**Supplemental Digital Content Legends**

Supplemental Digital Content 1 – Search Strategy Appendix.  
Supplemental Digital Content 2 – PRISMA Flowchart.

Supplemental Digital Content 3 – Table of Excluded Studies.

Supplemental Digital Content 4 – Risk of Bias Assessment. Studies were appraised using the Cochrane Risk of Bias Tool. Satisfactory performance, unclear performance, and unsatisfactory performance of each domain from the tool is denoted by green, clear, and red coloring, respectively.

Supplemental Digital Content 5 – Tables S1 and S2 that list the subgroup analyses performed for mortality for the critical care and surgical patient subgroups, respectively.

Supplemental Digital Content 6 – Trial sequential analysis for 30d mortality across critical care studies. This analysis demonstrates maturity of the evidence base for the critical care studies.

Supplemental Digital Content 7 – Trial sequential analysis for 30d mortality across perioperative studies. This analysis demonstrates immaturity of the evidence base for perioperative studies.

Supplemental Digital Content 8 –Funnel plot demonstrating asymmetry for the perioperative studies.

Supplemental Digital Content 9 –Funnel plot demonstrating symmetry for the critical care studies.

Supplemental Digital Content 10 – Summary of findings table with specific GRADE recommendations and support for authors’ judgement.