



A Guide to the **Nutritional Assessment and Treatment of the Critically Ill Patient**

2nd Ed.

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Foreword

The American Association for Respiratory Care (AARC) is excited to release its second edition of the Guide to the Nutritional Assessment and Treatment of the Critically Ill Patient. The goal of this guide is to assist respiratory therapists at the bedside as they provide a higher level of assessment and management to patients. Since its original introduction in 2014, the content in this guide has positively impacted the critical care community and we are pleased to provide updated information to continue to promote positive patient outcomes.

Proper nutritional assessment and treatment is essential to the successful management of critically ill patients. Unfortunately, these patients are often malnourished - especially those who require mechanical ventilation. Malnutrition can lengthen the time spent in the ICU and extend hospital length of stay. For the mechanically ventilated patient, it can delay or impede the weaning process - which adds another level of associated risks. All disciplines play an important role in managing the nutritional needs of the critically ill patient. All bedside clinicians have an obligation to ensure that critically ill patients are assessed for nutritional adequacy and intervention is taken when required.

This guide reviews the process of nutritional assessment and management of the adult critically ill patient, but also discusses specific patient populations where malnutrition is more prevalent. We hope that you find this guide helpful as you manage your patients' pulmonary and nutritional needs.



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Executive Summary

Introduction

The purpose of this guide is to provide an overview of the important considerations regarding nutritional assessment and treatment that the health care team must address to ensure patients are provided with appropriate nutritional support. The goal of this work is to review a broad list of topics that covers the nutritional support and care process to provide the health care team with a broad understanding of the nutrition assessment and treatment process for the hospitalized critically ill patient.

Overview

Appropriate nutrition is essential for improving outcomes in the health care environment. Hospitalized patients have high rates of malnutrition. Unmet nutritional needs and malnutrition lead to increased morbidity and mortality, decreased quality of life, prolonged duration of mechanical ventilation, and increased length of hospital stay, all of which contribute to the higher cost of health care. Critically ill patients and those patients with respiratory failure require special attention to prevent muscle wasting and to avoid overfeeding and complications associated with nutritional care. A functional nutrition support system should include an interdisciplinary team approach for assessment and treatment, which incorporates an evaluation of nutritional risk, standards for nutritional support, an appropriate assessment and reassessment process, proper implementation, route of support based on patient condition, and a means of measuring nutrient requirements to determine if target goals are being met.

Interdisciplinary Approach

The Society of Critical Care Medicine (SCCM) recognizes the value and importance of a multidisciplinary team approach to nutritional care as a means to improve clinical outcomes. Each discipline in an intensivist led interdisciplinary team, which includes dietitians, nurses, pharmacists, respiratory therapists, speech pathologists, and physical therapists, can contribute to improved outcomes and reduced health care costs.

Nutritional Risk and Assessment

Assessment of nutritional status is performed to identify patients at higher risk for malnutrition related complications. Patients with moderate or severe malnutrition are likely to have longer ICU and hospital length of stay and higher risk of death. After the initial assessment, the primary goals of nutritional support are to maintain lean body mass in at-risk patients and to provide continuous evaluation of the nutrition care plan. Minimized risk of malnutrition can be achieved by prompt initiation of nutritional support, proper targeting of appropriate nutrient quantities, and promotion of motility through the gastrointestinal tract.

A registered dietitian or other trained clinician gathers information to examine the patient's nutrition related history and physical findings, anthropometric physical measurements, biochemical data, and medical tests and procedures, and then screens the patient for other nutrition associated conditions such as malnutrition, obesity, and the risk of refeeding syndrome.

Route of Nutritional Support

Enteral nutrition (EN) is the preferred route of nutritional support. EN should be started within the first 24–48 hours after admission in patients who are incapable of volitional intake. Gastric or small bowel feeding is acceptable in the ICU setting. Enteral feeding tube placement in the small bowel should be done in patients at high risk for aspiration or whose intolerance to gastric feeding is demonstrated. Holding enteral feeding for high gastric residual volumes (GRV) in the absence of clear signs of intolerance and demonstrated risk of aspiration may result in an inappropriate cessation of EN and cause a calorie deficit over time. The definition for high GRV should be determined by individual institutional protocol; but use of GRV up to 500 mL has not been shown to increase the risks of regurgitation, aspiration, or pneumonia in adult patients.

The decision to initiate parenteral nutrition (PN) is influenced by the patient's nutritional risk, clinical diagnosis and condition, gastrointestinal tract function, and duration of anticipated need. PN in a previously healthy patient should be considered when EN is not feasible for the first 7–10 days after hospital admission. Patients with evidence of moderate to severe malnutrition where

EN is not an option should receive PN within the first few days following admission. Supplemental PN may be considered in adult and pediatric patients when nutritional requirements cannot be achieved with EN within the first week

Nutritional Considerations During Critical Illness

The general goals of nutritional care in all patients, including those with respiratory disorders and critical illness, are to provide adequate calories to support metabolic demands, to preserve lean body mass, and to prevent muscle wasting.

Nutritional support during critical illness attenuates the metabolic response to stress, prevents oxidative cellular injury, and modulates the immune system. The stress response to critical illness causes wide fluctuation in metabolic rate. The hypercatabolic phase can last for 7–10 days and is manifested by an increase in oxygen demands, cardiac output, and carbon dioxide production. Caloric needs may be increased by up to 100% during this phase. The goal is to provide ongoing monitoring and support with high protein feedings while avoiding overfeeding and underfeeding. Nutritional modulation of the stress response includes early EN, appropriate macro- and micronutrient delivery, and glycemic control.

Determination of Nutritional Requirements

Nutrient requirements can be calculated by over 200 different equations. Predictive equations use traditional factors for age, sex, height, weight, and additional factors for temperature, body surface area, diagnosis, and ventilation parameters. Additional data such as injury- stress, activity, medications received, and obesity have been added to improve accuracy. Several predictive equations were developed with a focus on specific patient populations and medical conditions.

Predictive equations have varying degrees of accuracy. Error rates can be significant and result in under- and overestimation of caloric needs that impact outcomes. Some equations are unsuitable for use in critically ill patients, while others have been validated with improved accuracy. Due to the extreme metabolic changes that can occur during critical illness, energy needs should be measured using indirect calorimetry (IC) in patients not responding to nutritional support, have complex medical conditions, and are ventilator dependent.

Indirect calorimetry relies on accurate determination of oxygen consumption (VO_2) and carbon dioxide production (VCO_2) using a metabolic analyzer for precise measurements of inspired and expired fractions of oxygen and carbon dioxide. The abbreviated

Weir equation uses the measured VO_2 and VCO_2 to determine resting energy expenditure (REE). The respiratory quotient (RQ), the ratio of VCO_2 to VO_2 , can then be calculated. The RQ was once thought to be a means to determine nutritional substrate use, but this assumption has never been substantiated and use of the RQ measurement is of limited clinical value. Measured values of RQ between the physiologic ranges of 0.67–1.3 should be used as a way to validate test quality. Values of RQ outside of this range invalidate the results due to technical measurement errors and should be repeated.

When a metabolic analyzer is unavailable, IC using other methods should be considered as an alternative to the utilization and dependence on predictive equations.

Clinical Practice Recommendations

Several clinical practice guidelines are available to guide nutritional support. The SCCM and the American Society of Parenteral and Enteral Nutrition (ASPEN), the European Society for Clinical Nutrition and Metabolism (ESPEN), the Academy of Nutrition and Dietetics (AND), the Canadian Clinical Practice Guidelines for Nutritional Support (CCPG), and the European Society of Pediatric and Neonatal Intensive Care (ESPNIC) have developed best practice recommendations based on the interpretation of available evidence, consensus agreement, and expert opinion.

The following present summaries of some of the best-practice recommendations for adult patients from the various organizations:

- Nutritional support should be initiated early, within the first 24–48 hours in critically ill patients.
- Primary goals of nutritional support and care are to: preserve and maintain lean muscle mass; provide continuous assessment, reassessment, and modification to optimize outcome; monitor the patient for tolerance and complications such as refeeding syndrome; prevent protein energy malnutrition by giving higher protein content while providing adequate total calories; monitor nutrition goals and target achievement rate of > 65% within the first week; and prevent accumulation of a caloric deficit.
- Indirect calorimetry should be used when available or when predictive equations are known to be inaccurate.
- Current EN practice recommendations are to: preferentially feed via the enteral route; initiate EN within 24–48 hours; reduce interruptions of EN for

nursing care and bedside procedures to prevent underfeeding; maintain head of bed (HOB) elevation to reduce aspiration risk; avoid routine monitoring of GRV or accept GRV up to 500 mL before reducing or stopping EN in the absence of clear signs of intolerance; use motility agents to improve tolerance; and promote post-pyloric feeding tube placement when feasible.

- Current PN practice recommendations are to: only use exclusive PN when enteral route is not feasible; use PN based on the patient's nutritional risk classification for malnutrition; delay PN up to seven days in low nutritional risk patients; initiate PN early in high nutritional risk patients; consider use of supplemental PN when indicated; convert to EN as soon as tolerated to reduce the risks associated with PN.
- Use of trophic or "trickle feeding" and permissive underfeeding may be beneficial.
- Use of pharmaconutrients and immunonutrition: utilize high omega-3 fatty acid to omega-6 fatty acid ratios. The use of arginine, glutamine, nucleotides, antioxidants, and probiotics may be beneficial in specific patients. The use of arginine should be avoided in patients with severe sepsis.

In pediatric patients:

- EN is the preferred route of nutrition.
- Initiate EN within 24-48 hours of admission to the PICU
- A stepwise algorithmic approach to advancing EN should be utilized.
- Goal is to achieve at least 2/3 of nutrient goal within the first week of admission.
- IC should be utilized when available to determine energy requirements.
- Routine measurement of GRV is not recommended
- Either gastric or post-pyloric feeding can be utilized in the majority of children.
- Measurement of anthropometrics should be obtained on admission and regularly throughout the hospital course.
- Immunonutrition is not recommended for the critically ill child.
- Timing for PN initiation should be individualized with the below general guidance:
 - In cases of severe malnutrition, supplemental PN may be provided in the first week of admission if unable to advance beyond low volumes of EN.

- When EN is unable to be initiated in the first week of admission, PN may be provided.

Appropriate nutritional support in hospitalized patients and the prevention of malnutrition can improve outcomes and reduce health care costs. The nutritional care plan should utilize the team approach and be supported by organizational standards with policies and procedures that are based on the best available evidence. The health care team's proper implementation, continuous assessment, and monitoring of the nutrition care plan are key elements for success.

Nutritional Overview

The Importance of Appropriate Nutrition

Appropriate nutrition is essential for health and healing. In hospitalized patients, malnutrition is a common and serious problem affecting both adult and pediatric populations.¹⁻⁴ Critically ill patients are at high risk for malnutrition related complications. The resulting detrimental effects of malnutrition include increased morbidity and mortality, decreased functional quality of life, prolonged duration of mechanical ventilation, and increased length of hospital stay, all which contribute to higher health care cost.⁵

Critical illness associated with respiratory failure requires special attention to prevent catabolic or destructive metabolism.⁶ Nutritional therapy in this setting requires maintenance of adequate calorie and protein intake to prevent muscle wasting and avoid overfeeding and complications associated with nutritional care.⁷ Malnutrition is a risk factor for the onset of respiratory failure and can worsen further after respiratory failure is established. Nutritional support can affect respiratory muscle strength, endurance and function, carbon dioxide production, and immune system response. To ensure successful support and recovery from respiratory failure, the nutritional care plan must also consider other important aspects, such as fluid and electrolyte balance, micronutrient requirements, and acid-base status. Recovery from respiratory failure requires a regimented nutritional support process that includes a comprehensive assessment of risk, proper implementation, ongoing reassessment of caloric requirements, tolerance of treatment monitoring, and avoiding the development of complications.⁶

Importance of Interdisciplinary Collaboration

The role of health care team members in providing expertise regarding nutritional support has evolved around interdisciplinary collaboration. Registered dietitians and physicians complete specialized training programs to attain the Certified Nutrition Support Clinician (CNSC) credential and are increasingly involved in nutrition support organizations such as the American Society of Parenteral and Enteral Nutrition (ASPEN).⁸

Respiratory therapists have traditionally maintained the responsibility and technical expertise in performing metabolic

measurements by indirect calorimetry assessments, especially in the mechanically ventilated critically ill patient. Clinical practice guidelines developed by the American Association for Respiratory Care (AARC) maintain an evidence-based framework for nutritional assessments using indirect calorimetry for patients receiving mechanical ventilation.⁹

Speech pathologists aid in the assessment of post-extubation dysphagia. Detection of swallowing dysfunction that is common after prolonged mechanical ventilation can help prevent the detrimental impact and risks associated with aspiration and poor nutrition among patients with or without neurologic dysfunction.^{10,11} Post-extubation dysphagia is associated with longer hospitalization in survivors of critical illness with neurologic impairment.

Critical care organizations such as the Society of Critical Care Medicine (SCCM) recognize the importance of an intensivist led multidisciplinary team consisting of nurses, dietitians, pharmacists, respiratory therapists, and physical therapists.¹² Each discipline provides expertise pertinent to nutritional support and care, contributes to improved outcomes, and reduces costs.

The future and ongoing challenge to the evolution of health care is to facilitate the team approach toward best practices and therapeutic efficacy. Appropriate nutritional assessment and treatment protocols require devoted resources toward diagnosis, intervention, and monitoring. The integrated health care delivery team trained in nutritional assessment and treatment will be better equipped to optimize and ensure health care resources are maximized.¹³

Importance of Adequate Nutritional Assessment and Treatment

Nutritional deficits related to chronic disease and acute illnesses are frequently found in patients admitted to the ICU. Many patients who cannot resume oral food ingestion within the first few days of admission are prone to losing body mass due to poor nutrient intake and are at risk for developing an acute and prolonged inflammatory process. Patients in the ICU for more than 48 hours need nutritional assessment and support maintained constantly throughout their period of critical illness

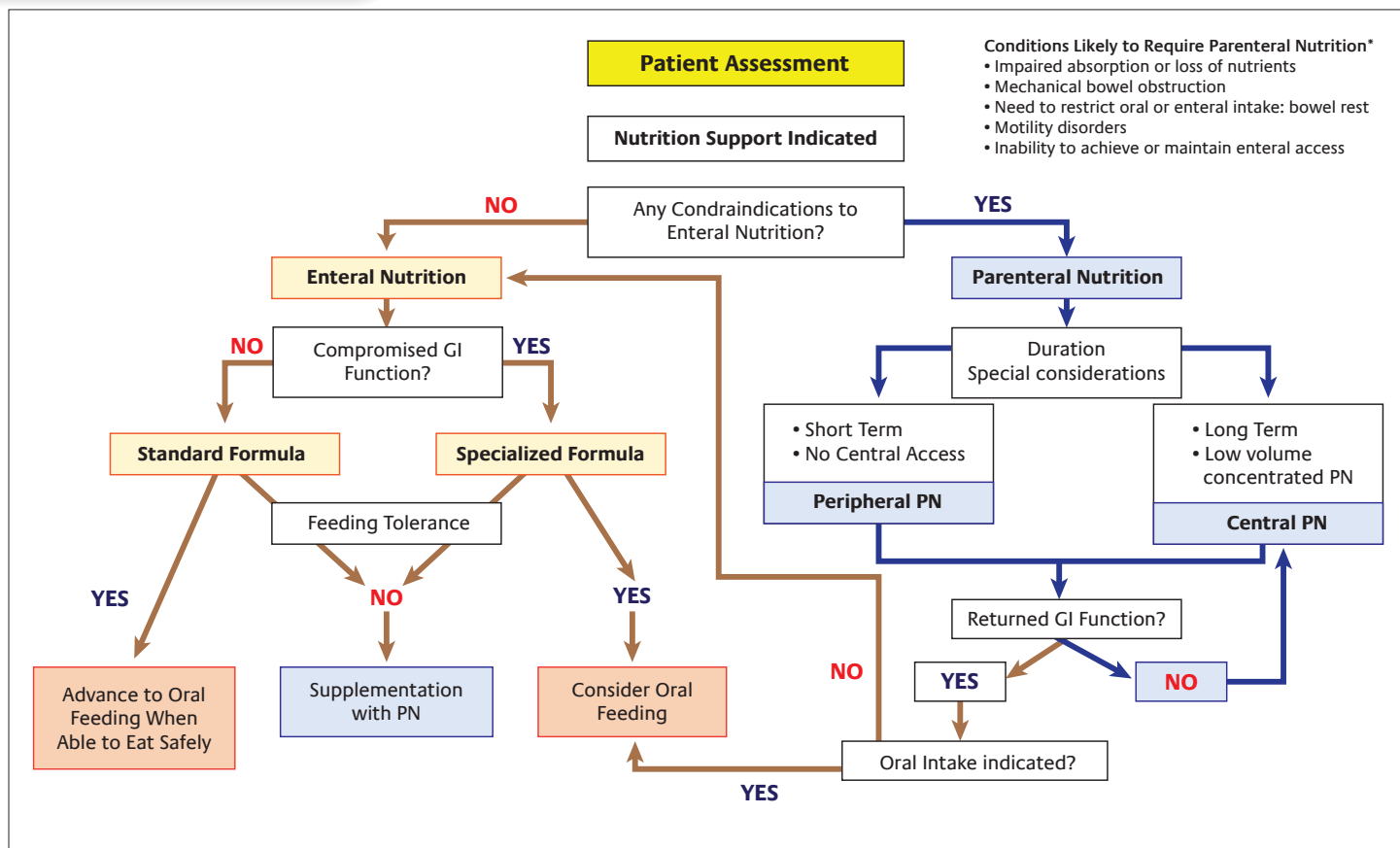


Figure 1. Example of a Nutrition Management Protocol¹⁵

and hospitalization. Many critically ill patients experience severe gastrointestinal motility disorders and can experience dysphagia following extubation, which may increase the risks for aspiration. Complications associated with critical illness can have serious consequences that can be diminished with early recognition and intervention. The promotion of effective nutrition can only be achieved with a standardized nutritional support protocol that incorporates regular assessments of gastrointestinal function and tolerance of parenteral and enteral feeding.¹⁴

In critically ill patients unable to take nutrition by mouth, enteral nutrition (EN) through the gastrointestinal tract is the preferred route. Parenteral nutrition (PN) by intravenous access is another alternative. Use of an evidence-based nutritional management protocol increases the likelihood that patients receive nutrition via the enteral route (Figure 1).

A standardized approach targeting gastric or post-pyloric feeding tube placement when indicated, gastric decompression,

and use of bowel motility agents can shorten the duration of mechanical ventilation and reduce the risk of death. Clinical outcome benefits from improving the rate of EN can be significant when adjusted for nutritional risk of moderate-to-severe malnutrition at baseline.¹⁶

Development and maintenance of a best-practice nutritional support program reduces costs and improves outcomes. Maintenance of nutritional support requires continuous monitoring of the appropriate route of administration and the adequacy of usage in order to minimize costs and reduce waste.¹⁷ Insufficient calorie intake is associated with an increase in mortality risk. The reasons for failure to achieve recommendations for best clinical practice include lack of sufficient nutritional support services to monitor adherence, inadequate training in nutritional support, and restricted use of nutrient formulations that show improved outcomes secondary to their higher cost, or disagreement about the supporting evidence.¹⁷

Nutritional Assessment

Nutritional Risk Assessment

The nutritional assessment process should begin at admission with a nutrition screening followed by a formal assessment when indicated.¹⁸ This process can be automatically triggered by electronic medical record (EMR) systems. Several nutrition screening and assessment tools are available to evaluate the risks of malnutrition in hospitalized patients.¹⁹ Parameters that are evaluated include BMI and weight loss, history of nutritional intake, severity of illness, and anthropometric measurements (Figure 2). Recent guidelines based on expert consensus, suggest the use of the Nutrition Risk Screening tool (NRS 2002) and the Nutrition Risk in the Critically Ill assessment tool (NUTRIC score) in all patients admitted to the ICU when insufficient volitional intake is anticipated.¹² Patients with a high risk of malnutrition are more likely to benefit from early initiation of EN.

Nutrition risk assessment should encompass two necessary elements. The initial assessment establishes the presence or

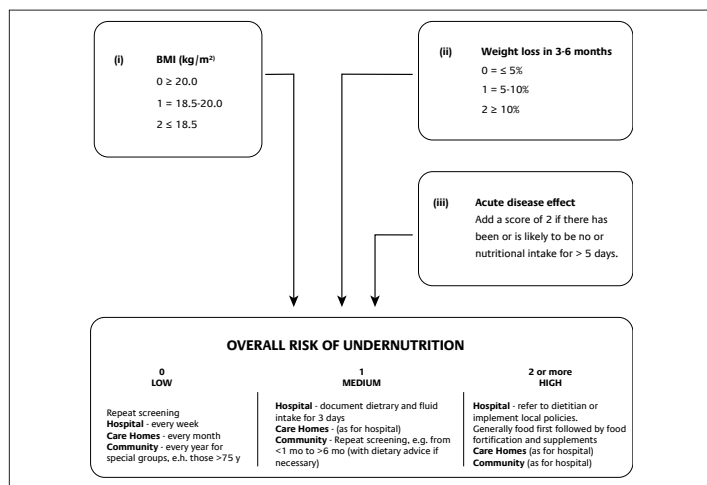


Figure 2. Malnutrition Universal Screening Tool (MUST) for Adults²⁰

estimate of lean body mass loss prior to ICU or hospital admission. The goal of preventing further loss of lean body mass can be achieved when acute illness is promptly controlled and with the formation of an adequate nutritional support process.

Additionally, the safe provision of nutritional support requires a continuous evaluation of the risks of nutritional care. Minimized risk can be achieved by prompt initiation of nutrition, targeting the appropriate nutrient quantities, promoting motility through the gastrointestinal tract, and averting serious life-threatening complications such as refeeding syndrome. Patients found to be at higher risk for nutrition-related problems should receive specialized nutritional support. Development of nutritional assessment and care protocols designed for the specific needs of critically ill patients are required to minimize the reduction of lean body mass until discharge. Nutritional care from admission to hospital discharge is essential to reducing risk of nutrition related complications and promoting recovery (Figure 3).

Standards for Nutritional Support

Nutritional support standards for acute care have been

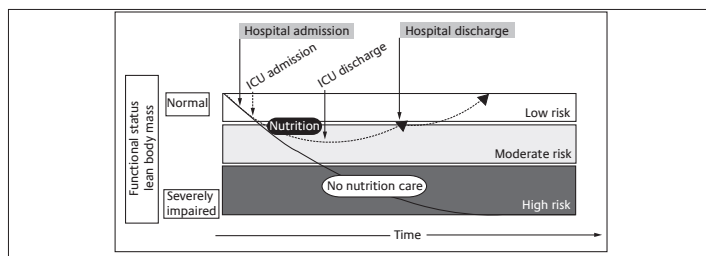


Figure 3. Time Course of Increasing Nutritional Risk Assessment and Support in Critical Illness¹⁴

developed to guide the nutrition support process. These standards are designed to optimize the development and performance of a competent nutritional care plan (Figure 4). Components of a nutritional support program should include the following.¹⁸

Organization

A nutritional support service or interdisciplinary team approach with established policies, procedures, and a performance improvement process should be initiated for each admitted patient.

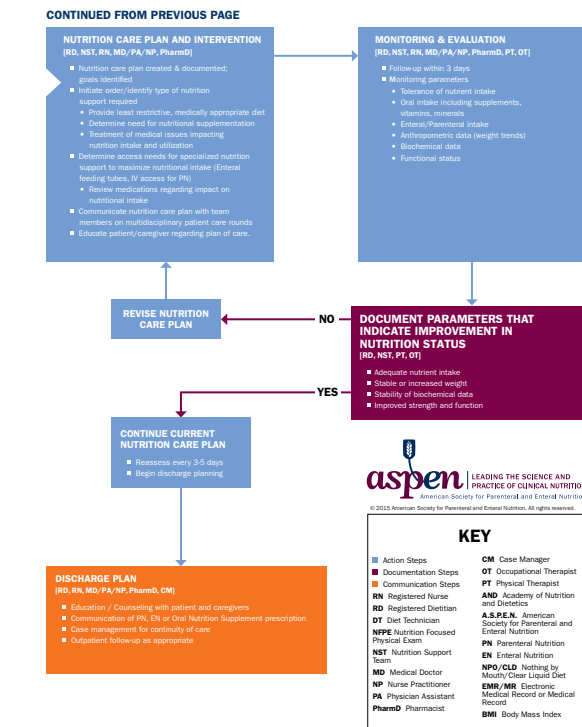
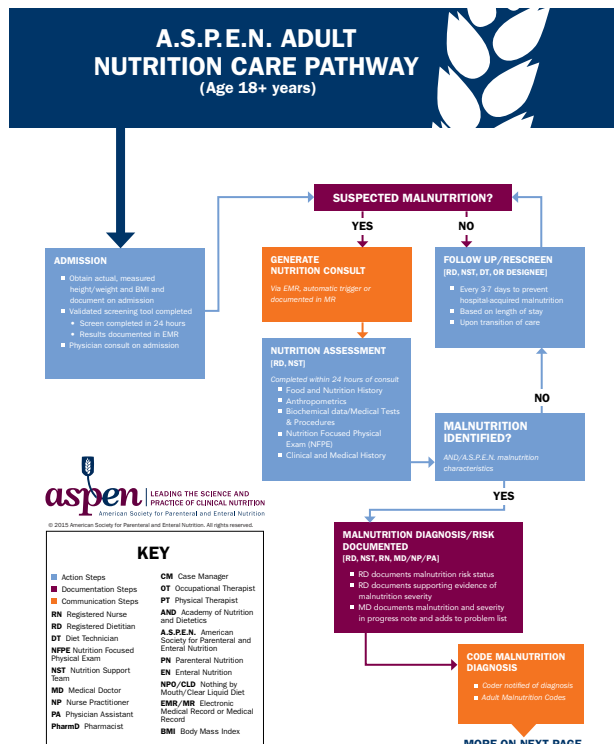


Figure 4. A.S.P.E.N. Adult Nutrition Care Pathway¹⁸

Nutritional Care Process

The process for nutritional care should identify at-risk patients using a screening process that is formalized and documented. Regulatory agencies such as The Joint Commission (PC.01.02.01 – EP 4) require that a nutritional screening be completed when the patient's condition warrants within the first 24 hours after admission. Identified nutritionally at-risk patients should undergo a formal nutritional assessment that includes subjective and objective criteria, classification of nutritional risk, requirements for treatment, and an assessment of appropriate route of nutrition intake.

Development of a Nutritional Care Plan

The nutritional care plan should include clear objectives, use a multidisciplinary approach, have defined goals, select the most appropriate route, select the least costly substrate formulation for the patient's disease process, and include a process for reassessment of adequacy and appropriateness.

Implementation Process

The ordering process for the nutritional care plan should be documented before administration occurs. The appropriate

nutritional access device should be inserted by a qualified health care professional using standardized procedures with appropriate placement confirmed and placement and/or adverse events documented. Enteral and parenteral formulations should be prepared accurately and safely using established policies and procedures. Parenteral formulation should be prepared in a sterile environment using aseptic techniques. Additives to formulations should be checked for incompatibilities and prepared under direct supervision of a pharmacist. All nutritional formulations should be labeled appropriately and administered as prescribed while monitoring patient tolerance. Protocols and procedures should be used to reduce and prevent the risks of regurgitation, aspiration and infection, and a process for Sentinel Event review should be established.

Monitoring and Re-evaluating the Nutritional Care Plan

Establish the frequency and parameters for monitoring the nutritional care plan based on the patient's degree of nutritional risk. Standard procedures for monitoring and re-evaluation should be established to determine whether progress toward short and long-term goals are met, or if realignment of goals is necessary.

Transition of Therapy Process

Assess achievement of targeted nutrient intake to ensure that estimated requirements are being met before nutritional support is transitioned between parenteral, enteral, and oral intake. Transitions should be based on clinical judgment and assessed and documented before nutrition support therapy is discontinued. Maintain continuity of care when transitioning between levels of care or changes in the care environment. Termination of nutritional support should follow protocols that take into account ethical and legal standards and the patient's advance directives.

Nutritional Assessment

The nutritional assessment process includes the collection of data to determine the nutritional status of an individual. A registered dietitian or physician trained in clinical nutrition gathers data to compare various social, pharmaceutical, environmental, physical, and medical factors to evaluate nutrient needs. The purpose of nutrition assessment is to obtain, verify, and interpret data needed to identify nutrition-related problems, their causes, and significance. This data is then used to ensure adequate nutrition is provided for the recovery of health and well-being.²¹

Food/Nutrition-related History

Past dietary behaviors can be identified in the nutritional assessment to determine the individual's pattern of food consumption. Assessment of dietary history should include:

- Appetite
- Weight history (loss, gain)
- Growth curves (pediatrics)
- Taste changes
- Nausea/vomiting
- Bowel pattern (constipation, diarrhea)
- Chewing, swallowing ability
- Substance abuse
- Usual meal pattern
- Diet restrictions
- Food allergies or intolerances
- Medications, herbal supplements
- Meal preparation, ability to buy/obtain food
- Activity level
- Knowledge/beliefs/attitudes
- Nutrient intake

The registered dietitian may use a 24-hour recall or a usual daily intake recall, a food diary or food record, or a food frequency questionnaire. The 24-hour recall or food frequency questionnaire employ retrospective data that can be easily used in a clinical setting. The 24-hour recall is a commonly used technique incorporated into the patient interview in which the individual states the foods and the amount of each food consumed in the previous 24 hours. Accuracy of the recall is dependent on the patient's memory, the perception of serving size, and the skill of the interviewer to elicit complete information. The 24-hour recall may underestimate usual energy intake. Food frequency questionnaires (FFQ) collect information on both the frequency and amount consumed of specific foods.²² The FFQ can help to identify eating patterns; however, intake of nutrients may be overestimated. In food diaries or food records, dietary intake is assessed by prospective information and contains dietary intake for three to seven days. These methods provide the most accurate data of actual intake but are very labor intensive and time consuming to analyze. Therefore, they are typically used in the research or outpatient setting.

Anthropometric Measurements

Anthropometrics refers to the physical measurements of the body. The measurements are used to assess the body habitus of an individual and include specific dimensions such as height, weight, and body composition (i.e., skin-fold thickness, body circumference including points at the waist, hips, chest, and arms).²³

Height and weight

Height and weight can be assessed by asking the patient or caregiver, or by taking a direct measurement. When recording data, note the date and whether the height and weight were stated or measured. Once these two measurements are obtained, a more useful number (the body mass index [BMI] or weight-for-length ratio in kg/cm for children under 2 years) can be calculated. BMI is defined by weight and height measurements where:

Using pounds and inches:

$$\text{BMI} = \text{Weight in pounds} / (\text{Height in inches})^2 \times 703$$

Using kilograms and meters:

$$\text{BMI} = \text{Weight in kilograms} / (\text{Height in meters})^2$$

BMI can have a strong correlation between body fat and risk of disease. This number is a useful tool for determining the BMI category: underweight, healthy weight, overweight, obese, or morbidly obese.

Table 1. BMI Classifications for Adults²⁴

BMI (kg / m ²)	Classification	Risk of Comorbidities
<16.0	Severe underweight	Severe
16.0-16.9	Moderate underweight	Moderate
17.0-18.5	Mild underweight	Average
<18.5	Underweight	Low, but risk of other clinical problems increases
18.5-24.9	Normal weight	Average
25.0-29.9	Overweight (pre-obese)	Increased
30.0-34.9	Obese Class I	Moderate
35.0-39.9	Obese Class II	Severe
≥ 40.0	Obese Class III	Very Severe

BMI categories

A healthy weight may be confirmed by a BMI of between 18.5 and 24.9 for adults or a BMI-for-age between the 5th and 85th percentiles for children. A BMI of 25.0 to 29.9 indicates excessive weight in the adult. BMI-for-age in children suggestive of excessive weight is between the 85th and 95th percentiles. Obesity is defined as a BMI greater than 30 in the adult and greater than the 95th percentile in boys and girls aged 2 to 20 years. Adults who are categorized as underweight have a BMI of less than 18.5, while underweight children score in the bottom 5th percentile for BMI-for-age (Table 1).

Body composition

Body composition measures body fat, muscle mass, and bone density. Body weight variations in individuals of similar height differ in the proportion of lean body mass, fat mass, and skeletal size. Several common measurements, which include skin-fold thickness, mid-upper arm circumference measurements, and more high tech measurements like bioelectrical impedance

analysis (BIA) or dual-energy X-ray absorptiometry (DXA) scans, can be used to determine body fat to body mass, intracellular water to extracellular water ratios, and bone density.

Skinfolds

Skinfold thickness measures subcutaneous fat with the assumption that it comprises 50% of total body fat. Usually, the triceps and subscapular skinfolds are the most useful for evaluation.²⁵ Skinfold thickness measurements are limited by reliability due to proper equipment and technique of the examiner and have limited practical application in the acute care setting.

Arm muscle area

The triceps skinfold (TSF) measurement, along with mid-upper arm circumference (MAC), is used to calculate the arm muscle area (AMA). The MAC is measured halfway between the acromion process of the scapula and the tip of the elbow. The results indicate muscle stores available for protein synthesis or energy

needs. Changes over time in AMA will show whether the patient has been deprived of protein or calories. AMA is one of the markers of nutritional status and can be a predictor of mortality.²⁶

Waist circumference

An alternative to BMI, waist circumference can be a more accurate predictor of excess body fat and risks associated with obesity.²⁷ The measurement of waist circumference has been correlated with visceral fat;²⁸ and the distribution of body fat, specifically as visceral fat, which is deposited in the abdominal region, is correlated with obesity related health risks (Figure 5).

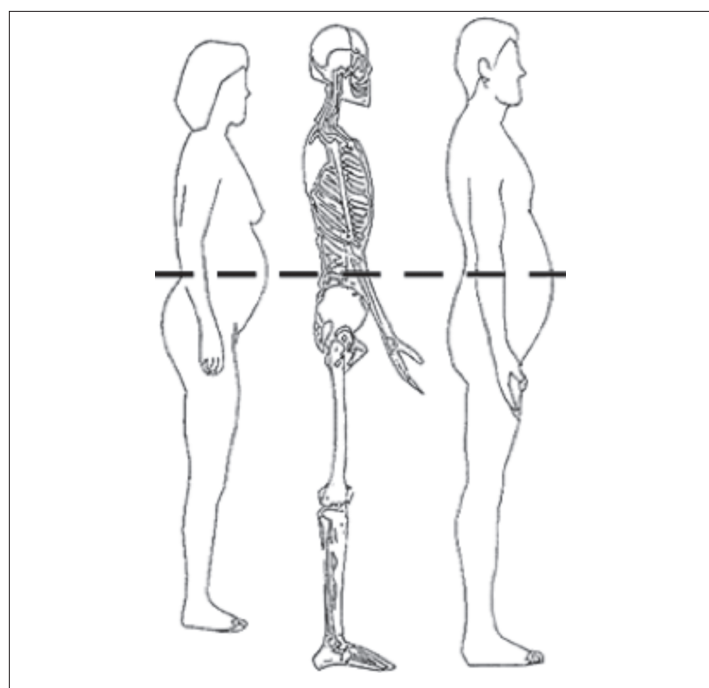


Figure 5. Measurement of Waist Circumference²⁹
Photo courtesy of the National Heart, Lung, and Blood Institute

According to the U.S. Department of Health and Human Services (HHS), the following individuals are at increased risk for developing chronic diseases:

- Women with a waist circumference of more than 35 inches
- Men with a waist circumference of more than 40 inches

However, the World Health Organization, due to recent research findings, has recommended lower thresholds for waist circumference for Asian populations.²⁹ Therefore, those at

increased risk for developing chronic disease include:

- Asian women with a waist circumference of more than 31 inches
- Asian men with a waist circumference of more than 35 inches

Other body assessment tools

More accurate measurements of body composition include the more advanced techniques of bioelectrical impedance analysis (BIA), low-density X-rays (DXA), computed tomography (CT) scan, and magnetic resonance imaging (MRI). These methods are very accurate and noninvasive; however, they are not necessarily ideal in the clinical setting, are expensive, and time consuming.

Biochemical Data

Laboratory values of particular significance used in assessing nutritional status include serum proteins and lymphocytes. An individual's protein stores may indicate the degree of nutritional risk. Protein-energy malnutrition (PEM) may be reflected in low values for albumin, transferrin, transthyretin (prealbumin), retinol-binding protein, and total lymphocyte count. Blood levels of these markers indicate the level of protein synthesis and thus yield information on overall nutritional status. However, inadequate intake may not be the cause of low protein values. Certain disease states, hydration level, liver and renal function, pregnancy, infection, and medical therapies may alter laboratory values of circulating proteins.³⁰ It is important to note that a nutritional disorder diagnosis cannot be made from one single laboratory value but should be utilized

Table 2. Common Biomarkers of Nutritional Status and Inflammation³¹

Biomarker	Normal range
Albumin	3.5-5g/dL
Transferrin	200-400 mg/dL
Prealbumin (Transthyretin)	18-50 mg/dL
Retinol-binding protein	3.0-8.0 mg/dL
C-reactive protein	0-1.0 mg/dL

with other assessment data to determine the nutritional status of the patient. The majority of laboratory values used in nutritional assessments lack sensitivity and specificity for malnutrition (Table 2).

Albumin

Comprising the majority of protein in plasma, albumin is commonly measured. The half-life of albumin is 14–20 days, which reduces its usefulness for monitoring the effectiveness of nutrition in the acute care setting. However, the general availability and stability of albumin levels from day to day make it one of the most common tests for assessing long-term trends and provides the clinician with a general idea of baseline nutritional status prior to a procedure, insult, or acute illness. Albumin levels often reflect the metabolic response and severity of disease, injury, or infection and can be a useful prognostic indicator. Albumin synthesis is affected by nutrition and also by inflammation. During an inflammatory state, the production of albumin diminishes. The effect of inflammation and hypoalbuminemia has been linked with increased morbidity, mortality, and longer hospitalization.³²

Transferrin

The transport protein for iron (transferrin) has a half-life of 8–10 days and, therefore, can be a better indicator of improved nutritional status than albumin. However, lack of iron influences its values along with a number of other factors, including hepatic and renal disease, inflammation, and congestive heart failure.³³

Transthyretin and retinol-binding protein

Transthyretin, also called prealbumin, and retinol-binding protein have a half-life of just 2–3 days and 12 hours, respectively. Each of these responds to nutritional changes much quicker than either albumin or transferrin. However, a number of metabolic conditions, diseases, therapies, and infectious states influence their values.³³ Levels of transthyretin and retinol-binding protein are influenced by many factors other than nutritional status. Similar to albumin, their use is limited in the setting of stress and inflammation. Because these conditions are so common among the critically ill, visceral protein markers are of limited usefulness for assessing nutritional deficiency but are of greater importance in assessing the severity of illness and the risk for future malnutrition.

Total lymphocyte count

The immune system may be compromised by a lack of protein. Two laboratory values, white blood cells and percentage of lymphocytes, have been used as measures of a compromised immune system. However, many non-nutritional variables influence lymphocyte count; therefore, their usefulness in assessing nutritional status is limited.³⁴

Biomarkers of inflammation

Biomarkers of inflammation are important values to measure along with serum proteins. The presence of inflammation affects the nutritional status of the patient. The inflammatory response increases the catabolic rate and causes albumin to leak out of the vascular compartment. Inflammation triggers a chemical cascade that causes a loss of appetite or anorexia, therefore decreasing dietary protein intake and further catabolism.³¹

One of the most common biomarkers of inflammation used in clinical practice is C-reactive protein (CRP). The production of CRP increases with infection and inflammation along with pro-inflammatory cytokines (i.e., IL-1a, IL-1b, IL-6, TNF) while the production of albumin and prealbumin decreases.³⁵ Other biomarkers of inflammation include prolactin, cholesterol, hyperglycemia, and ferritin.^{36–38}

Other Tests and Procedures

Creatinine-height index

Because the rate of creatinine formation in skeletal muscle is constant, the amount of creatinine excreted in the urine every 24 hours reflects skeletal muscle mass and can indicate muscle depletion. However, it requires an accurate urine collection and normal renal function. Other factors that influence creatinine excretion that can complicate interpretation of this index include age, diet, exercise, stress, trauma, fever, and sepsis.³³

Nitrogen balance (protein catabolism)

Nitrogen balance reflects skeletal muscle, visceral or organ, blood cell, and serum protein stores. Because nitrogen is a major byproduct of protein catabolism, its rate of urinary excretion can be used to assess protein adequacy. The amount of nitrogen excreted in the urine is typically measured as the 24-hour urinary urea nitrogen (UUN). If there is a positive urinary nitrogen balance, protein metabolizing is sufficient, and nitrogen is excreted in the urine. A UUN value less than zero indicates a negative nitrogen

balance, which indicates that the patient needs a higher protein intake. Theoretically, by increasing exogenous protein, loss of endogenous protein is reduced. However, because of invalid 24-hour urine collections, alterations in renal or liver function, large immeasurable insensible losses of protein from burns, high-output fistulas, wounds, ostomies, and inflammatory conditions, nitrogen balance calculations are generally negative and do not accurately reflect nutrition status.³³

Pulmonary function

Pulmonary function test results may change with malnutrition. Weakness of the diaphragm and other muscles of inspiration can lead to a reduced vital capacity and peak inspiratory pressures. The strength and endurance of respiratory muscles are affected, particularly the diaphragm. Respiratory muscle weakness can affect the ability to cough and clear secretions, which may impact rates of pulmonary complications. Dietary antioxidants are thought to protect tissue from oxidant injury or stress, due to their ability to stabilize reactive molecules. Oxidative stress contributes to airflow limitation; therefore, antioxidant vitamins provide pulmonary antioxidant defense.³⁹

Nutrition-focused physical findings

The nutritional-focused physical assessment is the evaluation of body systems, oral health, suck/swallow/breathing ability, and appetite, conducted by the Registered Dietitian or another member of the health care team as part of the nutritional assessment (IDNT 2009). Physical examination can reveal observable signs of nutrition deficiencies where high cell turnover occurs, like the hair, skin, mouth, and tongue.²³ Signs of weight loss, including loss of lean body mass and subcutaneous fat, should be investigated. Special attention should be given to fluid retention as this can mask weight loss.¹³ Other physical findings such as skeletal muscle depletion can be clinical indicators of inflammation or signs of systemic inflammatory response.

Patient History

Interviewing the patient or the caregiver to determine past and current eating practices can be helpful. The patient's medical record can also reveal additional information regarding social, pharmaceutical, environmental, and medical issues. Much of this data can give insight into a patient's nutritional status. The patient's social history indicates marital status, employment, education, and economic status. Drug-nutrient interactions may be identified from the prescribed medications that lead to

potential nutrient deficiencies. Environmental issues could shed light on the difficulties the patient has in procuring, storing, and/or preparing food. The patient's educational background could determine the potential for understanding and applying nutrition counseling. The economic status of the patient may drive certain food choices. Much of the information gained during the interview can be helpful to raise suspicion and guide the investigation further into revealing the nutritional status of the patient.

Malnutrition

Malnutrition is a serious and prevalent problem in adult and pediatric patients. In a recent study, severe malnutrition was diagnosed in 13% of critically ill adults.³ In various hospital settings, prevalence rates of malnourished or at-risk adults between 20% to 78% have been reported.⁴⁰⁻⁴³ Malnutrition rates of 15 to 53% have been reported in critically ill children.⁴⁴

Malnutrition can be characterized by deficient, excessive, or unbalanced nutrient intake. Malnutrition syndromes can be associated with acute or chronic inflammation. Etiology-based diagnosis of malnutrition falls into three categories: starvation-related malnutrition, when there is chronic starvation without inflammation (e.g., secondary to anorexia nervosa); chronic disease-related malnutrition, when inflammation is chronic and of mild-to-moderate degree (e.g., organ failure, pancreatic cancer, rheumatoid arthritis, or sarcopenic obesity); and acute disease or injury related malnutrition, when inflammation is acute and severe (e.g., major infection, burns, trauma, or closed-head injury)⁴⁵(Figure 6).

Standardized diagnostic characteristics for malnutrition have been created by the Academy of Nutrition and Dietetics (AND) and American Society for Parenteral and Enteral Nutrition (ASPEN).¹³ Identification of two or more of the following is recommended for a diagnosis of malnutrition based on degree of severity.

- Insufficient nutrient intake < 50–75% of estimated energy requirements over a defined time period
- Loss of weight (nonvolitional)
- Loss of muscle mass
- Loss of subcutaneous fat
- Localized or generalized fluid accumulation that may mask weight loss or loss of lean body mass
- Diminished functional status measured by hand- grip strength.³

Other criteria that can be used to assess the severity of malnutrition include low body mass index (BMI), underlying disease severity, and presence of acute or chronic inflammation.⁴⁶

Malnutrition in adults is a major contributor to increased morbidity and mortality, decreased functional quality of life, prolonged duration of mechanical ventilation, increased hospital length of stay, and higher health care costs.^{3,40}

Additionally, malnourished pediatric patients have been shown to have a higher risk of hospital acquired infections due to impaired immune function, delayed wound healing, and impaired GI function. Malnutrition has also been associated with poor or delayed growth, reduced or delayed mental and psychomotor development, childhood behavioral problems, and a suggested risk of developing adult onset conditions such as cardiovascular disease, hypertension, and type II diabetes.⁴

Undernutrition and Protein Energy Malnutrition

Undernutrition is a nutritional deficiency resulting from the lack of nutrient intake. Undernutrition suppresses immune function and is often a precursor of disease progression and/or worsening infection.⁴⁷ During critical illness, proteolysis (muscle protein breakdown) increases, which can cause dietary protein needs to more than double. Failure to meet this increased protein requirement can lead to a state of protein energy malnutrition, which can be characterized by weight loss and muscle wasting.⁴⁸

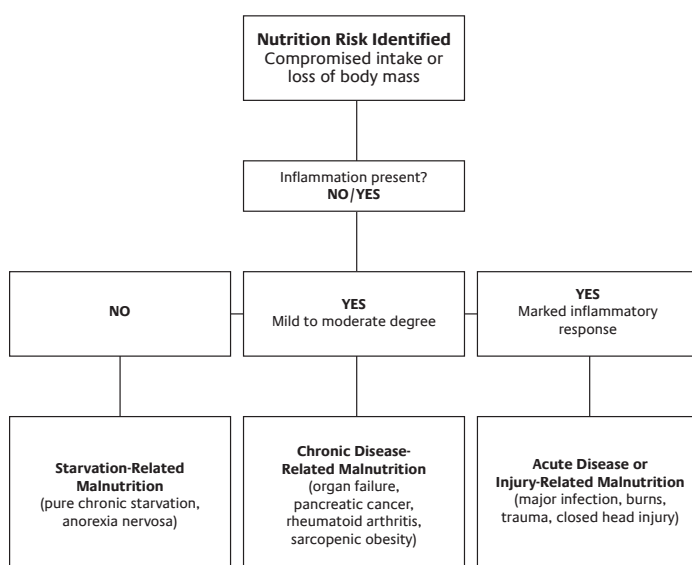


Figure 6. Etiology-Based Malnutrition¹³

Overnutrition, Obesity, and Metabolic Syndrome

Overnutrition in the obese patient can lead to fluid overload, hyperglycemia, fatty liver deposits and liver dysfunction, and the need for prolonged ventilator support.⁴⁹ Obese individuals have a higher incidence of inflammation associated chronic diseases, greater susceptibility to infection^{47,50} and have an increased risk of mortality.^{51,52} Obesity-induced inflammation is an important contributor to the development of insulin resistance and hyperglycemia.⁵³ Obesity increases the risk and prevalence of asthma in both adults and children.⁵⁴ Sarcopenic obesity is obesity associated with a decline in muscle strength and mass in elderly patients, which may further reduce physical activity and result in additional weight gain.⁵⁵ The additional weight loading of the chest wall increases the work of breathing, reduces lung volume, decreases functional residual capacity, and can result in atelectasis, hypoxemia, and hypercapnia. Obese patients have a high prevalence of obstructive sleep apnea and are prone to developing obesity hypoventilation syndrome.^{56,57}

The metabolic syndrome consists of a grouping of risk factors that have shown to be strongly associated with an increased risk for cardiovascular disease and the development of type-2 diabetes mellitus. Metabolic risk factors for metabolic syndrome consist of hyperlipidemia, hypertension, hyperglycemia, a proinflammatory state, and a prothrombotic state. The predominant underlying risk factors include abdominal obesity and insulin resistance.⁵⁸ Obesity hypoventilation syndrome, obstructive sleep apnea, and congestive heart failure are associated with the development of metabolic syndrome.⁵⁹⁻⁶⁴

Refeeding Syndrome

Refeeding Syndrome is a term used to describe the complex metabolic and clinical disturbances that occur after the reinstitution of nutrition to patients who are severely malnourished or starved.^{65,66} Clinical manifestations of refeeding syndrome are related to the resulting electrolyte and vitamin deficiencies caused by starvation and malnutrition, and the subsequent abnormalities

Table 3. Clinical Manifestations of Refeeding Syndrome⁶⁶⁻⁶⁸

Hypophosphatemia	Hypokalemia	Hypomagnesemia	Vitamin/Thiamine Deficiency	Sodium Retention
Impaired oxygen, transport and delivery, hypoxia Impaired cardiac function Impaired diaphragm contractility Respiratory failure Paresthesias Weakness Lethargy Somnolence Confusion Disorientation Restlessness Encephalopathy Areflexic paralysis Seizures Coma Death	Nausea Vomiting Constipation Weakness Paralysis Respiratory compromise Rhabdomyolysis Muscle necrosis Alterations in myocardial contraction Electrocardiograph changes ST-segment depression T-wave flattening T-wave inversion Presence of U-waves Cardiac arrhythmias Atrial tachycardia Bradycardia Atrioventricular block Premature ventricular contractions Ventricular tachycardia Ventricular fibrillation Sudden death	Weakness Muscle twitching Tremor Altered mental status Anorexia Nausea Vomiting Diarrhea Refractory hypokalemia and hypocalcemia Electrocardiograph changes Prolonged PR Widened QRS Prolonged QT ST depression Peaked T-wave T-wave flattening Cardiac arrhythmias Atrial fibrillation Torsade de pointes Ventricular arrhythmias Ventricular tachycardia Tetany Convulsions Seizures Coma Death	Encephalopathy (e.g., Wernicke-Korsakoff encephalopathy) Lactic acidosis Death	Fluid overload Pulmonary edema Cardiac decompensation

that develop once nutritional support is initiated. Refeeding after a period of malnutrition and starvation increases the basal metabolic rate, which results in major alterations in macronutrient metabolism. This leads to hypophosphatemia, hypomagnesemia, hypokalemia, and thiamine deficiency and can cause hyperglycemia during refeeding, decreased excretion of sodium and water, and an expansion of fluid compartments. The development of refeeding syndrome can result in severe cardiovascular and pulmonary complications. Cardiac arrhythmias and death have been seen in chronically malnourished patients receiving aggressive parenteral nutrition and early carbohydrate administration. Other significant complications include confusion, coma, and seizures. Congestive heart failure, pulmonary edema, diaphragm and intercostal muscle weakness, decreased tissue oxygen delivery, and increased carbon dioxide production can cause respiratory failure and can make weaning from mechanical ventilation more difficult (Table 3).

Factors that aid in the identification of patients at risk for refeeding syndrome include:

- BMI < 16–18.5 kg/m²
- Unintentional weight loss >10–15% within last 3–6 months
- Little or no nutritional intake for >5–10 days
- A history of alcohol abuse or drugs, including insulin, chemotherapy, antacids, or diuretics
- Low levels of phosphorous, potassium, or magnesium prior to feeding
- Uncontrolled diabetes mellitus (diabetic ketoacidosis)
- Abused/neglected/depressed elderly adults
- Bariatric surgery
- Dysphagia
- Malabsorption (short bowel syndrome [SBS], inflammatory bowel disease [IBD], cystic fibrosis [CF], persistent nausea/vomiting/diarrhea, chronic pancreatitis)
- Chronic disease conditions (tuberculosis, HIV, cancer)
- Prolonged hypocaloric feeding or fasting
- Unconventional/eccentric diets^{65,68,69}

Nutritional Support

The two routes of nutritional support are enteral and parenteral. Enteral nutrition (EN) is provided via the gastrointestinal tract, either by mouth or through a feeding tube. Parenteral nutrition (PN) is an intravenous solution composed of nutrients infused through an IV line that bypasses the gastrointestinal tract. Determination of the most appropriate route is influenced by the patient's nutritional risk, clinical diagnosis and condition, gastrointestinal tract function, and duration of anticipated need (Figure 1).

Parenteral Nutrition

PN provides nutrition to patients who are unable to digest or absorb sufficient nutrition via the gastrointestinal tract. These patients may include those with an obstruction, severe malabsorption, bowel hypo- motility (ileus), or bowel ischemia (Table 4). EN is the preferred modality over PN as it has been shown to have cost, safety, and physiologic benefits. EN may reduce disease severity, complications, and length of stay, and improve patient outcome.¹²

Administration of PN typically requires insertion of a central venous catheter or peripherally inserted central catheter (PICC). Due to the risks of catheter-related complications and infection, current recommendations suggest that PN be considered after 7-10 days if early EN is not feasible to meet >60% of nutrition needs following ICU admission.¹² In multicenter, randomized controlled trials, later PN was associated with lower ICU mortality, shorter ICU length of stay, fewer infections and other complications, and greater reduction of costs when compared with early initiation.⁷⁰⁻⁷²

In adult patients with evidence of moderate-to-severe malnutrition where EN is not an option should receive PN within the first few days following admission.¹² There is currently no consensus on the timing of supplemental PN initiation in pediatric patients.⁴⁴

Parenteral nutrition formulations

PN is customized to individual patient needs for nutrients, electrolytes, vitamins, and trace elements by specially certified

Table 4. Contraindications to Enteral Nutrition Support⁷⁰

- Severe short-bowel syndrome (< 100-150 cm small bowel remaining in the absence of the colon or 50-75 cm remaining small bowel in the presence of the colon)
- Other severe malabsorptive conditions
- Severe GI bleed
- Distal high-output GI fistula
- Paralytic ileus
- Intractable vomiting and/or diarrhea that does not improve with medical management
- Inoperable mechanical obstruction
- When the GI tract cannot be assessed—for example, when upper GI obstructions prevent feeding tube placement

GI: gastrointestinal.

pharmacists through a process called compounding.⁷³ Manual and automated compounding devices are available, but numerous cases of parenteral compounding errors in ordering, transcribing, compounding, and infectious complication have been reported.⁷⁴⁻⁷⁵ To address these problems, preparation of PN can be outsourced to specialized compounding pharmacies. Standardized, premixed, and commercial products are available. To improve the safe administration of parenteral nutrition, standardized procedures for ordering, labeling, nutrient dosing, screening orders, administering, and monitoring are recommended.⁷⁶

Enteral Nutrition

Short-term EN is typically administered via a nasally or orally inserted small-bore weighted tip feeding tube called a Dobhoff tube. The weighted tip helps the tube travel past the stomach and through the pyloric valve into the duodenum and jejunum. Initial placement is performed with a guide wire inserted into the tube. Complications during insertion can include soft-tissue

trauma and hemorrhage, esophageal perforation, and placement into the lungs. Definitive verification of tube placement is determined by chest radiograph.

Percutaneous endoscopic gastrostomy (PEG) or jejunostomy (PEJ) tubes placed surgically through the abdominal wall should be considered for long-term enteral feeding when nutritional support is expected for at least 6 weeks.⁷⁷

Enteral nutrition formulations

Numerous EN formulations are available with various products designed for specific disease states such as renal failure, gastrointestinal disease, diabetes and hyperglycemia, hepatic failure, acute and chronic pulmonary disease, and immunocompromised states (Table 5).

Unfortunately, most of these specialty products lack strong scientific evidence to promote use because of inconsistent, inconclusive, or unavailable clinical trial results.⁷⁷ Until clinical evidence becomes available, standard formulas should be used for the majority of patients requiring enteral feeding. In the critically ill, at-risk patient, evaluation of age, the nutritional needs, physical assessment, metabolic abnormalities, gastrointestinal (GI) function, fluid status, food allergies, diet preferences, and overall medical condition should be used to identify the enteral formula that will meet the individual patient requirements and determine product selection.^{79,80}

Feeding Tube Placement — Gastric versus Post-Pyloric

There is an ongoing controversy in clinical practice regarding post-pyloric versus gastric feeding tube placement. Generally, in the intensive care unit it is preferred to place the feeding tube in the post-pyloric position due to the assumption that delayed gastric emptying results in a predisposition to bleeding, regurgitation, reflux, and aspiration.⁸¹⁻⁸³ The clinical condition of the patient generally dictates the placement of the feeding tube. Patients who are at high risk for aspiration and delayed gut motility should be considered for post-pyloric small bowel access. Per ASPEN guidelines, these patients include those who have sustained severe blunt and penetrating torso and abdominal injuries, severe head injuries, major burns, undergone major intra-abdominal surgery, had a previous episode of aspiration or emesis, had persistent high gastric residuals, are unable to protect the airway, require prolonged supine or prone positioning, or are anticipated to have multiple surgical procedures.¹⁵

Post-pyloric feeding access can be difficult and may delay the introduction of EN. The repeated attempts of placement and using more advanced modalities such as fluoroscopy to determine placement can increase costs of providing care.⁸⁴ An electromagnetic sensor-guided enteral access system (Cortrak 2, Avanos Medical Devices, Alpharetta, Georgia) can provide real-time feeding tube location and help reduce the risk of complications and avert the need for additional procedures.⁸⁵

Multiple studies have not shown a significant difference in improved clinical outcomes with post-pyloric feeding tube placement. Meta-analysis of clinical outcomes of several small sample size studies have evaluated mortality, incidence of pneumonia, and reducing aspiration risk.⁸⁶⁻⁸⁸ Although numerous studies have demonstrated that EN via the small bowel decreases the risk of pneumonia, no difference in ICU and hospital LOS, duration of ventilation, or mortality were determined.^{12,80} The only clinical outcome that has been shown to have an improvement with post-pyloric feeding is an increase in the volume of targeted nutrient delivery.⁸⁸

Current recommended practice in adults is to target post-pyloric feeding tube placement but not to delay gastric feeding unless clear signs of intolerance, aspiration risk, and high gastric residual volume are evident. See table 6 for suggested enteral access route selection based on duration of anticipated need and gastric function.

Based on several observational studies, it is suggested that the gastric route is acceptable for patients in the pediatric ICU unless the patient has demonstrated intolerance to gastric feeding or a high risk for aspiration.⁴⁴

Gastric Residual Volume

The practice of measuring GRV is a standard nursing practice used to determine tolerance of gastric tube feedings. It is assumed that high GRV is correlated with an increased risk of reflux, aspiration, and pneumonia. However, little evidence exists in the literature correlating GRV with these risks.⁸⁹ GRV has not been shown to be a marker of aspiration.^{90,91} Aspiration occurs in critically ill patients whether GRV is low or high, but aspiration risk may increase with high GRV.⁹² It has also been shown that GRV does not correlate with gastric emptying in both the adult and pediatric populations.^{93,94}

The practice of checking GRV is time intensive, and small-bore feeding tubes often occlude during the process.⁹⁵ In fact, the practice of checking GRV may result in inappropriate cessation of

Table 5. Common Enteral Formulations Marketed for Patient and Disease Specific Conditions⁷⁸

Enteral Formula Type	Description	Products
Standard Formulas	• Meant to match nutrient requirements for healthy individuals • Concentrations vary from 1.0-2.0 kcals/mL • May or may not include soluble/insoluble fiber	Osmolite ⁺ Jevity ⁺ Promote ⁺ TwoCal HN ⁺ Nutren ⁺ Isosource ⁺ Fibersource ⁺ Replete ⁺
Diabetic	• Lower carbohydrate with increased fat • Contain more complex carbohydrates • Concentrations vary from 1.0-1.5 kcal/mL	Glucerna ⁺ Glytrol ⁺ Diabetisource AC ⁺
Renal	• Generally lower in protein, calorically dense, and lower in potassium, magnesium and phosphorus • May vary in protein, electrolytes, vitamin and minerals depending on renal replacement therapy	Nepro ⁺ Suplena ⁺ Novasource Renal ⁺ Renalcal ⁺
Liver	• Increased amounts of branched chain amino acids (BCAA) with decreased aromatic amino acids (AAA) • Calorically dense, low in total protein, sodium, and fat-soluble vitamins and minerals	Nutrihep ⁺
Pulmonary	• Calorically dense, low in carbohydrate and high in fat (COPD) • Calorically dense, high omega-3 to omega-6 fatty acid ratio, antioxidants, (ARDS)	Pulmocare (COPD) ⁺ Nutren Pulmonary (COPD) ⁺
Immune Modulating	• Key ingredients include arginine, glutamine, nucleotides, and omega-3 fatty acids	Vital High Protein ⁺ Peptamen Intense VHP ⁺
Bariatric	• Designed for critically ill morbidly obese patients • Very high in protein • Contains omega-3 fatty acids	Vital High Protein ⁺ Peptamen Intense VHP ⁺
Pediatric	• Formulated for pediatric nutrient needs and conditions	Pediasure ⁺ Elecare ⁺ Nutren Junior ⁺ Peptamen Junior ⁺ Renastart ⁺

Abbott Nutrition⁺, Nestle Health Science⁺

Table 6. Selection of Enteral Access Based on Gastric Tolerance and Anticipated Feeding Duration^{18,77}

Normal Gastric Motility	Duration	
	Short term (4-6 Weeks)	Long Term (Longer than 6 Weeks)
Yes	Nasogastric	Gastrostomy
No	Nasoduodenal Nasojejunal	Jejunostomy

EN and cause a decrease in nutrient delivery and accumulation of calorie deficit over time. Caloric deficit in already at-risk mechanically ventilated patients may increase complications and morbidity.⁹⁶

In the absence of other signs of intolerance (such as emesis and abdominal distension), the most recent ASPEN/SCCM clinical practice guidelines recommend not using GRV for routine monitoring in ICU patients. ICUs that continue this practice should use GRVs in the 200-500 mL range to trigger interventions toward minimizing the risk of aspiration and should avoid the mandatory interruption of EN.¹² This highly controversial recommendation is supported by several studies that show that a higher tolerable GRV was not associated with an increase in adverse events such as regurgitation, emesis, aspiration, and ventilator associated pneumonia.⁹⁷⁻¹⁰¹ Higher GRV in combination with prokinetic agents to promote bowel motility have been shown to improve nutrient volume administered and reduce the time to reach target goals without increasing complications.⁹⁹⁻¹⁰² Two prospective studies compared routine GRV monitoring to not checking GRV and also found no difference in adverse events.^{95,102} Therefore, the controversial practice of not routinely monitoring GRV is supported by current evidence.

Routine assessment of GRV in critically ill children is also not recommended in the recent European Society for Pediatric and Neonatal Intensive Care (ESPINIC) guidelines.¹⁰³

Regardless of the acceptable GRV used by an institution, the following practices have been proven to reduce the risk of aspiration.¹²

- Head of bed elevation to 30-45 degrees
- Use of bowel motility agents such as metoclopramide
- Post-pyloric or small bowel feeding tube placement when indicated.
- Continuous EN vs bolus EN

- Reducing the level of sedation/analgesia when possible
- Minimizing transport out of the ICU for diagnostic tests and procedures
- Use of chlorhexidine mouthwash

Trophic Feedings

Hypocaloric, low-dose, “trickle,” or trophic feeding is the practice of feeding minimal amounts (e.g. 10–30 mL/hr in adults, 1-10 mL/hr in pediatrics, and 15-20 mL/kg/day in neonates) of EN with the primary goal to maintain gut function and integrity despite not meeting daily caloric needs. It is most often used in preterm infants on PN¹⁰⁴ or in adult and pediatric patients with impaired enteral feeding tolerance or gut function. EN stimulates organs of digestion to function in their normal capacity and to assist in the digestion and absorption of nutrients. It also prevents passage of bacteria across the GI tract into the systemic circulation, reducing infection rates, enhancing immune function, and preserving GI mucosal structure and function.³³ Trophic feeding may also reduce the development of a postoperative ileus.^{12,105} Studies in mechanically ventilated patients with respiratory failure or ARDS show that trophic feedings resulted in fewer episodes of gastrointestinal intolerance but resulted in similar clinical outcomes compared to early advancement to full enteral feeding.^{106,107}

In extremely premature infants, a short vs extended duration of trophic enteral feeding was associated with earlier full EN support and not associated with an increased risk of necrotizing enterocolitis.¹⁰⁸

A meta-analysis¹⁰⁹ and a systematic review¹¹⁰ showed no outcome benefit in morbidity and mortality, acquired infections, duration of ventilation, and ICU length of stay of intentional underfeeding when compared to full feeding strategies. However, in patients with high nutritional risk, refeeding syndrome, shock, and acute gastrointestinal injury, early targeted full enteral nutrition maybe associated with worse prognosis.¹¹¹ Furthermore, it has been proposed that in the early acute phase of critical illness, overuse of the gut may worsen outcomes and that a gut protective, trophic feeding strategy should be initiated.¹¹²

Stress Ulcer Prophylaxis

Critically ill patients are at risk of GI bleeding from gastric or duodenal ulcers due to increased gastric acidity and decreases in the gastric mucosal barrier. EN can improve mucosal blood flow and reverse the production of inflammatory mediators that

cause gastropathy. EN may provide stress prophylaxis and help to reduce the use of acid-suppressive therapy in the ICU.¹¹³

Stress ulcer prophylaxis was investigated in a randomized controlled study and did not find additional benefit of gastric acid suppression versus early enteral nutrition in mechanically ventilated medical ICU patients. The authors concluded that these results further support the protective effect of early initiation of EN.¹¹⁴

Nutrient Requirements and Distribution

The purpose of a nutritional assessment is to determine a nutrition care plan with the primary goal of meeting the nutritional requirements of the patient. This includes determination of total energy, protein, carbohydrate, fat, and micronutrient needs.

Carbohydrate requirements

Carbohydrates are the primary fuel source for the body. It is recommended that approximately 45–65% of total calories come from carbohydrates. A minimum daily amount of 100–150g/day in adults is necessary to provide adequate glucose to the brain. If consumed in insufficient amounts, an accumulation of ketone bodies develops as a result of excessive fat and protein catabolism, and acidosis occurs.¹¹⁵

Protein requirement

Amino acids or proteins are essential to maintaining or restoring lean body mass. Because illness usually increases protein catabolism and protein requirements, the recommended dietary allowance (RDA) of 0.8 g/kg per day in adults or 0.8–1.5 g/kg per day in pediatrics is generally insufficient for critically ill patients. Based on the assessment of the protein catabolism rate, protein intake may need to be doubled or even tripled above the RDA (1.5 to 2.5 g/kg/day in adults or 1.5–4 g/kg/day in pediatrics). Ideally, approximately 20% of a patient's estimated calorie needs should be provided by protein. Higher percentages of protein may be needed in patients with “wasting syndrome” or cachexia, elderly persons, and persons with severe infections. However, whenever high protein intakes are given, the patient should be monitored for progressive uremia or azotemia (rising BUN > 100 mg/dl).¹¹⁵ Too much protein is harmful, especially for patients with limited pulmonary reserves. Excess protein can increase O₂ consumption, REE, minute ventilation, and central ventilatory drive.¹¹⁶ In addition, overzealous protein feeding may lead to symptoms such as dyspnea in patients with chronic pulmonary disease.

Fat requirements

The remaining calories (20–30%) should be provided from fat. A minimum of 2–4% is needed to prevent essential fatty acid deficiency. Fat intakes in excess of 50% of energy needs have been associated with fever, impaired immune function, liver dysfunction, and hypotension.¹¹⁵

Vitamins, minerals, and electrolytes

The dietary reference intakes (DRI) provide the recommended optimal level of intake for vitamins, minerals, and electrolytes. The primary goal is to prevent nutrient deficiencies as well as help reduce the risk of chronic diseases. Some nutrients may need to be supplemented above the DRI for certain disease states, therapies, or conditions.¹¹⁵

Fluid requirements

Fifty to sixty percent of body weight consists of water. Fluid requirements are estimated at 1 ml/kcal/day or 20–40 ml/kg/day in adults. In pediatric patients the modified Holliday Segar (4-2-1) method is generally utilized.¹¹⁷ There are some specific populations where practitioners deviate from utilizing this method to establish maintenance fluid requirements, however it is institution specific. Depending on a patient's medical condition, fluid restriction may be warranted. Additional fluid may be required for excessive fluid losses (urinary, fecal, blood, wound, emesis) and with excessive insensible losses (fever).¹¹⁵

Nutrition Support and Respiratory Function

Patients with acute and chronic respiratory failure may present with or have the potential to develop nutrition-related complications. Nutrition support plays a significant role in treatment as further deterioration can have a direct effect on respiratory function, further decline, and poor outcomes.¹¹⁸ Specific nutrition recommendations exist for intervention and treatment of acute and chronic respiratory failure.

Respiratory consequences of malnutrition may include the following:¹¹⁹

- Loss of diaphragmatic and accessory muscle mass and contractility
- Ineffective cough
- Decreased maximum expiratory pressure and maximum inspiratory pressure
- Decreased FVC or FEV₁

- Reduced production of surfactant¹²⁰
- Fluid imbalance
- Congestive heart failure
- Decreased lung compliance, atelectasis, and hypoxemia
- Decreased hypoxic and hypercapnic response¹²¹⁻¹²³
- Increased CO₂ production
- Increased incidence of hospital-acquired infections¹²⁴
- Decreased lung clearance mechanisms
- Increased bacterial colonization
- Emphysematous changes to lung parenchyma¹²⁵⁻¹²⁶

Chronic Obstructive Pulmonary Disease

Disease-related malnutrition is common in patients with chronic obstructive pulmonary disease (COPD). Between 30–60% of inpatients and 10–45% of outpatients with COPD are at risk for malnutrition.¹²⁷⁻¹²⁸ Malnourished COPD patients exhibit a higher degree of gas trapping, reduced diffusing capacity, and a diminished exercise tolerance when compared to patients with normal body weight, adequate nutrition, and comparable disease

severity.¹²⁹ The underlying mechanism between malnutrition and COPD is thought to be from a variety of contributing factors (Figure 7).

Malnutrition may be responsible for the respiratory muscle wasting, which intensifies the progression of COPD or may simply be a consequence of disease severity. Similarly, long-term caloric malnutrition is associated with the loss of body weight that includes an extensive loss of lung tissue and reduction in diffusion capacity. Emphysematous-like changes are found to occur in persons with chronic anorexia nervosa and those who die of starvation.^{125,126}

In COPD patients with acute respiratory failure, malnutrition may have detrimental effects, especially in weaning from ventilatory support.¹³⁰ Malnutrition is associated with a decrease in diaphragmatic muscle strength,^{131,132} a decrease in ventilatory drive,¹²² reduced surfactant production,¹²³ and an increased risk of nosocomial pneumonia.^{124,127} Protein energy malnutrition is common in COPD patients.¹³³ Early and aggressive nutritional support in COPD patients can produce significant improvements

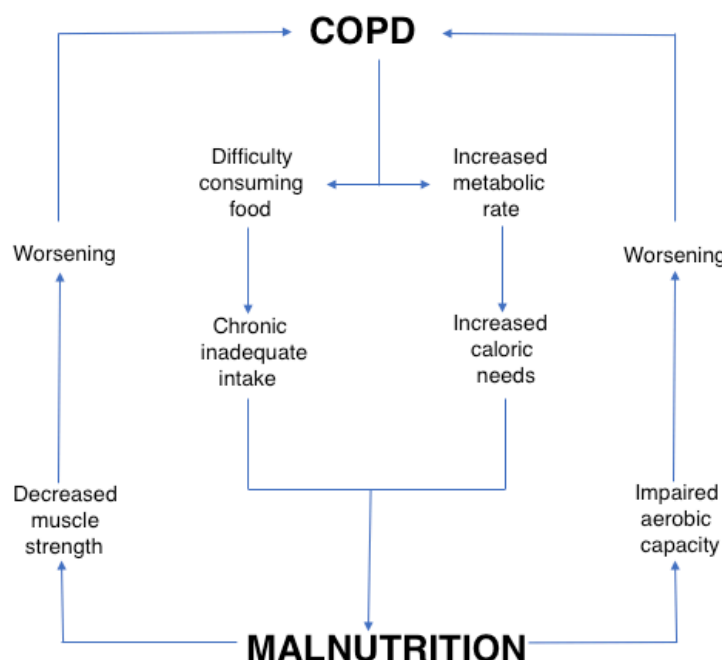


Figure 7 . Cycle of Malnutrition in COPD¹¹⁹

in several functional outcomes including respiratory and limb muscle strength.

Increased protein intake may improve ventilatory response to CO₂.¹²² Several meta-analyses of nutritional support studies have demonstrated improved nutrition related to anthropometric improvements,¹³⁴ inspiratory and expiratory muscle strength, exercise tolerance, and quality of life.¹²⁷ The most recent Cochrane systematic review found evidence of significant improvements in weight gain, indices of respiratory muscle strength, walking distance, and quality of life in malnourished COPD patients who received nutritional supplementation.¹³⁵

CO₂ is produced with the metabolism of all macronutrients, with the largest amount coming from carbohydrates. It is well known that overfeeding with an excess carbohydrate load increases CO₂ production. However, overfeeding with non-carbohydrate calories can be as detrimental in regard to CO₂ production and the increased work of breathing.¹³⁶ A high-fat, reduced carbohydrate nutrition formulation has been marketed in an effort to encourage the benefits of nutrition repletion and weight gain while reducing CO₂ production;¹³⁷ however, several studies have refuted this theoretical benefit, and the practice is not recommended.¹³⁸⁻¹⁴³

Underlying causes of malnutrition

Underlying causes of malnutrition in COPD patients include increased energy expenditure due to increased caloric cost of breathing, increased systemic inflammation, and the thermogenic effect of medications such as bronchodilators. Also, COPD patients have an inadequate caloric intake caused by dyspnea while eating, chewing and swallowing difficulties, taste alterations and suppressed appetite from medications, the use of a nasal cannula or tracheostomy, and early satiety.¹¹⁹

Psychosocial factors are also underlying causes that contribute to malnutrition in COPD patients. Depression, poverty, difficulty shopping, and tiring easily when preparing food often prevent good nutrition.¹¹⁹

Acute Respiratory Distress Syndrome

Acute respiratory distress syndrome (ARDS) is acute pulmonary failure that manifests from inflammatory conditions. Omega-3 fatty acids are metabolized to substances that reduce inflammation and inflammatory mediator production. Enteral supplementation with omega-3 fatty acids may have a beneficial effect in treatment for ARDS. Several studies observed reduced duration of mechanical ventilation, number of days in the ICU,

rates of organ failure, and mortality compared to use of standard enteral formulas.^{12,144,146}

Omega-6 fatty acids are metabolized to proinflammatory substances that influence cytokine production, platelet aggregation, vasodilation, and vascular permeability, and therefore may be harmful.¹⁴⁷⁻¹⁵¹

Nutritional support high in omega-6 fatty acids should be avoided.¹⁴⁷ Nutritional supplementation with higher omega-3 to omega-6 fatty acid ratios have been recommended to reduce the risks of inflammatory disorders such as coronary heart disease, diabetes, arthritis, cancer, osteoporosis, rheumatoid arthritis, and asthma.^{148,151} Due to conflicting results from recent trial, the practice of omega-3 supplementation, fish oils, borage oils, and antioxidants remains controversial in patients with ARDS.^{12,152,153}

Nutritional Support During Critical Illness

The general goals of nutritional support in the critically ill patient are to provide the energy and protein necessary to meet metabolic demands and to preserve lean body mass. Nutritional support is also an important therapy in critical illness as it attenuates the metabolic response to stress, prevents oxidative cellular injury, and modulates the immune response. Nutritional modulation of the stress response includes early enteral nutrition, appropriate macro and micronutrient delivery, and meticulous glycemic control.¹²

Stress Response in Critical Illness

Metabolic needs vary during critical illness. The metabolic response to critical illness occurs in three phases: the ebb phase, the early flow phase, and the late flow phase (Figure 8).

The ebb phase typically lasts for 24–48 hours and is associated with physiologic stress characterized by hemodynamic instability, hypotension, tissue hypoxia, and a decrease in oxygen consumption, body temperature, and metabolic rate. During this period of physiologic stress, insulin resistance and endogenous glucose production (EGP) increases. EGP can account for up to two-thirds of total energy requirements. In contrast to the healthy state, exogenous feeding does not attenuate EGP and can result in excessive energy availability. Exogenous feeding equivalent to the determined energy requirement during the first 24–48 hours of critical illness can result in overfeeding. During the ebb phase, the primary goal is resuscitation and hemodynamic stabilization.

After resuscitation and stabilization, the ebb phase is followed by a prolonged catabolic flow phase characterized by a hypermetabolic

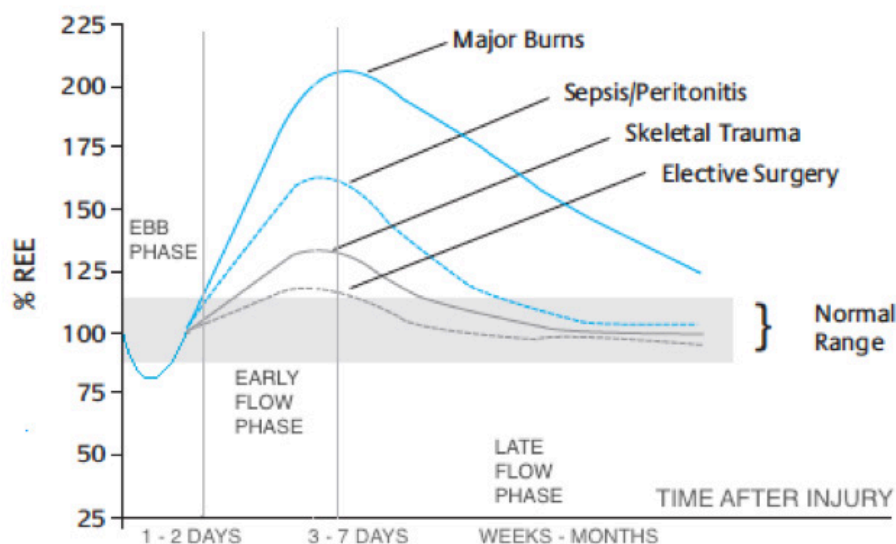


Figure 8. Stress Response in Critical Illness: EBB and Flow Phases¹⁵⁴⁻¹⁵⁷

state where a marked increase in cardiac output, oxygen consumption, carbon dioxide production, and REE are seen. The early flow phase usually lasts 7–10 days, and the goal is to provide ongoing metabolic support with high-protein feedings while avoiding overfeeding. Caloric needs may be increased by up to 100% during the early flow phase in patients with severe burns.^{154,158}

As the catabolic phase resolves, the late flow phase begins and can last for months. Caloric needs may remain elevated during the late flow phase for anabolic repletion of lean body mass and fat stores.

Under- and Overfeeding During Critical Illness

Providing inadequate provision of nutrients can have negative effects on the critically ill patient.¹⁶⁰⁻¹⁶² Underfeeding can result in a loss of lean body mass, immunosuppression, poor wound healing, and an increased risk of infection.¹⁶¹ This can also result in an inability to respond to hypoxemia and hypercapnia, and a diminished weaning capacity.¹⁶² Continual underfeeding in the ICU results in a cumulative caloric deficit, which increases length of stay, days of mechanical ventilation, and mortality.⁹⁶

Overfeeding patients can be equally detrimental as well.

Excess amounts of nutrients can exacerbate respiratory failure by increasing carbon dioxide production.¹⁶¹ Excess total calories (not excess carbohydrates) increase CO_2 production and, therefore, increase the work of breathing.¹³⁶ If under- or overfeeding is suspected, indirect calorimetry is an important tool to help determine accurate energy requirements.

Glycemic Control in Critical Illness

Control of serum glucose levels in non-diabetic patients during critical illness is important due to the adverse effect of hyperglycemia in patient outcomes. Control of hyperglycemia has been shown to reduce morbidity and mortality in hospitalized patients. Hyperglycemia is a normal response to physiologic stress and the inflammatory response related to critical illness. Since hyperglycemia can be caused by enteral and parenteral nutrition, control of hyperglycemia during nutritional support is of critical importance. The stress response to critical illness causes wide swings in nutrient requirements. Therefore, the nutritional support process needs to balance the potential detrimental effects of both under- and overfeeding with glycemic control. Current recommendations in adults are to maintain a target

Table 7. Consequences of Over–Underfeeding¹⁵⁹⁻¹⁶²

Overfeeding	Underfeeding
Physiologic stress	Increased complications
Respiratory compromise	Immune suppression
Prolonged mechanical ventilation	Prolonged hospitalization
Hyperosmolar state	Respiratory compromise
Hyperglycemia	Poor wound healing
Hepatic dysfunction	Nasocomial infection
Excessive cost	Prolonged mechanical ventilation
Immune suppression	
Fluid overload	
Axotemia	

blood glucose goal range of 140–180 mg/dL and to consider a blood glucose value of <70 mg/dL during nutritional support as a treatable hypoglycemia.¹⁶³⁻¹⁶⁶ Targeted glucose control in this range was associated with improved outcomes compared to tighter glucose control and resulted in fewer incidences of hypoglycemia in adult¹⁶⁷ and pediatric patients.¹⁶⁸

Nutritional Support of the Obese Patient

Hypocaloric feeding is recommended for the critically ill obese adult patient. Guidelines suggest the goal of EN should not exceed 65–70% of target energy requirements with a high protein goal of 2.0–2.5 g/kg of ideal body weight depending on the patients BMI.^{12,169}

It is essential to provide adequate protein in these patients to maintain nitrogen balance and lean body mass while encouraging

the use of adipose tissue for fuel. Morbidly obese patients receiving high protein through permissive underfeeding have reduced insulin resistance, lower insulin requirements, better glycemic control, decreased ICU stay, and reduced duration of mechanical ventilation.¹⁷⁰⁻¹⁷²

Pediatric Critical Illness

Optimizing nutritional therapy in pediatric patients can improve clinical outcomes. As in the adult, the goals of pediatric nutrition support encompass preservation of tissue stores and resolution of disease progress. However, the goals of pediatric nutrition are often more complicated than those for adult patients. In addition, needs that address the support for appropriate growth and development, the need for preservation of oral motor skills must also be considered. Components of energy requirements in pediatrics consist of basal or resting energy expenditure, activity, growth, gender, maintenance of normal body temperature, and stress factors. Requirements for vitamins and minerals vary based on age, medical status, and size of the child.^{44,103}

A multicenter international study demonstrated that the use of PN was associated with higher mortality in children. However, pediatric intensive care units that used protocols for starting and advancing EN support had a higher percent of target calorie goals administered, reduced 60-day mortality, and lower rates of acquired infections.¹⁷³ In another study, a stepwise algorithmic protocol for advancing EN was shown to significantly improve the time to reach nutrition delivery goals, minimize unnecessary interruptions, and reduce the utilization of PN.¹⁷⁴

Immunonutrition

Critical illness is often complicated by systemic inflammation and generalized immunosuppression. Malnutrition associated with critical illness leads to impeded wound healing, loss of lean muscle mass, delayed weaning from ventilatory support, longer hospital stays, and increased rates of infection. Immunonutrition using immune modulating nutrition formulations containing omega-3 fatty acids, arginine, glutamine, nucleotides, and antioxidants are used with the goal to modulate mucosal barrier function, cellular defense function, and local and systemic inflammatory response.¹⁷⁵

The use of these formulas in surgical patients has been shown to decrease risk for infections, reduce length of stay, and reduce mortality.¹⁷⁶ However, caution is advised within the use of arginine in critically ill severe septic patients.¹⁷⁷ Agreement concerning the use of specific immune-modulating substrates for different patient populations is lacking.¹⁷⁵

Due to the paucity of available studies, the use of immunonutrition is not currently recommended for the critically ill pediatric patient.⁴⁴

Determining Nutritional Requirements

Calculating, estimating, or measuring the number of calories required by an individual determines nutrient requirements. A calorie is a unit of energy equivalent to the amount of potential heat produced or contained in food when released during the metabolic oxidation processes of the body. A calorie is defined as the amount of heat needed to raise the temperature of 1 gram of water by 1 C° (also called a small calorie, abbreviated as cal). A Calorie (also called a large calorie, abbreviated as Cal) is defined as the amount of heat needed to raise the temperature of 1 kilogram of water by 1 C°, is equivalent to 1,000 calories and is also referred to as a kilocalorie. Kilocalories (kcal) are used to quantify the energy value of foods.

Calorie or energy needs are fundamental to the recommendations of the nutritional care plan. Macronutrients supply the body's energy requirements. The calorie contribution of the three major macronutrients are: protein = 4 kcal/g; carbohydrate = 4 kcal/g; and fat = 9 kcal/g. Alcohol is the only other calorie source with approximately 7 kcal/g.

Energy needs vary according to activity level and state of health. Energy needs of critically ill patients can be significantly different than normal values. Additional factors such as energy expended in catabolic states may be needed to adjust the estimate in patients with medical conditions such as injury, major wounds, and infection.¹⁷⁸ Energy needs for obese individuals are less because adipose tissue uses less energy than muscle uses.

The estimated energy requirement is the average dietary energy intake needed to maintain energy balance in an individual.¹⁷⁹ Estimating energy requirements for people according to their age, sex, weight, height, and level of physical activity is accomplished by the use of predictive equations.

Predictive Equations

Numerous equations have been developed to predict caloric requirements. The Harris Benedict Equation (HBE), the most well-known predictive equation, was developed in 1919 by comparing measured calories and their correlation to height, weight, age, and sex in normal subjects to estimate the basal metabolic rate (BMR). BMR is defined as the amount of heat produced in a state at rest with complete muscle inactivity during a post-absorptive

period 12–14 hours after the last meal. Since BMR as defined by Harris and Benedict is not necessarily reflective of the way nutritional requirements are determined in hospitalized patients, the more relevant terms – resting metabolic rate (RMR) or resting energy expenditure (REE) – are used in clinical practice to predict or measure caloric needs.

Predictive equations use factors validated by the original work by Harris and Benedict and incorporate additional factors such as temperature, body surface area, diagnosis, and ventilation parameters, as shown in Table 8.

Predictive equations have been modified as additional data (such as injury-stress, activity, medications received, and obesity) and have been added to the regression correlation equations. Several predictive equations were developed with a focus on specific patient populations and medical conditions. Predictive equations have varying degrees of agreement compared to measured calorie requirements.^{157,180} Error rates are not insignificant and, therefore, can have high degrees of under- and overestimation of caloric needs in adults¹⁸¹ and children.^{182,183}

This variability can result in errors large enough to impact outcomes.¹⁸⁴ Error rates with some equations make them unsuitable as assessment methods of energy expenditure in critically ill patients. Systematic reviews and meta-analyses comparing measured calorie requirements show that some predictive equations may be useful in critically ill non-obese patients while others might be useful in obese patients.^{181,184} Studies have compared accuracy reliability, and error ranges for predicting REE within 10% of measured.^{185,186} However, in both the adult and pediatric populations error rates can still be significant regardless of the prediction method used (Table 8).^{44,157,182,183} For example, in obese patients, there are no clinical features that can identify individual patients where a predictive equation is inaccurate.¹⁸⁷

There are more than 200 predictive equations in existence. Many were developed as long as 50–80 years ago and may not reflect body composition, nutritional risks, age, or ethnicity of the populations they are applied to. There is often no consensus on how a predictive equation is selected, and results can vary significantly between clinicians.¹⁸⁸ Furthermore, there are large segments of populations in whom predictive equations have no validation studies performed. These groups include the

Table 8. Examples of Commonly Used Predictive Equations¹⁵⁷

Equations	Parameters Used for Calculation	Accuracy Rate*
General Hospitalized Population		
25 kcal/kg	25 x WT	43% 10%
Harris & Benedict (1919)	M: $13.75 \times \text{WT} + 5.00 \times \text{HT} - 6.75 \times \text{age} + 66.47$ F: $9.56 \times \text{WT} + 1.85 \times \text{HT} - 0.67 \times \text{age} + 655.09$	43% 38%
Ireton-Jones (1992)	$1925 - 10 \times \text{age} + 5 \times \text{WT} + (281 \text{ if male})$ $+ (292 \text{ if trauma}) + (851 \text{ if burn})$	28%
Critically Ill Patients		
25 kcal/kg	25 x WT	12%
Harris & Benedict (1919)	M: $13.75 \times \text{WT} + 5.00 \times \text{HT} - 6.75 \times \text{age} + 66.47$ F: $9.56 \times \text{WT} + 1.85 \times \text{HT} - 0.67 \times \text{age} + 655.09$	37%
Ireton-Jones (1997)	$1784 - 11 \times \text{age} + 5 \times \text{WT} + (244 \text{ if male})$ $+ (239 \text{ if trauma}) + (804 \text{ if burn})$	37%
Mifflin-St. Jeor (1990)	M: $10 \times \text{WT} + 6.25 \times \text{HT} - 5 \times \text{age} + 5$ F: $10 \times \text{WT} + 6.25 \times \text{HT} - 5 \times \text{age} - 161$	35% 32%
Owen (1987)	M: $\text{WT} \times 10.2 + 879$ F: $\text{WT} \times 7.18 + 795$	12%
Penn State (2003)	$0.85 \times \text{HB} + 175 \times \text{Tmax} + 33 \times \text{Ve} - 6433$	43%
Swinamer (1990)	$945 \times \text{BSA} - 6.4 \times \text{age} + 108 \times \text{T} + 24.2 \times \text{RR}$ $+ 81.7 \times \text{V}_\text{T} - 4349$	55% 45%
*within +/- 10% of measured REE	M=Male F=Female HT=Height WT=Weight RR=Respiratory rate V _T =Tidal Volume BSA=Body Surface Area TMAX= Maximum Temperature (degrees Celsius)	

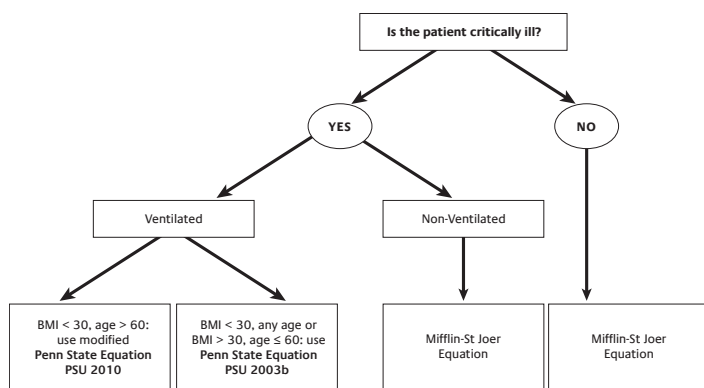


Figure 9. Predictive Equation Algorithm

elderly and many non-white racial groups. The limitations and variability of predictive equations when applied to an individual patient accentuates the need to use a regimented nutritional risk assessment process and sensible clinical judgment when deciding whether to use a predictive equation. Figure 9 provides an example algorithm for using predictive equations.

The American College of Chest Physicians' 1997 equation is a simple and prompt method to estimate daily energy needs of the average adult using a factor of 25–35 kcal/kg. This method is not necessarily as accurate as predictive equations, as it does not take into account gender, age, stature, and severity of illness. To rapidly estimate the energy needs of the average adult in kcal/day, identify the target goal for weight change and multiply the individual's actual body weight in kilograms times the factor listed as follows:¹⁸⁹

Goal	Energy Needs (kcal/kg)
Weight maintenance	25 to 30
Weight gain	30 to 35
Weight loss	20 to 25

A similar method is often utilized for calorie needs estimation in pediatrics. The daily recommended intake (DRI), which is presented in calories per kilogram, is often used as the target for calories goals.¹⁹⁰

To overcome the limitations of predictive equations and estimating formulas, energy needs can be measured at the bedside using indirect calorimetry to determine RMR or REE.

Calorimetry

Calorimeters measure heat released from chemical reactions or physical changes. Calorimetry has been used since the late 19th and early 20th centuries and was adopted as the major method of determining energy needs in individuals. Calculations of calorie requirements by mathematical equation were developed from the use of direct and indirect calorimetry.

Direct calorimetry

Direct calorimeters measure heat. A bomb calorimeter measures the energy value of food by measuring the precise amount of heat liberated as the food is burned in a closed chamber. Another type of direct calorimeter requires that the subject be enclosed in a sealed chamber for extended periods and a precise measurement of heat transfer conducted. The early experimentation conducted by nutrition scientists led to the development of the respiration chamber and IC.

Respiration chamber

The development of the respiration chamber combined the process of direct calorimetry with measurements of oxygen consumption (VO_2) and carbon dioxide production (VCO_2) (Figure 10).

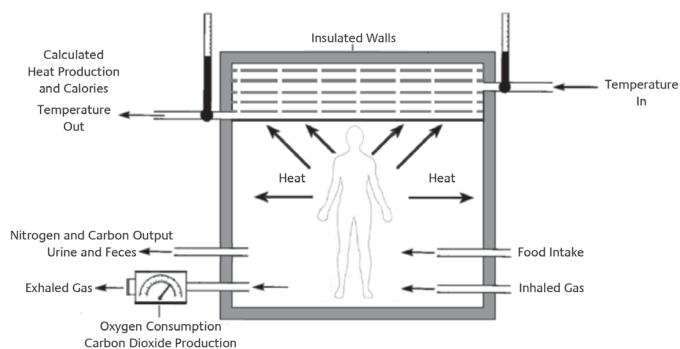


Figure 10. Respiration Chamber

Table 9. RQ Substrate Interpretation: Interpretation of Substrate Utilization Derived from the Respiratory Quotient¹⁹⁶⁻¹⁹⁸

Substrate Utilized	Respiratory Quotient
Ethanol	0.67
Ketones	0.67
Fat Oxidation	0.71
Protein oxidation	0.80-0.82
Mixed substrate oxidation	0.85-0.90
Carbohydrate oxidation	1.0
Lipogenesis	1.0-1.3

Around the turn of the century, the correlation of heat production in calories, the rate of VO_2 and VCO_2 , the quantity of nutrients consumed, and the mass of carbon and nitrogen excreted was used to derive the caloric value of oxygen and carbon dioxide.¹⁹¹⁻¹⁹⁵ By simultaneous measurement of the ratio VCO_2 to VO_2 , the respiratory quotient (Table 9) and caloric equivalent of each gas (Table 10) in relation to the oxidation of specific food substrates could be determined.¹⁹⁶⁻²⁰¹

It was observed that during short observational periods, the errors in computation of heat production and calories using indirect measurements of VO_2 and VCO_2 were less than the errors in computation when using direct calorimetry measurements.¹⁹³ Direct calorimetry and respiration chambers in relation to

metabolic testing to this day primarily remain as a research tool in animals.

Indirect calorimetry

Indirect calorimetry is the most accurate method for determining RMR and REE in various states of health and disease and, when available, is considered to be the gold standard for measuring energy expenditure in critically ill adult^{157,202,203} and pediatric patients.^{204,205} It relies on the determination of VO_2 and VCO_2 using precise measurements from a metabolic analyzer of the inspired and expired fractions of oxygen and carbon dioxide where:

$$\text{VO}_2 \text{ (mL/min)} = (\text{Vi} \times \text{FiO}_2) - (\text{Ve} \times \text{FeO}_2) \quad (1) \text{ and}$$

$$\text{VCO}_2 \text{ (mL/min)} = (\text{Ve} \times \text{FeCO}_2) - (\text{Vi} \times \text{FiCO}_2) \quad (2)$$

The abbreviated Weir equation uses the measured VO_2 and VCO_2 to determine REE where:

$$\text{REE} = (3.9 \times \text{VO}_2) + (1.1 \times \text{VCO}_2) \times 1.44 \quad (3)$$

The respiratory quotient, the ratio of VCO_2 to VO_2 , can then be calculated where:

$$\text{RQ} = \text{VCO}_2 / \text{VO}_2 \quad (4)$$

Since the normal $\text{RQ} = 0.85$, the volume of CO_2 produced is lower than the volume of O_2 consumed. Therefore, small differences in the inhaled versus exhaled volumes occur. In order to accurately calculate VO_2 and VCO_2 , the gas concentration measurements of a metabolic analyzer need to be within $\pm 0.01\%$. In regard to VO_2 measurements, elevated FiO_2 introduces error as the oxygen concentration approaches 1.0. As a result, the accuracy of IC diminishes as FiO_2 increases. Additionally, any error in gas concentration analysis or delivery is amplified at a higher FiO_2 . Due to this technical limitation, IC is not recommended or considered to be accurate at $\text{FiO}_2 > 0.60$.^{9,196,206}

Table 10. Caloric Equivalence¹⁹⁹⁻²⁰¹

Substrate	Respiratory Quotient	Oxygen Caloric Equivalent (kcal/L)	Carbon Dioxide Caloric Equivalent (kcal/L)
Carbohydrate	1.0	5.05	5.05
Mixed	0.90	4.83	5.52
Protein	0.80	4.46	5.57
Fat	0.71	4.74	6.67

Respiratory quotient was once thought to be useful as a means to determine nutritional substrate utilization. However, the accuracy of this assumption has never been substantiated. The large stores of CO_2 in the body can be mobilized with ventilation and, thus, would reflect an increase in CO_2 excretion but not necessarily production. Likewise, the effect of hypoventilation would have a similar yet opposite effect and result in a decrease in CO_2 excretion while CO_2 production remained constant. An increase or decrease in VCO_2 measured as a result of this mechanism would have an erroneous effect on the measured RQ (Figure 11).

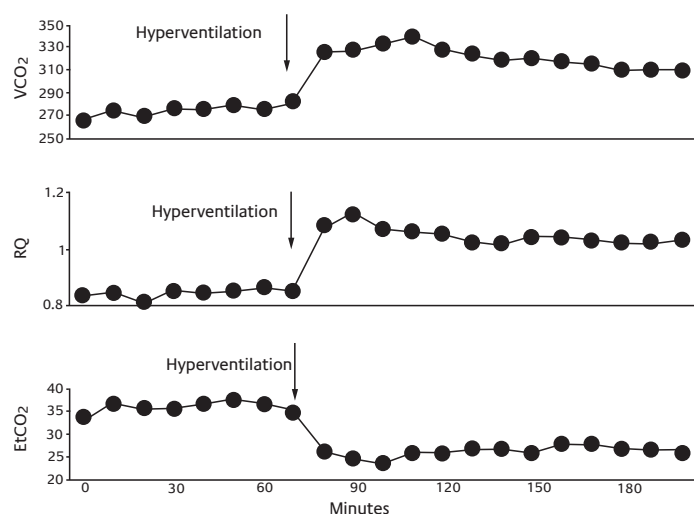


Figure 11. Effect of Acute Hyperventilation on VCO_2 , RQ, and EtCO_2 in a Patient with Head Injury¹⁹⁶

Therefore, the use of the RQ measurement is of limited clinical value in determining nutritional substrate requirements. Measured values of RQ between the physiologic ranges of 0.67–1.3 should be used as a means of quality control and a way to verify test validity. Values of RQ outside of this range obtained during IC testing invalidate the results due to technical measurement errors and should be repeated.¹⁹⁶

Indirect calorimetry is performed using a stand-alone metabolic cart by hood, face mask, and mouthpiece, or by connection to a ventilator (Figure 12).

Open circuit systems sample inspired gas concentrations, measured expired gas concentrations, and expired minute volume collected back into the analyzer to determine VCO_2 , VO_2 , and RQ. Indirect calorimetry has also been integrated into several ventilators (Table 11). Handheld calorimeters are also available (Figure 13). The MedGem handheld indirect calorimeter measures oxygen consumption to calculate resting energy expenditure using the caloric equivalent of oxygen.²⁰⁹⁻²¹¹

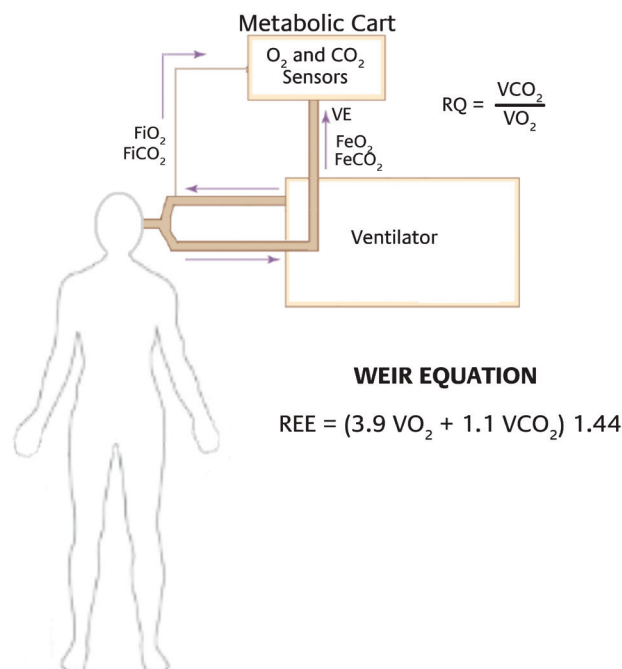


Figure 12. Metabolic Study During Mechanical Ventilation



Figure 13. MedGem Handheld Indirect Calorimeter
Courtesy of Microlife USA Inc., Clearwater, FL

Table 11. Advantages and Disadvantages of Ventilators with Integrated Indirect Calorimetry^{207,208}

Advantages	Disadvantages
• Enables real-time monitoring of metabolic stress during critical illness • Allows frequent repeatable measurements for closer monitoring of caloric balance • Disconnection from ventilator or when on high PEEP levels may be avoided • Reduces interference with ventilator function from attachment of an external device • Reduces infection risks from moving metabolic cart between patients • Fick cardiac output estimates are more accessible	• Limited use to a single patient attached to the ventilator • Cost of installation on several ventilators may be prohibitive • Difficult or unable to use on patients without an artificial airway (hood, face mask, and mouth piece studies)

Newer open-circuit breath-by-breath designs use a system where inspired and expired gases and volumes are measured at the airway, which simplify the measurement procedure (Figure 14). Accuracy of indirect calorimetry measurements are dependent on the technical aspects of test performance and patient care related variables.

Technical considerations when performing IC measurements during mechanical ventilation include:^{199,212}

- Warm-up time of 30 minutes for the indirect calorimeter
- Errors in calibration of flow, oxygen, and carbon dioxide sensors
- Presence of leaks (ventilator circuit, artificial airway, broncho-pleural fistulas)
- $\text{FiO}_2 > 60\%$
- Fluctuation of $\text{FiO}_2 > \pm 0.01\%$
- Changes in the ventilator setting within 1 to 2 hours of testing
- Acute hyperventilation or hypoventilation (changes body CO_2 stores)
- Moisture in the sampling system
- Bias flow through the ventilator may affect accuracy of indirect calorimeter
- Attachment of indirect calorimeter may affect ventilator function



Figure 14. CCM Express Indirect Calorimeter
Courtesy of Medical Graphics Corp., St Paul, MN

- Use of inhaled nitric oxide
- Presence of anesthetic gases

Recommendations for improving accuracy of IC include:²¹²

- Patient should be hemodynamically stable
- Patient should be in a comfortable resting position for 30 minutes before the study
- Avoid instability caused by disconnection from high levels of positive end-expiratory pressure (PEEP)
- Avoid voluntary activity for 30 minutes^{198,213-215}
- Avoid intermittent feedings or meals taken four hours before study^{198,215}
- Nutrient infusion should remain stable for at least 12 hours before and during the study^{198,215}
- Measurements are made in a quiet, neutral- thermal environment^{198,213-215}
- Limited voluntary skeletal muscle activity during the study^{198,213-215}
- Use of steady state data (coefficient of variation $\leq 10\%$)^{198,213-215}
- No general anesthesia within six to eight hours before the study¹⁹⁸
- Analgesics or sedatives for pain or agitation given at least 30 minutes prior to study^{198,213-216}
- Delay study for three to four hours after hemodialysis^{198,213,217}
- Delay study for one hour after painful procedures¹⁹⁸
- Delay study for unstable body temperature
- Routine care or activities avoided during the study^{198,214}

Indications for Indirect Calorimetry

Indirect calorimetry measurements are indicated when the use of predictive equations are inaccurate because of the patient's clinical condition, when patients fail to respond to nutrition support based on predictive equations, and when serial adjustments to the nutritional support plan are necessary as caloric requirements change during the stress response phases of critical illness. Use of this methodology and use of IC improves nutritional care and reduces complications associated with over or underfeeding.^{157, 212}

The conditions where caloric requirements estimated by predictive equations may be inadequate include: ^{157,198,207,212,214,218}

- Acute respiratory distress syndrome
- Chronic respiratory disease
- Large or multiple open wounds, burns

- Multiple trauma or neurologic trauma
- Multisystem organ failure
- Systemic inflammatory response syndrome, sepsis
- Postoperative organ transplantation
- Use of sedation and paralytic agents
- Altered body composition:
 - Limb amputation
 - Peripheral edema
 - BMI <18
 - BMI >30
 - Ascites

Use and Interpretation of Indirect Calorimetry Measurements

REE from indirect calorimetry is recognized as an accurate, objective, patient-specific reference standard for determining energy expenditure. Current recommendations are to decrease the reliance on predictive equations in critically ill patients.¹⁵⁷

REE measurements should be the targeted goal for nutritional calories in ICU patients without the use of correction factors for activity and metabolic stress. REE measures total caloric needs of the patient but does not distinguish protein from non-protein calories needed. Current practice recommendations are to provide adequate protein calories, as high as 1.5–2.5 g protein/kg/day in adults, balanced with carbohydrates and fats to meet >65-80% of goal calories predicted or measured by REE.¹²

Several preliminary studies show that preventing a cumulative nutrition deficit and a degree of tight calorie balance reduces mortality and may impact the consequences of overfeeding and underfeeding, such as increased ventilator days, infection rates, and length of stay.²¹⁸⁻²²⁴

Several studies have shown that a cumulative negative energy balance >10,000 kcal determined by IC measurements resulted in worse clinical outcomes. In a small multi-center study comparing nutritional support guided by predictive equations (control group) to measured energy requirements by IC (study group), the proportion of patients with a positive energy balance was higher in the study group and there was a significant difference in ventilator and ICU days when patients had a positive versus a negative energy balance (Table 12). Another study using IC reported a lower incidence of organ failure and mortality of 26% when the calorie deficit was < 10,000 kcal versus a mortality of 75% when the calorie deficit was > 10,000 kcal.²²² In a third study, there was a correlation with calorie deficit determined

Table 12. Impact of Energy Balance on Clinical Outcome²²¹

(*p < 0.005, ‡0.05)	Controls	Study	Vent Days	ICU days
Positive Energy Balance (PEB) (n=51)	24(69%)	27(84%)	10.6 ± 1.1	15.9 ± 1.6
Negative Energy Balance (NEB) (n=16)	11 (31%)	5(16%)	19.9 ± 3.9*	24.6 ± 4.0‡

Note: Negative energy balance was defined by > 10,000 kcal deficit.
Control patients received nutrition support per standard estimations of energy and protein (blinded to MREE and UUN). Study patients received nutrition support per daily MREE and UUN.

by IC and the development of pressure ulcers in nursing home patients. This correlation was stronger in patients where the negative energy balance exceeded 10,000 kcal.²²³ This data suggests that nutritional support guided by sequential monitoring and use of IC to maintain a positive energy balance may provide important clinical benefits (decreased mortality, ventilator and ICU days, infection rates, and complications rates).

Indirect Calorimetry Using Other Methods

Modifications to the Weir equation can be used to calculate REE. By substituting a calculated factor for either VO_2 or VCO_2 adjusted for a normal RQ, the REE based on VO_2 (REE- O_2) or VCO_2 (REE- CO_2) can be calculated where:

$$\text{REE-}\text{O}_2 = (3.9 \times \text{VO}_2) + (1.1 \times [\text{VO}_2 \times .85]) \times 1.44 \quad (5)$$

or

$$\text{REE-}\text{CO}_2 = (3.9 \times [\text{VCO}_2/.85]) + (1.1 \times \text{VCO}_2) \times 1.44 \quad (6)$$

When the actual RQ is equal to 0.85, both the REE- O_2 and REE- CO_2 equations will return a REE value equivalent to the value calculated by the standard Weir method. The CCM Express® metabolic analyzer (Medical Graphics Corporation, St. Paul, MN) uses Equation 6 to calculate REE- CO_2 with an accuracy of approximately +/- 10% compared to the REE.²²⁵

The REE measured by the CCM Express using the standard Weir equation was retrospectively compared to 116 calculated

REE- CO_2 measurements in 67 adult medical and surgical ICU patients.²²⁵ The correlation coefficient $r = 0.99$ and the coefficient of determination $r^2 = 0.98$, with bias and precision between measurements of 15 ± 126 kcal/day. When comparing the differences between REE to REE CO_2 to the measured RQ, there was a distinct pattern of agreement, whereby as RQ approached 0.70 (fat oxidation) the percent error (mean bias / mean REE for the range of RQ) became more positive, and as RQ approached 1.0 (carbohydrate oxidation) the percent error became more negative (Table 9). More importantly, when RQ was within the normal range of 0.80 to 0.90, the average error was approximately $\pm 5\%$ (Figure 15).

Other studies in mechanically ventilated adult patients have concluded that the REE CO_2 method when compared to IC measurements had a low level of agreement²²⁷ and were not sufficiently accurate.²²⁸ However, when VCO_2 -based calorimetry was compared to predictive equations, REE CO_2 methods were

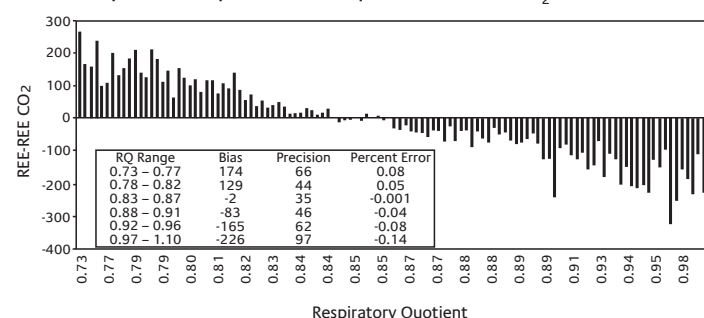


Figure 15. Comparison of REE and REE- CO_2 to RQ in 67 Patients²²⁶

more precise with higher accuracy rates within 10-15% of IC measurements^{229,230} and were associated with a lower rate of inaccuracy.²²⁹

In pediatric patients when the REE CO₂ method was compared to the IC measurements, it was concluded that IC was more accurate.^{231,232} Likewise, several studies confirmed that predictive equations are inaccurate in this population and have demonstrated that REE derived from VCO₂-based calorimetry can be more precise and superior to the reliance on estimations based on standard equations.^{232,233}

Use of REE CO₂ measurements relies on a choice of the RQ value used in the calculation. Investigators have used an RQ value based on the nutritional RQ (RQ_{macro}) of the 24-hr macronutrient composition delivered which included glucose and propofol.^{230,233} Substitution of the RQ_{macro} for RQ in Equations 5 and 6 where:

$$\text{REE-O}_2 = (3.9 \times \text{VO}_2) + (1.1 \times [\text{VO}_2 \times \text{RQ}_{\text{macro}}]) \times 1.44, \text{ and}$$

$$\text{REE-CO}_2 = (3.9 \times [\text{VCO}_2/\text{RQ}_{\text{macro}}]) + (1.1 \times \text{VCO}_2) \times 1.44$$

However, in both adults²³⁰ and children,²³³ use of the nutritional RQ failed to improve the accuracy of the REE CO₂ calculation.²²⁹

It has been suggested that the choice of RQ for VCO₂-based calorimetry of 0.85 is appropriate as this value is the midpoint of the normal physiological range (0.7– 1.0) between fat and carbohydrate oxidation.²²⁹ Furthermore, the RQ value of 0.85 is within the published cohort values of RQ in adults (0.76–0.89)²²⁹ and is close to the mean value of RQ (0.87) reported in study cohorts in children.²³³ In the study by Rousing, sensitivity analysis of different RQ values (0.70, 0.76, 0.85, 0.89 and 1.0) revealed that the RQ value of 0.85 had the lowest root mean square error (RMSE), (6%, range 6-17%) and the highest number of patients with accurate estimates within 10% of measured REE by IC (89%, range 22-89%).²²⁹

Additionally, the mean value of RQ for a given patient population can also be used in the REE CO₂ calculation.²³³ However in the Rousing study when the average RQ of the cohort (0.81) was compared a RQ of 0.85 to calculate REE CO₂, there was little difference in the mean REE from IC (-1% vs -4%), or RMSE (6% vs 7%), and no difference in the accuracy rate (89%).²²⁹

Recommendations for improving the accuracy REE CO₂ measurements include:

- Calibrating the CO₂ sensor at regular intervals and when necessary.
- Maintaining the CO₂ sensor interface free of secretions and condensation.
- Assessing the CO₂ waveform to confirm and validate accuracy of VCO₂ measurements.
- Allow adequate equilibration time between periods of instability and after adjustments in minute ventilation.
- Using 24 hour mean values of VCO₂ as opposed brief snapshot evaluation periods.

The ability to perform IC measurements using just a determination of VCO₂ has several important implications. VCO₂ determinations are technically easier to perform compared to VO₂. VCO₂ measurement capabilities are increasingly more available on stand-alone monitors and ventilators. This makes VCO₂-based calorimetry estimates of REE more accessible where metabolic analyzers are not available. Additionally, FiO₂ does not affect the accuracy of the REE-CO₂ calculation. Therefore, settings of FiO₂ > 0.60 may no longer be a limitation of measuring REE within a known range of acceptable error of ± 5–10%. This means that even the most severe critically ill patients on mechanical ventilation receiving 100% oxygen can have REE estimates performed to manage their complex nutritional needs using the REE-CO₂ method.²³⁴

Both the REE-O₂ and REE-CO₂ equations can be further simplified. By solving either equation with any combination of VCO₂ and VO₂ that equals an RQ of 0.85, and dividing the calculated REE by the measured VCO₂ or VO₂, a single factor can be derived for calculating REE-O₂ and REE-CO₂, whereby:

$$\text{REE-CO}_2 = 8.19 \times \text{VCO}_2 \quad (7)$$

$$\text{REE-O}_2 = 6.96 \times \text{VO}_2 \quad (8)$$

For example, when VCO₂ = 221 mL/min and VO₂ = 260 mL/min, RQ = 221/260 = 0.85, the Weir equation returns a calculate REE of 1810 kcal/day whereby:

$$\text{REE} = (3.9 \times 260) + (1.1 \times 221) \times 1.44 = 1810 \text{ kcal/day}$$

REE-CO₂ can be calculated as follows:

$$\text{REE-CO}_2 = (3.9 \times (221/0.85)) + (1.1 \times 221) \times 1.44 = 1810 \text{ kcal/day,}$$

$$\text{REE-CO}_2 \text{ Factor} = 1810/221 = 8.19, \text{ and}$$

$$\text{REE-CO}_2 = 8.19 \times 221 = 1810 \text{ kcal/day.}$$

Similarly, REE-O₂ becomes:

$$\text{REE-O}_2 = (3.9 \times 260) + (1.1 \times [260 \times .85]) \times 1.44 = 1810 \text{ kcal/day,}$$

$$\text{REE-O}_2 \text{ Factor} = 1810/260 = 6.96, \text{ and}$$

$$\text{REE-O}_2 = 6.96 \times 260 = 1810 \text{ kcal/day.}$$

The REE-O₂ and REE-CO₂ factors for any RQ value can also be determined where:

REE-CO₂ factor for RQ values of 0.80 and 0.90 are 8.60 and 7.82, and

REE-O₂ factor for RQ values of 0.80 and 0.90 are 6.88 and 7.04.

The MedGem® (Microlife USA Inc., Clearwater, FL) handheld calorimeter uses measurements of VO₂ and a calculation similar to Equation 8 above. The accuracy of this handheld device shares the same technical limitations and inaccuracies as traditional metabolic testing using IC and can only be performed on spontaneously breathing patients using a mouthpiece and nose clips.

The caloric equivalence of oxygen and carbon dioxide can also be used to indirectly calculate REE (Table 10). When the RQ = 0.90, the CO₂ or O₂ caloric equivalent factors equal 5.52 and 4.83 kcal/L respectively, where:

$$\text{REE-CO}_2 \text{ Equivalent} = 5.52 \times \text{VCO}_2 \times 1.44 \quad (9)$$

$$\text{REE-O}_2 \text{ Equivalent} = 4.83 \times \text{VO}_2 \times 1.44 \quad (10)$$

This technique has been compared to the Harris Benedict calculation and the Weir equation. The HBE significantly underestimated REE, but there was no significant difference between the Weir and REE-O₂ or CO₂ equivalent calculations.¹⁹⁹ Due to the technical complexity of measuring VO₂, the REE-CO₂ equivalent equation is simpler and should be the preferred method, especially when the FiO₂ is >0.60 or when small air leaks that would otherwise invalidate the VO₂ calculation are present.²⁰⁰

Additionally, VO₂ can be calculated using the reverse Fick equation when a pulmonary artery catheter is available and an accurate thermodilution cardiac output (CO), the oxygen

content of arterial (CaO₂), and mixed venous blood (CvO₂) can be determined.^{235,236}

The Fick equation is $\text{CO} = \text{VO}_2 / (\text{CaO}_2 - \text{CvO}_2)$ and therefore, VO₂ can be calculated where, $\text{VO}_2 = \text{CO} \times (\text{CaO}_2 - \text{CvO}_2)$.

The Weir equation is a superior method for IC measurements, especially when RQ is at or close to the physiologic extremes of 0.67 and 1.3. However, in the current ICU environment the physiologic extremes of RQ caused by prolonged starvation and excessive carbohydrate administration occur less frequently. Therefore, the use of a RQ value range between 0.85 and 0.90 to calculate REE using alternative methods is appropriate.

Since IC is not universally accessible, many ICUs still rely on predictive equations that have repeatedly demonstrated to be inaccurate. Despite its known limitations, when IC is unavailable, the REE CO₂ method is the best alternative for estimating REE than predictive equations.^{237,238}

The value of IC calculated using alternative methods is recognized by the recent *ESPEN Guideline on Clinical Nutrition in the Intensive Care Unit* recommendation 15, which states “If calorimetry is not available, using VO₂ (oxygen consumption) from pulmonary arterial catheter or VCO₂ (carbon dioxide production) derived from the ventilator will give a better evaluation on energy expenditure than predictive equations.”⁸⁰ Furthermore, the ASPEN guidelines for the pediatric critically ill patient acknowledge that the increased availability of bedside devices that provide measurements of VCO₂ may allow VCO₂-based calorimetry to replace predictive equations in mechanically ventilated patients.⁴⁴

Due to the cost and availability of metabolic analyzers or ventilators with an integrated function, and the technical problems associated with measuring VO₂, REE measurement based on VCO₂ is an attractive alternative. The accuracy of equations 6, 7, and 9 above compared to the Weir equation are¹⁹⁹⁻²⁰¹ within clinically acceptable limits needed for monitoring nutritional¹⁹⁹ interventions. Since VCO₂ monitoring is becoming more readily available, can be performed on any FiO₂, and is less costly than traditional metabolic testing, its use should be considered for incorporation into a standard nutrition assessment and treatment process where IC is unavailable or is restricted by limited resources. Additional validation studies and outcome measurements are needed to determine the true impact of these alternative methods of indirect calorimetry.

Clinical Practice Recommendations for Nutritional Support

Several clinical practice guidelines from different organizations for the various aspects of nutritional assessment and treatment have been developed. The Society of Critical Care Medicine and the American Society of Parenteral and Enteral Nutrition (SCCM/ASPEN), the European Society for Clinical Nutrition and Metabolism (ESPEN), the Academy of Nutrition and Dietetics (AND), the Canadian Clinical Practice Guideline for Nutritional Support (CCPG), and the European Society of Pediatric and Neonatal Intensive Care (ESPNIC), and SCCM/ASPEN Guidelines for the Provision and Assessment of Nutrition Support Therapy in the Pediatric Critically Ill Patient have developed best practice recommendations based on the interpretation of available evidence, consensus agreement, and expert opinion.^{12,44,80,104,239,240} The present concentration on evidence-based practice dictates that guidelines be supported by the available literature. The problem with multiple guidelines from different professional societies is that they often contradict one another (Table 13). Varying degrees of agreement, disagreement, and controversy over the strength of the evidence can be confusing to the clinician. The review and interpretation of practice recommendations, knowledge of the current available literature, clinical judgment, the specific patient population, and the needs of the individual patient should drive the translation of recommendations into clinical practice.

The following is a summary of some of the best practice recommendations from the various organizations:

- Nutritional support should be initiated early within the first 24–48 hours in critically ill patients. Primary goals of nutritional support and care are to:
 - Preserve and maintain lean muscle mass
 - Provide continuous assessment, reassessment, and modification to optimize outcome
 - Monitor the patient for tolerance and complications, such as refeeding syndrome
 - Prevent protein energy malnutrition by giving higher protein content while providing adequate total calories
 - Monitor nutrition goals and target achievement rate of >65% within the first week
 - Prevent accumulation of a caloric deficit
- Indirect calorimetry is the gold standard for determining energy requirements and should be used when available. Predictive equations are known to be inaccurate. Alternative methods of calculating REE are available.
- Current EN practice recommendations are to:
 - Consider using nutritional risk screening and assessment tools
 - Preferentially feed via the enteral route
 - Initiate EN within 24–48 hours
 - Reduce interruptions of EN for nursing care and procedures to prevent underfeeding
 - HOB elevation to reduce aspiration risk
 - Avoid routine measurement of GRV in adult and pediatric patients
 - Accept GRV up to 500 mL in adults before reducing or stopping EN in the absence of clear signs of intolerance
 - Use of motility agents to improve tolerance and reduce GRV when indicated
 - Promote post-pyloric feeding tube placement in adults when feasible
 - Gastric feeding is safe and is the preferred route for EN in most pediatric patients
- Current PN practice recommendations are to:
 - Only use PN when enteral route not feasible
 - Use PN based on the patient's nutritional risk classification for malnutrition
 - Initiate PN early if nutritional risk is high
 - Delay PN up to seven days if nutritional risk is low
 - Convert to EN as soon as tolerated to reduce the risks associated with PN
- Use of trophic or “trickle feeding” may be beneficial in adults, children, and infants
- Use of pharmaconutrients and immunonutrition:
 - Increase omega-3 fatty acids to omega-6 fatty acid ratios
 - Use of arginine, glutamine, nucleotides, antioxidants, and probiotics may be beneficial
 - Avoid using arginine in patients with severe sepsis

- In pediatric patients:^{44,103}
 - EN is the preferred route of nutrition
 - Initiate EN within 24-48 hours of admission to PICU
 - A stepwise algorithmic approach to advancing EN should be utilized
 - Goal is to achieve at least 2/3 of nutrient goal within the first week of admission
 - IC studies should be utilized when available to determine energy requirements
 - Routine measurement of GRV is not recommended
 - Either gastric or post-pyloric feeding can be utilized in the majority of children
 - Measurement of anthropometrics expressed in Z scores should be obtained on admission and regularly throughout the hospital course
 - Immunonutrition is not recommended for the critically ill child
 - Timing for PN initiation should be individualized with the below general guidance:
 - In cases of severe malnutrition, supplemental PN may be provided in the first week of admission if unable to advance beyond low volumes of EN
 - When EN is unable to be initiated in the first week of admission, PN may be provided

Table 13. Summary of Clinical Practice Guidelines for Nutrition Therapy in Critically Ill Adult Patients^{12,80,239,240}

Topics	ASPEN/SCCM	AND	Canadian	ESPEN
Nutritional Risk Screening/Assessment Tool	NRS 2002/ NUTRIC performed on all admissions when insufficient intake is anticipated	General clinical assessment	General clinical assessment	General clinical assessment for malnutrition until a specific tool has been validated.
Use and Timing of EN	24-48 hours following admission to ICU	24-48 hours following admission to ICU	24-48 hours following admission to ICU	Within 48 hours
Method for determining REE	IC when available or predictive equations or weight-based equation	IC when available or predictive equations for non-obese PSU (2003b), obese PSU(2010)	Insufficient data to make a recommendation on the use of indirect calorimetry vs. predictive equations	IC when available. If IC not available, using VO_2 or VCO_2 methods will give a better evaluation on REE than predictive equations.
Dose of Nutrition and Achieving Target (trophic or hypocaloric feedings)	Provide >50-65% of goal calories over first week. Low nutritional risk patients do not require specialized nutrition therapy over the first week of hospitalization in the ICU. High nutritional risk patients (severely malnourished) should be advanced toward goal over 24-48 hours to achieve >80% of goal energy and protein within 72 hours.	At least 60% of total estimated energy requirements within the first week.	No specific recommendation. Strategies to optimize delivery should be considered.	Hypocaloric nutrition <70% of REE in the early phase of acute illness. Increase up to 80-100% after day 3.

Protein target per day	1.2 – 2.0 g/kg actual body weight BMI 30-40 2.0 g/kg BMI > 40 2.5 g/kg	Consider hypocaloric, high protein feeding in obese patients	Insufficient data, No specific recommendation	1.3 g/kg adjusted body weight
EN: Arginine	Recommended in immune modulating formulas except in patients with sepsis	In patients without ARDS and sepsis immune modulating formulas containing arginine should be carefully evaluated	Not recommended	No recommendation
EN/PN: Fish Oil	EN: recommended in post-operative patients EN: cannot recommended for ARDS due to conflicting data	EN: in patients without ARDS and sepsis immune modulating formulas containing fish oil should be carefully evaluated	Should be considered for ARDS	PN: can be provided in patients receiving lipid emulsions.
EN/PN: Glutamine	EN/PN: should not be routinely used in critically ill patients	EN: in patients without ARDS and sepsis immune modulating formulas containing glutamine should be carefully evaluated PN: should be considered to reduce infectious complications	EN/PN: recommends that glutamine not be used in critically ill patients.	EN: use in burn patients, consider in trauma patients PN: should not be use in patients with liver and renal failure Consider in burn and trauma patients
EN: High Fat, Low CHO	Not recommended	No specific recommendation	Insufficient data	No specific recommendation
EN: Gastric Residual Volume (GRV)	Recommends not using GRV for routine monitoring. ICUs that continue this practice should the 200-500 mL range to trigger interventions toward minimizing the risk of aspiration and should avoid the mandatory interruption of EN	If GRV is used as an indicator for intolerance, EN should not be held if GRV is less than 500 ml in the absence of abdominal distention, nausea, vomiting.	GRVs 250 or 500 mLs (or somewhere in between) and checking q4 or q8 hours should be considered as a strategy to optimize delivery	No specific recommendation

EN: Motility agents	Use in patients at high risk of aspiration	If history of gastroparesis or high GRV	Recommended with EN intolerance	Recommended with EN intolerance
EN: Small bowel feeding	Recommends small bowel feeding tube placement in patients at risk for aspiration	Recommends small bowel feeding tube placement	Recommends the routine use of small bowel feeding tubes	Recommends small bowel feeding tube placement in patients with gastric feeding intolerance
EN: Body Position	HOB 30-45 degrees	HOB 30-45 degrees	HOB >45 degrees	No specific recommendation
EN: Prebiotics/Probiotics/Synbiotics	Probiotics recommended for patients with severe acute pancreatitis	No specific recommendation	Consider probiotics	No specific recommendation
EN: Continuous vs bolus	Continuous EN for high risk patients or those shown to be intolerant to bolus EN	No specific recommendation	Insufficient data. No recommendation	Continuous EN should be used
EN with PN	In low or high nutritional risk supplemental PN after 7-10 days	No specific recommendation	Supplemental PN should not be started until strategies to maximize EN delivery have been attempted	EN plus supplemental PN should not be instituted until 3-7 days to avoid overfeeding
Parenteral Nutrition	Low nutritional risk exclusive PN after 7 days High nutritional risk, severe malnutrition when EN not feasible start PN as soon as possible	No specific recommendation	PN should not be used routinely, but early PN should be considered in high nutritional risk patients with EN contraindications Not recommended	Low nutritional risk, EN is contraindicated, PN should be instituted in 3-7 days EN is contraindicated, patient is severely malnourished, early PN should be initiated
PN: Lipid Emulsions	Avoid omega-6 soy-based lipid in the first week	No specific recommendation	Reduce omega-6 load; insufficient data on type	Integral part of PN for energy; varying lipid emulsions available in Europe
Intensive Insulin Therapy	Glucose 140-180 mg/dl	Glucose 140-180 mg/dl	Glucose 140-180 mg/dl	Glucose >180 mg/dl

Summary

Nutritional support is important in the care of patients with acute and chronic illness. The prevention of malnutrition and the maintenance of appropriate nutritional care brings with it the potential for reducing morbidity and mortality, shortening the duration of mechanical ventilation and the length of hospital stay, and lowering health care costs while improving functional quality of life. Appropriate nutritional management is best achieved by using a comprehensively designed nutritional care process supported by the best available evidence. This process should include an interdisciplinary team approach and organizational standards of care with policies and procedures that ensure implementation, continuous assessment, and monitoring of the nutrition care plan.

Varying degrees of agreement, disagreement, and controversy from various organizations regarding nutritional support include the role and timing of enteral versus parenteral route, positioning of feeding tubes, thresholds for GRV, and use of predictive equations and indirect calorimetry. The use of immunonutrition and dietary supplements continue to evolve as practice changes develop when new evidence becomes available. All members of the integrated health care team should maintain awareness of the importance and continued evolution of best practices for nutritional assessment and treatment. Optimizing nutritional support and care of the critically ill and patients with acute and chronic respiratory disorders will contribute to improved outcomes and reduced health care costs.

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Acronyms

AARC	American Association for Respiratory Care	PICC	peripherally inserted central catheter
AMA	arm muscle area	PN	parenteral nutrition
AND	Academy of Nutrition and Dietetics	RDA	recommended dietary allowance
ARDS	acute respiratory distress syndrome	REE	resting energy expenditure
ASPEN	American Society of Parenteral and Enteral Nutrition	RMR	resting metabolic rate
		RQ	respiratory quotient
BIA	bioelectrical impedance analysis	RR	respiratory rate
BMI	body mass index	SBS	short bowel syndrome
BMR	basal metabolic rate	SCCM	Society of Critical Care Medicine
RQ	respiratory quotient	TMAX	temperature maximum (degrees Celsius)
BSA	body surface area	RQ	respiratory quotient
CCPG	Canadian Clinical Practice Guidelines for Nutritional Support	UUN	urinary urea nitrogen
		VCO₂	carbon dioxide production
CF	cystic fibrosis	VO₂	oxygen consumption
CNSC	Certified Nutrition Support Clinician	V_T	tidal Volume
CO₂	carbon dioxide		
COPD	chronic obstructive pulmonary disease		
CRP	C-reactive protein		
CT	computed tomography		
DRI	dietary reference intakes		
DXA	dual-energy X-ray absorptiometry		
EGP	endogenous glucose production		
EN	enteral nutrition		
ESPEN	European Society for Clinical Nutrition and Metabolism		
FFQ	food frequency questionnaires		
GI	gastrointestinal		
GRV	gastric residual volumes		
HBE	Harris Benedict Equation		
HHS	Health and Human Services		
HOB	head of bed		
IBD	inflammatory bowel disease		
IC	indirect calorimetry		
kcal	kilocalories		
MAC	mid-upper arm circumference		
MRI	magnetic resonance imaging		
PEEP	positive end-expiratory pressure		
PEG	percutaneous endoscopic gastrostomy		
PEM	protein-energy malnutrition		