an adequate sample size. Moreover, the design of the intervention itself hinders the execution of randomized clinical trials, as the strategy should be implemented at the entire unit level rather than at the patient level.

Methods

Objective

The objective of this study is to evaluate the impact of implementing SDD in a Burns Unit (BU) on the incidence of HAIs in critically ill patients with acute burn injuries.

We assessed the incidence of infections that were considered potentially preventable with SDD: bacteremia, pneumonia, and skin and soft tissue or burn wound infection. Bacteremia and pneumonia were defined according to the ENVIN-HELICS Manual of Definitions and Terms.¹⁶ Skin and soft tissue or burn wound infection were defined as the growth of $\geq 10^5$ colony-forming units (CFU) of a microorganism per gram of tissue biopsy.^{17,18}

Study design

We conducted a retrospective before-and-after cohort study in a BU, between January 1st, 2017 and June 30th, 2023. Patients were divided into two groups based on exposure to SDD, which was adopted on March 1st, 2019. The primary endpoint was the incidence of HAIs during the stay in the BU. Secondary outcomes included the incidence of specific types of infections by site and microorganism, and safety endpoints.

The study was approved by the Institutional Review Board (Comité de Ética de la Investigación con Medicamentos del Área de Salud Valladolid Oeste, reference 21-EO195) with a waiver for informed consent, according to local regulations. The investigators adhered to the STROBE statement to report the study.¹⁹

Setting

The study was conducted in a 4-bed BU based in a tertiary referral University Hospital in Valladolid, Spain. Our area of influence for major burn patients covers a population of over 2.39 million inhabitants in the region of Castilla y León, Spain.

Our BU is provided with positive pressure rooms for the patients and a dedicated operating room. Patient care is led by plastic surgeons and intensive care physicians, and supported by other specialists as needed.

Admission criteria to the BU included:

- 1) Second- or third-degree burn injury affecting >20% TBSA in patients aged 14–60.
- 2) Second- or third-degree burn injury affecting >10% TBSA in either patients aged ≥ 60 , chemical or electric burns, in the context of major trauma, or in patients with comorbidities that may influence the prognosis.
- 3) Second- or third-degree burn injury affecting >5% TBSA in patients with suspected inhalation injury.

The management of the patients includes a primary and secondary assessment following the Advanced Trauma Life Support (ATLS) and Advanced Burn Life Support (ABLS) recommendations.^{20,21} This involves a comprehensive examination for an accurate evaluation of the extent²² and depth²³ of the burn, formal resuscitation with a protocol including albumin (Biological Engineering Technology [BET] formula),²⁴ and systematic screening for inhalation injury.^{25,26}

The treatment of burn wounds is based on either early surgical excision,^{27,28} early bromelain-based enzymatic debridement,²⁹ or a combination of both, along with early coverage using autografts.²⁸

General supportive measures, including thermoregulation, nutritional support, analgesia, and control of the hypermetabolic and inflammatory response, fall within the standard of care for the critically ill burned patient.³⁰

Participants

All the patients admitted to the BU during the study period were eligible for analysis, regardless of age and TBSA. We

excluded the patients who died or were discharged within the first 48 hours of admission, as HAI require more than 48 hours to develop. Additionally, patients with an exceedingly poor prognosis upon admission (Abbreviated Burn Severity Index [ABSI] >12 points, corresponding to a predicted survival rate on admission of <10%³¹) were excluded, unless they were deemed suitable for full escalation of treatment at the time of admission.

Interventions

SDD comprised three elements. Firstly, intravenous administration of cefotaxime 1 g every 8 hours for 4 days or, alternatively, levofloxacin 500 mg every 24 hours for 4 days in allergic patients. Secondly, oral or via nasogastric tube administration of a suspension containing 4% vancomycin, 1% colistin, 3% nystatin, and 1.2% tobramycin. Patients received 10 ml of the solution, previously shaken and diluted into 30 ml of water, every 8 hours until discharge. When administered via a nasogastric tube, it was flushed with water before and after the procedure. Thirdly, topical application of a paste containing 4% vancomycin, 2% colistin, 2% nystatin, and 3% tobramycin. Before application, the oral cavity was cleaned with an antiseptic (chlorhexidine 0.1%). The paste was then applied to the oral mucosa, gums, palate, and tracheal stoma in patients with tracheostomies and maintained in place for 20 minutes every 8 hours until discharge from the BU.

The multimodal approach of the Spanish ICU Zero Programs, endorsed by our national scientific society (*Sociedad Española de Medicina Intensiva, Crítica y Unidades Coronarias*, SEMICYUC) and other organizations, represents the basic framework in our unit and has been described elsewhere.^{11,32–34} The BU also participates in the Antimicrobial Stewardship Program (ASP) established in our institution, based on an *audit and feedback* strategy.³⁵ These strategies and policies did not change throughout the study time.

Variables and data sources

Age, sex and type of admission were retrieved from the electronic medical record (EMR) (IntelliSpace Critical Care and Anesthesia, Philips). APACHE II was calculated as described by Knaus WA et al.³⁶ ABSI was calculated as described by Tobiasen J et al.³¹ The percentage of burned TBSA was calculated as described by Lund CC and Browder NC.²² Inhalation injury was defined based on clinical evaluation, laboratory tests, bronchoscopy findings, and chest X-ray.^{25,26} Relevant comorbidities and potential risk exposures were defined as described in the Spanish surveillance program for ICU-acquired infections (ENVIN-HELICS registry).¹⁶ Previous history of smoking, alcohol abuse or psychiatric disorders was reported given their prognostic implications. HAIs were classified by type of microorganism and source. Acquisition of multidrug-resistant bacteria (MDRB) was extracted from the ENVIN-HELICS registry.¹⁶ Withdrawal/withholding of life support, duration of mechanical ventilation (MV), length of stay and mortality were retrieved from the EMR.

Statistical methods

The study was powered on the primary endpoint, which was the incidence of HAIs during the stay in the BU. We anticipated an incidence of HAI of 70% and the potential of SDD to achieve a 50% decrease in this rate. Assuming an enrollment ratio of 1:1, an alpha of 0.05, and a power of 0.8, the required sample size was 62 patients (31 in each group).

The quantitative variables were expressed as mean and 95% confidence interval (95% CI) or median (interquartile range, IQR). Qualitative variables were presented as absolute frequencies or percentages (%). Comparisons between quantitative variables were conducted using either the Student's T-test or the Mann-Whitney U test, depending on whether they exhibited a normal distribution or not. The Chi-square test was employed for comparing qualitative variables. The level of statistical significance was set at <0.05.

A multivariable logistic regression analysis was performed to ascertain the effects of explanatory predictor variables on the likelihood of HAI. Potential variables were identified based on previous literature.³⁷ Variable selection was conducted using the backward elimination method at a p-value of 0.2. Odds ratios (OR) and corresponding 95% CIs were calculated.

Statistical calculations were conducted using Stata 16 statistical software (StataCorp LLC, Texas).

Results

Patients and descriptive data

During the study period, 86 patients were admitted to the BU. Six patients were excluded from the analysis due to a hospital stay of <48 hours: three died during this period (burned surface areas of 15%, 28%, and 60%), and three were discharged alive (burned surface areas of 7%, 8%, and 17%). Additionally, four patients with ABSI >12 points were considered unsuitable for full escalation of treatment and, thus, were excluded from the analysis.

A total of 72 patients were analyzed: 27 did not receive SDD, and 45 received SDD. None of the patients discontinued the SDD regimen during the BU stay. The clinical characteristics of the patients during the two study periods are summarized in [Table 1](#).

Outcome data and main results

The incidence of HAIs in both the non-SDD and SDD groups, categorized by site and microorganism, is illustrated in [Table 2](#). The mean number of hospital-acquired infectious episodes and the median time to BU-acquired infection are depicted in [Table 3](#). Mortality, withdrawal or withholding of life support, duration of MV, and length of stay are reported in [Table 4](#). SDD was not independently associated with mortality. Logistic regression analysis was performed to examine the influence of predictor variables on the incidence of HAIs ([Table 5](#)).

Table 1 Baseline Characteristics.

Characteristic	Regimen		p
	Non-SDD group (n = 27)	SDD group (n = 45)	
Age at time of BU, years, Mean (95% CI)	61.8 (54.7–68.9)	58.1 (52.2–64)	0.43
Male sex, No. (%)	20 (74.1)	30 (66.7)	0.51
Burn characteristics			
Inhalation injury, No. (%)	8 (29.6)	9 (20)	0.35
Flame injury, No. (%)	23 (85.2)	39 (86.7)	0.51
%TBSA, Median (IQR)	20 (11–30)	20 (12–30)	0.94
ABSI, Mean (95% CI)	7.6 (6.9–8.3)	7.6 (7–8.1)	1
APACHE II, Mean (95% CI)	18.3 (15.8–20.7)	14.5 (12.9–16.2)	<0.01
Type of admission			
Community or emergency room, No. (%)	21 (77.8)	30 (66.7)	0.03
Referral from non-specialized ICU, No. (%)	0 (0)	10 (22.2)	
Other, No. (%)	6 (22.2)	5 (11.1)	
Previous or preexistent condition, No. (%)			
Diabetes mellitus	5 (18.5)	7 (15.6)	0.74
Chronic kidney disease	1 (3.7)	3 (6.7)	0.60
Neoplasia	3 (11.1)	2 (4.4)	0.28
Poor nutritional status	3 (11.1)	5 (11.1)	1
Smoking	11 (40.7)	14 (31.1)	0.41
Alcohol abuse	8 (29.6)	5 (11.1)	0.05
Psychiatric disorder	5 (18.5)	7 (15.6)	0.74
Risk factors for infections, No. (%)			
Central venous catheterization	25 (92.6)	45 (100)	0.06
Mechanical ventilation	16 (59.3)	34 (75.6)	0.15
Parenteral nutrition	3 (11.1)	3 (6.7)	0.51
Renal replacement therapy	2 (7.4)	6 (13.3)	0.44
Urgent surgery	5 (18.5)	8 (17.8)	0.94

Table 2 Incidence of Hospital-Acquired Infections for Patients With a Length of BU Stay More Than 2 Days.

	Regimen		OR (95% CI) SDD vs no SDD	p
	Non-SDD group (n = 27)	SDD group (n = 45)		
Source of Infection, No. (%)				
Any	21 (77.8)	21 (46.7)	0.25 (0.09–0.72)	0.009
Bacteremia	7 (25.9)	8 (17.8)	0.62 (0.2–1.89)	0.409
Pneumonia	12 (44.4)	10 (22.2)	0.36 (0.13–0.99)	0.048
Skin or soft tissue	11 (40.7)	13 (28.9)	0.59 (0.22–1.59)	0.302
Gram-positive Infections, No. (%)				
Any	15 (55.6)	11 (24.4)	0.26 (0.09–0.71)	0.008
<i>Staphylococcus aureus</i>	2 (7.4)	5 (11.1)	1.56 (0.32–17.49)	0.608
Other <i>Staphylococci</i>	12 (44.4)	6 (13.3)	0.19 (0.06–0.59)	0.003
<i>Streptococcus pneumoniae</i>	3 (11.1)	0 (0)	0 (0–0.72)	0.022
Gram-negative Infections, No. (%)				
Any	15 (55.6)	11 (24.4)	0.26 (0.09–0.71)	0.008
<i>Enterobacteriaceae</i>	10 (37)	3 (6.7)	0.12 (0.03–0.47)	0.001
Nonfermenting Gram-negative bacilli	7 (25.9)	7 (15.6)	0.53 (0.17–1.65)	0.282
Other Gram-negative bacteria	5 (18.5)	1 (2.2)	0.1 (0–0.7)	0.015
Fungal Infections, No. (%)				
Any	5 (18.5)	7 (15.6)	0.81 (0.24–2.72)	0.744

Table 3 Number of Hospital-Acquired Infectious Episodes Per Patient and Time to Infection for Patients with a Length of BU Stay More Than 2 Days.

	Regimen		p
	Non-SDD group (n = 27)	SDD group (n = 45)	
Any BU-acquired infection			
Number of episodes per patient, Mean (95% CI)	2.52 (1.21–3.82)	1.13 (0.54–1.73)	0.029
Time to infection (days), Median (IQR)	7 (5–13)	16 (6–29)	0.040
Bacteremia			
Number of episodes per patient, Mean (95% CI)	0.37 (0.08–0.66)	0.22 (0.07–0.38)	0.322
Time to infection (days), Median (IQR)	22 (1–37)	17 (12–49)	0.807
Pneumonia			
Number of episodes per patient, Mean (95% CI)	1 (0.36–1.64)	0.33 (0.12–0.55)	0.018
Time to infection (days), Median (IQR)	5 (4–10)	9 (5–17)	0.220
Skin or soft tissue			
Number of episodes per patient, Mean (95% CI)	1.15 (0.37–1.92)	0.58 (0.22–0.93)	0.126
Time to infection (days), Median (IQR)	15 (10–31)	25 (14–30)	0.505

Table 4 Safety Endpoints.

	Regimen		p
	Non-SDD group (n = 27)	SDD group (n = 45)	
Acquisition of MDRB during BU stay, No. (%)	8 (26.6)	10 (22.2)	0.482
HAI produced by MDRB during BU stay, No. (%)	5 (18.5)	3 (6.7)	0.121
Duration of MV (days), Median (IQR)	13.5 (6.5–29)	11.5 (4–31)	0.755
Length of stay (days), Median (IQR)			
BU	42 (17–54)	29 (13–58)	0.762
Hospital	42 (20–55)	38 (21–64)	0.830
Withdrawal/withholding of life support, No. (%)	4 (14.8)	5 (11.1)	0.645
Mortality, No. (%)			
Day 28	3 (11.1)	6 (13.3)	0.783
BU	4 (14.8)	7 (15.6)	0.942
Hospital	5 (18.5)	7 (15.6)	0.744

Table 5 Logistic regression analysis examining the influence of predictor variables on the incidence of HAIs. The model as a whole was significant (n = 72, LR $\chi^2(3) = 35.02$, pseudo- $R^2 = 0.36$, $p < 0.001$, Akaike's information criterion [AIC] = 70.79).

	OR (95% CI)	p
Female sex	3.89 (0.96–15.78)	0.057
Duration of mechanical ventilation, per day	1.15 (1.06–1.24)	<0.001
Selective decontamination of the digestive tract	0.09 (0.02–0.39)	0.001

Discussion

Key findings

We found SDD was associated with a lower incidence of HAIs during the BU stay. SDD acted as an independent pro-

TECTIVE factor against BU-acquired infection, even after adjusting for confounding factors. This finding is probably explained by a significant reduction in the incidence of pneumonia and a downtrend in the incidence of other infections. SDD showed a significant decrease in Gram-positive and Gram-negative infections, with no observed effect on fungal

infections. Moreover, patients who received SDD developed fewer episodes of infection per patient compared to those who did not, and these infections occurred significantly later. Finally, SDD was not associated with improved survival, decreased duration of MV, or reduced length of stay.

Relationship to previous studies

Infections remain among the leading causes of mortality in burn patients, but they may be reduced by adopting prevention strategies.³⁸ In this regard, the only single-center randomized, controlled trial (RCT) conducted in a Spanish BU showed SDD was associated with a reduction in both BU and hospital mortality, and a lower incidence of pneumonia.³⁹ Besides, observational studies have shown a possible mortality benefit of SDD in critically ill burn patients.^{15,40,41} In addition, SDD has proved to reduce the burden of organ dysfunction in the burn patient.⁴² According to a recent systematic review of RCTs and observational studies,¹⁴ SDD may improve the survival of critically ill burn patients by reducing the incidence of infections such as pneumonia and bloodstream infections. Despite these findings, implementation of SDD in BUs is scarce. Although some experts have postulated a link between SDD and an increased risk of MDRB, neither contemporary literature¹⁴ nor our study provide results that support this hypothesis in burn patients.

While the strengths of the RCT conducted by De la Cal et al.³⁹ are undeniable, the study did not address the benefits of implementing SDD as a unit-wide infection prevention policy. Therefore, not including all patients in the burn unit may have resulted in selection bias. In contrast, our before-and-after study exhibits a more pragmatic approach, as it investigates the effect of SDD on all the patients admitted during the study period. Besides, the implementation of the "Zero Programs", aiming to reduce the incidence of catheter-related bloodstream infections, VAP, and multidrug resistance in Spanish ICUs,^{11,32–34} as well as the participation in ASPs,³⁵ may affect the external validity and extrapolation of results from studies conducted a few years ago to the present moment.

Preventing pneumonia is a primary goal of SDD.¹⁰ De la Cal et al.³⁹ observed a decreasing trend in pneumonia incidence among burn patients receiving SDD (48% vs. 34%, $p=0.14$) and a reduction in the mean number of pneumonia episodes per patient from 0.67 (0.54–0.79) to 0.41 (0.28–0.55), $p<0.01$. Similarly, Mackie et al.¹⁵ reported a substantial decrease in pneumonia incidence among burn patients on SDD (27% vs. 7%, $p=0.03$). However, their study's generalizability may be limited as follow-up was censored at 28 days and only five categories of microorganisms were examined. In contrast, our study offers additional insights, as patients were monitored until discharge and all microorganisms were documented. We found a lower pneumonia incidence in the SDD group (44.4% vs. 22.2%, $p=0.048$) and a decrease in the mean number of pneumonia episodes per patient from 1 (0.36–1.64) to 0.33 (0.12–0.55), $p=0.018$. These findings hold particular relevance, especially in light of the implementation of a national multimodal intervention designed to prevent VAP in our BU in 2011.³³ Our logistic regression analysis showed that both SDD and the duration

of MV were independently associated with the risk of developing infection, consistent with previous studies.⁴³

The effect of SDD on bloodstream infections in critically ill burn patients is controversial, with observational studies suggesting benefit,^{15,44–46} but with no effect in the only RCT.³⁹ In line with the findings of the before-mentioned RCT, we did not find differences in the incidence of bacteremia or the mean number of episodes of bacteremia per patient between both groups. However, the incidence of bacteremia in our non-SDD group was only 25.9%, compared to 26–80% in the other studies.^{15,39,44–46} The potential benefit of SDD in reducing bloodstream infections may have been weakened by the introduction of a national multimodal intervention aimed at preventing catheter-related bloodstream infections in our BU in 2014.³² Of note, we found a downtrend in the incidence of bacteremia in our SDD group, despite a higher rate of central venous catheterization, close to statistical significance.

Some studies have hypothesized that SDD may influence the prevention of skin and soft tissue infections or burn wound infections,¹⁵ while others have demonstrated no discernible effect.^{39,47} Consistent with the latter, we observed comparable rates in both the non-SDD (40.7%) and SDD groups (28.9%).

A reduction in mortality has been linked to SDD in burn patients, decreasing from 28% to 9% in an RCT³⁹ and from 23% to 3% in an observational study.¹⁵ However, we found similar hospital mortality rates in the non-SDD group (18.5%) and the SDD group (15.6%). Our study was underpowered to detect differences in mortality because of its low sample size and the low mortality observed in both groups.

Clinical implications

This study implies that SDD reduces the incidence of HAIs and the number of infectious episodes per patient during BU stay, together with an increase in the time to the development of infections. Therefore, SDD represents a reasonable strategy to reduce the burden of infections in this population. Since infectious complications are common in severely ill burn patients, our finding has an impact on the cost of burn care⁹ and the potential to reduce its associated morbidity and disability.

Strengths and limitations

This study evaluated the effect of SDD as a unit-wide infection prevention policy applied to all the patients admitted to a BU. Thus, the patient population was less likely to be subject to selection bias than in previous studies. Furthermore, our study is the first to address the effect of SDD in the setting of modern BU care, where further infection prevention strategies and ASPs are commonly implemented. Finally, the pragmatic approach of our work makes it more generalizable.

We acknowledge several limitations. Although the before-and-after study design utilized in this research implies a quasi-experimental design, it also introduces acknowledged biases. The absence of a concurrent control group presents challenges in establishing a direct

causal relationship between the intervention and outcomes, potentially allowing confounding by external factors and temporal trends. Randomization was not employed due to the unit-wide implementation of SDD, making a control group unfeasible. Concerns about selection bias arise from the non-random assignment of participants, which is mitigated by reporting baseline characteristics. We attribute the significantly lower APACHE II score in the SDD group to the higher number of patients referred from a non-specialized ICU, leading to a lower score calculation within the first 24 hours of BU admission. Despite the potential for information bias, data collection methods and surveillance practices were kept consistent across the before and after periods.

Conclusions

Selective decontamination of the digestive tract in a Burns Unit is associated with a reduced incidence of hospital-acquired bacterial infections and a decrease in the number of infectious episodes per patient, which is not associated with reduced mortality.

Authors' contributions

DPT, AIML and JABG designed the study. DPT, AIML, JABG, MLFR, JSB, PBS and JMPP participated in the management of the patients during the study period. DPT, AIML, CCR, JABG, CDR, ICP, MLFR, CCC, JSB, PBS and TGTE extracted the data from the patients from the electronic medical record. DPT, AIML, JABG, PBS and JMPP critically interpreted the data. DPT drafted the manuscript. All authors read and approved the final text.

Ethics approval and consent to participate

The study was approved by the Institutional Review Board (Comité de Ética de la Investigación con Medicamentos del Área de Salud Valladolid Oeste, reference 21-EO195) with a waiver for informed consent, according to local regulations.

Consent for publication

Not applicable.

Availability of data and materials

The datasets analyzed during the current study are available from the corresponding author on reasonable request.

Competing interests

The authors declare that they have no competing interests.

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References

- Hansbrough JF, Field TO, Gadd MA, Soderberg C. Immune response modulation after burn injury: T cells and antibodies. *J Burn Care Rehabil.* 1987;8(6):509–12, <http://dx.doi.org/10.1097/00004630-198708060-00011>.
- Alexander JW. Mechanism of immunologic suppression in burn injury. *J Trauma.* 1990;30 12 Suppl, <http://dx.doi.org/10.1097/00005373-199012001-00017>.
- Heideman M, Bengtsson A. The immunologic response to thermal injury. *World J Surg.* 1992;16(1):53–6, <http://dx.doi.org/10.1007/BF02067115>.
- Kiley JL, Greenhalgh DG. Infections in burn patients. *Surg Clin North Am.* 2023;103(3):427–37, <http://dx.doi.org/10.1016/J.SUC.2023.02.005>.
- Lachiewicz AM, Hauck CG, Weber DJ, Cairns BA, Van Duin D. Bacterial infections after burn injuries: impact of multidrug resistance. *Clin Infect Dis.* 2017;65(12):2130–6, <http://dx.doi.org/10.1093/CID/CIX682>.
- Fitzwater J, Purdue GF, Hunt JL, O'keefe GE. The risk factors and time course of sepsis and organ dysfunction after burn trauma. *J Trauma.* 2003;54(5):959–66, <http://dx.doi.org/10.1097/01.TA.0000029382.26295.AB>.
- Stanojic M, Vainik R, Jeschke MG. Status and challenges of predicting and diagnosing sepsis in burn patients. *Surg Infect (Larchmt).* 2018;19(2):168–75, <http://dx.doi.org/10.1089/SUR.2017.288>.
- Dvorak JE, Ladhani HA, Claridge JA. Review of sepsis in burn patients in 2020. *Surg Infect (Larchmt).* 2021;22(1):37–43, <http://dx.doi.org/10.1089/SUR.2020.367>.
- Tsai SY, Lio CF, Yao WC, Liu CP, Shih SC, Wang TYT, et al. Cost-drivers of medical expenses in burn care management. *Burns.* 2020;46(4):817–24, <http://dx.doi.org/10.1016/J.BURNS.2020.01.004>.
- Wittekamp BHJ, Oostdijk EAN, Cuthbertson BH, Brun-Buisson C, Bonten MJM. Selective decontamination of the digestive tract (SDD) in critically ill patients: a narrative review. *Intensive Care Med.* 2020;46(2):343–9, <http://dx.doi.org/10.1007/S00134-019-05883-9>.
- Arias-Rivera S, Jam-Gatell R, Nuvials-Casals X, Vázquez-Calatayud M. Update of the recommendations of the Pneumonia Zero project. *Enferm Intensiva.* 2022;33:517–30, <http://dx.doi.org/10.1016/J.ENFI.2022.05.005>.
- Klompas M, Branson R, Cawcutt K, Crist M, Eichenwald EC, Greene LR, et al. Strategies to prevent ventilator-associated pneumonia, ventilator-associated events, and nonventilator hospital-acquired pneumonia in acute-care hospitals: 2022 Update. *Infect Control Hosp Epidemiol.* 2022;43(6):687–713, <http://dx.doi.org/10.1017/ICE.2022.88>.
- Hammond NE, Myburgh J, Seppelt I, Garside T, Vlok R, Mahendran S, et al. Association between selective decontamination of the digestive tract and in-hospital mortality in intensive care unit patients receiving mechanical ventilation: a systematic review and meta-analysis. *JAMA.* 2022;328(19):1922–34, <http://dx.doi.org/10.1001/JAMA.2022.19709>.
- Rubio-Regidor M, Martín-Pellicer A, Silvestri L, van Saene HKF, Lorente JA, de la Cal MA. Digestive decontamination in burn patients: a systematic review of randomized clinical trials and observational studies. *Burns.* 2018;44(1):16–23, <http://dx.doi.org/10.1016/J.BURNS.2017.04.001>.
- Mackie D, van Hertum W, Schumburg T, Kuijper E, Knappe P. Prevention of infection in burns: preliminary experience with selective decontamination of the digestive tract in patients with extensive injuries. *J Trauma.* 1992;32(5):570–5, <http://dx.doi.org/10.1097/00005373-199205000-00006>.
- SEMICYUC, Grupo de Trabajo de Enfermedades Infecciosas y Sepsis (GTEIS). Manual de Definiciones y Términos, Estudio

- Nacional de Vigilancia de Infección Nosocomial en UCI (ENVIN-HELICS) [Internet]. 2018 [Accessed 2023 Feb 7].
17. Bowler PG, Duerden BI, Armstrong DG. Wound microbiology and associated approaches to wound management. *Clin Microbiol Rev.* 2001;14(2):244–69, <http://dx.doi.org/10.1128/CMR.14.2.244-269.2001>.
 18. Bonham PA. Swab cultures for diagnosing wound infections: a literature review and clinical guideline. *J Wound Ostomy Cont Nurs Off Publ Wound Ostomy Cont Nurses Soc.* 2009;36(4):389–95, <http://dx.doi.org/10.1097/WON.0B013E3181AAEF7F>.
 19. von Elm E, Altman DG, Egger M, Pocock SJ, Gøtzsche PC, Vandenbroucke JP. The strengthening of reporting of observational studies in epidemiology (STROBE) statement: guidelines for reporting observational studies. *Lancet (London, England).* 2007;370(9596):1453–7, [http://dx.doi.org/10.1016/S0140-6736\(07\)61602-X](http://dx.doi.org/10.1016/S0140-6736(07)61602-X).
 20. Henry S, editor. *ATLS Advanced Trauma Life Support, Student Course Manual.* 10th Edition Chicago: ACS American College of Surgeons; 2018.
 21. Pham TN, Bettencourt AP, Bozinko GM, Chang PH, Chung KK, Craig CK, et al. *Advanced Burn Life Support Course. Provider Manual.* 2018 Update. Chicago: American Burn Association; 2018.
 22. Lund CC, Browder NC. The estimation of area of burns. *Surg Gynecol Obs.* 1944;79(79):352–8.
 23. Hettiaratchy S, Papini R. Initial management of a major burn: II-assessment and resuscitation. *BMJ.* 2004;329(7457):101–3, <http://dx.doi.org/10.1136/BMJ.329.7457.101>.
 24. Blanco-Schweizer P, Sánchez-Ballesteros J, Bendito B, Martín AI, Fernández L, Piqueras JM, et al. Resuscitation with albumin using BET formula keeps at bay fluid administration in burned patients. An observational study. *Burns.* 2020;46(4):860–7, <http://dx.doi.org/10.1016/J.BURNS.2019.10.024>.
 25. Sheridan RL. Fire-related inhalation injury. *N Engl J Med.* 2016;375(5):464–9, <http://dx.doi.org/10.1056/NEJMRA1601128>.
 26. Canas-Pérez I, Pérez-Torres D, Martín-Luengo AI. Inhalation injury. *Med Intensiva.* 2023;47(9):558–9, <http://dx.doi.org/10.1016/J.MEDINE.2023.02.002>.
 27. Still JJ, Law E. Primary excision of the burn wound. *Clin Plast Surg.* 2000;27(1):23–47.
 28. Jeschke MG, van Baar ME, Choudhry MA, Chung KK, Gibran NS, Logsetty S. Burn injury. *Nat Rev Dis Prim.* 2020;6(1), <http://dx.doi.org/10.1038/S41572-020-0145-5>.
 29. Serracanta J, Baena J, Martínez-Mendez JR, Sánchez-Sánchez M, López-Suso E, Galeiras R, et al. Bromelain-based enzymatic burn debridement: Spanish multidisciplinary consensus. *Eur J Plast Surg.* 2023;46(2):271–9, <http://dx.doi.org/10.1007/S00238-022-01999-2>.
 30. McGovern C, Puxty K, Paton L. Major burns: part 2. Anaesthesia, intensive care and pain management. *BJA Educ.* 2022;22(4):138–45, <http://dx.doi.org/10.1016/J.BJAE.2022.01.001>.
 31. Tobiasen J, Hiebert JM, Edlich RF. The abbreviated burn severity index. *Ann Emerg Med.* 1982;11(5):260–2, [http://dx.doi.org/10.1016/S0196-0644\(82\)80096-6](http://dx.doi.org/10.1016/S0196-0644(82)80096-6).
 32. Palomar M, Álvarez-Lerma F, Riera A, Díaz MT, Torres F, Agra Y, et al. Impact of a national multimodal intervention to prevent catheter-related bloodstream infection in the ICU: the Spanish experience. *Crit Care Med.* 2013;41(10):2364–72, <http://dx.doi.org/10.1097/CCM.0B013E3182923622>.
 33. Álvarez Lerma F, Sánchez García M, Lorente L, Gordo F, Añón JM, Álvarez J, et al. Guidelines for the prevention of ventilator-associated pneumonia and their implementation. The Spanish 'Zero-VAP' bundle. *Med Intensiva.* 2014;38(4):226–36, <http://dx.doi.org/10.1016/J.MEDIN.2013.12.007>.
 34. Garnacho Montero J, Lerma FÁ, Gallego PR, Martínez MP, Rocha LÁ, Gaité FB, et al. Combatting resistance in intensive care: the multimodal approach of the Spanish ICU 'Zero Resistance' program. *Crit Care.* 2015;19(1), <http://dx.doi.org/10.1186/S13054-015-0800-5>.
 35. Pérez-Torres D, Tamayo-Lomas LM, González MDG, Almendros-Muñoz R, Sacristán-Salgado MA, González-González E, et al. Antimicrobial stewardship program in an Intensive Care Unit: a retrospective observational analysis of the results 15 months after its implementation. *Rev Esp Quimioter.* 2023;36(5):477–85, <http://dx.doi.org/10.37201/REQ/142.2022>.
 36. Knaus WA, Draper EA, Wagner DP, Zimmerman JE. APACHE II: a severity of disease classification system. *Crit Care Med.* 1985;13(10):818–29, <http://dx.doi.org/10.1097/00003246-198510000-00009>.
 37. Stoltzfus JC. Logistic regression: a brief primer. *Acad Emerg Med.* 2011;18(10):1099–104, <http://dx.doi.org/10.1111/J.1553-2712.2011.01185.X>.
 38. Vainik R, Barayan D, Shahrokhi S, Jeschke MG. Management and prevention of drug resistant infections in burn patients. *Expert Rev Anti Infect Ther.* 2019;17(8):607–19, <http://dx.doi.org/10.1080/14787210.2019.1648208>.
 39. De La Cal MA, Cerdá E, García-Hierro P, Van Saene HKF, Gómez-Santos D, Negro E, et al. Survival benefit in critically ill burned patients receiving selective decontamination of the digestive tract: a randomized, placebo-controlled, double-blind trial. *Ann Surg.* 2005;241(3):424–30, <http://dx.doi.org/10.1097/01.SLA.0000154148.58154.D5>.
 40. Jarrett F, Balish E, Moylan JA, Ellerbe S. Clinical experience with prophylactic antibiotic bowel suppression in burn patients. *Surgery.* 1978;83(5):523–7, <http://dx.doi.org/10.5555/URI:PII:003960607890154X>.
 41. Cerdá E, Abella A, De La Cal MA, Lorente JA, García-Hierro P, Van Saene HKF, et al. Enteral vancomycin controls methicillin-resistant *Staphylococcus aureus* endemicity in an intensive care burn unit: a 9-year prospective study. *Ann Surg.* 2007;245(3):397–407, <http://dx.doi.org/10.1097/01.SLA.0000250418.14359.31>.
 42. López-Rodríguez L, De La Cal MA, García-Hierro P, Herrero R, Martins J, Van Saene HKF, et al. Selective digestive decontamination attenuates organ dysfunction in critically ill burn patients. *Shock.* 2016;46(5):492–7, <http://dx.doi.org/10.1097/SHK.0000000000000664>.
 43. McCusker ME, Périssé ARS, Roghmann MC. Severity-of-illness markers as predictors of nosocomial infection in adult intensive care unit patients. *Am J Infect Control.* 2002;30(3):139–44, <http://dx.doi.org/10.1067/mic.2002.121662>.
 44. Jarrett F, Balish E, Moylan J, Ellerbe S. Clinical experience with prophylactic antibiotic bowel suppression in burn patients. *Surgery.* 1978;83(5):523–7.
 45. Shalaby H, Higazi M, El Far N. Selective gastrointestinal decontamination and burn wound sepsis. *Ann Burns Fire Disasters.* 1998;11(1):1–6.
 46. Aboelatta Y, Abd-Elsalam A, Omar A, Abdelaal M, Farid A. Selective digestive decontamination (SDD) as a tool in the management of bacterial translocation following major burns. *Ann Burns Fire Disasters.* 2013;26(4):182–8.
 47. Barret JP, Jeschke MG, Herndon DN. Selective decontamination of the digestive tract in severely burned pediatric patients. *Burns.* 2001;27(5):439–45, [http://dx.doi.org/10.1016/S0305-4179\(00\)00147-9](http://dx.doi.org/10.1016/S0305-4179(00)00147-9).