

# Role of interleukin-3 as a prognostic marker in septic patients

## Avaliação da interleucina 3 como marcador prognóstico na sepse

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### Sample size determination

There is only one study published in the medical literature regarding the use of IL-3 as a prognostic marker in sepsis.<sup>(9)</sup> In that study, analysis of IL-3 levels in 97 patients (retrospective data of a previously published prospective cohort with 60 patients and prospective data of a new cohort with 37 patients) revealed that IL-3 was associated with 28-day follow-up mortality in severe sepsis or septic shock patients. Considering a cut-off value of 89.4pg/mL, 38% of deceased patients had IL-3 levels above that value, compared with 11% of the surviving patients, with a 4.9 odds ratio in the Kaplan-Meier survival curve. Global 28-day mortality was 37.1%. In the retrospective phase of the present study, we observed a

27.8% 28-day mortality rate. Considering the prevalence of positive IL-3 (> 89.4pg/mL) in surviving and deceased patients up to the 28th day of the aforementioned study and the allocation ratio of 0.38 in both patient subgroups in our retrospective cohort (i.e., 1 death for every 2.6 survivors), 95 survivors and 36 deaths would be required for a statistical significance to be found between the two groups. Because the primary outcome analyzed in our study was hospital mortality (as opposed to 28-day mortality), the measurement was adjusted considering a 32.9% hospital mortality rate in the first phase of our cohort. Thus, 40 deaths and 81 survivors during hospital stay (yielding a total of 121) would be required for a statistically significant difference to be found between these two subgroups, with a 95% power and 5%  $\alpha$ -error.



**Figure 1S** - Circulating levels of IL-3 on days 1, 3 and 7 according to in-hospital survival. Confidence interval bars with median levels of interleukin-3 (IL-3) and 75<sup>th</sup> percentiles on days 1 (D1), 3 (D3), and 7 (D7). \* p statistically significant.

**Table 1S** - Hospital mortality adjusted for the covariables age and sequential SOFA - Cox survival analysis

Variable	Hazard ratio	95%CI	p value
IL-3 D1	1.032	1.010 - 1.055	0.005
SOFA D1, D3, D7	1.154	1.068 - 1.248	< 0.001
Age	1.035	1.010 - 1.060	0.5

95%CI - 95% confidence interval; IL-3 - Interleukin-3; D1 - Day 1; SOFA - Sequential Organ Failure Assessment; D3 - Day 3; D7 - Day 7.

**Table 2S** - Model performance measures - hospital mortality prediction

Model	Interclass correlation	Coefficient of determination (R <sup>2</sup> )	AIC
IL-3 D1 + SOFA + AGE	0.777	0.063 (0.486)	234.20
SOFA + AGE	0.743	0.052 (0.483)	236.52

IL-3 - Interleukin-3; D1 - Day 1; SOFA - Sequential Organ Failure Assessment; AIC - Akaike Information Criteria.