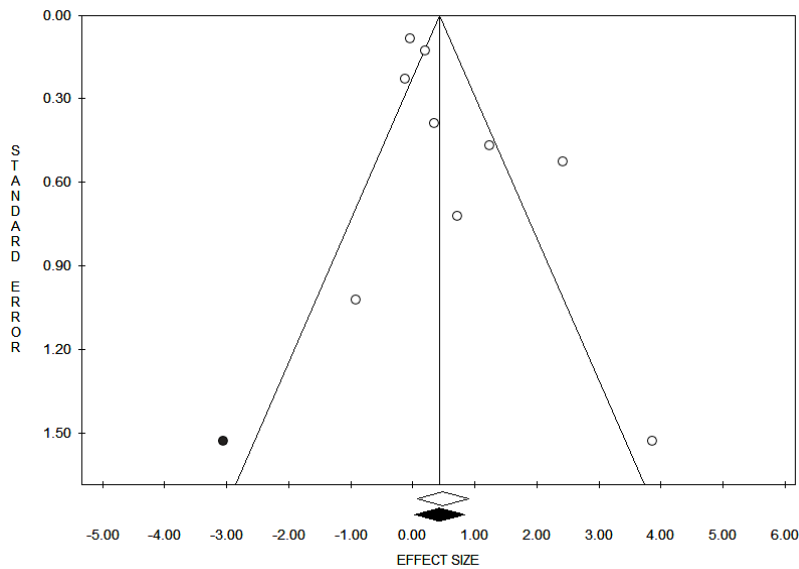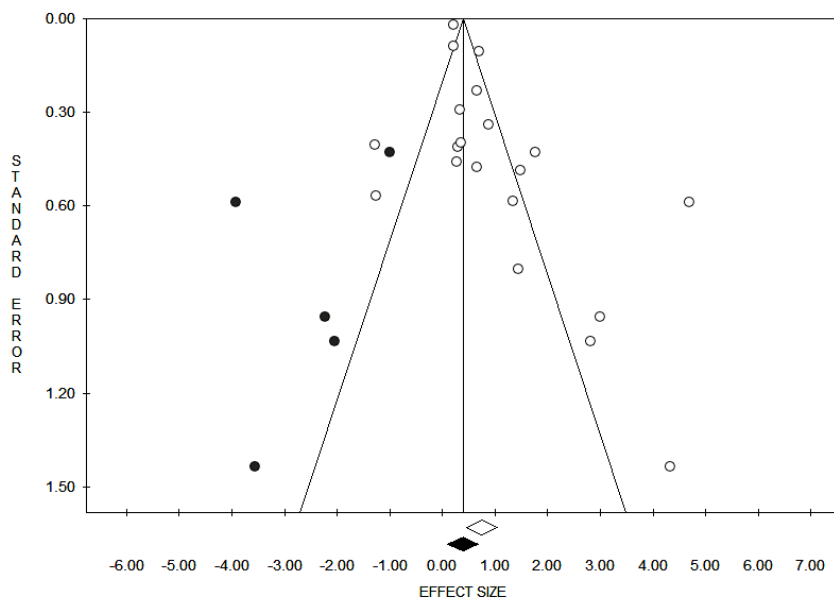**S4 Fig.**

Funnel plot and trim-and-fill analysis of studies evaluating compliance with individual 24-hour bundle targets. Open circles indicate the analyzed studies, full circles indicate the trimmed studies.
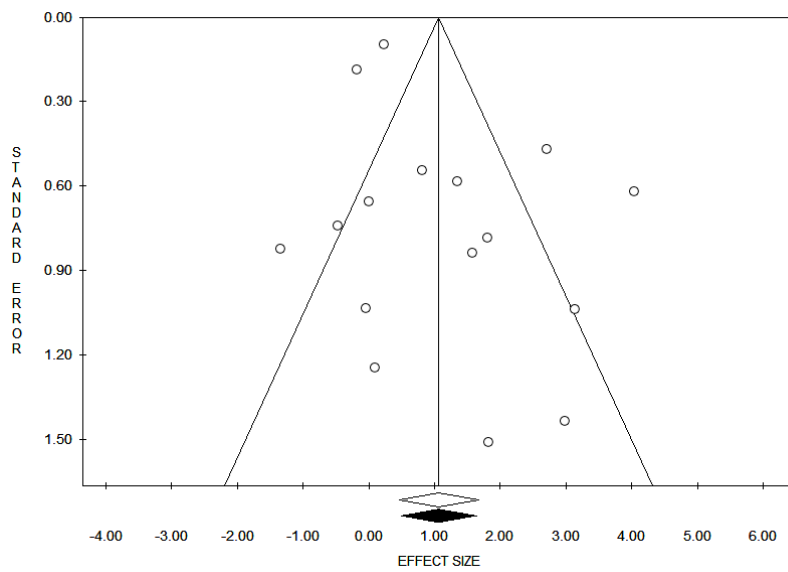
(A) Lung protective ventilation. The trim-and-fill analysis revealed an asymmetry of the funnel plot (estimated ES = 1.54 [1.00-2.37] versus observed ES = 1.63 [1.07-2.47]; number of trimmed studies: 1). The Egger's linear regression test showed a t = 2.19, p = 0.065.

(B) Low-dose steroids. The trim-and-fill analysis revealed an asymmetry of the funnel plot (estimated ES = 1.48 [1.09-2.01] versus observed ES = 1.77 [1.37-2.29]; number of trimmed studies: 5). The Egger's linear regression test confirmed the possible presence of a publication bias (t = 2.66, p = 0.016).

(C) Drotrecogin alfa (activated). Trim-and-fill analysis and Egger's linear regression test (t = 2.00, p = 0.066) did not show any asymmetry of the funnel plot.

(D) Glucose control. Trim-and-fill analysis and Egger's linear regression test (t = 0.51, p = 0.621) did not show any asymmetry of the funnel plot.