

Surviving Sepsis Campaign[®]



SURVIVING SEPSIS CAMPAIGN INTERNATIONAL GUIDELINES FOR THE MANAGEMENT OF SEPTIC SHOCK AND SEPSIS-ASSOCIATED ORGAN DYSFUNCTION IN CHILDREN

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SCREENING, DIAGNOSIS, AND SYSTEMATIC MANAGEMENT RECOMMENDATIONS TABLE

| RECOMMENDATION #1 | STRENGTH & QUALITY OF EVIDENCE |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|
| <p>In children who present as acutely unwell, we suggest implementing systematic screening for timely recognition of septic shock and other sepsis-associated organ dysfunction.</p> <p>Remarks: Systematic screening needs to be tailored to the type of patients, resources, and procedures within each institution. Evaluation for the effectiveness and sustainability of screening should be incorporated as part of this process.</p> | <ul style="list-style-type: none">• Weak• Very Low-Quality of Evidence |
| RECOMMENDATION #2 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We were unable to issue a recommendation about using blood lactate values to stratify children with suspected septic shock or other sepsis-associated organ dysfunction into low- versus high-risk of having septic shock or sepsis. However, in our practice, if lactate levels can be rapidly obtained, we often measure blood lactate in children when evaluating for septic shock and other sepsis-associated organ dysfunction.</p> | <p>Insufficient</p> |
| RECOMMENDATION #3 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We recommend implementing a protocol/guideline for management of children with septic shock or other sepsis-associated organ dysfunction.</p> | <p>Best Practice Statement</p> |

RECOMMENDATION #4

**STRENGTH &
QUALITY OF EVIDENCE**

We *recommend obtaining* blood cultures before initiating antimicrobial therapy in situations where this does not substantially delay antimicrobial administration.

Best Practice
Statement

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ANTIMICROBIAL THERAPY RECOMMENDATIONS TABLE

| RECOMMENDATION #5 | STRENGTH & QUALITY OF EVIDENCE |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------|
| In children with septic shock, we recommend starting antimicrobial therapy as soon as possible, within one (1) hour of recognition. | <ul style="list-style-type: none"> • Strong • Very Low-Quality of Evidence |
| RECOMMENDATION #6 | STRENGTH & QUALITY OF EVIDENCE |
| In children with sepsis-associated organ dysfunction but without shock, we suggest starting antimicrobial therapy as soon as possible after appropriate evaluation, within three (3) hours of recognition. | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |
| RECOMMENDATION #7 | STRENGTH & QUALITY OF EVIDENCE |
| We recommend empiric broad-spectrum therapy with one or more antimicrobials to cover all likely pathogens. | Best Practice Statement |
| RECOMMENDATION #8 | STRENGTH & QUALITY OF EVIDENCE |
| Once the pathogen(s) and sensitivities are available, we recommend narrowing empiric antimicrobial therapy coverage. | Best Practice Statement |

RECOMMENDATION #9**STRENGTH &
QUALITY OF EVIDENCE**

If no pathogen is identified, we **recommend** narrowing or stopping empiric antimicrobial therapy according to clinical presentation, site of infection, host risk factors, and adequacy of clinical improvement in discussion with infectious disease and/or microbiological expert advice.

Best Practice
Statement

RECOMMENDATION #10**STRENGTH &
QUALITY OF EVIDENCE**

In children without immune compromise and without high risk for multidrug-resistant pathogens, we **suggest against** the routine use of empiric multiple antimicrobials directed against the same pathogen for the purpose of synergy. **Remarks:** In certain situations, such as confirmed or strongly suspected group B streptococcal sepsis, use of empiric multiple antimicrobials directed against the same pathogen for the purpose of synergy may be indicated.

- Weak
- Very Low-Quality of Evidence

RECOMMENDATION #11**STRENGTH &
QUALITY OF EVIDENCE**

In children with immune compromise and/or at high risk for multidrug-resistant pathogens, we **suggest** using empiric multi-drug therapy when septic shock or other sepsis-associated organ dysfunction is present/suspected.

- Weak
- Very Low-Quality of Evidence

RECOMMENDATION #12**STRENGTH &
QUALITY OF EVIDENCE**

We **recommend** using antimicrobial dosing strategies that have been optimized based on published pharmacokinetic/pharmacodynamic principles and with consideration of specific drug properties.

Best Practice
Statement

RECOMMENDATION #13

STRENGTH & QUALITY OF EVIDENCE

In children with septic shock or sepsis-associated organ dysfunction who are receiving antimicrobials, we **recommend** daily assessment (e.g., clinical, laboratory assessment) for de-escalation of antimicrobial therapy. **Remarks:** This assessment should include a review of the ongoing indication for empiric antimicrobial therapy after the first 48 hours that is guided by microbiologic results and in response to clinical improvement and/or evidence of infection resolution. This recommendation applies to patients being treated with empiric, targeted, and combination therapy.

Best Practice
Statement

RECOMMENDATION #14

STRENGTH & QUALITY OF EVIDENCE

We **recommend** determining the duration of antimicrobial therapy according to the site of infection, microbial etiology, response to treatment, and ability to achieve source control.

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SOURCE CONTROL RECOMMENDATIONS TABLE

| RECOMMENDATION #15 | STRENGTH & QUALITY OF EVIDENCE |
|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------|
| <p>We recommend that emergent source control intervention be implemented as soon possible after a diagnosis of an infection amenable to a source control procedure is made. Remarks: Appropriate diagnostic testing to identify the site of infection and microbial etiology should be performed, and advice from specialist teams (e.g., infectious diseases, surgery) should be sought, as appropriate, in order to prioritize interventions needed to achieve source control.</p> | <p>Best Practice Statement</p> |
| RECOMMENDATION #16 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We recommend removal of intravascular access devices that are confirmed to be the source of sepsis or septic shock after other vascular access has been established and depending on the pathogen and the risks/benefits of a surgical procedure.</p> | <ul style="list-style-type: none">• Strong• Low-Quality of Evidence |

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FLUID THERAPY RECOMMENDATIONS TABLE

| RECOMMENDATION #17 | STRENGTH & QUALITY OF EVIDENCE |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------|
| In healthcare systems with availability of intensive care, we suggest administering up to 40–60mL/kg in bolus fluid (10–20mL/kg per bolus) over the first hour, titrated to clinical markers of cardiac output and discontinued if signs of fluid overload develop, for the initial resuscitation of children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |
| RECOMMENDATION #18 | STRENGTH & QUALITY OF EVIDENCE |
| In healthcare systems with no availability of intensive care and in the <i>absence of hypotension</i> , we recommend against bolus fluid administration while starting maintenance fluids. | <ul style="list-style-type: none">• Strong• High-Quality of Evidence |
| RECOMMENDATION #19 | STRENGTH & QUALITY OF EVIDENCE |
| In healthcare systems with no availability of intensive care, if hypotension is present, we suggest administering up to 40mL/kg in bolus fluid (10–20mL/kg per bolus) over the first hour with titration to clinical markers of cardiac output and discontinued if signs of fluid overload develop. Remarks: Clinical markers of cardiac output may include heart rate, blood pressure, capillary refill time, level of consciousness, and urine output. In all settings, the need for fluid administration should be guided by frequent reassessment of clinical markers of | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |

cardiac output, serial blood lactate measurement and advanced monitoring, when available. Signs of fluid overload that should limit further fluid bolus therapy may include clinical signs of pulmonary edema or new or worsening hepatomegaly.

| RECOMMENDATION #20 | STRENGTH & QUALITY OF EVIDENCE |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|
| We suggest using crystalloids, rather than albumin, for the initial resuscitation of children with septic shock or other sepsis-associated organ dysfunction. Remarks: Although there is no difference in outcomes, this recommendation takes into consideration cost and other barriers of administering albumin compared with crystalloids. | <ul style="list-style-type: none">• Weak• Moderate-Quality of Evidence |

| RECOMMENDATION #21 | STRENGTH & QUALITY OF EVIDENCE |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|
| We suggest using balanced/buffered crystalloids, rather than 0.9% saline, for the initial resuscitation of children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none">• Weak• Very Low-Quality of Evidence |

| RECOMMENDATION #22 | STRENGTH & QUALITY OF EVIDENCE |
|---------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------|
| We recommend against using starches in the acute resuscitation of children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none">• Strong• Moderate-Quality of Evidence |

| RECOMMENDATION #23 | STRENGTH & QUALITY OF EVIDENCE |
|------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------|
| We suggest against using gelatin in the resuscitation of children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |

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HEMODYNAMIC MONITORING RECOMMENDATIONS TABLE

| RECOMMENDATION #24 | STRENGTH & QUALITY OF EVIDENCE |
|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|
| We were <i>unable to issue a recommendation</i> about whether to target mean arterial blood pressure (MAP) at the 5th or 50th percentile for age in children with septic shock and other sepsis-associated organ dysfunction. However, <i>in our practice</i> , we target MAP to between the 5th and 50th percentile or greater than 50th percentile for age. | Insufficient In Our Practice |
| RECOMMENDATION #25 | STRENGTH & QUALITY OF EVIDENCE |
| We <i>suggest not using</i> bedside clinical signs in isolation to categorize septic shock in children as “warm” or “cold”. | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |
| RECOMMENDATION #26 | STRENGTH & QUALITY OF EVIDENCE |
| We <i>suggest</i> using advanced hemodynamic variables, when available, in addition to bedside clinical variables to guide the resuscitation of children with septic shock or other sepsis-associated organ dysfunction. Remarks: Advanced hemodynamic monitoring may include cardiac output/cardiac index, systemic vascular resistance, or central venous oxygen saturation (Scvo2). | <ul style="list-style-type: none"> • Weak • Low-Quality of Evidence |

RECOMMENDATION #27

STRENGTH & QUALITY OF EVIDENCE

We **suggest** using trends in blood lactate levels, in addition to clinical assessment, to guide resuscitation of children with septic shock and other sepsis-associated organ dysfunction.

Remarks: In children with an elevated blood lactate, repeat testing that reveals a persistent elevation in blood lactate may indicate incomplete hemodynamic resuscitation and should prompt efforts, as needed, to further promote hemodynamic stability.

- Weak
- Very Low-Quality of Evidence

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VASOACTIVE MEDICATIONS RECOMMENDATIONS TABLE

| RECOMMENDATION #28 | STRENGTH & QUALITY OF EVIDENCE |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|
| We suggest using epinephrine, rather than dopamine, in children with septic shock. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |
| RECOMMENDATION #29 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest using norepinephrine, rather than dopamine, in children with septic shock. | <ul style="list-style-type: none">• Weak• Very Low-Quality of Evidence |
| RECOMMENDATION #30 | STRENGTH & QUALITY OF EVIDENCE |
| We were unable to issue a recommendation for a specific first-line vasoactive infusion for children with septic shock. However, in our practice, we select either epinephrine or norepinephrine as the first-line vasoactive infusion guided by clinician preference, individual patient physiology, and local system factors. | Insufficient |
| RECOMMENDATION #31 | STRENGTH & QUALITY OF EVIDENCE |
| We were unable to issue a recommendation about initiating vasoactive agents through peripheral access in children with septic shock. However, in our practice, we often or sometimes administer a dilute concentration of the initial vasoactive medication through a peripheral vein if central venous access is not readily accessible. | Insufficient |

RECOMMENDATION #32

STRENGTH & QUALITY OF EVIDENCE

We **suggest** either adding vasopressin or further titrating catecholamines in children with septic shock who require high-dose catecholamines. **Remarks:** No consensus was achieved on the optimal threshold for initiating vasopressin. Therefore, this decision should be made according to individual clinician preference.

- Weak
- Low-Quality of Evidence

RECOMMENDATION #33

STRENGTH & QUALITY OF EVIDENCE

We were unable to issue a recommendation about adding an inodilator in children with septic shock and cardiac dysfunction despite other vasoactive agents. However, in our practice, we sometimes use inodilators in children with septic shock and evidence of persistent hypoperfusion and cardiac dysfunction despite other vasoactive agents.

Insufficient

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VENTILATION RECOMMENDATIONS TABLE

| RECOMMENDATION #34 | STRENGTH & QUALITY OF EVIDENCE |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|
| We were unable to issue a recommendation about whether to intubate children with fluid-refractory, catecholamine-resistant septic shock. However, in our practice, we commonly intubate children with fluid-refractory, catecholamine-resistant septic shock without respiratory failure. | Insufficient |
| RECOMMENDATION #35 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest not to use etomidate when intubating children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |
| RECOMMENDATION #36 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest a trial of noninvasive mechanical ventilation (over invasive mechanical ventilation) in children with sepsis-induced pediatric ARDS (PARDS) without a clear indication for intubation and who are responding to initial resuscitation. | <ul style="list-style-type: none">• Weak• Very Low-Quality of Evidence |

| RECOMMENDATION #37 | STRENGTH & QUALITY OF EVIDENCE |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|
| <p>We suggest using high positive end-expiratory pressure (PEEP) in children with sepsis-induced PARDS. Remarks: The exact level of high PEEP has not been tested or determined in PARDS patients. Some RCTs and observational studies in PARDS have used and advocated for use of the ARDS-network PEEP to Fio2 grid though adverse hemodynamic effects of high PEEP may be more prominent in children with septic shock.</p> | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |
| RECOMMENDATION #38 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We cannot suggest for or against the use of recruitment maneuvers in children with sepsis-induced PARDS and refractory hypoxemia. Remarks: If a recruitment maneuver is considered, the use of a stepwise, incremental and decremental PEEP titration maneuver is preferred over sustained inflation techniques that have not been optimized through direct testing in PARDS patients. All PARDS patients must be carefully monitored for tolerance of the maneuver.</p> | <p>Insufficient</p> |
| RECOMMENDATION #39 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We suggest a trial of prone positioning in children with sepsis and severe PARDS. Remarks: Research trials in adults with ARDS and children with PARDS have emphasized prone positioning for at least 12 hours per day, as tolerated.</p> | <ul style="list-style-type: none"> • Weak • Low-Quality of Evidence |
| RECOMMENDATION #40 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We recommend against the routine use of inhaled nitric oxide (iNO) in all children with sepsis-induced PARDS.</p> | <ul style="list-style-type: none"> • Strong • Low-Quality of Evidence |
| RECOMMENDATION #41 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We suggest using iNO as a rescue therapy in children with sepsis-induced PARDS and refractory hypoxemia after other oxygenation strategies have been optimized.</p> | <ul style="list-style-type: none"> • Weak • Moderate-Quality of Evidence |



RECOMMENDATION #42

STRENGTH & QUALITY OF EVIDENCE

We were **unable to issue a recommendation** to use high-frequency oscillatory ventilation (HFOV) versus conventional ventilation in children with sepsis-induced PARDS.

Insufficient

However, **in our practice**, there is no preference to use or not use HFOV in patients with severe PARDS and refractory hypoxia.

In Our Practice

RECOMMENDATION #43

STRENGTH & QUALITY OF EVIDENCE

We suggest using neuromuscular blockade in children with sepsis and severe PARDS. **Remarks:** The exact duration of neuromuscular blockade to use in severe PARDS patients has not been determined to date. Most of the adult RCT data and pediatric observational data support treatment for 24–48 hours after ARDS onset.

- Weak
- Very Low-Quality of Evidence

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CORTICOSTEROIDS RECOMMENDATIONS TABLE

| RECOMMENDATION #44 | STRENGTH & QUALITY OF EVIDENCE |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------|
| We suggest against using IV hydrocortisone to treat children with septic shock if fluid resuscitation and vasopressor therapy are able to restore hemodynamic stability. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |
| RECOMMENDATION #45 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest that either IV hydrocortisone or no hydrocortisone may be used if adequate fluid resuscitation and vasopressor therapy are not able to restore hemodynamic stability. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |

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ENDOCRINE & METABOLIC RECOMMENDATIONS TABLE

| RECOMMENDATION #46 | STRENGTH & QUALITY OF EVIDENCE |
|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------|
| We recommend against insulin therapy to maintain a blood glucose target at or below 140mg/dL (7.8 mmol/L). | <ul style="list-style-type: none"> • Strong • Moderate-Quality of Evidence |
| RECOMMENDATION #47 | STRENGTH & QUALITY OF EVIDENCE |
| We were unable to issue a recommendation regarding what blood glucose range to target for children with septic shock or other sepsis-associated organ dysfunction. However, in our practice , there was consensus to target blood glucose levels below 180mg/dL (10 mmol/L) but there was not consensus about the lower limit of the target range. | <p>Insufficient</p> <p>In Our Practice</p> |
| RECOMMENDATION #48 | STRENGTH & QUALITY OF EVIDENCE |
| We were unable to issue a recommendation as to whether to target normal blood calcium levels in children with septic shock or sepsis-associated organ dysfunction. However, in our practice , we often target normal calcium levels for children with septic shock requiring vasoactive infusion support. | <p>Insufficient</p> <p>In Our Practice</p> |

RECOMMENDATION #49

We **suggest against** the routine use of levothyroxine in children with septic shock and other sepsis-associated organ dysfunction in a sick euthyroid state.

STRENGTH & QUALITY OF EVIDENCE

- Weak
- Low-Quality of Evidence

RECOMMENDATION #50

We **suggest** either antipyretic therapy or a permissive approach to fever in children with septic shock or other sepsis-associated organ dysfunction.

STRENGTH & QUALITY OF EVIDENCE

- Weak
- Moderate-Quality of Evidence



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NUTRITION RECOMMENDATIONS TABLE

| RECOMMENDATION #51 | STRENGTH & QUALITY OF EVIDENCE |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------|
| <p>We were unable to issue a recommendation regarding early hypocaloric/trophic enteral feeding followed by slow increase to full enteral feeding versus early full enteral feeding in children with septic shock or sepsis-associated organ dysfunction without contraindications to enteral feeding. However, in our practice, there is a preference to commence early enteral nutrition within 48 hours of admission in children with septic shock or sepsis-associated organ dysfunction who have no contraindications to enteral nutrition and to increase enteral nutrition in a stepwise fashion until nutritional goals are met.</p> | Insufficient |
| RECOMMENDATION #52 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We suggest not withholding enteral feeding solely on the basis of vasoactive-inotropic medication administration.</p> <p>Remarks: Enteral feeding is not contraindicated in children with septic shock after adequate hemodynamic resuscitation who no longer require escalating doses of vasoactive agents or in whom weaning of vasoactive agents has started.</p> | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |

RECOMMENDATION #53**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest** enteral nutrition as the preferred method of feeding and that parenteral nutrition may be withheld in the first 7 days of PICU admission in children with septic shock or other sepsis-associated organ dysfunction.

- Weak
- Moderate-Quality of Evidence

RECOMMENDATION #54**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest against** supplementation with specialized lipid emulsions in children with septic shock or other sepsis-associated organ dysfunction.

- Weak
- Very Low-Quality of Evidence

RECOMMENDATION #55**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest against** the routine measurements of gastric residual volumes (GRVs) in children with septic shock or other sepsis-associated organ dysfunction.

- Weak
- Low-Quality of Evidence

RECOMMENDATION #56**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest** administering enteral feeds through a gastric tube, rather than a postpyloric feeding tube, to children with septic shock or other sepsis-associated organ dysfunction who have no contraindications to enteral feeding.

- Weak
- Low-Quality of Evidence

RECOMMENDATION #57**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest against** the routine use of prokinetic agents for the treatment of feeding intolerance in children with septic shock or other sepsis-associated organ dysfunction.

- Weak
- Low-Quality of Evidence

RECOMMENDATION #58**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest against** the use of selenium in children with septic shock or other sepsis-associated organ dysfunction.

- Weak
- Low-Quality of Evidence

| | |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|
| RECOMMENDATION #59 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest against the use of glutamine supplementation in children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none"> • Weak • Low-Quality of Evidence |
| RECOMMENDATION #60 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest against the use of arginine in the treatment of children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |
| RECOMMENDATION #61 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest against using zinc supplementation in children with septic shock and other sepsis-associated organ dysfunction. | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |
| RECOMMENDATION #62 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest against the use of ascorbic acid (vitamin C) in the treatment of children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |
| RECOMMENDATION #63 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest against the use of thiamine to treat children with sepsis-associated organ dysfunction. | <ul style="list-style-type: none"> • Weak • Low-Quality of Evidence |
| RECOMMENDATION #64 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest against the acute repletion of vitamin D deficiency (VDD) for treatment of septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |



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BLOOD PRODUCTS RECOMMENDATIONS TABLE

| RECOMMENDATION #65 | STRENGTH & QUALITY OF EVIDENCE |
|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|
| <p>We suggest against transfusion of RBCs if the blood hemoglobin concentration is greater than or equal to 7 g/dL in hemodynamically stabilized children with septic shock or other sepsis-associated organ dysfunction. Remarks: According to the 2018 Transfusion and Anemia Expertise Initiative (TAXI) guidelines, for the purposes of RBC transfusion, “hemodynamically stabilized” is defined as a MAP higher than 2 sds below normal for age and no increase in vasoactive medications for at least 2 hours.</p> | <ul style="list-style-type: none"> • Weak • Low-Quality of Evidence |
| RECOMMENDATION #66 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We cannot make a recommendation regarding hemoglobin transfusion thresholds for critically ill children with unstable septic shock.</p> | <p>Insufficient</p> |
| RECOMMENDATION #67 | STRENGTH & QUALITY OF EVIDENCE |
| <p>We suggest against prophylactic platelet transfusion based solely on platelet levels in nonbleeding children with septic shock or other sepsis-associated organ dysfunction and thrombocytopenia.</p> | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |

RECOMMENDATION #68

STRENGTH & QUALITY OF EVIDENCE

We **suggest against** prophylactic plasma transfusion in nonbleeding children with septic shock or other sepsis-associated organ dysfunction and coagulation abnormalities.

Remarks: Prophylactic plasma transfusion refers to situations in which there is an abnormality in laboratory coagulation testing but no active bleeding.

- Weak
- Very Low-Quality of Evidence



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PLASMA EXCHANGE, RENAL REPLACEMENT AND EXTRACORPOREAL SUPPORT RECOMMENDATIONS TABLE

| RECOMMENDATION #69 | STRENGTH & QUALITY OF EVIDENCE |
|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|
| We suggest against using plasma exchange (PLEX) in children with septic shock or other sepsis-associated organ dysfunction without thrombocytopenia-associated multiple organ failure (TAMOF). | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |
| RECOMMENDATION #70 | STRENGTH & QUALITY OF EVIDENCE |
| We cannot suggest for or against the use of PLEX in children with septic shock or other sepsis-associated organ dysfunction with TAMOF. | Insufficient |
| RECOMMENDATION #71 | STRENGTH & QUALITY OF EVIDENCE |
| We suggest using renal replacement therapy to prevent or treat fluid overload in children with septic shock or other sepsis-associated organ dysfunction who are unresponsive to fluid restriction and diuretic therapy. | <ul style="list-style-type: none"> • Weak • Very Low-Quality of Evidence |

RECOMMENDATION #72**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest against** high-volume hemofiltration (HVHF) over standard hemofiltration in children with septic shock or other sepsis-associated organ dysfunction who are treated with renal replacement therapy.

- Weak
- Low-Quality of Evidence

RECOMMENDATION #73**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest** using venovenous ECMO in children with sepsis-induced PARDS and refractory hypoxia.

- Weak
- Very Low-Quality of Evidence

RECOMMENDATION #74**STRENGTH &
QUALITY OF EVIDENCE**

We **suggest** using venoarterial ECMO as a rescue therapy in children with septic shock only if refractory to all other treatments.

- Weak
- Very Low-Quality of Evidence

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IMMUNOGLOBULINS AND PROPHYLAXIS RECOMMENDATIONS TABLE

IMMUNOGLOBULINS

| RECOMMENDATION #75 | STRENGTH & QUALITY OF EVIDENCE |
|----------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------|
| We suggest against the routine use of IV immune globulin (IVIG) in children with septic shock or other sepsis-associated organ dysfunction. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |

PROPHYLAXIS

| RECOMMENDATION #76 | STRENGTH & QUALITY OF EVIDENCE |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|
| We suggest against the routine use of stress ulcer prophylaxis in critically ill children with septic shock or other sepsis-associated organ dysfunction, except for high-risk patients. | <ul style="list-style-type: none">• Weak• Very Low-Quality of Evidence |

| RECOMMENDATION #77 | STRENGTH & QUALITY OF EVIDENCE |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------|
| We suggest against routine deep vein thrombosis (DVT) prophylaxis (mechanical or pharmacologic) in critically ill children with septic shock or other sepsis-associated organ dysfunction, but potential benefits may outweigh risks and costs in specific populations. | <ul style="list-style-type: none">• Weak• Low-Quality of Evidence |