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| National Clinical Guidelines Centre |

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| Sepsis |

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| Search strategies appendix |

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| Clinical guideline |

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| Appendices |

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| 26 November 2015 |

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| Draft |
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| Commissioned by the National Institute for Health and Care Excellence |

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| Guideline short title |
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| |  | | --- | | Disclaimer  Healthcare professionals are expected to take NICE clinical guidelines fully into account when exercising their clinical judgement. However, the guidance does not override the responsibility of healthcare professionals to make decisions appropriate to the circumstances of each patient, in consultation with the patient and, where appropriate, their guardian or carer. | | Copyright | | National Clinical Guideline Centre, 2015. Confidential | | Funding | | National Institute for Health and Care Excellence | |

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1. Literature search strategies

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Search strategies used for the Sepsis guideline are outlined below and were run in accordance with the methodology in the NICE guidelines manual.{NICE2012} All searches were run up to 9 October 2015unless otherwise stated. Any studies added to the databases after this date (even those published prior to this date) were not included unless specifically stated in the text. We do not routinely search for electronic, ahead of print or ‘online early’ publications. Where possible searches were limited to retrieve material published in English.

Table : Database date parameters

| Database | Dates searched |
| --- | --- |
| Medline | 1946 – 9 October 2015 |
| Embase | 1974 – 9 October 2015 |
| The Cochrane Library | Cochrane Reviews to 2015 Issue 10 of 12  CENTRAL to 2015 Issue 9 of 12  DARE to 2013 Issue 2 of 4 |
|  | HTA to 2013 Issue 3 of 4 |
|  | NHSEED to 2013 Issue 2 of 4 |
| PsycINFO (Ovid)  PsycINFO (ProQuest) | 1806 - March Week 1 2015  1806 – 6 October 2015 |
| CINAHL | 1981 – 6 October 2015 |

Searches for the **clinical reviews** were run in Medline (OVID), Embase (OVID) and the Cochrane Library (Wiley). Additional searches were run in CINAHL and PsycINFO for some questions see Table 2.

Table 2: Databases searched

|  | Question | Databases |
| --- | --- | --- |
| A.4.7 | Acid-base pH | Medline/Embase/Cochrane Library |
| A.4.6 | Antimicrobials | Medline/Embase/Cochrane Library |
| A.4.3 | Blood tests | Medline/Embase/Cochrane Library |
| A.4.13 | Educational programmes/Identification protocols | Medline/Embase/Cochrane Library |
| A.4.10 | Escalation of care | Medline/Embase/Cochrane Library |
| A.5.1 | Health economic reviews | Medline/Embase/CRD/HEED |
| A.4.12 | Information support | Medline/Embase/Cochrane Library /CINAHL/PsycINFO |
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| A.4.5 | IV Fluids | Medline/Embase/Cochrane Library |
| A.4.11 | Monitoring | Medline/Embase/Cochrane Library |
| A.4.8 | Oxygen | Medline/Embase/Cochrane Library |
| A.5.2 | Quality of life reviews | Medline/Embase |
| A.4.2 | Scoring systems/Prognostic tools | Medline/Embase/Cochrane Library |
| A.4.1 | Signs and symptoms | Medline/Embase/Cochrane Library |
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Searches for **intervention and diagnostic studies** were usually constructed using a PICO formatwhere population (P) terms were combined with Intervention (I) and sometimes Comparison (C) terms. An intervention can be a drug, a procedure or a diagnostic test. Outcomes (O) are rarely used in search strategies for interventions. Search filters were also added to the search where appropriate.

Searches for **prognostic studies** were usually constructed combining population terms with prognostic variable terms and sometimes outcomes. Search filters were added to the search where appropriate.

Searches for the **health economic reviews** were run in Medline, Embase, the NHS Economic Evaluations Database (NHS EED), the Health Technology Assessment (HTA) database and the Health Economic Evaluation Database (HEED). The Health Economic Evaluation Database (HEED) ceased production in 2014 with access ceasing in January 2015. For the final dates of HEED searches, please see individual economic questions.

For Medline and Embase an economic filter (instead of a study type filter) was added to the same clinical search strategy. Searches in CRD and HEED were constructed using population terms only.

* 1. Population search strategies
     1. Standard sepsis population

The standard sepsis population was used in all questions except question 11 [A.4.11]

**Medline search terms**

|  |  |
| --- | --- |
|  | exp sepsis/ |
|  | sepsis.ti,ab. |
|  | blood-borne pathogens/ |
|  | (blood adj2 (pathogen\* or poison\*)).ti,ab. |
|  | exp systemic inflammatory response syndrome/ |
|  | 'systemic inflammatory response syndrome'.ti,ab. |
|  | sirs.ti,ab. |
|  | (septicaemi\* or septicemi\*).ti,ab. |
|  | (septic adj2 shock).ti,ab. |
|  | (pyaemi\* or pyemi\* or pyohemi\*).ti,ab. |
|  | (bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*).ti,ab. |
|  | or/1-11 |

**Embase search terms**

|  |  |
| --- | --- |
|  | exp \*sepsis/ |
|  | sepsis.ti,ab. |
|  | \*bloodborne bacterium/ |
|  | (blood adj2 (pathogen\* or poison\*)).ti,ab. |
|  | exp \*systemic inflammatory response syndrome/ |
|  | sirs.ti,ab. |
|  | 'systemic inflammatory response syndrome'.ti,ab. |
|  | \*septicemia/ |
|  | (septicaemi\* or septicemi\*).ti,ab. |
|  | \*septic shock/ |
|  | (septic adj2 shock).ti,ab. |
|  | (pyaemi\* or pyemi\* or pyohemi\*).ti,ab. |
|  | (bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*).ti,ab. |
|  | or/1-13 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | MeSH descriptor: [sepsis] explode all trees |
|  | MeSH descriptor: [blood-borne pathogens] explode all trees |
|  | MeSH descriptor: [systemic inflammatory response syndrome] explode all trees |
|  | ((systemic inflammatory response syndrome) or (sirs) or (sepsis)):ti,ab |
|  | (septicaemi\* or septicemi\* or pyaemi\* or pyemi\* or pyohemi\* or bacteremi\* or fungemi\* or parasitemi\* or viremi\* or bacteraemi\* or fungaemi\* or parasitaemi\* or viraemi\*):ti,ab |
|  | (septic near/2 shock):ti,ab |
|  | (blood near/2 (pathogen\* or poison\*)):ti,ab |
|  | {or #1-#7} |

**CINAHL search terms**

|  |  |
| --- | --- |
|  | (MH "sepsis+") |
|  | (MH "systemic inflammatory response syndrome+") |
|  | (MH "bloodborne pathogens") |
|  | (bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*) |
|  | (pyaemi\* or pyemi\* or pyohemi\*) |
|  | (septic n2 shock) |
|  | (septicaemi\* or septicemi\*) |
|  | sirs or sepsis |
|  | (blood n2 (pathogen\* or poison\*)) |
|  | S1 orS2 orS3 orS4 orS5 orS6 orS7 orS8 or S9 |

**PsycINFO (Ovid) search terms**

|  |  |
| --- | --- |
|  | sepsis.ti,ab. |
|  | (blood adj2 (pathogen\* or poison\*)).ti,ab. |
|  | systemic inflammatory response syndrome'.ti,ab. |
|  | sirs.ti,ab. |
|  | (septicaemi\* or septicemi\*).ti,ab. |
|  | (septic adj2 shock).ti,ab. |
|  | (pyaemi\* or pyemi\* or pyohemi\*).ti,ab. |
|  | (bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*).ti,ab. |
|  | or/1-8 |

**PsycINFO (ProQUEST) search terms**

|  |  |
| --- | --- |
|  | (ti,ab(sepsis) or ti,ab(blood near/2 (pathogen\* or poison\*)) or ti,ab(systemic inflammatory response syndrome) or ti,ab(sirs) or ti,ab(septicaemi\* or septicemi\*) or ti,ab(septic near/2 shock) or ti,ab(bacter\*emi\* or fung\*emi\* or parasite\*emi\* or vir\*emi\*) |

**CRD search terms**

|  |  |
| --- | --- |
|  | (MeSH descriptor sepsis explode all trees) |
|  | (MeSH descriptor blood-borne pathogens explode all trees) |
|  | (MeSH descriptor systemic inflammatory response syndrome explode all trees) |
|  | ((systemic inflammatory response syndrome) or (sirs) or (sepsis)) |
|  | ((septicaemi\* or septicemi\* or pyaemi\* or pyemi\* or pyohemi\* or bacteremi\* or fungemi\* or parasitemi\* or viremi\* or bacteraemi\* or fungaemi\* or parasitaemi\* or viraemi\*)) |
|  | ((septic adj2 shock)) |
|  | ((blood adj2 (pathogen\* or poison\*))) |
|  | (#1 or #2 or #3 or #4 or #5 or #6 or #7) |

**HEED search terms**

|  |  |
| --- | --- |
|  | ax=sepsis or septicaemi\* or septicemi\* or systemic inflammatory response syndrome or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacteremi\* or fungemi\* or parasitemi\* or viremi\* or bacteraemi\* or fungaemi\* or parasitaemi\* or viraemi\* |
|  | ax=septic shock |
|  | ax=blood pathogen or blood pathogens or blood borne pathogen or blood borne pathogens |
|  | ax=blood poisoning |
|  | cs=1 or 2 or 3 or 4 |

* + 1. Standard bacterial meningitis population

The standard bacterial meningitis population was used in questions A.4.4, A.4.10, A.4.13 and A.5.1 only.

**Medline search terms**

|  |  |
| --- | --- |
|  | exp bacterial meningitis/ |
|  | ((bacterial\* or infect\*) adj3 (meningitis or meningitides)).ti,ab. |
|  | ((bacterial\* or infect\*) adj3 meninges).ti,ab. |
|  | (infect\* adj3 (leptomeninges or subarachnoid space?)).ti,ab. |
|  | ((meningitis or meningitides) adj3 (e coli or escherichia coli)).ti,ab. |
|  | ((meningitis or meningitides) adj3 (haemophilus or hemophilus)).ti,ab. |
|  | ((meningitis or meningitides) adj3 listeria).ti,ab. |
|  | ((meningitis or meningitides or meningeal or pachymeningitis) adj3 (tuberculosis or tuberculous or tubercular)).ti,ab. |
|  | meningoencephalitis.ti,ab. |
|  | meningoencephalitis/ |
|  | meningitis/ |
|  | exp pneumococcal infections/ |
|  | pneumococc\*.ti,ab. |
|  | exp neisseria meningitidis/ |
|  | (neisseria adj1 meningitidis).ti,ab. |
|  | exp meningococcal infections/ |
|  | (meningococcosis or meningococc\* or meningococcemia).ti,ab. |
|  | exp streptococcal infections/ |
|  | streptococc\*.ti,ab. |
|  | or/1-19 |

**Embase search terms**

|  |  |
| --- | --- |
|  | exp bacterial meningitis/ |
|  | ((bacterial\* or infect\*) adj3 (meningitis or meningitides)).ti,ab. |
|  | ((bacterial\* or infect\*) adj3 meninges).ti,ab. |
|  | (infect\* adj3 (leptomeninges or subarachnoid space?)).ti,ab. |
|  | ((meningitis or meningitides) adj3 (e coli or escherichia coli)).ti,ab. |
|  | ((meningitis or meningitides) adj3 (haemophilus or hemophilus)).ti,ab. |
|  | ((meningitis or meningitides) adj3 listeria).ti,ab. |
|  | ((meningitis or meningitides or meningeal or pachymeningitis) adj3 (tuberculosis or tuberculous or tubercular)).ti,ab. |
|  | meningoencephalitis.ti,ab. |
|  | meningoencephalitis/ |
|  | meningitis/ |
|  | exp pneumococcal infections/ |
|  | pneumococc\*.ti,ab. |
|  | exp neisseria meningitidis/ |
|  | (neisseria adj1 meningitidis).ti,ab. |
|  | exp meningococcal infections/ |
|  | (meningococcosis or meningococc\* or meningococcemia).ti,ab. |
|  | exp streptococcal infections/ |
|  | streptococc\*.ti,ab. |
|  | or/1-19 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | MeSH descriptor: [meningitis, bacterial] explode all trees |
|  | ((bacterial\* or infect\*) near/3 (meningitis or meningitides)):ti,ab |
|  | ((bacterial\* or infect\*) near/3 meninges):ti,ab |
|  | (infect\* near/3 (leptomeninges or subarachnoid space?)):ti,ab |
|  | ((meningitis or meningitides) near/3 (e coli or escherichia coli)):ti,ab |
|  | ((meningitis or meningitides) near/3 (haemophilus or hemophilus)):ti,ab |
|  | ((meningitis or meningitides) near/3 listeria):ti,ab |
|  | ((meningitis or meningitides or meningeal or Pachymeningitis) near/3 (tuberculosis or tuberculous or tubercular)):ti,ab |
|  | meningoencephalitis:ti,ab |
|  | MeSH descriptor: [meningoencephalitis] this term only |
|  | MeSH descriptor: [meningitis] this term only |
|  | MeSH descriptor: [pneumococcal infections] explode all trees |
|  | pneumococc\*:ti,ab |
|  | MeSH descriptor: [neisseria meningitidis] explode all trees |
|  | (neisseria near/1 meningitidis):ti,ab |
|  | MeSH descriptor: [meningococcal infections] explode all trees |
|  | (meningococcosis or meningococc\* or meningococcemia):ti,ab |
|  | MeSH descriptor: [streptococcal infections] explode all trees |
|  | streptococc\*:ti,ab |
|  | {or #1-#19} |

**CRD search terms**

|  |  |
| --- | --- |
|  | ( MeSH descriptor meningitis, bacterial explode all trees) |
|  | ((mening\*)) |
|  | (MeSH descriptor meningoencephalitis explode all trees) |
|  | (MeSH descriptor meningitis explode all trees) |
|  | (MeSH descriptor meningitis explode all trees) |
|  | ((pneumococc\*)) |
|  | (MeSH descriptor neisseria meningitidis explode all trees) |
|  | (MeSH descriptor meningococcal infections explode all trees) |
|  | (MeSH descriptor streptococcal infections explode all trees) |
|  | ((streptococc\*)) |
|  | (#1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10) |

**HEED search terms**

|  |  |
| --- | --- |
|  | ax=mening\* or pneumococc\* or streptococc\* |

* 1. Study filter search terms
     1. Excluded study designs and publication types

The following study designs and publication types were removed from retrieved results using the NOT operator.

**Medline search terms**

|  |  |
| --- | --- |
|  | letter/ |
|  | editorial/ |
|  | news/ |
|  | exp historical article/ |
|  | anecdotes as topic/ |
|  | comment/ |
|  | case report/ |
|  | (letter or comment\*).ti. |
|  | or/1-8 |
|  | randomized controlled trial/ or random\*.ti,ab. |
|  | 9 not 10 |
|  | animals/ not humans/ |
|  | exp animals, laboratory/ |
|  | exp animal experimentation/ |
|  | exp models, animal/ |
|  | exp rodentia/ |
|  | (rat or rats or mouse or mice).ti. |
|  | or/11-17 |

**Embase search terms**

|  |  |
| --- | --- |
|  | letter.pt. or letter/ |
|  | note.pt. |
|  | editorial.pt. |
|  | case report/ or case study/ |
|  | (letter or comment\*).ti. |
|  | or/1-5 |
|  | randomized controlled trial/ or random\*.ti,ab. |
|  | 6 not 7 |
|  | animal/ not human/ |
|  | nonhuman/ |
|  | exp animal experiment/ |
|  | exp experimental animal/ |
|  | animal model/ |
|  | exp rodent/ |
|  | (rat or rats or mouse or mice).ti. |
|  | or/8-15 |

**CINAHL search terms**

|  |  |
| --- | --- |
|  | pt anecdote or pt audiovisual or pt bibliography or pt biography or pt book or pt book review or pt brief item or pt cartoon or pt commentary or pt computer program or pt editorial or pt games or pt glossary or pt historical material or pt interview or pt letter or pt listservs or pt masters thesis or pt obituary or pt pamphlet or pt pamphlet chapter or pt pictorial or pt poetry or pt proceedings or pt “questions and answers” or pt response or pt software or pt teaching materials or pt website |

* + 1. Randomised controlled trials (RCT) search terms

**Medline search terms**

|  |  |
| --- | --- |
|  | randomized controlled trial.pt. |
|  | controlled clinical trial.pt. |
|  | randomi#ed.ab. |
|  | placebo.ab. |
|  | drug therapy.fs. |
|  | randomly.ab. |
|  | trial.ab. |
|  | groups.ab. |
|  | or/1-8 |

**Embase search terms**

|  |  |
| --- | --- |
|  | random\*.ti,ab. |
|  | factorial\*.ti,ab. |
|  | (crossover\* or cross over\*).ti,ab. |
|  | ((doubl\* or singl\*) adj blind\*).ti,ab. |
|  | (assign\* or allocat\* or volunteer\* or placebo\*).ti,ab. |
|  | crossover procedure/ |
|  | double blind procedure/ |
|  | single blind procedure/ |
|  | randomized controlled trial/ |
| 10. | or/1-9 |

* + 1. Systematic review (SR) search terms

**Medline search terms**

|  |  |
| --- | --- |
|  | meta-analysis/ |
|  | meta-analysis as topic/ |
|  | (meta analy\* or metanaly\* or metaanaly\*).ti,ab. |
|  | ((systematic\* or evidence\*) adj3 (review\* or overview\*)).ti,ab. |
|  | (reference list\* or bibliograph\* or hand search\* or manual search\* or relevant journals).ab. |
|  | (search strategy or search criteria or systematic search or study selection or data extraction).ab. |
|  | (search\* adj4 literature).ab. |
|  | (medline or pubmed or cochrane or embase or psychlit or psyclit or psychinfo or psycinfo or cinahl or science citation index or bids or cancerlit).ab. |
|  | cochrane.jw. |
|  | ((multiple treatment\* or indirect or mixed) adj2 comparison\*).ti,ab. |
|  | or/1-10 |

**Embase search terms**

|  |  |
| --- | --- |
|  | systematic review/ |
|  | meta-analysis/ |
|  | (meta analy\* or metanaly\* or metaanaly\*).ti,ab. |
|  | ((systematic or evidence) adj3 (review\* or overview\*)).ti,ab. |
|  | (reference list\* or bibliograph\* or hand search\* or manual search\* or relevant journals).ab. |
|  | (search strategy or search criteria or systematic search or study selection or data extraction).ab. |
|  | (search\* adj4 literature).ab. |
|  | (medline or pubmed or cochrane or embase or psychlit or psyclit or psychinfo or psycinfo or cinahl or science citation index or bids or cancerlit).ab. |
|  | cochrane.jw. |
|  | ((multiple treatment\* or indirect or mixed) adj2 comparison\*).ti,ab. |
|  | or/1-10 |

* + 1. Health economics (HE) search terms

**Medline search terms**

|  |  |
| --- | --- |
|  | economics/ |
|  | value of life/ |
|  | exp "costs and cost analysis"/ |
|  | exp economics, hospital/ |
|  | exp economics, medical/ |
|  | economics, nursing/ |
|  | economics, pharmaceutical/ |
|  | exp "fees and charges"/ |
|  | exp budgets/ |
|  | budget\*.ti,ab. |
|  | cost\*.ti. |
|  | (economic\* or pharmaco?economic\*).ti. |
|  | (price\* or pricing\*).ti,ab. |
|  | (cost\* adj2 (effective\* or utilit\* or benefit\* or minimi\* or unit\* or estimat\* or variable\*)).ab. |
|  | (financ\* or fee or fees).ti,ab. |
|  | (value adj2 (money or monetary)).ti,ab. |
|  | or/1-16 |

**Embase search terms**

|  |  |
| --- | --- |
|  | health economics/ |
|  | exp economic evaluation/ |
|  | exp health care cost/ |
|  | exp fee/ |
|  | budget/ |
|  | funding/ |
|  | budget\*.ti,ab. |
|  | cost\*.ti. |
|  | (economic\* or pharmaco?economic\*).ti. |
|  | (price\* or pricing\*).ti,ab. |
|  | (cost\* adj2 (effective\* or utilit\* or benefit\* or minimi\* or unit\* or estimat\* or variable\*)).ab. |
|  | (financ\* or fee or fees).ti,ab. |
|  | (value adj2 (money or monetary)).ti,ab. |
|  | or/1-13 |

* + 1. Quality of life (QOL) search terms

**Medline search terms**

|  |  |
| --- | --- |
|  | quality-adjusted life years/ |
|  | sickness impact profile/ |
|  | (quality adj2 (wellbeing or well-being)).ti,ab. |
|  | sickness impact profile.ti,ab. |
|  | disability adjusted life.ti,ab. |
|  | (qal\* or qtime\* or qwb\* or daly\*).ti,ab. |
|  | (euroqol\* or eq5d\* or eq 5d\*).ti,ab. |
|  | (qol\* or hql\* or hqol\* or h qol\* or hrqol\* or hr qol\*).ti,ab. |
|  | (health utility\* or utility score\* or disutilit\*).ti,ab. |
|  | (hui or hui1 or hui2 or hui3).ti,ab. |
|  | health\* year\* equivalent\*.ti,ab. |
|  | (hye or hyes).ti,ab. |
|  | rosser.ti,ab. |
|  | (willingness to pay or time tradeoff or time trade off or tto or standard gamble\*).ti,ab. |
|  | (sf36 or sf 36 or short form 36 or shortform 36 or shortform36).ti,ab. |
|  | (sf20 or sf 20 or short form 20 or shortform 20 or shortform20).ti,ab. |
|  | (sf12 or sf 12 or short form 12 or shortform 12 or shortform12).ti,ab. |
|  | (sf8 or sf 8 or short form 8 or shortform 8 or shortform8).ti,ab. |
|  | (sf6 or sf 6 or short form 6 or shortform 6 or shortform6).ti,ab. |
|  | or/1-19 |

**Embase search terms**

|  |  |
| --- | --- |
|  | quality adjusted life year/ |
|  | "quality of life index"/ |
|  | short form 12/ or short form 20/ or short form 36/ or short form 8/ |
|  | sickness impact profile/ |
|  | (quality adj2 (wellbeing or well-being)).ti,ab. |
|  | sickness impact profile.ti,ab. |
|  | disability adjusted life.ti,ab. |
|  | (qal\* or qtime\* or qwb\* or daly\*).ti,ab. |
|  | (euroqol\* or eq5d\* or eq 5d\*).ti,ab. |
|  | (qol\* or hql\* or hqol\* or h qol\* or hrqol\* or hr qol\*).ti,ab. |
|  | (health utility\* or utility score\* or disutilit\*).ti,ab. |
|  | (hui or hui1 or hui2 or hui3).ti,ab. |
|  | health\* year\* equivalent\*.ti,ab. |
|  | (hye or hyes).ti,ab. |
|  | rosser.ti,ab. |
|  | (willingness to pay or time tradeoff or time trade off or tto or standard gamble\*).ti,ab. |
|  | (sf36 or sf 36 or short form 36 or shortform 36 or shortform36).ti,ab. |
|  | (sf20 or sf 20 or short form 20 or shortform 20 or shortform20).ti,ab. |
|  | (sf12 or sf 12 or short form 12 or shortform 12 or shortform12).ti,ab. |
|  | (sf8 or sf 8 or short form 8 or shortform 8 or shortform8).ti,ab. |
|  | (sf6 or sf 6 or short form 6 or shortform 6 or shortform6).ti,ab. |
|  | or/1-21 |

* + 1. Diagnostic test accuracy (DIAG) search terms

**Medline search terms**

|  |  |
| --- | --- |
|  | exp "sensitivity and specificity"/ |
|  | (sensitivity or specificity).ti,ab. |
|  | ((pre test or pretest or post test) adj probability).ti,ab. |
|  | (predictive value\* or ppv or npv).ti,ab. |
|  | likelihood ratio\*.ti,ab. |
|  | likelihood function/ |
|  | (roc curve\* or auc).ti,ab. |
|  | (diagnos\* adj3 (performance\* or accurac\* or utilit\* or value\* or efficien\* or effectiveness)).ti,ab. |
|  | gold standard.ab. |
|  | or/1-9 |

**Embase search terms**

|  |  |
| --- | --- |
|  | exp "sensitivity and specificity"/ |
|  | (sensitivity or specificity).ti,ab. |
|  | ((pre test or pretest or post test) adj probability).ti,ab. |
|  | (predictive value\* or ppv or npv).ti,ab. |
|  | likelihood ratio\*.ti,ab. |
|  | (roc curve\* or auc).ti,ab. |
|  | (diagnos\* adj3 (performance\* or accurac\* or utilit\* or value\* or efficien\* or effectiveness)).ti,ab. |
|  | diagnostic accuracy/ |
|  | diagnostic test accuracy study/ |
|  | gold standard.ab. |
|  | or/1-10 |

* + 1. **Observational studies (OBS) search terms**

**Medline search terms**

|  |  |
| --- | --- |
|  | epidemiologic studies/ |
|  | exp case control studies/ |
|  | exp cohort studies/ |
|  | cross-sectional studies/ |
|  | case control.ti,ab. |
|  | (cohort adj (study or studies or analys\*)).ti,ab. |
|  | ((follow up or observational or uncontrolled or non randomi#ed or nonrandomi#ed or epidemiologic\*) adj (study or studies)).ti,ab. |
|  | ((longitudinal or retrospective or prospective or cross sectional) and (study or studies or review or analys\* or cohort\*)).ti,ab. |
|  | or/1-8 |

**Embase search terms**

|  |  |
| --- | --- |
|  | clinical study/ |
|  | exp case control study/ |
|  | family study/ |
|  | longitudinal study/ |
|  | retrospective study/ |
|  | prospective study/ |
|  | cross-sectional study/ |
|  | cohort analysis/ |
|  | follow-up/ |
|  | cohort\*.ti,ab. |
|  | 9 and 10 |
|  | case control.ti,ab. |
|  | (cohort adj (study or studies or analys\*)).ti,ab. |
|  | ((follow up or observational or uncontrolled or non randomi#ed or nonrandomi#ed or epidemiologic\*) adj (study or studies)).ti,ab. |
|  | ((longitudinal or retrospective or prospective or cross sectional) and (study or studies or review or analys\* or cohort\*)).ti,ab. |
|  | or/1-8,11-15 |

* + 1. Prognostic/prediction rule (PROG) search terms

**Medline search terms**

|  |  |
| --- | --- |
|  | predict.ti. |
|  | (validat\* or rule\*).ti,ab. |
|  | (predict\* and (outcome\* or risk\* or model\*)).ti,ab. |
|  | ((history or variable\* or criteria or scor\* or characteristic\* or finding\* or factor\*) and (predict\* or model\* or decision\* or identif\* or prognos\*)).ti,ab. |
|  | decision\*.ti,ab. and Logistic models/ |
|  | (decision\* and (model\* or clinical\*)).ti,ab. |
|  | (prognostic and (history or variable\* or criteria or scor\* or characteristic\* or finding\* or factor\* or model\*)).ti,ab. |
|  | (stratification or discrimination or discriminate or c statistic or "area under the curve" or AUC or calibration or indices or algorithm or multivariable).ti,ab. |
|  | ROC curve/ |
|  | or/1-9 |

**Embase search terms**

|  |  |
| --- | --- |
|  | predict\*.ti. |
|  | (validat\* or rule\*).ti,ab. |
|  | (predict\* and (outcome\* or risk\* or model\*)).ti,ab. |
|  | ((history or variable\* or criteria or scor\* or characteristic\* or finding\* or factor\*) and (predict\* or model\* or decision\* or identif\* or prognos\*)).ti,ab. |
|  | decision\*.ti,ab. and Statistical model/ |
|  | (decision\* and (model\* or clinical\*)).ti,ab. |
|  | (prognostic and (history or variable\* or criteria or scor\* or characteristic\* or finding\* or factor\* or model\*)).ti,ab. |
|  | (stratification or discrimination or discriminate or c statistic or "area under the curve" or AUC or calibration or indices or algorithm or multivariable).ti,ab. |
|  | receiver operating characteristic/ |
|  | or/1-9 |

* 1. Searches for specific questions
     1. Signs and symptoms
* What is the predictive value/utility of different signs and symptoms, alone or in combination, for the diagnosis of sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*heart rate/ |
|  | \*respiratory rate/ |
|  | ((heart or pulse or respirat\* or breath\*) adj2 rate).ti,ab. |
|  | \*blood pressure/ |
|  | ((systolic or blood) adj2 pressure).ti,ab. |
|  | exp \*consciousness disorders/ |
|  | (consciousness or unconsciousness or semiconsciousness).ti,ab. |
|  | \*delirium/ |
|  | ((alter\* or chang\*) adj2 mental state\*).ti,ab. |
|  | (oxygen saturation or blood oxygen or sats).ti,ab. |
|  | \*anoxia/ |
|  | (anoxia or anoxemia or anoxaemia or hypoxia or hypoxaemia or hypoxemia or "oxygen deficien\*").ti,ab. |
|  | \*body temperature/ |
|  | exp \*body temperature changes/ |
|  | (fever\* or hypothermi\* or hyperthermi\* or pyrex\* or febri\* or hyper-therm\* or hyper-pyrex\* or hypo-therm\* or body temperature?).ti,ab. |
|  | or/5-19 |
|  | \*accidental falls/ |
|  | (fall or falls or falling).ti,ab. |
|  | \*dysuria/ |
|  | \*oliguria/ |
|  | (oliguria or hypouresis or dysuria).ti,ab. |
|  | \*pallor/ |
|  | \*exanthema/ |
|  | (pallor or cyanosis or exanthem\* or rash\*).ti,ab. |
|  | \*cyanosis/ |
|  | ((pale or mottled or pallid or wan or ashen or blue) adj2 (skin or lips or tongue or lip)).ti,ab. |
|  | ((ill\* or sick\* or unwell) adj3 (look\* or appear\*)).ti,ab. |
|  | ((reduc\* or low\*) adj2 urin\* adj2 (volume? or output? or level?)).ti,ab. |
|  | \*chills/ |
|  | \*shivering/ |
|  | (chill? or rigor? or shiver\*).ti,ab. |
|  | exp \*pain/ |
|  | pain?.ti,ab. |
|  | exp \*vaginal discharge/ |
|  | (vagina\* adj2 (discharge\* or secret\*)).ti,ab. |
|  | (capill?ary refill time? or crt).ti,ab. |
|  | \*capillary resistance/ |
|  | \*microcirculation/ |
|  | ((cold or chill\*) adj3 (hand? or feet or foot)).ti,ab. |
|  | \*respiratory sounds/ |
|  | (rales or crackles or rhonchi or grunt\*).ti,ab. |
|  | ((chest or sternal or sternum or intercostal) adj3 (in drawing or in?drawing or recess\* or retract\*)).ti,ab. |
|  | ((respirat\* or breath\*) adj3 (distress\* or disorder? or alter\*)).ti,ab. |
|  | ((nose or nasal or nostril? or alar) adj3 flar\*).ti,ab. |
|  | \*Cranial fontanelles/ |
|  | (fontanel\* adj3 (bulg\* or tens\*)).ti,ab. |
|  | (pulse adj2 pressure).ti,ab. |
|  | \*arterial pressure/ |
|  | (mean arterial adj2 pressure).ti,ab. |
|  | (confus\* adj2 mental state\*).ti,ab. |
|  | \*confusion/ |
|  | exp \*diarrhea/ |
|  | diarrh\*.ti,ab. |
|  | (water\* adj3 (bowel movement\* or stool\* or feces or faeces or fecal or foecal)).ti,ab. |
|  | \*vomiting/ |
|  | (vomit\* or emesis or emeses).ti,ab. |
|  | \*crying/ |
|  | ((weak\* or continu\* or high-pitch\* or high pitch\*) adj2 (cry\* or voice\* or sound\* or articulat\*)).ti,ab. |
|  | \*postpartum period/ |
|  | \*pregnant women/ |
|  | (pregnant or pregnancy).ti,ab. |
|  | or/21-65 |
|  | exp "signs and symptoms"/ |
|  | symptom assessment/ |
|  | diagnosis/ or prognosis/ |
|  | (clinical adj3 (manifestation? or feature? or finding? or aspect? or marker?)).ti,ab. |
|  | (presenting adj3 (feature? or finding? or factor?)).ti,ab. |
|  | presentation?.ti,ab. |
|  | (physical adj3 (manifestaion? or characteristic? or feature? or finding?)).ti,ab. |
|  | (sign or signs or symptom\* or recogni\* or identif\* or complain\*).ti,ab. |
|  | (diagnos\* or prognos\* or assess\* or criteria\* or predict\*).ti,ab. |
|  | or/67-75 |
|  | (20 or 66) and 76 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 77 and 78 |
|  | limit 80 to yr="1990 -Current" |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*heart rate/ |
|  | \*breathing rate/ |
|  | ((heart or pulse or respirat\* or breath\*) adj2 rate).ti,ab. |
|  | \*systolic blood pressure/ |
|  | ((systolic or blood) adj2 pressure).ti,ab. |
|  | \*consciousness level/ |
|  | exp \*consciousness disorder/ |
|  | (consciousness or unconsciousness or semiconsciousness).ti,ab. |
|  | ((alter\* or chang\*) adj2 mental state\*).ti,ab. |
|  | \*delirium/ |
|  | (oxygen saturation or blood oxygen or sats).ti,ab. |
|  | \*oxygen saturation/ |
|  | \*anoxia/ |
|  | (anoxia or anoxemia or anoxaemia or hypoxia or hypoxaemia or hypoxemia or "oxygen deficien\*").ti,ab. |
|  | \*body temperature/ |
|  | exp \*body temperature disorder/ |
|  | (fever\* or hypothermi\* or hyperthermi\* or pyrex\* or febri\* or hyper-therm\* or hyper-pyrex\* or hypo-therm\* or body temperature?).ti,ab. |
|  | or/5-21 |
|  | \*falling/ |
|  | (fall or falls or falling).ti,ab. |
|  | \*dysuria/ |
|  | \*oliguria/ |
|  | (oliguria or hypouresis or dysuria).ti,ab. |
|  | \*pallor/ |
|  | \*rash/ |
|  | (pallor or cyanosis or exanthem\* or rash\*).ti,ab. |
|  | \*cyanosis/ |
|  | ((pale or mottled or pallid or wan or ashen or blue) adj2 (skin or lips or tongue or lip)).ti,ab. |
|  | ((ill\* or sick\* or unwell) adj3 (look\* or appear\*)).ti,ab. |
|  | ((reduc\* or low\*) adj2 urin\* adj2 (volume? or output? or level?)).ti,ab. |
|  | \*chill/ |
|  | \*shivering/ |
|  | (chill? or rigor? or shiver\*).ti,ab. |
|  | exp \*pain/ |
|  | pain?.ti,ab. |
|  | \*vagina discharge/ |
|  | (vagina\* adj2 (discharge\* or secret\*)).ti,ab. |
|  | (capill?ary refill time? or crt).ti,ab. |
|  | \*capillary resistance/ |
|  | \*microcirculation/ |
|  | ((cold or chill\*) adj3 (hand? or feet or foot)).ti,ab. |
|  | \*abnormal respiratory sound/ |
|  | (rales or crackles or rhonchi or grunt\*).ti,ab. |
|  | ((chest or sternal or sternum or intercostal) adj3 (in drawing or in?drawing or recess\* or retract\*)).ti,ab. |
|  | ((respirat\* or breath\*) adj3 (distress\* or disorder? or alter\*)).ti,ab. |
|  | ((nose or nasal or nostril? or alar) adj3 flar\*).ti,ab. |
|  | \*fontanel/ |
|  | (fontanel\* adj3 (bulg\* or tens\*)).ti,ab. |
|  | \*pathological crying/ or \*crying/ |
|  | ((weak\* or continu\* or high-pitch\* or high pitch\*) adj2 (cry\* or voice\* or sound\* or articulat\*)).ti,ab. |
|  | (pulse adj2 pressure).ti,ab. |
|  | \*arterial pressure/ |
|  | (mean arterial adj2 pressure).ti,ab. |
|  | (confus\* adj2 mental state\*).ti,ab. |
|  | \*confusion/ or \*acute confusion/ or \*"confusion (uncertainty)"/ |
|  | exp \*diarrhea/ |
|  | diarrh\*.ti,ab. |
|  | (water\* adj3 (bowel movement\* or stool\* or feces or faeces or fecal or foecal)).ti,ab. |
|  | \*vomiting/ |
|  | (vomit\* or emesis or emeses).ti,ab. |
|  | \*puerperium/ |
|  | \*pregnant woman/ |
|  | (pregnant or pregnancy).ti,ab. |
|  | or/23-67 |
|  | symptom assessment/ |
|  | diagnosis/ |
|  | prognosis/ |
|  | (clinical adj3 (manifestation? or feature? or finding? or aspect? or marker?)).ti,ab. |
|  | (presenting adj3 (feature? or finding? or factor?)).ti,ab. |
|  | presentation?.ti,ab. |
|  | (physical adj3 (manifestaion? or characteristic? or feature? or finding?)).ti,ab. |
|  | (sign or signs or symptom\* or recogni\* or identif\* or complain\*).ti,ab. |
|  | (diagnos\* or prognos\* or assess\* or criteria\* or predict\*).ti,ab. |
|  | exp symptomatology/ |
|  | or/69-78 |
|  | (22 or 68) and 79 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 80 and 81 |
|  | limit 82 to yr="1990 -Current" |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | [mh "heart rate"] |
|  | [mh "respiratory rate"] |
|  | [mh "body temperature changes"] |
|  | [mh "body temperature"] |
|  | [mh anoxia] |
|  | [mh delirium] |
|  | [mh "blood pressure"] |
|  | [mh "consciousness disorders"] |
|  | ((heart or pulse or respirat\* or breath\*) near/2 rate):ti,ab |
|  | ((systolic or blood) near/2 pressure):ti,ab |
|  | (consciousness or unconsciousness or semiconsciousness):ti,ab |
|  | ((alter\* or chang\*) near/2 mental state\*):ti,ab |
|  | (oxygen saturation or blood oxygen or sats):ti,ab |
|  | (anoxia or anoxemia or anoxaemia or hypoxia or hypoxaemia or hypoxemia or "oxygen deficien\*"):ti,ab |
|  | (fever\* or hypothermi\* or hyperthermi\* or pyrex\* or febri\* or hyper-therm\* or hyper-pyrex\* or hypo-therm\* or body temperature\*):ti,ab |
|  | {or #2-#16} |
|  | MeSH descriptor: [accidental falls] explode all trees |
|  | (fall or falls or falling):ti,ab |
|  | MeSH descriptor: [dysuria] explode all trees |
|  | MeSH descriptor: [oliguria] explode all trees |
|  | (oliguria or hypouresis or dysuria):ti,ab |
|  | MeSH descriptor: [pallor] explode all trees |
|  | MeSH descriptor: [exanthema] explode all trees |
|  | (pallor or cyanosis or exanthem\* or rash\*):ti,ab |
|  | MeSH descriptor: [cyanosis] explode all trees |
|  | ((pale or mottled or pallid or wan or ashen or blue) near/2 (skin or lips or tongue or lip)):ti,ab |
|  | ((ill\* or sick\* or unwell) adj3 (look\* or appear\*)):ti,ab |
|  | ((reduc\* or low\*) near/2 urin\* near/2 (volume? or output? or level?)):ti,ab |
|  | MeSH descriptor: [chills] explode all trees |
|  | MeSH descriptor: [shivering] explode all trees |
|  | (chill? or rigor? or shiver\*):ti,ab |
|  | MeSH descriptor: [pain] explode all trees |
|  | pain?.ti,ab |
|  | MeSH descriptor: [vaginal discharge] explode all trees |
|  | (vagina\* near/2 (discharge\* or secret\*)):ti,ab |
|  | (capill?ary refill time? or crt):ti,ab |
|  | MeSH descriptor: [capillary resistance] explode all trees |
|  | MeSH descriptor: [microcirculation] explode all trees |
|  | ((cold or chill\*) near/3 (hand? or feet or foot)):ti,ab |
|  | MeSH descriptor: [respiratory sounds] explode all trees |
|  | (rales or crackles or rhonchi or grunt\*):ti,ab |
|  | ((chest or sternal or sternum or intercostal) adj3 (in drawing or in?drawing or recess\* or retract\*)):ti,ab |
|  | ((respirat\* or breath\*) near/3 (distress\* or disorder? or alter\*)):ti,ab |
|  | ((nose or nasal or nostril? or alar) near/3 flar\*):ti,ab |
|  | MeSH descriptor: [cranial fontanelles] explode all trees |
|  | (fontanel\* near/3 (bulg\* or tens\*)):ti,ab |
|  | MeSH descriptor: [crying] explode all trees |
|  | ((weak\* or continu\* or high-pitch\* or high pitch\*) near/2 (cry\* or voice\* or sound\* or articulat\*)):ti,ab |
|  | (pulse adj2 pressure):ti,ab |
|  | MeSH descriptor: [arterial pressure] explode all trees |
|  | (mean arterial adj2 pressure):ti,ab |
|  | (confus\* adj2 mental state\*):ti,ab |
|  | MeSH descriptor: [confusion] explode all trees |
|  | MeSH descriptor: [diarrhea] explode all trees |
|  | diarrh\*:ti,ab |
|  | (water\* near/3 (bowel movement\* or stool\* or feces or faeces or fecal or foecal)):ti,ab |
|  | MeSH descriptor: [vomiting] explode all trees |
|  | (vomit\* or emesis or emeses):ti,ab |
|  | MeSH descriptor: [postpartum period] explode all trees |
|  | MeSH descriptor: [pregnant women] explode all trees |
|  | (pregnant or pregnancy):ti,ab |
|  | {or #18-#62} |
|  | [mh "signs and symptoms"] |
|  | [mh "symptom assessment"] |
|  | [mh ^diagnosis] |
|  | [mh ^prognosis] |
|  | (clinical near/3 (manifestation\* or feature\* or finding\* or aspect\* or marker\*)):ti,ab |
|  | (presenting near/3 (feature\* or finding\* or factor\*)):ti,ab |
|  | presentation\*:ti,ab |
|  | (physical near/3 (manifestaion\* or characteristic\* or feature\* or finding\*)):ti,ab |
|  | (sign or signs or symptom\* or recogni\* or identif\* or complain\*):ti,ab |
|  | (diagnos\* or prognos\* or assess\* or criteria\* or predict\*):ti,ab |
|  | {or #64-#73} |
|  | #1 and (#17 or #63) and #74 Publication Year from 1990 to 2015 |

* + 1. Scoring systems/Prognostic tools
* What is the most accurate and cost effective assessment tool to identify patients with sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*"predictive value of tests"/ |
|  | \*"severity of illness index"/ |
|  | \*decision support techniques/ |
|  | \*triage/mt [methods] |
|  | scor\* system\*.ti,ab. |
|  | (risk adj2 calculation\*).ti,ab. |
|  | ((risk or warn\* or triage\* or observation\* or observ\*) adj2 (system\* or scor\*)).ti,ab. |
|  | (severity adj2 index).ti,ab. |
|  | (curb65 or crb65).ti,ab. |
|  | exp \*health status indicators/ |
|  | "track and trigger".ti,ab. |
|  | ((trigger or calling or alert) adj5 criteria).ti,ab. |
|  | or/5-16 |
|  | ((paediatric or pediatric or child\*) adj4 (observation\* or observ\* or scor\*)).ti,ab. |
|  | (tool kit or toolkit).ti,ab. |
|  | "sequential organ failure assessment".ti,ab. |
|  | "acute physiology and chronic health evaluation".ti,ab. |
|  | "mortality probability model\*".ti,ab. |
|  | "predisposition infection response and organ dysfunction".ti,ab. |
|  | "mortality in emergency department sepsis".ti,ab. |
|  | "rapid emergency medicine score".ti,ab. |
|  | "emergency severity index".ti,ab. |
|  | "simplified acute physiology score".ti,ab. |
|  | "pediatric logistic organ dysfunction".ti,ab. |
|  | "pediatric risk of mortality score".ti,ab. |
|  | "glasgow meningococcal sepsis prognostic score".ti,ab. |
|  | "pediatric index of mortality ".ti,ab. |
|  | (pews or mews or mts or pops or sofa or apache or mpm or piro or meds or rems or esi or saps or saps ii or pelod or prism or gmsps or pim).ti,ab. |
|  | or/18-32 |
|  | intensive care units/ |
|  | 33 not 34 |
|  | 4 and (17 or 35) |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*"predictive value"/ |
|  | \*"severity of illness index"/ |
|  | \*decision support system/ |
|  | scor\* system\*.ti,ab. |
|  | (risk adj2 calculation\*).ti,ab. |
|  | ((risk or warn\* or triage\* or observation\* or observ\*) adj2 (system\* or scor\*)).ti,ab. |
|  | (severity adj2 index).ti,ab. |
|  | (curb65 or crb65).ti,ab. |
|  | exp \*health status indicators/ |
|  | "track and trigger".ti,ab. |
|  | ((trigger or calling or alert) adj5 criteria).ti,ab. |
|  | or/5-15 |
|  | ((paediatric or pediatric or child\*) adj4 (observation\* or observ\* or scor\*)).ti,ab. |
|  | (tool kit or toolkit).ti,ab. |
|  | "sequential organ failure assessment".ti,ab. |
|  | "acute physiology and chronic health evaluation".ti,ab. |
|  | "mortality probability model\*".ti,ab. |
|  | "predisposition infection response and organ dysfunction".ti,ab. |
|  | "mortality in emergency department sepsis".ti,ab. |
|  | "rapid emergency medicine score".ti,ab. |
|  | "emergency severity index".ti,ab. |
|  | "simplified acute physiology score".ti,ab. |
|  | "pediatric logistic organ dysfunction".ti,ab. |
|  | "pediatric risk of mortality score".ti,ab. |
|  | "glasgow meningococcal sepsis prognostic score".ti,ab. |
|  | "pediatric index of mortality ".ti,ab. |
|  | (pews or mews or mts or pops or sofa or apache or mpm or piro or meds or rems or esi or saps or saps ii or pelod or prism or gmsps or pim).ti,ab. |
|  | apache/ |
|  | \*"named inventories, questionnaires and rating scales"/ |
|  | sequential organ failure assessment score/ or simplified acute physiology score/ |
|  | \*scoring system/ |
|  | or/17-35 |
|  | intensive care unit/ |
|  | 36 not 37 |
|  | 4 and (16 or 38) |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | MeSH descriptor: [predictive value of tests] this term only |
|  | MeSH descriptor: [severity of illness index] this term only |
|  | MeSH descriptor: [decision support techniques] this term only |
|  | MeSH descriptor: [triage] explode all trees and with qualifier(s): [Methods - MT] |
|  | scor\* system\*:ti,ab |
|  | (risk near/2 calculation\*):ti,ab |
|  | ((risk or warn\* or triage\* or observation\* or observ\*) near/2 (system\* or scor\*)):ti,ab |
|  | (severity near/2 index):ti,ab |
|  | (curb65 or crb65):ti,ab |
|  | MeSH descriptor: [health status indicators] this term only |
|  | track and trigger:ti,ab |
|  | ((trigger or calling or alert) near/5 criteria):ti,ab |
|  | {or #2-#13} |
|  | ((paediatric or pediatric or child\*) near/4 (observation\* or observ\* or scor\*)):ti,ab |
|  | (tool kit or toolkit):ti,ab |
|  | sequential organ failure assessment:ti,ab |
|  | acute physiology and chronic health evaluation:ti,ab |
|  | mortality probability model\*:ti,ab |
|  | predisposition infection response and organ dysfunction:ti,ab |
|  | mortality in emergency department sepsis:ti,ab |
|  | rapid emergency medicine score:ti,ab |
|  | emergency severity index:ti,ab |
|  | simplified acute physiology score:ti,ab |
|  | pediatric logistic organ dysfunction:ti,ab |
|  | pediatric risk of mortality score:ti,ab |
|  | (pews or mews or mts or pops or sofa or apache or mpm or piro or meds or rems or esi or saps or saps ii or pelod or prism or gmsps or pim):ti,ab |
|  | {or #15-#27} |
|  | MeSH descriptor: [intensive Care] explode all trees |
|  | #28 not #29 |
|  | #1 and (#14 or #30) |

* + 1. Blood tests
* What is the predictive value/usefulness of blood tests for the recognition and early assessment of sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | biological markers/ |
|  | (blood adj6 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | blood gas analysis/ |
|  | (abg or vbg or cbg).ti,ab. |
|  | blood glucose/an, bl, du [analysis, blood, diagnostic use] |
|  | lactic acid/an, bl, du [analysis, blood, diagnostic use] |
|  | ((lactate or lactic) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | exp blood cell count/ |
|  | ((blood or leukocyte\* or leucocyte\* or erythrocyte\* or thrombocyte\* or platelet or wbc\* or rbc\*) adj2 (differential or count\*)).ti,ab. |
|  | (fbc or cbc or fbe).ti,ab. |
|  | (polymorph\* or polymorphonucleocyte\* or neutrophil\*).ti,ab. |
|  | ((polymorph\* or polymorphonucleocyte\* or neutrophil\*) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | leukocytes/an, bl, di, du [analysis, blood, diagnosis, diagnostic use] |
|  | neutrophils/an, bl, bs, di [analysis, blood, blood supply, diagnosis] |
|  | blood platelets/an [analysis] |
|  | urea/an, bl, du [analysis, blood, diagnostic use] |
|  | (urea adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | electrolytes/bl, du [blood, diagnostic use] |
|  | (electrolyte\* adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | u&e.ti,ab. |
|  | (blood urea nitrogen adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | bun.ti,ab. |
|  | ((kidney or renal) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | creatinine/bl, du [blood, diagnostic use] |
|  | (creatinine adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | ((liver or hepatic) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | limax.ti,ab. |
|  | ((coagul\* or anticoagul\* or act) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | (partial thromboplastin time or ptt or aptt or pt or aptr).ti,ab. |
|  | ((prothrombin or bleed\* or clot\* or thrombin or blood) adj2 time\*).ti,ab. |
|  | fibrinogen/bl, di, du [blood, diagnosis, diagnostic use] |
|  | (fibrinogen\* adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | c-reactive protein/bl, du [blood, diagnostic use] |
|  | (c-reactive protein\* adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | or/5-38 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or DIAG [A.3.6] |
|  | 4 and 39 and 40 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*biological marker/ |
|  | (blood adj6 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | \*blood gas analysis/ |
|  | (abg or vbg or cbg).ti,ab. |
|  | \*glucose blood level/ |
|  | \*lactic acid/ |
|  | ((lactate or lactic) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | exp \*blood cell count/ |
|  | ((blood or leukocyte\* or leucocyte\* or erythrocyte\* or thrombocyte\* or platelet or wbc\* or rbc\*) adj2 (differential or count\*)).ti,ab. |
|  | (fbc or cbc or fbe).ti,ab. |
|  | (polymorph\* or polymorphonucleocyte\* or neutrophil\*).ti,ab. |
|  | ((polymorph\* or polymorphonucleocyte\* or neutrophil\*) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | \*leukocyte/ |
|  | \*neutrophil/ |
|  | \*thrombocyte/an [drug analysis] |
|  | \*urea/ |
|  | (urea adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | \*electrolyte/ |
|  | (electrolyte\* adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | u&e.ti,ab. |
|  | (blood urea nitrogen adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | bun.ti,ab. |
|  | ((kidney or renal) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | \*creatinine/ |
|  | (creatinine adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | ((liver or hepatic) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | limax.ti,ab. |
|  | ((coagul\* or anticoagul\* or act) adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | (partial thromboplastin time or ptt or aptt or pt or aptr).ti,ab. |
|  | ((prothrombin or bleed\* or clot\* or thrombin or blood) adj2 time\*).ti,ab. |
|  | \*fibrinogen/ |
|  | (fibrinogen\* adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | \*c reactive protein/ |
|  | (c-reactive protein\* adj3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)).ti,ab. |
|  | or/5-38 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or DIAG [A.3.6] |
|  | 4 and 39 and 40 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | MeSH descriptor: [biological markers] explode all trees |
|  | (blood near/6 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | MeSH descriptor: [blood gas analysis] explode all trees |
|  | (abg or vbg or cbg):ti,ab |
|  | MeSH descriptor: [blood glucose] explode all trees |
|  | MeSH descriptor: [lactic acid] explode all trees and with qualifier(s): [analysis - an, blood - bl, diagnostic use - du] |
|  | ((lactate or lactic) near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | MeSH descriptor: [blood cell count] explode all trees |
|  | ((blood or leukocyte\* or leucocyte\* or erythrocyte\* or thrombocyte\* or platelet or wbc\* or rbc\*) near/2 (differential or count\*)):ti,ab |
|  | (fbc or cbc or fbe):ti,ab |
|  | (polymorph\* or polymorphonucleocyte\* or neutrophil\*):ti,ab |
|  | ((polymorph\* or polymorphonucleocyte\* or neutrophil\*) near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | MeSH descriptor: [leukocytes] explode all trees |
|  | MeSH descriptor: [neutrophils] explode all trees |
|  | MeSH descriptor: [blood platelets] explode all trees |
|  | MeSH descriptor: [urea] explode all trees and with qualifier(s): [analysis - an, blood - bl, diagnostic use - du] |
|  | (urea near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)) ;ti,ab |
|  | MeSH descriptor: [electrolytes] explode all trees and with qualifier(s): [blood - bl, diagnostic use - du] |
|  | (electrolyte\* near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | u&e:ti,ab |
|  | (blood urea nitrogen near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | bun:ti,ab |
|  | ((kidney or renal) near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | MeSH descriptor: [creatinine] explode all trees and with qualifier(s): [blood - bl, diagnostic use - du] |
|  | (creatinine near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | ((liver or hepatic) near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | limax:ti,ab |
|  | ((coagul\* or anticoagul\* or act) near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | (partial thromboplastin time or ptt or aptt or pt or aptr):ti,ab |
|  | ((prothrombin or bleed\* or clot\* or thrombin or blood) near/2 time\*):ti,ab |
|  | MeSH descriptor: [fibrinogen] explode all trees |
|  | (fibrinogen\* near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | MeSH descriptor: [c-reactive protein] explode all trees |
|  | (c-reactive protein\* near/3 (analys\* or analyze\* or test\* or investigat\* or evaluat\* or examin\* or check\* or assess\* or measur\* or diagnos\* or identif\* or verif\* or assay)):ti,ab |
|  | {or #2-#35} |
|  | #1 and #36 |

* + 1. Supplementary blood tests

Searches for the following three questions were were run as one search:

* What is the predictive value of lactate in people with sepsis for the recognition and early assessment of worsening sepsis?
* What is the predictive value of serum creatinine in people with sepsis for the recognition and early assessment of worsening sepsis?
* What is the predictive value of disseminated intravascular coagulation in people with sepsis for the recognition and early assessment of worsening sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | exp arthritis, infectious/ or exp bone diseases, infectious/ or exp community-acquired infections/ or exp respiratory tract infections/ or exp skin diseases, infectious/ or exp soft tissue infections/ or exp urinary tract infections/ or "meningitis".mp. or "serious infections".mp. or exp gastroenteritis/ [mp=title, abstract, original title, name of substance word, subject heading word, keyword heading word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier] |
|  | or/1-3 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 4 not 5 |
|  | Limit 6 to English language |
|  | exp lactic acid/ |
|  | (lactate\* or lactic acid).ti,ab. |
|  | 8 or 9 |
|  | exp disseminated intravascular coagulation/ |
|  | (disseminated adj2 intravascular adj2 (coagulat\* or coagulopath\* or clot\*)).ti,ab. |
|  | ((consumption or consumptive) adj1 coagulopath\*).ti,ab. |
|  | dic.ti,ab. |
|  | or/11-14 |
|  | creatinine/ |
|  | (creatinine or cystatin c or acr or kreatinine).ti,ab. |
|  | 16 or 17 |
|  | acute kidney injury/ |
|  | kidney tubular necrosis, acute/ |
|  | (aki or acute kidney necrosis or acute kidney tubul\* necrosis or acute kidney injury).ti,ab. |
|  | ((acute or early) adj (kidney or renal) adj (failure\* or injur\* or insufficien\* or dysfunction\* or impair\*)).ti,ab. |
|  | or/19-22 |
|  | 15 or (18 and 23) |
|  | Study filters PROG [A.3.8] |
|  | 7 and 24 and 25 |
|  | 7 and 10 |
|  | 26 or 27 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | exp infectious arthritis/ or exp hematogenous osteomyelitis/ or exp communicable disease/ or exp respiratory tract infection/ or exp skin infection/ or exp soft tissue infection/ or exp urinary tract infection/ or exp gastroenteritis/ or (serious and infections).mp. [mp=title, abstract, heading word, drug trade name, original title, device manufacturer, drug manufacturer, device trade name, keyword] |
|  | or/1-3 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 4 not 5 |
|  | Limit 6 to English language |
|  | exp lactic acid/ |
|  | (lactate\* or lactic acid).ti,ab. |
|  | 8 or 9 |
|  | exp disseminated intravascular clotting/ |
|  | (disseminated adj2 intravascular adj2 (coagulat\* or coagulopath\* or clot\*)).ti,ab. |
|  | ((consumption or consumptive) adj1 coagulopath\*).ti,ab. |
|  | DIC.ti,ab. |
|  | or/11-14 |
|  | creatinine/ |
|  | (creatinine or cystatin c or acr or kreatinine).ti,ab. |
|  | 16 or 17 |
|  | acute kidney failure/ or acute kidney tubule necrosis/ |
|  | (aki or acute kidney necrosis or acute kidney tubul\* necrosis or acute kidney injury).ti,ab. |
|  | ((acute or early) adj (kidney or renal) adj (failure\* or injur\* or insufficien\* or dysfunction\* or impair\*)).ti,ab. |
|  | or/19-21 |
|  | 15 or (18 and 22) |
|  | Study filters PROG [A.3.8] |
|  | 7 and 23 and 24 |
|  | 7 and 10 |
|  | 25 or 26 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | MeSH descriptor: [arthritis, infectious] explode all trees |
|  | MeSH descriptor: [bone diseases, infectious] explode all trees |
|  | MeSH descriptor: [community-acquired infections] explode all trees |
|  | MeSH descriptor: [respiratory tract infections] explode all trees |
|  | MeSH descriptor: [skin diseases, infectious] explode all trees |
|  | MeSH descriptor: [soft tissue infections] explode all trees |
|  | MeSH descriptor: [urinary tract infections] explode all trees |
|  | MeSH descriptor: [gastroenteritis] explode all trees |
|  | serious infections |
|  | or/1-11 |
|  | MeSH descriptor: [lactic acid] explode all trees |
|  | (lactate\* or lactic acid):ti,ab |
|  | #13 or #14 |
|  | MeSH descriptor: [disseminated intravascular coagulation] explode all trees |
|  | (disseminated near/2 intravascular near/2 (coagulat\* or coagulopath\* or clot\*)):ti,ab |
|  | ((consumption or consumptive) next coagulopath\*):ti,ab |
|  | dic:ti,ab |
|  | {or #16-#19} |
|  | MeSH descriptor: [creatinine] this term only |
|  | (creatinine or cystatin c or acr or kreatinine):ti,ab |
|  | #21 or #22 |
|  | MeSH descriptor: [acute kidney injury] this term only |
|  | MeSH descriptor: [kidney tubular necrosis, acute] this term only |
|  | (aki or acute kidney necrosis or acute kidney tubul\* necrosis or acute kidney injury):ti,ab |
|  | ((acute or early) adj (kidney or renal) adj (failure\* or injur\* or insufficien\* or dysfunction\* or impair\*)):ti,ab |
|  | {or #24-#27} |
|  | #23 and #28 |
|  | #15 or #20 or #29 |
|  | #12 or #30 |

* + 1. IV Fluids

Searches for the following three questions were run as one search:

* What is the most clinical and cost effective immediate/bolus IV fluid for resuscitation of patients with sepsis?
* What is the clinical and cost effectiveness of different volumes/dosages of immediate/bolus IV fluid resuscitation in patients with sepsis?
* What is the most clinically and cost effective rate of administration of immediate/bolus IV fluids in patients with sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | fluid therapy/ |
|  | exp water-electrolyte balance/ |
|  | exp water-electrolyte imbalance/ |
|  | ((fluid\* or electrolyte\*) adj3 (document\* or chart\* or strateg\* or regimen\* or load\*)).ti,ab. |
|  | ((fluid\* or electrolye\* or water) adj3 (requir\* or need\* or prescri\* or intravenous or iv or infusion\* or drip or drips or maint\* or volume\* or therap\* or administrat\* or manag\* or balance\* or imbalance\* or overload\* or loss\* or status or monit\* or assess\* or reassess\* or evaluat\* or re-evaluat\* or reevaluat\*)).ti,ab. |
|  | ((fluid\* or volum\*) adj3 (restor\* or resus\* or replac\* or deplet\* or deficien\* or replenish\* or therap\* or substitut\* or rehydrat\*)).ti,ab. |
|  | (fluid\* adj3 (challenge or bolus)).ti,ab. |
|  | (volume adj2 (expand\* or expansion\* or substitut\*)).ti,ab. |
|  | ((perioperativ\* or intraoperativ\* or postoperativ\*) adj3 fluid\*).ti,ab. |
|  | (euvol?emi\* or normovol?emi\*).ti,ab. |
|  | insensible loss\*.ti,ab. |
|  | hyponatr?emi\*.ti,ab. |
|  | exp hemoglobins/ |
|  | h?emoglobin\*.ti,ab. |
|  | ((red blood cell\* or rbc or prbc or red cell\* or blood or packed cell\* or erythrocyte\*or fluid\* or volum\* or plasma\*) adj5 (therap\* or transfus\* or replac\* or resuscita\* or substitut\* or restor\* or deficien\* or replenish\*)).ti,ab. |
|  | exp plasma/ |
|  | (ffp or ((frozen or thawed or tp or fresh) adj3 plasma)).ti,ab. |
|  | ((lyophili?ed or freeze-dried or liquid or "not frozen" or "never frozen") adj3 plasma).ti,ab. |
|  | (fdsp or fdp or lqp or lhp).ti,ab. |
|  | exp freeze drying/ and plasma.ti,ab,sh. |
|  | ((platelet\* or thrombocyte\*) adj3 (transfus\* or prophyla\* or therap\* or infus\* or administ\*)).ti,ab. |
|  | blood platelets/ |
|  | or/5-26 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 27 and 28 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*fluid therapy/ |
|  | exp \*electrolyte balance/ |
|  | exp \*electrolyte disturbance/ |
|  | fluid resuscitation/ |
|  | ((fluid\* or electrolyte\*) adj3 (document\* or chart\* or strateg\* or regimen\* or load\*)).ti,ab. |
|  | ((fluid\* or electrolye\* or water) adj3 (requir\* or need\* or prescri\* or intravenous or iv or infusion\* or drip or drips or maint\* or volume\* or therap\* or administrat\* or manag\* or balance\* or imbalance\* or overload\* or loss\* or status or monit\* or assess\* or reassess\* or evaluat\* or re-evaluat\* or reevaluat\*)).ti,ab. |
|  | ((fluid\* or volum\*) adj3 (restor\* or resus\* or replac\* or deplet\* or deficien\* or replenish\* or therap\* or substitut\* or rehydrat\*)).ti,ab. |
|  | (fluid\* adj3 (challenge or bolus)).ti,ab. |
|  | (volume adj2 (expand\* or expansion\* or substitut\*)).ti,ab. |
|  | ((perioperativ\* or intraoperativ\* or postoperativ\*) adj3 fluid\*).ti,ab. |
|  | (euvol?emi\* or normovol?emi\*).ti,ab. |
|  | insensible loss\*.ti,ab. |
|  | hyponatr?emi\*.ti,ab. |
|  | exp \*hemoglobin/ |
|  | h?emoglobin\*.ti,ab. |
|  | ((red blood cell\* or rbc or prbc or red cell\* or blood or packed cell\* or erythrocyte\* or fluid\* or volum\*) adj3 (therap\* or transfus\* replac\* or resuscita\* or substitut\* or restor\* or deficien\* or replenish\*)).ti,ab. |
|  | exp \*plasma/ |
|  | ((ffp or frozen or fresh or thawed or tp) adj3 plasma).ti,ab. |
|  | ((lyophili?ed or freeze-dried or liquid or "not frozen" or "never frozen") adj3 plasma).ti,ab. |
|  | (fdsp or fdp or lqp or lhp).ti,ab. |
|  | exp \*freeze drying/ and plasma.ti,ab,sh. |
|  | ((platelet\* or thrombocyte\*) adj3 (transfus\* or prophyla\* or therap\* or infus\* or administ\*)).ti,ab. |
|  | \*thrombocyte/ |
|  | or/5-28 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 28 and 29 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | MeSH descriptor: [fluid therapy] explode all trees |
|  | MeSH descriptor: [water-electrolyte balance] explode all trees |
|  | MeSH descriptor: [water-electrolyte imbalance] explode all trees |
|  | ((fluid\* or electrolyte\*) near/3 (document\* or chart\* or strateg\* or regimen\* or load\*)):ti,ab |
|  | ((fluid\* or electrolye\* or water) near/3 (requir\* or need\* or prescri\* or intravenous or iv or infusion\* or drip or drips or maint\* or volume\* or therap\* or administrat\* or manag\* or balance\* or imbalance\* or overload\* or loss\* or status or monit\* or assess\* or reassess\* or evaluat\* or re-evaluat\* or reevaluat\*)):ti,ab |
|  | ((fluid\* or volum\*) near/3 (restor\* or resus\* or replac\* or deplet\* or deficien\* or replenish\* or therap\* or substitut\* or rehydrat\*)):ti,ab |
|  | (fluid\* near/3 (challenge or bolus)):ti,ab |
|  | (volume near/2 (expand\* or expansion\* or substitut\*)):ti,ab |
|  | ((perioperativ\* or intraoperativ\* or postoperativ\*) near/3 fluid\*):ti,ab |
|  | (euvol\*emi\* or normovol\*emi\*):ti,ab |
|  | insensible loss\*:ti,ab |
|  | hyponatr\*emi\*:ti,ab |
|  | mesh descriptor: [hemoglobins] explode all trees |
|  | h\*emoglobin\*:ti,ab |
|  | ((red blood cell\* or rbc or prbc or red cell\* or blood or packed cell\* or erythrocyte\*or fluid\* or volum\* or plasma\*) near/5 (therap\* or transfus\* or replac\* or resuscita\* or substitut\* or restor\* or deficien\* or replenish\*)):ti,ab |
|  | mesh descriptor: [plasma] explode all trees |
|  | ((ffp or frozen or thawed or tp or fresh) near/3 plasma):ti,ab |
|  | ((lyophili?ed or freeze-dried or liquid or "not frozen" or "never frozen") near/3 plasma):ti,ab |
|  | (fdsp or fdp or lqp or lhp):ti,ab |
|  | MeSH descriptor: [freeze drying] explode all trees |
|  | ((platelet\* or thrombocyte\*) near/3 (transfus\* or prophyla\* or therap\* or infus\* or administ\*)):ti,ab |
|  | MeSH descriptor: [blood platelets] explode all trees |
|  | {or #2-#23} |
|  | #1 and #24 |

* + 1. Antimicrobials

Searches for the following two questions were run as one search:

* What are the most clinically and cost effective timings of IV or IM empiric antimicrobial treatments in patients with (a) septic shock, (b) severe sepsis without shock or (c) sepsis?
* What is the most clinically and cost effective IV or IM empiric antimicrobial treatment in patients with sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*anti-infective agents/ |
|  | exp anti-bacterial agents/ |
|  | ((antiinfect\* or anti infect\* or anti-infect\*) adj3 (agent\* or therap\* or treatment\*)).ti,ab. |
|  | ((anti-mycobacterial\* or antimycobacterial\* or anti-mycobacterial\*) adj3 (agent\* or therap\* or treatment\*)).ti,ab. |
|  | (microbicid\* or bacteriocid\* or bactericid\*).ti,ab. |
|  | ((antibacterial\* or anti bacterial\* or anti-bacterial\* or bacteriocidal or anti-microbial\* or antimicrobial\*) adj3 (agent\* or therap\* or treatment\*)).ti,ab. |
|  | (antibiotic\* or anti-biotic\* or anti biotic\*).ti,ab. |
|  | exp \*beta-lactams/ |
|  | (co-amoxiclav or amoxicillin-clavulanic or piperacillin or tazobactam or tazocin or piptazobatam or ampicillin or sulbactam).ti,ab. |
|  | (benzylpenicillin or flucloxacillin or amoxicillin).ti,ab. |
|  | (cephalosporin\* or cefuroxime or cefotaxime or ceftriaxone or ceftazidime or ceftaroline or ceftobiprole or cefipime).ti,ab. |
|  | aztreonam.ti,ab. |
|  | (carbapenem\* or imipenem or meropenem or ertapenem).ti,ab. |
|  | aminoglycosides/ or gentamicins/ |
|  | amikacin/ |
|  | tobramycin/ |
|  | (gentamicin or amikacin or tobramycin).ti,ab. |
|  | ciprofloxacin/ or levofloxacin/ |
|  | (ciprofloxacin or levofloxacin).ti,ab. |
|  | teicoplanin/ or vancomycin/ |
|  | (vancomycin or teicoplanin).ti,ab. |
|  | daptomycin/ |
|  | daptomycin.ti,ab. |
|  | clindamycin/ |
|  | tigecycline.ti,ab. |
|  | rifampin/ |
|  | rifampicin.ti,ab. |
|  | chloramphenicol/ |
|  | chloramphenicol.ti,ab. |
|  | co-trimoxazole.ti,ab. |
|  | colistin/ |
|  | colistin.ti,ab. |
|  | metronidazole/ |
|  | metronidazole.ti,ab. |
|  | exp erythromycin/ |
|  | (clar#thromycin or clindamycin or linezolid).ti,ab. |
|  | (azithromycin or erythromycin).ti,ab. |
|  | doxycycline/ |
|  | doxycycline.ti,ab. |
|  | or/5-43 |
|  | time factors/ |
|  | ((early or earlier or time or timing or late or later or delay\*) adj4 (initiat\* or start\* or treat\* or therap\* or administ\* or prescri\* or antibiotic\*)).ti,ab. |
|  | 45 or 46 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 44 and 47 and 48 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*antiinfective agent/ |
|  | ((antiinfect\* or anti infect\* or anti-infect\*) adj3 (agent\* or therap\* or treatment\*)).ti,ab. |
|  | ((anti-mycobacterial\* or antimycobacterial\* or anti-mycobacterial\*) adj3 (agent\* or therap\* or treatment\*)).ti,ab. |
|  | (microbicid\* or bacteriocid\* or bactericid\*).ti,ab. |
|  | ((antibacterial\* or anti bacterial\* or anti-bacterial\* or bacteriocidal or anti-microbial\* or antimicrobial\*) adj3 (agent\* or therap\* or treatment\*)).ti,ab. |
|  | (antibiotic\* or anti-biotic\* or anti biotic\*).ti,ab. |
|  | exp \*beta lactam antibiotic/ |
|  | exp \*beta lactamase inhibitor/ |
|  | exp \*penicillin derivative/ |
|  | exp \*cephalosporin derivative/ |
|  | \*aztreonam/ |
|  | exp \*aminoglycoside antibiotic agent/ |
|  | exp \*quinoline derived antiinfective agent/ |
|  | exp \*polypeptide antibiotic agent/ |
|  | \*rifampicin/ |
|  | \*chloramphenicol/ |
|  | \*cotrimoxazole/ |
|  | \*linezolid/ |
|  | exp \*macrolide/ |
|  | exp \*tetracycline derivative/ |
|  | (co-amoxiclav or amoxicillin-clavulanic or piperacillin or tazobactam or tazocin or piptazobatam or ampicillin or sulbactam).ti,ab. |
|  | (benzylpenicillin or flucloxacillin or amoxicillin).ti,ab. |
|  | (cephalosporin\* or cefuroxime or cefotaxime or ceftriaxone or ceftazidime or ceftaroline or ceftobiprole or cefipime).ti,ab. |
|  | aztreonam.ti,ab. |
|  | (carbapenem\* or imipenem or meropenem or ertapenem).ti,ab. |
|  | (gentamicin or amikacin or tobramycin).ti,ab. |
|  | (ciprofloxacin or levofloxacin).ti,ab. |
|  | (vancomycin or teicoplanin).ti,ab. |
|  | daptomycin.ti,ab. |
|  | tigecycline.ti,ab. |
|  | rifampicin.ti,ab. |
|  | chloramphenicol.ti,ab. |
|  | co-trimoxazole.ti,ab. |
|  | colistin.ti,ab. |
|  | metronidazole.ti,ab. |
|  | (clar#thromycin or clindamycin or linezolid).ti,ab. |
|  | (azithromycin or erythromycin).ti,ab. |
|  | doxycycline.ti,ab. |
|  | or/5-42 |
|  | \*time/ |
|  | therapy delay/ |
|  | ((prompt or hour\* or rapid or within or early or earlier or late\* or time\* or late or later or delay\*) adj8 (initiat\* or start\* or treat\* or therap\* or administ\* or prescri\* or antibiotic\*)).ti,ab. |
|  | or/44-46 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 43 and 47 and 48 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | MeSH descriptor: [anti-infective agents] explode all trees |
|  | ((antiinfect\* or anti infect\* or anti-infect\*) near/3 (agent\* or therap\* or treatment\*)):ti,ab |
|  | ((anti-mycobacterial\* or antimycobacterial\* or anti-mycobacterial\*) near/3 (agent\* or therap\* or treatment\*)):ti,ab |
|  | (microbicid\* or bacteriocid\* or bactericid\*):ti,ab |
|  | ((antibacterial\* or anti bacterial\* or anti-bacterial\* or bacteriocidal or anti-microbial\* or antimicrobial\*) near/3 (agent\* or therap\* or treatment\*)):ti,ab |
|  | (antibiotic\* or anti-biotic\* or anti biotic\*):ti,ab |
|  | MeSH descriptor: [beta-lactams] explode all trees |
|  | (co-amoxiclav or amoxicillin-clavulanic or piperacillin or tazobactam or tazocin or piptazobatam or ampicillin or sulbactam):ti,ab |
|  | (benzylpenicillin or flucloxacillin or amoxicillin):ti,ab |
|  | (cephalosporin\* or cefuroxime or cefotaxime or ceftriaxone or ceftazidime or ceftaroline or ceftobiprole or cefipime):ti,ab |
|  | aztreonam:ti,ab |
|  | (carbapenem\* or imipenem or meropenem or ertapenem):ti,ab |
|  | MeSH descriptor: [aminoglycosides] explode all trees |
|  | (gentamicin or amikacin or tobramycin):ti,ab |
|  | MeSH descriptor: [fluoroquinolones] explode all trees |
|  | (ciprofloxacin or levofloxacin):ti,ab |
|  | MeSH descriptor: [glycopeptides] explode all trees |
|  | (vancomycin or teicoplanin):ti,ab |
|  | MeSH descriptor: [daptomycin] explode all trees |
|  | daptomycin:ti,ab |
|  | MeSH descriptor: [clindamycin] explode all trees |
|  | tigecycline:ti,ab |
|  | MeSH descriptor: [rifampin] explode all trees |
|  | rifampicin:ti,ab |
|  | MeSH descriptor: [chloramphenicol] explode all trees |
|  | chloramphenicol:ti,ab |
|  | MeSH descriptor: [trimethoprim-sulfamethoxazole combination] explode all trees |
|  | co-trimoxazole:ti,ab |
|  | MeSH descriptor: [colistin] explode all trees |
|  | colistin:ti,ab |
|  | MeSH descriptor: [metronidazole] explode all trees |
|  | metronidazole:ti,ab |
|  | MeSH descriptor: [erythromycin] explode all trees |
|  | MeSH descriptor: [doxycycline] explode all trees |
|  | (clar?thromycin or clindamycin or linezolid):ti,ab |
|  | (azithromycin or erythromycin):ti,ab |
|  | doxycycline:ti,ab |
|  | {or #2-#38} |
|  | [mh "time factors"] |
|  | ((prompt or hour\* or rapid or within or early or earlier or late\* or time\* or late or later or delay\*) near/8 (initiat\* or start\* or treat\* or therap\* or administ\* or prescri\* or antibiotic\*)):ti,ab |
|  | #40 and #41 |
|  | #1 and #39 and #42 |

* + 1. Acid-base pH
* Is acid-base balance (that is, the use of bicarbonate) clinically and cost effective in people with sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | acid-base equilibrium/ or acid-base imbalance/ |
|  | (acid base or acid-base or ph).ti,ab. |
|  | exp bicarbonates/ |
|  | bicarbonate\*.ti,ab. |
|  | anion gap\*.ti,ab. |
|  | or/5-9 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 10 and 11 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*ph/ |
|  | \*bicarbonate blood level/ or exp \*bicarbonate/ |
|  | (acid base or acid-base or ph).ti,ab. |
|  | bicarbonate\*.ti,ab. |
|  | anion gap\*.ti,ab. |
|  | or/5-9 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 10 and 11 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | MeSH descriptor: [acid-base equilibrium] explode all trees |
|  | MeSH descriptor: [acid-base imbalance] explode all trees |
|  | MeSH descriptor: [bicarbonates] explode all trees |
|  | (acid base or acid-base or ph):ti,ab |
|  | anion gap\*:ti,ab |
|  | bicarbonate\*:ti,ab |
|  | {or #2-#7} |
|  | #1 and #8 |

* + 1. Oxygen
* Is the use of supplemental oxygen clinically and cost effective in patients with sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | oxygen inhalation therapy/ |
|  | oxygen/ |
|  | oxygen.ti,ab. |
|  | noninvasive ventilation/ |
|  | oximetry/ |
|  | hyperoxia/ |
|  | exp oxygen consumption/ |
|  | ((non invasive or noninvasive) adj ventilat\*).ti,ab. |
|  | (oximetry or hyperoxia).ti,ab. |
|  | or/5-13 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 14 and 15 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | \*oxygen therapy/ |
|  | \*oxygen/ |
|  | oxygen.ti,ab. |
|  | noninvasive ventilation/ |
|  | oximetry/ |
|  | hyperoxia/ |
|  | exp oxygen consumption/ |
|  | ((non invasive or noninvasive) adj ventilat\*).ti,ab. |
|  | (oximetry or hyperoxia).ti,ab. |
|  | or/5-13 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 14 and 15 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | MeSH descriptor: [oxygen inhalation therapy] explode all trees |
|  | MeSH descriptor: [oxygen] explode all trees |
|  | oxygen:ti,ab |
|  | MeSH descriptor: [noninvasive ventilation] explode all trees |
|  | MeSH descriptor: [oximetry] explode all trees |
|  | MeSH descriptor: [hyperoxia] explode all trees |
|  | MeSH descriptor: [oxygen consumption] explode all trees |
|  | (non invasive or noninvasive) near/1 ventilat\*:ti,ab |
|  | (oximetry or hyperoxia):ti,ab |
|  | {or #2-#10} |
|  | #1 and #11 |

* + 1. Inotropes

Searches for the following two questions were run as one search:

* What is the most clinical and cost effective inotropic agent and vasopressor for early management of people with severe sepsis?
* What are the most clinically and cost effective timings of inotropic agents and vasopressors in patients with sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | inotrope\*.ti,ab. |
|  | inotropic\*.ti,ab. |
|  | milrinone/ |
|  | enoximone/ |
|  | dobutamine/ |
|  | exp dopamine/ |
|  | soproterenol/ |
|  | (milrinone or primacor).ti,ab. |
|  | (enoximone or perfan).ti,ab. |
|  | (dobutamine or dopamine).ti,ab. |
|  | (dopexamine or isoprenaline or dopacard or dobutrex or isuprel).ti,ab. |
|  | (vasoactive agent\* or vasopressor\* or vasopressin\*).ti,ab. |
|  | (adrenalin or epinephrine).ti,ab. |
|  | epinephrine/ |
|  | metaraminol/ |
|  | metaraminol.ti,ab. |
|  | norepinephrine/ |
|  | (noradrenalin\* or norepinrphrine).ti,ab. |
|  | or/5-22 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 23 and 24 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | inotrope\*.ti,ab. |
|  | inotropic\*.ti,ab. |
|  | milrinone/ |
|  | enoximone/ |
|  | dobutamine/ |
|  | exp dopamine/ |
|  | (milrinone or primacor).ti,ab. |
|  | (enoximone or perfan).ti,ab. |
|  | (dobutamine or dopamine).ti,ab. |
|  | (dopexamine or isoprenaline or dopacard or dobutrex or isuprel).ti,ab. |
|  | (vasoactive agent\* or vasopressor\* or vasopressin\*).ti,ab. |
|  | (adrenalin or epinephrine).ti,ab. |
|  | adrenalin/ |
|  | isoprenaline/ |
|  | noradrenalin/ |
|  | metaraminol/ |
|  | metaraminol.ti,ab. |
|  | (noradrenalin\* or norepinephrine).ti,ab. |
|  | or/5-22 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 4 and 23 and 24 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | inotrope\*:ti,ab |
|  | inotropic\*:ti,ab |
|  | MeSH descriptor: [milrinone] this term only |
|  | MeSH descriptor: [enoximone] this term only |
|  | MeSH descriptor: [dobutamine] this term only |
|  | MeSH descriptor: [dopamine] explode all trees |
|  | (milrinone or primacor):ti,ab |
|  | (enoximone or perfan):ti,ab |
|  | (dobutamine or dopamine):ti,ab |
|  | (dopexamine or isoprenaline or dopacard or dobutrex or isuprel):ti,ab |
|  | (vasoactive agent\* or vasopressor\* or vasopressin\*):ti,ab |
|  | (adrenalin or epinephrine):ti,ab |
|  | MeSH descriptor: [epinephrine] this term only |
|  | MeSH descriptor: [isoproterenol] this term only |
|  | MeSH descriptor: [metaraminol] this term only |
|  | metaraminol:ti,ab |
|  | MeSH descriptor: [norepinephrine] this term only |
|  | (noradrenalin\* or norepinrphrine):ti,ab |
|  | {or #2-#19} |
|  | #1 and #20 |

* + 1. Escalation of care

Searches for the following two questions were run as one search:

* When is the most appropriate time for care of people with sepsis to be directed to (a) a senior healthcare professional, and (b) critical care providers?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | 1 or 2 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 3 not 4 |
|  | Limit 5 to English language |
|  | "delivery of health care"/ |
|  | exp critical care/ |
|  | ((intensive or critical) adj2 care).ti,ab. |
|  | (intensivist\* or consultant\* or specialist\* or senior\*1 or junior\*1 or sho or registrar\* or spr or house officer\* or houseofficer\* or housestaff\* or physician\* or intern\*1 or internship or resident\*1 or fellow\*1 or foundation doctor or nurs\*).ti,ab. |
|  | or/7-10 |
|  | \*time factors/ |
|  | (time or times or timing or referral or refer or refers or referring).ti,ab. |
|  | ((early or earlie\* or late or later or schedul\* or hour\* or rapid\* or fast\* or slow\* or delay\* or immediate\* or escalat\* or manage\* or managing or hospital) adj2 care).ti,ab. |
|  | or/12-14 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 6 and 11 and 15 and 16 |
|  | limit 17 to yr="1999 -Current" |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | 1 or 2 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 3 not 4 |
|  | Limit 5 to English language |
|  | \*health care delivery/ |
|  | \*exp intensive care/ |
|  | ((intensive or critical) adj2 care).ti,ab. |
|  | (intensivist\* or consultant\* or specialist\* or senior\*1 or junior\*1 or sho or registrar\* or spr or house officer\* or houseofficer\* or housestaff\* or physician\* or intern\*1 or internship or resident\*1 or fellow\*1 or foundation doctor or nurs\*).ti,ab. |
|  | or/7-10 |
|  | \*time factors/ |
|  | (time or times or timing or referral or refer or refers or referring).ti,ab. |
|  | ((early or earlie\* or late or later or schedul\* or hour\* or rapid\* or fast\* or slow\* or delay\* or immediate\* or escalat\* or manage\* or managing or hospital) adj2 care).ti,ab. |
|  | or/12-14 |
|  | Study filters RCT [A.3.2] or SR [A.3.3] or OBS [A.3.7] |
|  | 6 and 11 and 15 and 16 |
|  | limit 17 to yr="1999 -Current" |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | #1 or #2 |
|  | MeSH descriptor: [delivery of health care] this term only |
|  | MeSH descriptor: [critical care] explode all trees |
|  | ((intensive or critical) near/2 care):ti,ab |
|  | (intensivist\* or consultant\* or specialist\* or senior\*1 or junior\*1 or SHO or registrar\* or SPR or house officer\* or houseofficer\* or housestaff\* or physician\* or intern\*1 or internship or resident\*1 or fellow\*1 or foundation doctor or nurs\*):ti,ab |
|  | {or #4-#7} |
|  | MeSH descriptor: [time factors] explode all trees |
|  | (time or times or timing or referral or refer or refers or referring):ti,ab |
|  | ((early or earlie\* or late or later or schedul\* or hour\* or rapid\* or fast\* or slow\* or delay\* or immediate\* or escalat\* or manage\* or managing or hospital) near/2 care):ti,ab |
|  | {or #9-#11} |
|  | #3 and #8 and #12 Publication Year from 1999 to 2015 |

* + 1. Monitoring
* In people with sepsis or severe sepsis, what is the clinical and cost effectiveness of scoring systems, and specified blood markers in monitoring response to treatment?

**Medline search terms**

|  |  |
| --- | --- |
|  | (early warn\* adj4 (system\* or scor\* or criteri\* or tool\*)).ti,ab. |
|  | (pews or mews).ti,ab. |
|  | 1 or 2 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 3 not 4 |
|  | Limit 5 to English language |

**Embase search terms**

|  |  |
| --- | --- |
|  | (early warn\* adj4 (system\* or scor\* or criteri\* or tool\*)).ti,ab. |
|  | (pews or mews).ti,ab. |
|  | 1 or 2 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 3 not 4 |
|  | Limit 5 to English language |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | (early warn\* near/4 (system\* or scor\* or criteri\* or tool\*)):ti,ab |
|  | (pews or mews):ti,ab |
|  | #1 or #2 |

* + 1. Information support

Searches for the following four questions were run as one search:

* What information, education and support would be useful for people assessed for possible sepsis, but discharged from medical care
* What information, education and support would be useful for people at high risk of sepsis
* What information, education and support would be useful for people who have sepsis or severe sepsis, families and carers
* What information, education and support would be useful for people who survived episodes of severe sepsis

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | patient education as topic/ |
|  | patient acceptance of health care/ |
|  | patient satisfaction/ |
|  | patient education handout/ |
|  | consumer health information/ |
|  | (information adj (need\* or requirement\* or support\*)).ti,ab. |
|  | (discharg\* adj2 (information\* or advice)).ti,ab. |
|  | ((patient\* or carer\* or famil\* or parent\*1 or father\*1 or mother\*1 or caregiver\* or next of kin) adj3 (inform\* or educat\* or support\* or advice\* or advise\*)).ti,ab. |
|  | ((patient\* or carer\* or famil\* or parent\*1 or father\*1 or mother\*1 or caregiver\* or next of kin) adj2 (pamphlet\* or leaflet\* or booklet\* or manual\* or brochure\* or publication\* or handout\* or website\* or web site\* or web page\* or webpage\* or video\* or dvd\* or education or educate or educating or literature or information or internet or computer\* or program\* or interactive or email\* or e-mail\* or wireless or bluetooth or telephone or phone or sms or text\*)).ti,ab. |
|  | or/5-13 |
|  | 4 and 14 |
|  | limit 15 to yr="1990 -Current" |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | patient education/ |
|  | patient attitude/ |
|  | patient satisfaction/ |
|  | consumer health information/ |
|  | (information adj (need\* or requirement\* or support\*)).ti,ab. |
|  | (discharg\* adj2 (information\* or advice)).ti,ab. |
|  | ((patient\* or carer\* or famil\* or parent\*1 or father\*1 or mother\*1 or caregiver\* or next of kin) adj3 (inform\* or educat\* or support\* or advice\* or advise\*)).ti,ab. |
|  | ((patient\* or carer\* or famil\* or parent\*1 or father\*1 or mother\*1 or caregiver\* or next of kin) adj2 (pamphlet\* or leaflet\* or booklet\* or manual\* or brochure\* or publication\* or handout\* or website\* or web site\* or web page\* or webpage\* or video\* or dvd\* or education or educate or educating or literature or information or internet or computer\* or program\* or interactive or email\* or e-mail\* or wireless or bluetooth or telephone or phone or sms or text\*)).ti,ab. |
|  | or/5-12 |
|  | 4 and 13 |
|  | limit 14 to yr="1990 -Current" |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | [mh "patient education as topic"] |
|  | [mh "patient acceptance of health care"] |
|  | [mh "patient satisfaction"] |
|  | [mh "patient education handout"] |
|  | [mh "consumer health information"] |
|  | (information near/1 (need\* or requirement\* or support\*)):ti,ab |
|  | (discharg\* near/2 (information\* or advice)):ti,ab |
|  | ((patient\* or carer\* or famil\* or parent or parents or father\* or mother\* or caregiver\* or "next of kin") near/3 (inform\* or educat\* or support\* or advice\* or advise\*)):ti,ab |
|  | ((patient\* or carer\* or famil\* or parent or parents or father\* or mother\* or caregiver\* or "next of kin") near/2 (pamphlet\* or leaflet\* or booklet\* or manual\* or brochure\* or publication\* or handout\* or website\* or web site\* or web page\* or webpage\* or video\* or dvd\* or education or educate or educating or literature or information or internet or computer\* or program\* or interactive or email\* or e-mail\* or wireless or bluetooth or telephone or phone or sms or text\*)):ti,ab |
|  | {or #2-#10} |
|  | #1 and #11 Publication Year from 1990 to 2015 |

**CINAHL search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | (MH "patient satisfaction") |
|  | (MH "consumer health information") |
|  | (MH "patient discharge education") |
|  | (MH "patient education") |
|  | (discharg\* n2 (information\* or advice)) |
|  | (information n1 (need\* or requirement\* or support\*)) |
|  | ((patient\* or carer\* or famil\* or parent or parents or father\* or mother\* or caregiver\* or next of kin) n3 (inform\* or educat\* or support\* or advice\* or advise\*)) |
|  | ((patient\* or carer\* or famil\* or parent or parents or father\* or mother\* or caregiver\* or next of kin) n2 (pamphlet\* or leaflet\* or booklet\* or manual\* or brochure\* or publication\* or handout\* or website\* or web site\* or web page\* or webpage\* or video\* or dvd\* or education or educate or educating or literature or information or internet or computer\* or program\* or interactive or email\* or e-mail\* or wireless or bluetooth or telephone or phone or sms or text\*)) |
|  | S5 or S6 or S7 or S8 or S9 or S10 or S11 or S12 |
|  | S4 and S13 Limiters - Published Date: 19900101-20151231; Exclude MEDLINE records |

**PsycINFO (Ovid) search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | limit 1 to (English language) |
|  | client education/ |
|  | client attitudes/ |
|  | client satisfaction/ |
|  | (information adj (need\* or requirement\* or support\*)).ti,ab. |
|  | (discharg\* adj2 (information\* or advice)).ti,ab. |
|  | ((patient\* or carer\* or famil\* or parent\*1 or father\*1 or mother\*1 or caregiver\* or next of kin) adj3 (inform\* or educat\* or support\* or advice\* or advise\*)).ti,ab. |
|  | ((patient\* or carer\* or famil\* or parent\*1 or father\*1 or mother\*1 or caregiver\* or next of kin) adj2 (pamphlet\* or leaflet\* or booklet\* or manual\* or brochure\* or publication\* or handout\* or website\* or web site\* or web page\* or webpage\* or video\* or dvd\* or education or educate or educating or literature or information or internet or computer\* or program\* or interactive or email\* or e-mail\* or wireless or bluetooth or telephone or phone or SMS or text\*)).ti,ab. |
|  | or/3-9 |
|  | 2 and 10 |
|  | limit 11 to yr="1990 -Current" |

**PsycINFO (ProQUEST) search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | ((su.exact("client education") or su.exact("client attitudes") or su.exact("client satisfaction") or (ti,ab(information near (need\* or requirement\* or support\*)) or ti,ab(discharg\* near/2 (information\* or advice)))) or ti,ab((patient\* or carer\* or famil\* or parent\* or father\* or mother\* or caregiver\* or "next of kin") near/3 (inform\* or educat\* or support\* or advice\* or advise\*)) or ti,ab((patient\* or carer\* or famil\* or parent\* or father\* or mother\* or caregiver\* or "next of kin") near/2 (pamphlet\* or leaflet\* or booklet\* or manual\* or brochure\* or publication\* or handout\* or website\* or "web site\*" or "web page\*" or webpage\* or video\* or dvd\* or education or educate or educating or literature or information or internet or computer\* or program\* or interactive or email\* or e-mail\* or wireless or bluetooth or telephone or phone or sms or text\*)) |
|  | 1 and 2 additional limits - Date: From 1990 to 2015; Language: English |

* + 1. Educational programmes/Identification protocols
* What education and training programmes are available, effective and cost effective for the early recognition, diagnosis and management of sepsis and severe sepsis?

**Medline search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | 1 or 2 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 3 not 4 |
|  | Limit 5 to English language |
|  | exp health personnel/ |
|  | (clinician\* or doctor\* or physician\* or nurse or nurses\* or specialist\* or registrar\* or gp or gps or general practitioner\* or hca or hcas or health care assistant\* or consultant\* or trainee\* or team or teams or personnel or staff or professional\* or healthcare).ti,ab. |
|  | exp primary health care/ |
|  | exp physician's practice patterns/ |
|  | exp family practice/ |
|  | exp physicians, primary care/ |
|  | exp general practice/ |
|  | exp physicians, family/ |
|  | exp general practitioners/ |
|  | ((primary or communit\*) adj5 care).ti,ab. |
|  | (family practi\* or family doctor\* or family physician\* or gp\* or general practi\*).ti,ab. |
|  | exp community health services/ or exp community health centers/ |
|  | exp ambulatory care/ |
|  | casualty\*.tw. |
|  | paramedic\*.ti,ab. |
|  | ((ambulance or ambulatory) adj2 (care or caring)).ti,ab. |
|  | emergency medical services/ |
|  | "pre hospital\*".ti,ab. |
|  | or/7-24 |
|  | early diagnosis/ |
|  | education/ |
|  | exp education, professional/ |
|  | exp inservice training/ |
|  | ((program\* or system or systems) adj4 (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ti,ab. |
|  | (manag\* and (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ti. |
|  | (manag\* adj3 (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ti,ab. |
|  | (awareness or identif\* or recogni\* or detect\* or alert\* or warn\*).ti. |
|  | ((awareness or identif\* or recogni\* or detect\* or alert\* or warn\*) adj3 (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ab. |
|  | (education\* or train\* or program\* or learn\*).ti. |
|  | ((program\* or system or systems) adj4 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)).ti,ab. |
|  | manage\*.ti. |
|  | (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*).ti. |
|  | 37 and 38 |
|  | (manag\* adj3 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)).ti,ab. |
|  | ((awareness or identif\* or recogni\* or detect\* or alert\* or warn\*) adj3 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)).ab. |
|  | or/26-36,39-41 |
|  | 6 and 25 and 42 |

**Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | 1 or 2 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 3 not 4 |
|  | Limit 5 to English language |
|  | exp health personnel/ |
|  | (clinician\* or doctor\* or physician\* or nurse or nurses\* or specialist\* or registrar\* or gp or gps or general practitioner\* or hca or hcas or health care assistant\* or consultant\* or trainee\* or team or teams or personnel or staff or professional\* or healthcare).ti,ab. |
|  | exp primary health care/ |
|  | exp professional practice/ |
|  | exp clinical practice/ |
|  | exp general practice/ |
|  | general practitioner/ |
|  | ((primary or communit\*) adj5 care).ti,ab. |
|  | exp ambulatory care/ |
|  | exp community care/ |
|  | health center/ |
|  | paramedic\*.ti,ab. |
|  | ((ambulance or ambulatory) adj2 (care or caring)).ti,ab. |
|  | emergency health service/ |
|  | "pre hospital\*".ti,ab. |
|  | or/7-21 |
|  | early diagnosis/ |
|  | education/ |
|  | exp education, professional/ |
|  | exp inservice training/ |
|  | ((program\* or system or systems) adj4 (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ti,ab. |
|  | (manag\* and (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ti. |
|  | (manag\* adj3 (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ti,ab. |
|  | (awareness or identif\* or recogni\* or detect\* or alert\* or warn\*).ti. |
|  | ((awareness or identif\* or recogni\* or detect\* or alert\* or warn\*) adj3 (sepsis or septic\* or (blood adj2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)).ab. |
|  | (education\* or train\* or program\* or learn\*).ti. |
|  | ((program\* or system or systems) adj4 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)).ti,ab. |
|  | manage\*.ti. |
|  | (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*).ti. |
|  | 34 and 35 |
|  | (manag\* adj3 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)).ti,ab. |
|  | ((awareness or identif\* or recogni\* or detect\* or alert\* or warn\*) adj3 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)).ab. |
|  | or/23-33,36-38 |
|  | 6 and 22 and 39 |

**Cochrane search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | #1 or #2 |
|  | MeSH descriptor: [health personnel] explode all trees |
|  | (clinician\* or doctor\* or physician\* or nurse or nurses\* or specialist\* or registrar\* or gp or gps or general practitioner\* or hca or hcas or health care assistant\* or consultant\* or trainee\* or team or teams or personnel or staff or professional\* or healthcare):ti,ab |
|  | MeSH descriptor: [primary health care] explode all trees |
|  | MeSH descriptor: [physician's practice patterns] explode all trees |
|  | MeSH descriptor: [family practice] explode all trees |
|  | MeSH descriptor: [physicians, primary care] explode all trees |
|  | MeSH descriptor: [general practice] explode all trees |
|  | MeSH descriptor: [general practitioners] explode all trees |
|  | ((primary or communit\*) near/5 care):ti,ab |
|  | (family practi\* or family doctor\* or family physician\* or gp\* or general practi\*):ti,ab |
|  | MeSH descriptor: [community health services] explode all trees |
|  | MeSH descriptor: [community health centers] explode all trees |
|  | MeSH descriptor: [ambulatory care] explode all trees |
|  | casualty\*:ti,ab,kw |
|  | paramedic\*:ti,ab |
|  | ((ambulance or ambulatory) near/2 (care or caring)):ti,ab |
|  | MeSH descriptor: [emergency medical services] this term only |
|  | pre hospital\*:ti,ab |
|  | {or #4-#21} |
|  | MeSH descriptor: [early diagnosis] this term only |
|  | MeSH descriptor: [education] this term only |
|  | MeSH descriptor: [education, professional] explode all trees |
|  | MeSH descriptor: [inservice training] explode all trees |
|  | ((program\* or system or systems) near/4 (sepsis or septic\* or (blood near/2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)):ti,ab |
|  | (manag\* and (sepsis or septic\* or (blood near/2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)):ti |
|  | (manag\* near/j3 (sepsis or septic\* or (blood near/2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)):ti,ab |
|  | (awareness or identif\* or recogni\* or detect\* or alert\* or warn\*):ti |
|  | ((awareness or identif\* or recogni\* or detect\* or alert\* or warn\*) near/3 (sepsis or septic\* or (blood near/2 (pathogen\* or poison\*)) or systemic inflammatory response syndrome' or sirs or pyaemi\* or pyemi\* or pyohemi\* or bacter?emi\* or fung?emi\* or parasit?emi\* or vir?emi\*)):ab |
|  | (education\* or train\* or program\* or learn\*):ti |
|  | ((program\* or system or systems) near/4 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)) .ti,ab |
|  | manage\*:ti |
|  | (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*):ti |
|  | #34 and #35 |
|  | (manag\* near/3 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)):ti,ab |
|  | ((awareness or identif\* or recogni\* or detect\* or alert\* or warn\*) near/3 (meningitis or meningitides or meningococc\* or meningococcemia or pneumoccoc\* or streptococc\*)):ab |
|  | {or #23-#33,#36-#38} |
|  | #3 and #22 and #39 |

* 1. Health economics search
     1. Health economic (HE) reviews

Economic searches were conducted in Medline, Embase, CRD and HEED.

**Medline & Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | 1 or 2 |
|  | Excluded study designs and publication types [A.3.1] |
|  | 3 not 4 |
|  | Limit 5 to English language |
|  | Study filter HE [A.3.4] |
|  | 6 and 7 |
|  | limit 8 to yr="2012 -Current" |

**CRD search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | (#1 or #2) |
|  | ((#3) from 1999 to 2015) in NHSEED |
|  | ((#3) from 1999 to 2015) in HTA |

**HEED search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Standard bacterial meningitis population [A.2.2] |
|  | cs=1 or 2 jd=2012-2014 |

* + 1. Quality of life (QOL) reviews

Quality of life searches were conducted in Medline and Embase only

**Medline & Embase search terms**

|  |  |
| --- | --- |
|  | Standard sepsis population [A.2.1] |
|  | Excluded study designs and publication types [A.3.1] |
|  | 1 not 2 |
|  | Limit 3 to English language |
|  | exp child/ not (exp child/ and (exp adult/ or adolescent/)) |
|  | exp infant/ not (exp infant/ and (exp adult/ or adolescent/)) |
|  | 5 or 6 |
|  | 4 not 7 |
|  | Study filter QOL [A.3.5] |
|  | 8 and 9 |

* 1. References