



The spectrum and outcome of surgical sepsis in Pietermaritzburg, South Africa

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Background. Sepsis is a leading cause of morbidity and mortality worldwide, and the incidence appears to be increasing. In the resource-limited environment in low- and middle-income countries, the management of surgical sepsis (SS) continues to represent a significant portion of the workload for most general surgeons.

Objective. To describe the spectrum of SS seen at a busy emergency department, and categorise the outcomes.

Methods. The Pietermaritzburg Metropolitan Trauma Service (PMTS) and Pietermaritzburg Metropolitan Surgical Service (PMSS) in KwaZulu-Natal Province, South Africa (SA), maintain a prospective electronic registry. All patients with features of sepsis among emergency general surgical patients >15 years of age admitted to the PMSS over the period January 2012 - January 2015 were identified. From this cohort, all patients with sepsis that required surgical source control or who had a documented surgical source of sepsis (i.e. had SS) were selected for analysis.

Results. Of a total of 6 020 adult surgical patients on the database, a cohort of 1 240 acute surgical patients with features of sepsis were identified, and 675 with SS were then analysed further. Of the 675 patients, 49.2% were male, and the mean age was 46 years (standard deviation (SD) 19); 47.0% presented to the PMSS directly from within the metropolitan area, while the remaining 53.0% were referred from hospitals outside the area. Physiological parameters (mean values) on presentation were as follows: systolic blood pressure 123 mmHg (standard deviation (SD) 23), respiratory rate 22 breaths/min (SD 5.2), heart rate 107 bpm (SD 19), temperature 37°C (SD 2) and white cell count $20 \times 10^9/L$ (SD 8). Of the patients, 21.6% were known to be HIV-positive, 13.5% (91/675) were negative and 64.9% were of unknown status; 57.6% had intra-abdominal sepsis, 26.1% diabetes-related limb sepsis and the remaining 16.3% soft-tissue infections; 17.5% required intensive care unit admission, with a mean length of stay of 4 days (SD 4), and 30.7% developed complications. In this last group ($n=207$), a total of 313 morbidities were identified. The overall mortality rate was 12.7% (86/675). The mortality rate for intra-abdominal sepsis was 13.1%, for diabetic foot sepsis 14.2% and for necrotising fasciitis 27.3%.

Conclusions. The spectrum of SS in SA is different to that seen in the developed world. Intra-abdominal sepsis is the most common SS and is overwhelmingly caused by acute appendicitis. Diabetic foot infection is a major cause of SS, reflecting the increasing burden of non-communicable chronic diseases in SA.

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Sepsis is a leading cause of morbidity and mortality worldwide, and the incidence appears to be increasing.^[1-4] Surgical conditions are responsible for up to one-third of the overall disease burden of sepsis. The annual cost of hospital care of sepsis is estimated to be over USD14 billion in the USA alone. Surgical sepsis (SS) is a broad term covering a spectrum of conditions in which the patient has a septic process that requires surgical source control. In addition to operative management, these patients require aggressive resuscitation and organ support.^[5-11] In the resource-limited environment in low- and middle-income countries (LMICs), the management of SS continues to represent a significant portion of the workload for most general surgeons.

Objective

To review the spectrum of SS disease and outcome among patients who presented to a major tertiary surgical service in South Africa (SA).

Methods

This retrospective study was undertaken at Grey's Hospital in Pietermaritzburg, the capital of KwaZulu-Natal Province, SA. The Pietermaritzburg Metropolitan Surgical Service (PMSS) covers the entire western part of the province, with a total catchment population of over 3 million, most of whom are black Africans.

The electronic clinical registry^[12] at the PMSS was retrospectively reviewed from January 2012 to January 2015. The electronic registry is prospectively maintained in 'real time', where patients' clinical information is entered directly into an electronic database. On review of the prospectively maintained database, all emergency general surgical patients aged >15 years presenting to the emergency room who fulfilled the standard Surviving Sepsis Campaign (SSC) criteria^[1] for systemic inflammatory response syndrome (SIRS), sepsis, severe sepsis or septic shock from January 2012 to January 2015 were identified. The following criteria (two or more had to be present) were used to identify patients with sepsis: fever (temperature >38.3°C

or hypothermia (core temperature $<36^{\circ}\text{C}$), heart rate >90 bpm or more than two standard deviations (SDs) above the normal value for age, tachypnoea, altered mental status, and hyperglycaemia (plasma glucose >7.7 mmol/L) in the absence of diabetes. Once this cohort had been identified, the registry was reviewed to identify all patients with sepsis that required surgical source control or who had a documented surgical source of sepsis (i.e. had SS). Those with incomplete records or without subsequent surgical confirmation of the source of sepsis were excluded. Data reviewed from the registry included basic demographic information, admission physiological parameters, diagnosis, requirement for intensive care unit (ICU) admission, length of hospital stay and mortality. Morbidities were classified according to organ system, e.g. hospital-acquired pneumonia was classified as a respiratory complication. The surgical pathologies were classified into the following categories: intra-abdominal sepsis, limb sepsis and soft-tissue sepsis.

Statistical analysis

Data were processed and analysed using Stata version 13.0 (StataCorp, USA). Continuous variables were summarised using means and SDs. If there was evidence of skewing or asymmetrical outliers, medians and interquartile ranges were presented instead.

Ethics approval

Ethics approval for this study was formally granted by the Biomedical Research Ethics Committee of the University of KwaZulu-Natal (ref. no. BCA221/13) and the Knowledge and Epidemiology Unit of the Department of Health (no reference number).

Table 1. The spectrum of SS according to anatomical region (N=675)

	<i>n</i> (%)
Abdominal sepsis	389/675 (57.6)
Appendicitis	156/389 (40.1)
Perforated peptic ulcer	60/389 (15.4)
Biliary sepsis	47/389 (12.1)
Intra-abdominal abscess	25/389 (6.4)
Tuberculosis of the abdomen	21/389 (5.3)
Perforated viscus (non-peptic ulcers)	31/389 (7.9)
Pelvic inflammatory disease	15/389 (3.8)
Complicated bowel ischaemia	14/389 (3.5)
Diverticulitis	5/389 (1.2)
Anastomotic leak	5/389 (1.2)
Infective colitis	4/389 (1.0)
Peritoneal dialysis catheter sepsis	4/389 (1.0)
Primary peritonitis	2/389 (0.5)
Diabetic foot sepsis	176/675 (26.1)
Soft-tissue sepsis	110/675 (16.3)
Necrotising fasciitis	33/110 (30.0)
Soft-tissue abscesses	77/110 (70.0)
Head/neck	24/77 (31.2)
Perianal	15/77 (19.5)
Limbs	11/77 (14.3)
Trunk	11/77 (14.3)
Other	16/77 (20.8)

Results

Demographics

Of a total of 6 020 adult surgical patients on the database, we identified 1 240 acute surgical patients with SIRS. Of these, 675 patients with sepsis that required surgical source control or who had a documented surgical source of sepsis (i.e. had SS) were analysed further; 49.2% (332/675) were male, the mean age was 46 years (SD 19), and 47.0% (317/675) presented to the PMSS directly from within the metropolitan area, while the remaining 53.0% (358/675) were referred from other hospitals outside the area.

Clinical presentation

Physiological parameters (mean values) on presentation were as follows: systolic blood pressure 123 mmHg (SD 23), respiratory rate 22 breaths/min (SD 5.2), heart rate 107 bpm (SD 19), temperature 37°C (SD 2), and white cell count $20 \times 10^9/\text{L}$ (SD 8). Of the patients, 21.6% were known to be HIV-positive, 13.5% (91/675) were negative and 64.9% were of unknown status.

Surgical disease spectrum

Of the 675 patients, 57.6% ($n=389$) had intra-abdominal sepsis, 26.1% ($n=176$) had diabetes-related limb sepsis and the remaining 16.3% ($n=110$) had soft-tissue infections. Table 1 summarises the spectrum of surgical pathologies in each category. A total of 585 operative procedures were performed for source control. These are summarised in Table 2.

Outcome

Of all the patients, 17.5% (118/675) required ICU admission, with a mean length of stay of 4 days (SD 4), and 30.7% (207/675) developed

Table 2. Procedures to obtain source control (N=585)

Procedures	<i>n</i> (%)
Laparotomy	281 (48.0)
Amputation	148 (25.3)
Incision and drainage	55 (9.4)
Debridement	39 (6.7)
Appendectomy	18 (3.1)
Radiological procedure	15 (2.6)
Endoscopic retrograde cholangiopancreatography	13 (2.2)
Laparoscopic cholecystectomy	8 (1.4)
Open cholecystectomy	8 (1.4)

Table 3. Organ system-related complications (N=313)

Complications (organ system)	<i>n</i> (%)
Neurological	74 (23.6)
Respiratory	73 (23.3)
Wound	70 (22.3)
Renal	47 (15.1)
Gastrointestinal	30 (9.5)
Cardiac	17 (5.4)
Urinary	2 (0.6)

Table 4. Mortality according to site of sepsis

Site of sepsis	Outcome		Total, N
	Died, n (%)	Survived, n (%)	
Intra-abdominal	51 (13.1)	338 (86.9)	389
Diabetic foot sepsis	25 (14.2)	151 (85.8)	176
Soft-tissue sepsis			
Abscess	1 (1.6)	60 (98.4)	61
Necrotising fasciitis	9 (27.3)	24 (72.7)	33
Other	-	16 (100)	16
Total	86 (12.7)	589 (87.3)	675

complications during the course of their illness. A total of 313 morbidities were identified in these patients (Table 3). The overall mortality rate was 12.7% (86/675). The mortality rate for intra-abdominal sepsis was 13.1%, for diabetic foot sepsis 14.2% and for necrotising fasciitis 27.3%. Table 4 summarises mortality according to type of sepsis.

Discussion

Sepsis remains a leading cause of death and a significant health problem worldwide.^[1-5] Despite the considerable amount of resources invested in managing and researching sepsis, the incidence continues to increase with little change in overall mortality. Most of the current management strategies originated from literature in the developed world (high-income countries, HICs), and literature focusing specifically on the spectrum of disease in LMICs is limited.^[1-5]

Our study highlighted the fact that several aspects of SS in SA differ from the experience in HICs.^[12-15] The patient cohort is generally much younger than in HICs, as mean life expectancy is lower in SA. The most common SS is intra-abdominal sepsis, which is mainly secondary to acute appendicitis, followed by complicated peptic ulcer disease. In HICs, colonic perforation secondary to diverticular disease or colonic malignancies are generally the main sources of intra-abdominal SS. Although there is evidence to suggest that the incidence of diverticular disease in Africa is rising, it remains an uncommon cause of sepsis in our environment.^[16] The severity of acute appendicitis in SA has been well described in the literature, and appendicitis in SA carries significantly higher morbidity and mortality than in the developed world, primarily as a result of delayed presentation and poor access to healthcare.^[17-19] Patients generally present with more advanced disease than in HICs. Similarly, complicated peptic ulcer disease is rarely seen in HICs, where there is ready access to drug therapy and endoscopy services.

Non-communicable diseases (NCDs) are an increasing contributor to the burden of disease in SA. This was reflected in our study by the fact that over 25% of the surgical workload due to sepsis was for diabetic limb sepsis. A recent review of the burden of diabetes mellitus in KwaZulu-Natal showed that the number of patients initiated on treatment was increasing.^[20] The authors also showed a direct correlation between defaulting and the development of diabetic foot sepsis, and that diabetics are more likely than non-diabetics to require lower limb amputations (LLAs). In SA, diabetes

is responsible for two-thirds of all LLAs. Poverty, poor sanitation, inadequate footwear and poor glycaemic control all contribute to this situation.

The overall mortality rate in our cohort was high. This is in keeping with studies from HICs, which report mortality rates ranging from 18% to 34%. The mortality rate for intra-abdominal sepsis is similar to that for diabetic foot sepsis, once again emphasising the growing problem of NCDs in our environment. The mortality rate for soft-tissue sepsis varies dramatically between different types of soft-tissue infections. Subcutaneous abscesses have a negligible mortality rate, but once the infection involves the fascial planes the mortality rate increases dramatically. Advanced necrotising fasciitis reflects inadequate early recognition and referral by primary healthcare services.

Conclusions

The spectrum of SS presenting to an acute surgical service in SA is different to that seen in HICs. Intra-abdominal sepsis is the most common SS and is overwhelmingly caused by severe acute appendicitis and complicated peptic ulcer disease. Diabetic foot infection is a major cause of SS, reflecting the growing burden of NCDs in SA.

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