Mortality Factors for Severe Septic States in Intensive Care Unit

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Introduction

Sepsis is a life-threatening organ dysfunction caused by an inappropriate host response to an infection. There is no more distinction between sepsis and severe sepsis [1]. Severe sepsis is a major source of morbidity and mortality in intensive care units. The objectives of our study were to analyze mortality factors associated with severe sepsis and to describe the epidemiological, clinical and therapeutic aspects.

Patients and Methods

This is a monocentric, observational and retrospective study from January 1st, 2015 to June 30th, 2017, dealing with mortality factors for severe sepsis in 40 adult patients in the intensive care unit at Aristide Le Dantec Teaching Hospital in Dakar. We included all adult patients who died from severe sepsis whose records were complete. The data was collected from the records and the register of the resuscitation unit.

The studied parameters concerned:

a) Sociodemographic and anamnestic data: age, sex, time of care if diagnosed before admission, reason for hospitalization, number of patients hospitalized during the period;

b) Clinical data: signs of sepsis and septic shock, signs of organ failure, infectious focus;

c) Microbiological data;

d) Therapeutic data: vascular filling modalities, vasopressor amines used, probabilistic and adapted antibiotherapy, supportive therapies (corticosteroids, insulin therapy, mechanical ventilation, extrarenal treatment ...);

e) Evolving data: duration of infection, length of stay, mortality factors and causes of death.

The collected data were captured and analyzed with the Sphinx version 2 software.
The descriptive study was carried out by calculating frequencies, proportions with confidence intervals for qualitative and quantitative variables, and by calculating averages with their standard deviation.

The analytical study was done with the crossed tables. The p value below 0.05 was considered statistically significant.

Results

During the study period, out of 1242 admitted, 47 patients died from complications of severe sepsis, i.e an incidence of 3.78%. Forty files could be exploited. The average age of patients was 55.75 years, with extremes of 21 and 86 years. The sex-ratio was 0.9% (19M / 21F). Diabetes (32.5%) was the most common site in our patients. Postoperative follow-up was the most common reason for hospitalization with 45% (27.5% related to gas gangrene and 17.5% related to peritonitis), followed by septic shock accounted for 25%. The most common infectious focus was cutaneous in 16 patients (40%), followed by peritoneal foci in 9 patients (22.5%) and pulmonary patients in 8 patients (17.5%). Bacteriological findings were positive in 14 patients with a predominance of gram-negative bacillus. Vascular filling was performed in all patients on admission with crystalloids alone in 87.5% of the patients and associated with the colloids for the rest. The most used vasopressor was norepinephrine in the majority of patients (29 cases), i.e 72.5%. All patients received antibiotic therapy, probabilistic in 80% (32 patients) and adapted in only 20% of cases (8 patients). The supportive treatments and various treatments given to our patients are summarized in the table below (Table 1).

**Table 1**: Overall representativeness of each of the therapies established in relation to the overall population.

| Treatment                  | Number (n) | Percentage (%) |
|----------------------------|------------|----------------|
| Dialysis                   | 2          | 5              |
| Corticotherapy             | 6          | 15             |
| Insulin therapy            | 14         | 35             |
| Mechanical ventilation     | 22         | 55             |
| Preventive anticoagulation | 27         | 67.5           |
| Vasopressive amines        | 34         | 85             |
| Antibiotics                | 40         | 100            |

For evolution data, the causes of death are listed in Table 2.

**Table 2**: Distribution of patients according to the causes of death.

| Causes of Death             | Number (n) | Percentage (%) |
|----------------------------|------------|----------------|
| Cardiovascular causes      | 2          | 5              |
| Neurological causes         | 4          | 10             |
| Respiratory causes (ARDS)  | 7          | 17.5           |
| Renal causes                | 8          | 20             |
| Multiple organ Failure      | 19         | 47.5           |

ARDS: acute respiratory distress syndrome

The average length of stay of our patients was 6.13 days with extremes ranging from 1 to 55 days. The analytical study showed that: septic shock was an early mortality factor with a significant p value at 0.024. But there was no correlation between early mortality and other reasons for hospitalization. There was a relationship between mortality and hyperthermia (p=0.038), duration of infection before admission (p=0.007), impairment of renal function (p=0.019), and hyperkalemia with p significant at 0.017.

Discussion

The epidemiology of severe septic syndromes in intensive care is now well known, as are their implications for morbidity and management difficulties. The incidence of severe sepsis is 3.78% in our study. In France, the “Episepsis” survey, conducted in 2001 in a large number of intensive care units, showed that about 15% of patients hospitalized in intensive care, had a severe septic syndrome [2]. It is accepted that the risk of developing a severe infection increases significantly from the age of 60, so that septic shock can be seen at any age, but it is particularly common in the elderly [3]. In our study, the average age was 55.75 years, confirming data from the literature that states a link between mortality and age in septic shock. With regard to gender, the analysis of published data does not find a causal link between sex and mortality [4]. Diabetes was the main field found in our patients. This high frequency could be explained by the delay in patient management and is thought to be responsible for the higher number of admissions for gas gangrene. This delay in management was also reflected in the frequent occurrence of septic shock at admission with a significant relationship with mortality, as well as the duration of infection before admission. The presence of hyperthermia at admission was also correlated with the occurrence of death. Other mortality factors noted in our study were acute renal failure and hyperkalemia. In the SOAP study conducted in Europe on septic patients at resuscitation department, the frequency of acute renal failure was 36%, occurring in 72% of cases during the first 2 days of hospitalization. The overall mortality at day 60 was 35.7%, with an increase depending on the severity of the acute renal failure: 36% in forms without extrarenal treatment, 41% if need of extrarenal treatment and 52% in case of anuria [5]. There is consensus among experts that prompt extrarenal treatment should be instituted in the following life-threatening situations: hyperkalemia, lysis syndrome, acute pulmonary edema, metabolic acidosis [6,7]. In our series, 29 patients had acute renal failure with only 2 patients under extrarenal treatment, and a statistically significant value was found at p = 0.019 with mortality. Therapeutically, only 20% of the patients were able to benefit from an adapted antibiotherapy.
whereas all the patients had had an initial infectious assessment as soon as admitted. This is due to the long delay in receiving results, which poses a real problem in adapting antibiotic therapy to bacteriological results. However, early and adapted antibiotic therapy is imperative and guarantees better progression during severe sepsis [8]. Inadequate initial antibiotic therapy is a source of delay in treatment, lengthening the duration of stay and significant excess mortality [9,10].

Supportive therapies are of great importance in the management of severe sepsis, aiming at the same time at improving the state of health of the patient, but above all at preventing the deleterious complications that may occur during the stay in intensive care; whose lethal potential would only worsen the state of these patients with already poor prognosis. With this in mind, the study on the Sepsis Survival Campaign conducted in 2012 also defined a number of recommendations to be applied in order to put the maximum potential for the patient’s survival, but also to improve their end of life where appropriate [8]. A number of factors may be intimately associated with poor prognosis in septic shock. A detailed analysis of the prognostic factors associated with infections and septic states was carried out in the European “Euro sepsis” survey. The variables remaining significantly associated with death are of three types: chronic associated pathologies, the importance of physiological alterations and organ dysfunctions and the characteristics of the infection [11]. Among organ failures, renal (77%), cardiovascular (44%) and hematological (15%) failures dominate the prognosis. Among the characteristics of infection, those acquired in the hospital (34%) and in intensive care (21%) are associated with a higher risk of death, as well as intra-abdominal infections (15%) and those of indeterminate origin (2%); infections with strict aerobic germs (20%) and yeasts (12%) are also associated with an increased risk of death, unlike Enterobacteriaceae (35%), associated with a comparatively more favorable prognosis. In the same French multi-centric study, mechanical ventilation was the prognostic factor associated with the highest mortality rate [12]. In our study, we had 20% renal failure and 55% ventilated patients. In our series, the prognostic factors of mortality in univariate analysis approached this study by several parameters: demographic (age, direct admission to intensive care, severity of the disease: acute respiratory insufficiency, acute respiratory distress syndrome ARDS, acute renal failure) and therapeutic interventions: mechanical ventilation, use of vasoactive amines.

Conclusion
Severe sepsis is associated with significant morbidity and mortality. In our context, the prognostic factors are essentially the duration of the infection, the state of shock and the severe renal failure. The prognosis can be improved thanks to the early and effective management of identified mortality factors.

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