Metabolic assessment and enteral tube feeding usage in children with acute neurological diseases

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Objective: To report on acquired experience of metabolic support for children with acute neurological diseases, emphasizing enteral tube feeding usage and metabolic assessment, and also to recommend policies aimed towards improving its implementation.

Design: Retrospective analysis. Setting: Pediatric Intensive Care Unit of Hospital do Servidor Público Estadual de São Paulo. Subjects: 44 patients consecutively admitted to the Pediatric ICU over a period of 3 years who were given nutrition and metabolic support for at least 72 hours. Head trauma, CNS infections and craniotomy post-operative period following tumor exeresis were the main diagnoses. Measurements: Records of protein-energy intake, nutrient supply route, nitrogen balance and length of therapy.

Results: From a total of 527 days of therapy, single parenteral nutrition was utilized for 34.3% and single enteral tube feeding for 79.1% of that period. 61.4% of the children were fed exclusively via enteral tube feeding, 9.1% via parenteral and 39.5% by both routes. The enteral tube feeding was introduced upon admission and transpyloric placement was successful in 90% of the cases. Feeding was started 48 hours after ICU admission. The caloric goal was achieved on the 7th day after admission, and thereafter parenteral nutrition was interrupted. The maximum energy supply was 104.2 ± 23.15 kcal/kg. The median length of therapy was 11 days (range 4-38). None of the patients on tube feeding developed GI tract bleeding, pneumonia or bronchoaspiration episodes and, of the 4 patients who were given exclusive TPN, 2 developed peptic ulcer. The initial urinary urea nitrogen was 7.11 g/m² and at discharge 6.44 g/m². The protein supply increased from 1.49 g/kg to 3.65 g/kg (p< 0.01). The nitrogen balance increased from -7.05 to 2.2 g (p< 0.01).

Conclusions: Children with acute neurological diseases are hypercatabolic and have high urinary nitrogen losses. The initial negative nitrogen balance can be increased by more aggressive feeding regimes than the usual ones. Early tube feeding was well tolerated, which permits the conclusion that it is a safe and effective method for nutrition support. Recommendations of basic rules for metabolic support are made.


INTRODUCTION

In spite of the general knowledge that central nervous system (CNS) diseases can lead to impairment of nutritional status, up until the last decade only a few studies had demonstrated the need for implementation of early nutritional therapy in these cases.1

Until then, the recommended method was the maintenance of fasting until full recovery of gastrointestinal function, which occurs at around one week after the onset of the injury. The nutritional support has to be instituted early, since these patients are considered at risk of malnutrition. In addition to the autophagic process resulting from the metabolic stress,2 there is the difficulty in delivering enteral nutrients to patients following head injury.

Impossibility of feeding by mouth and hypercatabolism are indications for nutritional support which, in its turn, can be a determining factor in the prognosis.3 Patients with CNS acute diseases are frequently in coma or have their swallowing reflexes impaired and need parenteral nutrition or enteral tube feeding.4,5 Even

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though the evidence favors enteral nutrition, due to its lower risk of complications and low cost, problems such as alteration of consciousness level and gastric motility abnormalities can delay the beginning of enteral tube feeding, leading to the more frequent use of parenteral nutrition in the first week following the neurological injury. In view of the inherent difficulties in nutrient administration and with the aim of contributing to the study of problems related to nutritional and metabolic therapy in children with CNS acute diseases, this study was developed to describe experience obtained with nutritional therapy in children with acute neurological diseases in a Pediatric Intensive Care Unit (PICU), with emphasis on the metabolic assessment and pattern of enteral nutrition, as well as recommending policies aimed at improving its implementation.

METHODS

Medical records of all patients with acute neurological diseases admitted to the PICU of Hospital do Servidor Público Estadual de São Paulo between June 1992 and January 1996 were analyzed. Of the total of 63 patients admitted during this period, the study included 44 who received nutritional therapy for at least 4 days. The remaining patients were excluded on the grounds of being discharged after less than 72 hours (11 patients) or having incomplete records. The age range was from 3 to 144 months (median: 68 months). Mechanical lung ventilation was used in 28 patients. The median Glasgow score of the patients with a diagnosis of head trauma and CNS infection at admission was 7 (range: 4 to 13). Table 1 shows the diagnosis at admission to the PICU.

The daily records of protein-energy delivery, initial nutritional assessment and metabolic monitoring were analyzed, from PICU admission until discharge. The nutritional assessment, performed within the first 72 hours after PICU admission, considered the weight/height measurements which were compared to NCHS standards. Nutritional status was obtained according to Waterlow’s criteria. The metabolic assessment was performed by determination of urinary urea nitrogen and nitrogen balance in those patients from whom it was possible to obtain 24 hour urine at admission and discharge from the PICU.

Urinary urea nitrogen determination was performed by the automated urease method. Nitrogen balance, obtained from the difference between nitrogen delivery and urinary losses, plus the estimated cutaneous and fecal losses, was calculated according to Wilmore. The excretion of urinary urea nitrogen by the subjects with acute neurologic disease was compared to that obtained from other patients admitted with a diagnosis of sepsis who were followed up during the same period.

All patients received nutritional support, administered according to the unit’s routine, following the standards of the American Society for Parenteral and Enteral Nutrition. Nutritional support is given to every child admitted into the unit, as of the moment they are hydrated and are in hemodynamic stability, initially given by the parenteral route. Parenteral nutrition is administered when enteral access cannot be obtained, when enteral nutrition support fails to meet nutritional requirements, and when feeding into the gastrointestinal tract is contraindicated. Solutions with dextrose concentration up to 12.5% are administered via peripheral vein, and those containing higher concentrations are infused via the central vein. Solutions are administered with lipid emulsions mixed in the same vial. When the digestive route cannot be used (due to presence of ileum, digestive hemorrhage or abdominal bloating) the nasoduodenal route is associated to the parenteral. Complete polymeric iso-osmolar diets are employed, with a nitrogen to non-protein calorie ratio varying from 1:150 to 1:100, and a caloric content of 1.0 kcal/ml. When there is intolerance, predigested commercial formulae with glucose polymers and protein hydrolyzates are used.

The placement of polyurethane tubing is as follows. The patient is placed in elevated decubitus at 30°, with prior administration of metoclopramide at 0.2 mg per kg of body weight. After the tube is placed in the stomach, 20-30 ml of air are injected, with the patient maintained in lateral right decubitus to facilitate tube migration. Adequate post-pyloric position is confirmed by abdominal roentgenogram showing the tip of the tube placed between the second duodenal segment and the jejunum. The diet is administered by infusion pump, with the patient maintained

<table>
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<tr>
<th>Diagnosis</th>
<th>Frequency (%)</th>
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<tbody>
<tr>
<td>Head injury</td>
<td>18 (40.9%)</td>
</tr>
<tr>
<td>CNS infection</td>
<td>12 (27.3%)</td>
</tr>
<tr>
<td>CNS Tumor</td>
<td>8 (18.2%)</td>
</tr>
<tr>
<td>Miscellanea *</td>
<td>6 (13.6%)</td>
</tr>
<tr>
<td><strong>Total</strong></td>
<td><strong>44</strong></td>
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*Guillain Barré syndrome (4), spinal cord injury by gunfire (1), intracranial hypertension (1).
in a right lateral position or raised at 30° decubitus. The energy delivery is calculated according to Holliday & Segar’s rule.\textsuperscript{11} Delivery is initiated at ¼ of the total daily volume of estimated energy needs, increasing to half on the second day, ¾ on the third, and reaching the full provision by the fourth day. This scheme may be modified according to the patient’s tolerance.

Statistical analysis

Considering the skewed distribution of the variable values, nonparametric methods were applied. The Kruskal-Wallis variance analysis\textsuperscript{12} was used for comparing urinary urea nitrogen in the various disease groups. The Wilcoxon test\textsuperscript{12} was used for comparing values of nitrogen balance, urinary nitrogen losses and protein-caloric delivery at admission and discharge of the PICU. The Mann-Whitney test was used for comparing values of energy delivery according to the use of mechanical lung ventilation. Data analysis was performed on a CSS computer package (Statsoft, Inc., 1991). The level of significance was set at 5\% for all analyses. Data were presented as median values and range and as mean ± SD.

RESULTS

Table 2 shows the nutritional status of the patients at time of admission to the PICU. Of the 44 patients studied, 27 received exclusively enteral tube feeding, 4 received exclusively parenteral nutrition and 13 received both forms of therapy. The enteral route was used for 79.9\% and the parenteral route for 34.3\% of a total period of 527 days of nutritional therapy.

The maximum energy delivery during hospital stay was 104.2 ± 23.15 kcal/kg, without significant variation between the different groups of neurological diseases. The planned energy amount was administered in its totality to 32 patients. No differences in energy supply was observed when patients who were under pulmonary mechanical ventilation (92.1 ± 21.2 kcal/kg) were compared to those who were not (105.2 ± 25.9 kcal/kg; z calculated = 1.5, z critical = 1.96).

The median period for nutritional support was 11 days, with variation from 4 to 38 days. The enteral tube was inserted on the admission day, with successful migration to the duodenum in 90\% of the cases. The median length of time between admission and the beginning of postpyloric enteral feeding was 2 days, with one day being the necessary time for adequate tube placement. The estimated energy needs were satisfied by the seventh day after admission (Figure 2), when parenteral nutrition was suspended. No return to parenteral nutrition was needed in any of the cases and the transition was to the oral route in all cases.

The complications observed in the course of enteral tube feeding are shown in Table 3. None of the children submitted to enteral nutrition developed pneumonia, bronchoaspiration episodes, nor gastrointestinal tract bleeding. Of the four patients that received exclusively parenteral nutrition, two developed peptic ulcer, confirmed by endoscopy.

The comparison between excretion of urinary urea nitrogen by patients with different CNS diseases at admission and by patients with a diagnosis of sepsis is shown in Figure 3. Although the median value for septic patients (10.12 g/m\textsuperscript{2}) was higher than for patients with head trauma (6.91 g/m\textsuperscript{2}), with CNS infection (8.5 g/m\textsuperscript{2}) and those in the craniotomy post-operative period following tumor exeresis (8.15 g/m\textsuperscript{2}) there was no significant difference between the groups (calculated H=3.7; critical X\textsuperscript{2} =7.81; p = 0.28). There was a significant increase in nitrogen balance, which went from -7.05 g/m\textsuperscript{2} at admission to 2.2 g/m\textsuperscript{2} at the time of discharge from the unit (p < 0.01), as shown in Figure 4.

The urinary excretion of urea nitrogen at admission had a median value of 7.11 g/m\textsuperscript{2} and of 6.44 g/m\textsuperscript{2} at discharge, which was not a significant variation. On the other hand, protein delivery changed from 1.49 g/kg/day at admission (range zero to 2.31 g/kg/day) to 3.65 g/kg/day (range 0.43 to 6.87 g/kg/day), thus showing a significant increase (p < 0.01).

Table 2

<table>
<thead>
<tr>
<th>Nutritional status at admission.</th>
<th>Frequency</th>
<th>%</th>
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<tbody>
<tr>
<td>Eutrophy</td>
<td>33</td>
<td>75%</td>
</tr>
<tr>
<td>Wasted</td>
<td>3</td>
<td>6.8%</td>
</tr>
<tr>
<td>Wasted and stunted</td>
<td>5</td>
<td>11.4%</td>
</tr>
<tr>
<td>Stunted</td>
<td>3</td>
<td>6.8%</td>
</tr>
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Table 3

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<thead>
<tr>
<th>Complications of enteral tube feeding.</th>
<th>Frequency</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Diarrhea</td>
<td>6</td>
<td>15%</td>
</tr>
<tr>
<td>Tube obstruction</td>
<td>4</td>
<td>10%</td>
</tr>
<tr>
<td>Abdominal bloating</td>
<td>4</td>
<td>10%</td>
</tr>
<tr>
<td>Metabolic disturbance *</td>
<td>2</td>
<td>5%</td>
</tr>
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</table>

* Hypernatremia
DISCUSSION

Patients with acute CNS disease present problems that, although not being specific, make nutritional metabolic therapy more difficult. CNS injury initiates a metabolic response similar to that which occurs with patients with sepsis or trauma. This response is characterized by hypermetabolism, hypercatabolism, altered vascular permeability, and altered GI function.

Hypercatabolism may lead to acute malnutrition which, depending on the degree and period of stress and on the previous conditions of the patient, can result in a significant loss of body mass. This metabolic response may vary according to the specific clinical situation. In the present study, the patients presented initial urinary urea nitrogen values of variable although generally high amplitude, which resulted in negative nitrogen balances at admission. This was observed in children with traumatic, infectious or inflammatory lesions.

There is great variability of metabolic response according to the clinical situation. Patients with similar Glasgow scores may present different nitrogen losses. Even in the case of identical CNS lesions, the complications and therapeutical approaches may interfere with the magnitude of the metabolic response. Sepsis, therapy using steroids, sedatives and neuromuscular blocking agents may also affect nitrogen losses and energy expenditure. As in other situations, there are individual variations in energy and oxygen consumption. Patients with decerebration and decortication have greater energy needs, whilst the needs are smaller in patients in barbituric coma or who are tetraplegic.

Studies in adult patients with head trauma have shown increases of over 50% in basal energy expenditure. Weekes and Elia observed energy expenditure increased by 30-35% above the estimate and that this increase was directly related to temperature and interruption in sedation. It has been suggested that energy needs do not necessarily increase, since the decrease in physical activity that accompanies the disease is sufficient to counterbalance this effect and, thus, the increase does not really occur.
In the present study, the increased loss of urinary urea nitrogen observed in children with Guillain-Barré syndrome suggests the existence of a hypercatabolic state, probably triggered by endocrine and inflammatory components of the disease. An increase in urinary urea nitrogen excretion was also observed by Roubenoff et al in adults with Guillain-Barré syndrome. In their study, the authors drew attention to the false impression that patients suffering from this disease were maintaining good nutritional status, a fact that often delays the implementation of adequate nutritional support.

The nitrogen balance may be a useful tool for tailoring protein supply to the individual needs of each patient, as has been shown in our study. In fact, the increase in nitrogen balance was made possible by the significant increase in protein delivery, since no variation in nitrogen loss was detected between the start and end points of the assessment. The increase in protein delivery is aimed at minimizing nitrogen losses and at partially compensating the hypercatabolism. Even though the supply of aminoacids may increase protein synthesis, it is not sufficient to reach protein anabolism in the short run. In children with head trauma or with severe CNS infection, a nitrogen to non-protein calorie ratio of 1:150 to 1:100 is recommended.

Patients with acute CNS disease are frequently comatose or have impairment of swallowing reflexes, generally receiving parenteral or enteral tube feeding. Enteral nutrition presents several obvious advantages over parenteral nutrition, since it is more physiological, its costs are lower and it presents a lower incidence of complications. Patients with acute neurological disease present clinical conditions than clearly, more than in any other disease, justify the indication of enteral tube feeding: 1) the presence or risk of malnutrition, in which the oral route is insufficient to avoid weight loss; and 2) neuromuscular impairment with a high probability of bronchoaspiration of gastric contents.

Clinical studies have demonstrated improvement in immunocompetence and survival in neurological patients who were given early nutritional support, and that nutrient delivery by the enteral route is safe when there is viability of the digestive tract. Sachs et al observed improvement in immunological function, assessed by the CD-4/CD-8 relationship and tests stimulating T-lymphocyte proliferation in patients with brain injury who had received early enteral nutrition.

Enteral tube feeding must be an obligatory component of the therapeutic plan for patients with acute neurological diseases and it may be administered by an intragastric or post-pyloric route. The intragastric route requires that the swallowing and coughing reflexes be preserved, that the esophageal sphincter be functional and that gastric motility be maintained. The post-pyloric route must be employed in those patients who present impaired protection of the respiratory tract or delayed gastric emptying, situations which give rise to greater probability of bronchoaspiration. Patients under metabolic stress frequently present some degree of gastric atony. Factors such as sepsis, increase in intracranial pressure and the use of drugs such as opiates and dopamine, amongst others, affect the hormonal control of GI tract motility, abolishing the activity of migratory motor complexes in the gastric antrum.

Saxe et al, studying head-injured patients, observed dysfunction of the lower esophageal sphincter, suggesting that intra-gastric enteral nutrition would not be a safe method for nutrient delivery. It has also been suggested that regurgitation is a common event in head-injured patients who receive intra-gastric tube feeding and that the use of the post-pyloric route presents advantages over the intra-gastric route in these patients. Thus, in situations where there is a high probability of delayed gastric emptying, feeding tubes placed beyond the pylorus allow for the provision of higher volumes of the diet at an earlier time, with a lower risk of aspiration or bronchopneumonia, when compared to the intragastric route.

In the present study, enteral tube feeding was initiated two days after patient admission, constituting early beginning as defined in the literature. The authors took into account that the median period of seven days needed to reach the planned energy delivery may be deemed short, considering the high proportion of patients with head injury or in the craniotomy post-operative period following brain tumor exeresis, situations in which sedatives for decreasing digestive tract motility are frequently used. A study with adult head-injured patients showed a median time of 11.5 days for achieving the planned energy delivery. There are no references regarding this particular aspect in pediatric studies.

Polymeric enteral diets are generally well tolerated, although the use of pre-digested ones, based on hydrolyzed protein, may be indicated in an initial phase of the treatment. This type of diet is preferred when there is persistent diarrhea or sepsis, situations in which there is reduction in absorptive surface and increase in permeability of the intestinal mucosa. The frequency of diarrhea varies from 5 to 32% and it presents multifactorial etiology. Osmolality and diet infusion mode, bacterial contamination, hypoalbuminemia, use of antiacids and antibiotics and intestinal mucosa lesions are the most frequently reported factors.

The low frequency of GI tract complications in this study could be attributed to the early implementation of...
feeding by the digestive route, which may improve
tolerance to the nutrients and, by strengthening the
trophism of the intestinal epithelium, may act as a
prophylactic against mucosa lesions. It is significant that
none of the patients fed by the enteral route developed
peptic ulcer, a complication that occurred in two of the
four patients on exclusively parenteral nutrition. The
association of stress and starvation promotes alterations
in the function and trophism of the intestinal mucosa,
characterized by decrease in secretory IgA and bacterial
adherence of inflammatory macromolecules. Nutrition
by the digestive route thus acquires great importance as a
protective measure for preventing alterations to the
integrity of the intestinal mucosa, and to the barrier
function it performs during stress.

Metabolic complications like hypermetatremia can occur
in infants who have received diets formulated for adult
patients which, due to their greater concentration of
electrolytes, are not appropriate for use in pediatric patients.
Severely undernourished patients may undergo
hypophosphatemia and hypopotassemia when they reach
nutritional recovery, needing supplementation of these ions.

Based on evidence presented in the literature and on
acquired experience, the authors suggest that early
nutritional regimens using small bowel feeding should be
attempted whenever possible. If there is hemodynamic
stability and if the digestive tract is functioning, enteral
nutrition can be beneficial with the use of adequate diets,
administered by continuous infusion. When fluid
restriction is indicated, formulae with higher energy
content may be used. Since nutrient delivery by the enteral
route does not initially fulfill the needs of acutely ill
patients, the concomitant administration of parenteral
nutrition is generally needed in this phase, while the enteral
nutrition promotes the stimulation of the GI tract. The
nursing staff should be educated regarding the risk of
aspiration and poor gastric emptying in enteral tube-fed
patients.

Monitoring of metabolic stress and response to the
administration of nutrients must be performed at regular
intervals, providing means of verifying the efficacy of the
therapy and the degree of intolerance to nutrient delivery,
adjusting it according to the patient’s needs. The use of
formulae based on the determination of urea nitrogen in
the 24 hour urine is useful for estimating the total nitrogen
loss. Urea nitrogen has a good correlation with total
nitrogen and it has been used in evaluating the degree
of metabolic stress and planning the protein offer. The
amount of nitrogen excreted in the urine reflects the
magnitude of the breakdown of muscular protein and the
degree of hypercatabolism.

The nitrogen balance, obtained from the difference
between the losses and the administration of nitrogen, in
spite of the difficulties inherent in serial collection of 24
hour urine, is still the simplest and quickest method for
the assessment of protein metabolism in the short run, as
well as for evaluating the efficacy of nutritional support
in hypercatabolic patients. Glycemic levels must be
routinely monitored, due to the risk of neurological lesions
and possible association with worsened prognosis in head
injuries. The serum levels of triglycerides in critically
ill children receiving intravenous lipid emulsions must also
be monitored, since in this situation there may be a
reduction in lipid clearance, with deposition in the vascular
endothelium and reticulo-endothelial system. The serum
levels of triglycerides must be determined before the start
of infusion and whenever there is an increase in lipid
delivery, maintaining levels below 200 mg/dl.

The results of this study have demonstrated that
patients with acute neurological diseases are
hypercatabolic and that the negative nitrogen balance can
be reversed by more aggressive nutritional support than
that commonly used. Early enteral nutrition was well
tolerated, demonstrating that it is a safe and effective
method for the administration of nutrients. It should be
emphasized, however, that the success of enteral nutrition
is directly related to the motivation and performance of
the medical, nursing and nutritional team. Awareness of
the advantages of early use of enteral tube feeding, its
administration using correct techniques and appropriate
diets, accompanied by continuous monitoring and care by
nursing personnel, are essential factors in improving the
quality of nutritional support in children with acute
neurological diseases.

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Resumo

Objetivos: Relatar a experiência com a terapia nutricional e metabólica em crianças com neuropatias agudas - com ênfase na avaliação metabólica e no uso da nutrição enteral - e propor medidas mínimas para a sua implementação. Pacientes e Métodos: Análise retrospectiva do padrão de terapia nutricional e metabólica de todos os pacientes com neuropatia aguda consecutivamente admitidos na UTI durante um período de três anos. Traumatismo craniano, infecção do SNC e pós-operatório de retirada de tumor cerebral foram os principais diagnósticos. Resultados: Dos 44 pacientes estudados, 61,4% receberam nutrição enteral exclusiva, 9,1% nutrição parenteral exclusiva e 29,5% por ambas as vias. A via enteral foi utilizada em 79,1% e a via parenteral em 34,3% do tempo total de 527 dias de terapia. A sonda enteral era introduzida na admissão, ocorrendo migração para o duodeno em 90% dos casos. A nutrição enteral pós-pilórica iniciou-se dois dias após a internação. As necessidades energéticas estimadas foram satisfeitas no sétimo dia de admissão quando era então suspensa a nutrição parenteral. A oferta energética máxima foi 104,2 ± 23,15 kcal/kg. O tempo mediano de terapia foi de 11 dias (variação de quatro a 38 dias). Nenhuma criança em nutrição enteral desenvolveu hemorragia digestiva, pneumonia ou episódio de broncoaspiração e, dos quatro pacientes que receberam nutrição parenteral exclusiva, dois desenvolveram úlcera péptica. A excreção urinária de nitrogênio uréico foi elevada, sendo 7,11 g/m² na admissão e 6,44 g/m² na alta. A oferta protéica passou de 1,49 g/kg/dia para 3,65 g/kg/dia (p< 0,01) e o balanço nitrogenado de -7,05 g na admissão para 2,2 g quando da alta da unidade (p< 0,01). Conclusões: As neuropatias agudas levam a um estado ao hipercatabólico, com elevadas perdas nitrogenadas. O balanço nitrogenado negativo pode ser revertido por regimes mais agressivos do que os habitualmente utilizados. A nutrição enteral iniciada precocemente foi bem tolerada, demonstrando ser um método seguro e efetivo para a administração de nutrientes.