

Fluid therapy

See initial resuscitation timing recommendations.

Use crystalloids or colloids.

Give fluid challenge to patients with suspected inadequate tissue perfusion at a rate of 500–1000 ml of crystalloids or 300–500 ml of colloids over 30 minutes and repeat if blood pressure and urine output do not increase and there is no evidence of intravascular volume overload.

Blood product administration

Following resolution of tissue hypoperfusion, and in the absence of

significant coronary artery disease or acute haemorrhage, transfuse red blood cells when haemoglobin decreases to <7.0 g/dl (<70 g/L) to target a haemoglobin of 7.0 - 9.0 g/dl.

Do not use erythropoietin to treat sepsis-related anaemia.

Erythropoietin may be used for other accepted reasons.

Do not use fresh frozen plasma to correct laboratory clotting abnormalities unless there is bleeding or planned invasive procedures.

Do not use antithrombin therapy.

Administer platelets when counts are $<5000/\text{mm}^3$ ($5 \times 10^9/\text{L}$) regardless of bleeding. Transfuse platelets when counts are 5000 to 30,000/ mm^3 ($5\text{--}30 \times 10^9/\text{L}$) and there is significant bleeding risk.

Higher platelet counts ($\geq 50,000/\text{mm}^3$ [$50 \times 10^9/\text{L}$]) are required for surgery or invasive procedures.

Sedation, analgesia, and neuromuscular blockade in sepsis

Use sedation protocols for critically ill mechanically

ventilated patients. Measure the sedation goal with a standardized subjective sedation scale.

Target sedation to predetermined end-points (sedation score).

Use either intermittent bolus sedation or continuous infusion sedation with daily interruption/lightening to produce awakening. Re-titrate if necessary.

Avoid neuromuscular blockers (NMBs), if at all possible.

If NMBs must be utilized for longer than the first 2 to 3 hours of mechanical ventilation, use either intermittent bolus as required or continuous infusion with monitoring of depth of block with train of four monitoring.

Mechanical ventilation of sepsis-induced acute lung injury (ALI)/ARDS

Avoid high tidal volumes coupled

with high plateau pressures. Reduce tidal volumes over 1-2 hours to a low tidal volume (6 ml per kilogram of lean body weight) as a goal in conjunction with the goal of maintaining end-inspiratory plateau pressures <30 cm H_2O .

If necessary, minimize plateau pressures and tidal volumes by allowing PaCO_2 to increase above normal.

Set a minimum amount of positive end-expiratory pressure (PEEP) to prevent lung collapse at end expiration. Set PEEP based on severity of oxygenation deficit and guided by the FiO_2 required to maintain adequate oxygenation (ARDSnet guidelines) or titrate PEEP according to bedside measurements of thoracopulmonary compliance.

Prone ARDS patients requiring potentially injurious levels of FiO_2 or plateau pressure. Only prone patients not at high risk from positional changes.

To prevent ventilator-associated pneumonia maintain mechanically ventilated patients in a semi-recumbent position (head of bed raised 45 degrees), unless contraindicated.

Use a weaning protocol and have mechanically ventilated patients undergo a spontaneous breathing trial (SBT), at least daily, to evaluate for ventilation discontinuation.

SBT options include a low level of pressure support with continuous positive airway pressure 5 cm H_2O or a T-piece. Prior to SBT, patients should:

- ◆ be arousable;
- ◆ be haemodynamically stable without vasopressors;
- ◆ have no new potentially serious conditions;
- ◆ have low ventilatory and end-expiratory pressure requirement; and
- ◆ require FiO_2 levels that can be safely delivered with a face mask or nasal cannula.

Consider extubation if SBT is successful.

Glucose control

Maintain blood glucose <150 mg/dl (8.3mmol/L) following initial stabilization. Use continuous insulin and glucose infusion. Monitor blood glucose every 30 - 60 minutes until stabilized, then monitor every 4 hours.

Include a nutritional protocol for glycaemic control.

Recombinant human activated protein C (rhAPC)

rhAPC is recommended in patients at high risk of death

(APACHE II ≥ 25 , sepsis-induced multiple organ failure, septic shock, or sepsis-induced acute respiratory distress syndrome) and with no absolute contraindication related to bleeding risk or relative contraindication that outweighs the potential benefit of rhAPC.

rhAPC is indicated for the treatment of adult patients with severe sepsis with multiple organ failure when added to best standard care.

Renal replacement

Intermittent haemodialysis and continuous veno venous haemofiltration (CVVH) are considered equivalent. CVVH offers easier management in haemodynamically unstable patients.

Bicarbonate therapy

Do not use bicarbonate therapy for the purpose of improving haemodynamics or reducing vasopressor requirements when treating hypoperfusion induced lactic acidemia with $\text{pH} \geq 7.15$.

Deep vein thrombosis (DVT) prophylaxis

Use either low-dose unfractionated heparin or low-molecular weight

heparin. Use a mechanical prophylactic device, such as compression stockings or an intermittent compression device, when heparin is contraindicated.

Use a combination of pharmacologic and mechanical therapy for patients who are at very high risk for DVT.

Stress ulcer prophylaxis

Provide stress ulcer prophylaxis.

The preferred agents are H_2 receptor inhibitors.

Consideration for limitation of support

Discuss advance care planning with patients and families. Describe likely outcomes and set realistic expectations.