Supplementary Online Content

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eFigure. PRISMA Flow Chart for Search Record

eTable. Details of the 15 Studies Included in the Systematic Review

This supplementary material has been provided by the authors to give readers additional information about their work.
Figure. PRISMA flow chart for Search record

Identification

Records identified through database searching (n = 347)
Additional records identified through other sources (n = 0)

Screening

Records after duplicates removed (n = 345)

Records screened (n = 345) → Records excluded (n = 309)

Eligibility

Full-text articles assessed for eligibility (n = 36) → Full-text articles excluded because inclusion/exclusion criteria not met (n = 21)

Included

Studies included in qualitative synthesis (n = 15)

Studies included in quantitative synthesis (meta-analysis) (n = 15)
**eTable 1. Details of the 15 studies included in the systematic review**

| Author (Year)          | Country      | Study type                                      | Acute illness                                                                 | Population size |
|------------------------|--------------|------------------------------------------------|-------------------------------------------------------------------------------|-----------------|
| Joosten (2000) 10      | Netherlands  | Prospective observational, 2 groups            | Meningococcal septic shock survived, nonsurvivors                             | 26              |
| van Woensel (2001) 11  | Netherlands  | Prospective observational, 3 subgroups         | Meningococcal meningitis without and with sepsis, and fulminant sepsis       | 32              |
| Bone (2002) 12         | UK           | Prospective, single group                      | Meningococcal disease                                                         | 65              |
| de Kleijn (2002) 13     | Netherlands  | Prospective, part of a trial, 3 subgroups      | Meningococcal sepsis with no shock, shock and survived, nonsurvivors          | 62              |
| Lichtarowicz-Krynska (2004) 14 | UK       | Prospective, single group                      | Meningococcal disease                                                         | 31              |
| Tasker (2004) 15        | UK           | Prospective controlled observational, two subgroups during acute illness and at recovery | Moderate and severe bronchiolitis                                             | 32              |
| Neville (2005) 16       | Australia    | Prospective observational, single group        | Acute gastroenteritis requiring intravenous fluid rehydration                | 52              |
| Pinto (2006) 17         | Chile         | Prospective controlled observational, two subgroups and at recovery; controls before elective minor surgery | Mild and severe bronchiolitis                                                | 42              |
| Singhi (2006) 18        | India         | Prospective observational, 2 groups            | Aseptic meningitis; bacterial meningitis                                       | 30              |
| Casartelli (2007) 19    | Brazil        | Prospective observational, 2 subgroups         | Septic shock survived and nonsurvivors                                       | 22              |
| Sarthi (2007) 20        | India         | Prospective, single group                      | Septic shock                                                                  | 30              |
| den Brinker (2008) 21   | Netherlands   | Prospective observational, 2 subgroups         | Meningococcal sepsis not intubated, intubated without etomidate              | 37              |
| Study                  | Country | Study Design                      | Conditions                                                                 | Participants |
|-----------------------|---------|-----------------------------------|-----------------------------------------------------------------------------|--------------|
| Diaz (2012) 22        | Chile   | Prospective controlled observational, two subgroups and controls before elective minor surgery | Mild/moderate and severe bronchiolitis                                      | 49           |
| Karaguzel (2012) 23   | Turkey  | Prospective controlled observational, 2 groups; age matched healthy controls | Severe sepsis; acute critical illness without sepsis                         | 50           |
| Alder (2018) 24       | USA     | Prospective controlled observational, 3 subgroups and controls before elective hernia surgery | Sepsis and septic shock; Systemic inflammatory response syndrome             | 129          |