Carol Rees Parrish, M.S., R.D., Series Editor

Pancreatic Exocrine Insufficiency and Enteral Feeding: A Practical Guide with Case Studies



Mary E. Phillips



Amy Berry



Lucy S. Gettle

Patients with pancreatic disease can develop severe malnutrition. Many factors contribute to the malnutrition seen in this patient population, one of the most problematic being pancreatic exocrine insufficiency (PEI) with resultant malabsorption. Pancreatic enzyme replacement therapies (PERT) are predominantly designed for oral administration, but this can be challenging in patients requiring enteral nutrition (EN). This review explores the use of PERT in complex patients who are on EN. Case studies will be utilized to demonstrate different methods available in order to provide practical guidance on administration, both in the United States (U.S.) and the United Kingdom (U.K.).

INTRODUCTION

Patients with pancreatic disease, pancreatic resection, and cystic fibrosis often develop significant malnutrition as a result of the numerous gastrointestinal (GI) complications preventing adequate oral intake. Malabsorption, as well as side effects of medications such as antibiotics and opiates, adds an additional challenge to meeting nutritional requirements. In patients requiring EN, standard elemental formulas may not be tolerated or absorbed. Semi-elemental (or peptide-based) formulas are often utilized, providing less long-chain triglycerides and more

medium-chain triglycerides, thereby decreasing the dependence of pancreatic lipase for absorption. However, some patients will continue to malasborb, even with semi-elemental products, worsening the malnutrition present.

Traditional signs and symptoms of malabsorption include the presence of pale stools (often described as yellow or clay colored), floating or oily stools, stools with an unusually foul and offensive odor, abdominal bloating, cramping or gas, urgency and/or frequent stools.¹ The use of gut slowing medications, such as opiates, or the use

Mary E. Phillips BSc (Hons) RD DipADP Advanced Specialist Dietitian (Hepato-pancreatico-biliary Surgery), Royal Surrey County Hospital NHS Foundation Trust, Egerton Road, Guildford, Surrey, UK Amy Berry MS, RD, CNSC Nutrition Support Specialist (GI Surgery at the Emily Couric Cancer Center), University of Virginia Medical Center, Charlottesville VA Lucy S. Gettle, RDN CNSC Clinical Dietitian Specialist (Adult Cystic Fibrosis), University of Virginia Medical Center, Charlottesville VA

of low fat enteral feedings may result in reduction of obvious clinical signs and mask ongoing malabsorption. In these patients, constipated or infrequent stools (vs. diarrhea) may be reported, as well as abnormal color, unusually large volume stools, abnormal cramping and/or gas etc. The addition of antibiotics or prokinetics, can further confuse the clinical picture. Hence, sometimes it is the presence of more subtle signs of malnutrition such as difficulty with wound healing, worsening functional status, ongoing weight loss, or recurrent hypoglycemia/reduction in insulin requirement despite adequate caloric intake, that should trigger the consideration to consider an empiric trial of PERT.^{2,3}

EN and PERT products vary internationally. The aim of this paper is to provide a guide in the management of this complex patient group, combining the experience of dietitians specializing in patients with PEI, both in the U.S. and the U.K. We will also provide practical guidance in implementing the various modalities that can be utilized when it is determined that a patient is malabsorbing on their current EN regimen, as direction provided in the literature is sparse.⁴⁻¹⁰

CASE #1

A 62 year old female with pancreatic cancer and poorly controlled diabetes (HbA1c = 8.3%) completed a neo-adjuvant course of chemo-radiation prior to surgery. She reported a recent reduction in her insulin dose due to multiple episodes of severe hypoglycemia. Her oncologist also recently started her on enteric coated PERT due to reported yellow, oily stools and continued weight loss. She underwent a classic Whipple procedure, combined with a percutaneous gastro-jejunostomy (PEG-J) tube placement. Semi-elemental EN (Perative[®], Abbott, U.S.) was chosen due to the presence of malabsorption preoperatively. Her recovery was complicated by delayed gastric emptying requiring NPO (nil per os) status, and she was treated with a prokinetic and high dose proton pump inhibitor (PPI) given via her J-port. Her gastric port was vented as needed for gastric decompression. She was experiencing diarrhea, but was receiving enteral electrolyte replacement therapy (including magnesium oxide) and a prokinetic. On discharge,

she was tolerating G-tube clamping around the clock. She transitioned to a nocturnal feeding regimen that met her nutritional needs and was kept NPO.

Once home, her diarrhea increased. Her prokinetic dose was successfully weaned, with no worsening of her nausea. The patient reported improvement in her diarrhea, yet, eventually frequent loose stools associated with cramps and urgency were again reported. Her glycemic control was erratic; she began holding her long acting insulin due to episodes of hypoglycemia.

Two weeks post discharge her weight was down 5 pounds (2.25 kg), and an area of her midline wound opened, which required packing. At this time, it was determined an EN change was required. The elemental formula Vivonex[®] (Nestle, U.S.) was considered for its very low fat profile, however the patient was quite volume sensitive and would not have tolerated the rate increase needed. She was to be tried on a non-enteric coated PERT (Viokace[®]; Aptalis Pharma, U.S.), that could be crushed and added directly to her EN, but her insurance denied coverage for this product. We were hesitant to attempt enteric coated granules to mix with EN (described elsewhere^{6,7}) for fear of tube clogging, as the patient was completely dependent on her jejunal EN.

Fortunately, she advanced to oral intake. Full liquids were tolerated and she was able to take medications by mouth. She began to take her enteric coated PERT (2 capsules of Zenpep® 25,000 USP; Aptalis Pharma, U.S.) by mouth at the start of feedings, ~ 3 hours into her feeding cycle, and again before turning her EN off in the morning. Her stools firmed and improved; however, she still reported urgency and pale stools in the middle of the night. We increased her oral PERT administration to four times during her nocturnal cycle (she reported being awake and ambulatory at night, and agreeable to medication at this time), with one loperamide at the start of EN. This finally resulted in normalization of stooling and weight maintenance. She also resumed her intermediate acting insulin due to rising blood glucose with improved absorption; her blood glucoses now remaining in the 80-200mg/dL range. Over the next 2 months she increased her weight by 12 pounds (5.5 kg) and her abdominal wound began to heal.

Table 1. Clinical Parameters Pre- and Post-PERT

Parameter	Case 1		Case 2		
Diagnosis	Pancreatic cancer, Post Whipple (classic)		Cystic fibrosis related PEI, no anatomic abnormalities		
IBW	52.3	52.3 kg/ 115 lbs		75 kg/ 166 lbs	
Goal wt	62.7 kg/ 138 lbs		72 kg/ 160 lbs		
EN Access	PEG-J		Low profile G-tube		
Timing	Pre-PERT	Post-PERT	Pre-PERT	Post-PERT	
PERT	N/A	3 Zenpep [®] 25,000 USP units. Taken orally at start of EN, mid-EN x 2, end of EN. Alternatively, crush and add 1-2 tablets, 10,440 USP units of Viokace [®] to EN.	Viokace® crushed and mixed directly to EN	2 Relizorb® cartridges/day	
Weight	62.7 kg/138 lbs	70 kg/154 lbs	54 kg/119 lbs	64 kg/141 lbs	
Stool Output	Constant loose stools w/ EN; clay colored	Color normalization, formed, decreased frequency	Loose stools, fecal urgency in middle of night	2-3 formed stools daily	
EN	Perative®	Perative®	TwoCal HN®	Nutren 2.0®	
Glycemic Control	Poor; insulin requirement variable, glucoses labile	Increased insulin requirement with stabilization of glucoses	Normal	Impaired glucose tolerance, required insulin w/ EN	
Additional Changes	Wound breakdown, weight loss	Wound healing, weight increase	Abdominal discomfort, fullness with EN, difficulty with complex enzyme regimen	Improved adherence to EN regimen, increased appetite	

Case 3		Case 4		Case 5	
Total gastrectomy, total colectomy; severe chronic pancreatitis		Pancreatic cancer, gastric outlet obstruction		Total pancreatectomy; duodenectomy, hepatico- jejunostomy on roux loop	
UK does not use		64.5 kg/ 142 lbs		UK does not use	
65 kg/ 143 lbs		64.5 kg/ 142 lbs		84 kg/ 185 lbs	
Jejunostomy		PEG-J		Naso-jejunal	
Pre-PERT	Post-PERT	Pre-PERT	Post-PERT	Pre-PERT	Post-PERT
N/A	Pancrex V® powder 4g (100,000 units) /400mL EN	N/A	Creon [®] 24,000 USP units; 2 capsules/can of EN; opened and mixed with Nabicarb tabs and 60mL water for 30 minutes; flush via tube q 4 hours during EN infusion	N/A	Pancrex V® powder 4g (100,000 units)/ 400mL EN
52 kg/114 lbs	57 kg/125 lbs	59.1 kg/130 lbs	64.5 kg/142 lbs	84 kg/185 lbs	Fluid overloaded
High volume stoma output >2000mL/day	Stoma output <600mL/day	Frequent, yellow, oily; alternating w/ constipation d/t use of opiates	Constipated, requiring miralax	Brown formed stool every 1-2 days	Brown formed stool every 1-2 days
Peptisorb [®] ; Peptamen HN [®] ; Peptamen [®]	Vital 1.5 [®]	Vital 1.5®	Vital 1.5 [®]	Peptisorb [®]	Vital 1.5 [®]
Normal	Normal	Well- controlled, HbA1C= 6.3	Well-controlled	2-32mmol/L (36 – 576 mg/dL)	5-11mmol/L (90-198mg/dL)
Weight stabilization with use of supplemental PN	Improved functional status, d/c PN	Inability to gain weight	Weight regain, improved strength	Wound breakdown, recurrent hypoglycemia, peripheral edema, poor functional status	Wound healing, improved functional status, glucose stabilization with no further incidence of hypoglycemia

Table 2. Advantages and Disadvantages of Diagnostic Tests for PEI 16,12,15

lest	Advantages	LIMITATIONS			
Quantitative Fecal Fat (24, 48, 72 hour collection)	 Demonstrates extent of malabsorption, comparing fat intake vs. output. Traditionally 100g fat diet prescribed, but results can still be used if less fat consumed when careful record of how much fat patient consumes for comparison with output. Can demonstrate adequacy of therapy for patients already on PERT or semi-elemental EN. 	 May require stopping PERT. Unpleasant for patients. Requires 100g daily fat ingestion. Cumbersome, requiring stool collection from 1-3 days. Any stool loss during the collection can skew results. Careful monitoring of fat intake essential for result interpretation. Not available in the UK. 			
Qualitative Fecal Fat or Sudan Stain ("spot check")	 Easy, only requires one small stool sample. Positive result showing fat in the stool indicates malabsorption. Must ingest fat; no particular amount. 	 Positive result is unable to inform the extent of malabsorption. Negative result does not definitively rule out malabsorption (based on one sample at one time, may not catch what would be in the next stool sample). Not available in the UK. 			
Fecal Elastase (FE-1) fecal elastase <200 µg/g = pancreatic insufficiency; < 100 µg/g = severe pancreatic insufficiency	 Easy, only requires one small stool sample. Do not have to stop PERT for test.* No special diet required. 	 Reduced accuracy if used post- pancreatic resection. Watery stools cause falsely low result due to dilution. Unable to evaluate for continued malabsorption on current PERT. 			

* Though the manufacturer of the polyclonal fecal elastase test issued a disclaimer to discontinue PERT prior to the test, evidence reveals that the antibodies in the ELISA are to human elastase and are unaffected by the porcine elastase found in PERT.¹⁵

CASE #2

A 26 year old male with cystic fibrosis (CF) related PEI was struggling to gain weight and adhere to his EN regimen. He had been taking oral enteric coated PERT with his meals since being diagnosed with PEI at a young age, and was automatically started on PERT with a polymeric EN formula via G-tube. His original EN regimen consisted of a 2.0 kcal/mL formula (TwoCalHN[®], Abbott, U.S.) with enteric coated PERT taken by mouth at the beginning and end of an 8-hour nocturnal EN schedule. He continually endorsed abdominal discomfort and stool urgency in the middle of the night during his EN. Though taking oral enzymes at the beginning and end of an EN cycle tends to be the easiest regimen for CF patients, this

(continued on page 68)

(continued from page 66)

method does not provide enzymatic coverage for the entire duration of nocturnal EN. Due to continued malabsorptive symptoms and failure to gain weight, a non-enteric coated PERT product (Viokace®) was crushed and mixed with his EN prior to administration. This helped relieve some of the abdominal discomfort, but he continued to have difficulty consuming breakfast due to postfeeding appetite suppression. This patient also struggled with EN adherence due to the reported burden of effort to crush and mix PERT into the EN formula. Furthermore, the patient did not have a consistent living situation and did not always have EN supplies with him. We then trialed an in-line digestive enzyme cartridge (Relizorb[®], Alcresta, U.S.), as this would greatly reduce the burden of EN preparation, and discontinued Viokace[®]. Due to poor fat hydrolysis data of TwoCal HN[®] using Relizorb[®], his EN was changed to the highest caloric density compatible formula available at our facility (Ösmolite 1.5®, Abbott, U.S.). The patient had a dramatic improvement in tolerance and adherence to his EN regimen. His abdominal discomfort was much improved and he was able to consume a large breakfast due to reduced post-EN appetite suppression (consistent with results of a recent study¹⁰). He gained 10 pounds (4.5 kg) in the first three months after this therapy change, but was not able to maintain the rate of weight gain. In order to increase caloric provision, his outpatient EN was changed to Nutren 2.0® (Nestle, U.S.). He continued to tolerate the new EN regimen with 2 cartridges of Relizorb[®] nightly and gained 22 pounds (10 kg; 17.5% body weight) over the following year. It should be noted that the patient's appetite and oral intake also improved, contributing to weight gain.

CASE #3

A 54 year old male with chronic pancreatitis suffered a splenic bleed which resulted in extensive ischaemia of the bowel. A total gastrectomy, hemihepatectomy, total colectomy with end ileostomy, and feeding jejunostomy was performed; high stool output followed. The oesophagus was left disconnected (with a drain in situ, or "spit fistula") for 18 months prior to reconstruction.

Short bowel syndrome was presumed, as

there had been concern over the perfusion of his small bowel, and we utilized a full range of trials with peptide-based EN. In addition, loperamide up to 16mg QID and 60mg of codeine TID was administered. An electrolyte mix was used to flush his jejunostomy instead of water. Additional sodium (50mL 30%NaCl solution which provides 50mmol sodium, increasing the total sodium to 90-100mmol/l) was added to the EN to optimize water absorption. His stoma output remained >2000mL per day, and he continued to require supplemental parenteral nutrition (PN) to prevent weight loss. As this management was ineffective, his ongoing high stoma output was presumed to be not only due to loss of compensatory mechanisms (saliva and gastric secretions; colonic fermentation of malabsorbed carbohydrate, intestinal hurry), but also to severe PEI.

PERT was administered as 2g Pancrex V[®] powder (Essential Pharmaceuticals, U.K.) dissolved in 50mL sterile water and flushed by the nursing staff every 2 hours through his 24 hour EN. No benefit was observed and the decision was made to administer PERT to the EN directly. The 4 g Pancrex V[®] powder (100,000 units lipase) was added directly to 400 mL of Vital 1.5® (Abbott, U.K.), the mixture was shaken well and run at 100mL/hr over 4 hours. UK guidelines limit hang time for EN with any additives to 4 hours.¹¹ This was repeated 4 times during the day, and provided a total of 2400 calories. His stoma output reduced to less than 600mL per day, and he was weaned off PN and began to gain weight. This method of EN was continued until he underwent reconstructive surgery and was able to recommence oral intake.

CASE #4

A 64 year old man status post chemotherapy, radiation, and distal pancreatectomy/splenectomy for pancreatic cancer suffered many post-operative complications including a chyle leak, GI bleed, recurrent abscesses, bacteremia, and delayed gastric emptying requiring placement of a PEG-J. He required multiple readmissions and continued to struggle with nausea, vomiting, abdominal pain and weight loss. He eventually requested a second opinion and was admitted to our facility. His esophagogastroduodenoscopy (EGD), upper

(continued on page 70)

(continued from page 68)

Table 3. Factors Decreasing Response to Oral or Enteral PERT 5-6, 16

- Poor adherence to therapy.
- Outdated prescription.
- Storing enzymes in a hot environment (ex: in the car, in a clothing pocket) or taking with hot drinks, which can denature enzymes.
- Not taking enzymes with fat-containing beverages, milk products, or snacks.
- "Grazing" eating behaviors which can be challenging to cover with enzymes.
- Not increasing enzyme dosage for very high fat diet.
- Excessively acidic gastric and intestinal environment as the optimal pH for enzyme activation is 5.5 or above; may need acid blocker.
- Delayed gastric emptying causing poor mixing of enzyme with food, and prolonged exposure to gastric acid.
- Inappropriate timing of enzymes such as taking at completion of meal instead of at beginning/during.
- Chewing enzymes.
- Gastric (or oral) Feeding: Opening PERT capsule, adding enzyme beads/granules onto alkaline foods causing premature activation of enzymes. *When using this route, may be advisable to add to acidic puree (such as applesauce) to protect the enteric coating.*
- Jejunal Feeding: Opening the capsule to add enzyme beads/ granules without first activating the enzyme by utilizing a bicarbonate slurry (See Figure 1. Section titled: Powdered or crushed nonenteric coated PERT).

Adapted with permission from University of Virginia Health System, Nutrition Support Traineeship Manual¹⁶

GI series and computed tomography (CT) scan showed gastric stenosis that would require surgical revision. Due to his malnourished state, nutritional optimization prior to surgery was necessary. He came to us receiving 6 cans of Vital 1.5[®] (Abbott, U.S.). Each can of Vital 1.5 was mixed with 2 capsules of Creon 24,000® (AbbVie, U.S.), which were opened and mixed with 2 tabs of sodium bicarbonate (650 mg) crushed and mixed with 60mL of water and sat as a slurry for 30 minutes prior to administration. This slurry was flushed via his feeding tube every 4 hours. He had lost 21% of his usual body weight over the last year, and presented to us with a weight of 130 pounds (59 kg). He was 5 feet, 6 inches (168cm); preoperative weight goal was 140-145 pounds (63-65 kg). He was switched to 8 cans of Vivonex[®], allowing him to stop the PERT slurry flushes. However, after 2-3 weeks, he had not gained any weight. His EN regimen was then changed to a mixture of 1 can of Vital 1.5[®] mixed with 3 cans of Vivonex[®] twice

daily for a total of 8 cans per day. This provided increased calories while keeping fat content low, avoiding the use of enzymes. However, his lack of weight gain persisted and the patient reported a decline in quality of life due to the continuous EN infusion regimen. At this time his EN was adjusted to 7 cans of Vital 1.5[®], and PERT slurries were restarted every 4 hours. Over the next 4 weeks EN was increased to 8 cans, then 9 cans of Vital 1.5[®]. One month after his 9th can was added, he reached his goal weight of 142 pounds (64.5 kg) and was able to undergo a surgical gastro-jejunostomy with eventual progression to an oral diet.

CASE #5

A 68 year old male underwent a pancreaticoduodenectomy for adenocarcinoma of the pancreas with subsequent completion pancreatectomy following the development of a high volume pancreatic fistula. A nasojejunal (NJ) feeding tube

(continued on page 72)

(continued from page 70)

was inserted intra-operatively and he was initially fed with a semi-elemental EN (Peptisorb[®], Nutricia Clinical Care, U.K.). He had a protracted length of stay and ongoing delayed gastric emptying, thus he remained dependent on NJ feedings for his nutrition. He had normal bowel function, yet continued to lose weight, developed a hospital acquired pneumonia and an abdominal wound breakdown requiring the use of a vacuum wound management system.

His glycemic control was erratic despite continuous EN, with frequent episodes of hypoglycemia. Given his complete pancreatectomy, malabsorption was presumed, and 4g of Pancrex V[®] powder (100,000 units lipase) was added to each bottle of his peptide EN, which was changed to a 1.5kcal/mL product (400mL Vital 1.5[®]) to allow a rest period for physiotherapy. This was repeated 4 times a day, to provide 2400 calories. Within a

Table 4. Medicare Coverage for PERT 2018¹⁹

week, his insulin requirements increased and he no longer experienced hypoglycemic episodes. Over the next few weeks his grip strength and weight started to increase. There was no change in his bowel habits, and he continued with brown formed stools every 24-36 hours. His wound began to heal, and he began to eat and drink. On day 92 he discontinued NJ feedings and was discharged home on day 94 on an oral diet, PERT and insulin. Table 1 provides an overview of all 5 cases including nutritional parameters, PERT, and outcomes.

Not Everyone Needs PERT

The cases above highlight the need to identify patients with PEI who would benefit from the use of PERT. However, it is worth noting that not all patients with pancreatic disease and frequent stooling require PERT. The use of antibiotics, prokinetics, diagnosis of *Clostridium difficile*, and occurrence of dumping syndrome among other

Pancreatic enzymes are a Tier II drug, covered under Medicare Part D. Initially the patient will payout-of-pocket for PERT until they reach their deductible (currently, no Medicare plan deductible can exceed \$405). After the deductible is met, the patient will pay a copayment or coinsurance for the drug (copay depends on particular plan). Once total expenses (the amount the patient pays for the medication PLUS what Medicare pays for the medication) reaches \$3750.00, the patient lands in the "coverage gap" or "donut hole". Once there, the patient will pay 35% of name brand drug cost until total costs (the 35% patient paid PLUS 50% manufacturer discount payment, which will get the patient past the coverage gap faster) of drug payments exceed \$5000.00. Once this amount is met, "catastrophic coverage" begins and the patient only pays a small flat co-pay for the remainder of the year. The plan the patient chooses at enrollment will determine this co-pay amount. Plans are available to help with the "donut hole" coverage that can be signed up for during Medicare's open enrollment. However, these plans will have a higher out of pocket monthly cost. Additionally, there is the "Extra Help" program Medicare offers, which is income based and further helps drug costs (to qualify, double income has to be less than \$24,960 with less than \$28,150 in countable resources).

At the time this article was written, the authors found the average cost was \$7-11 per 25,000 USP units of an enteric coated PERT. Therefore, if a patient consumes 8 pills in a day, the out of pocket cost of PERT could be \$56- \$88 per day or \$1680- \$2640 per month.

Relizorb[®] cartridges are classified by the FDA as a device and not a medication, which can affect coverage. It is currently covered by Medicaid and various private insurance plans, but not Medicare.

Of note, most PERT companies have financial assistance available for patients with commercial insurance or no insurance, but not for those on governmental assistance programs such as Medicaid/Medicare.

causative factors should be considered prior to adjusting EN or initiating PERT for suspected malabsorption from PEI.

A 76 year old patient post Whipple procedure for ampullary adenocarcinoma was readmitted with severe diarrhea, nausea, vomiting, dehydration and failure to thrive. He was also on a prokinetic due to symptoms of delayed gastric emptying, and receiving enteral electrolytes, both of which could potentially affect stool output. In attempts to rule out malabsorption, the surgical team ordered a fecal elastase, which came back low; the watery stool sample as well as recent pancreatic surgery invalidated the low result.¹² A fecal fat check (qualitative) was obtained, however, he had little to no enteral nutrition (hence fat) for days. As a result, the fecal fat came back negative, adding little to the clinical assessment. His CT scan showed colitis (stool sample was negative for Clostridium difficile). The team ordered PERT, with no change in stool output. Eventually the enzymes were stopped. Ultimately, the patient required a PEG-J, tolerating standard EN, and his stools improved over time as his colitis resolved based on his repeat CT scan.

DISCUSSION

Assessment of nutritional adequacy and feeding tolerance is complex in patients who have pancreatic disease. There are no standard guidelines in the administration of PERT with EN, and clinical practice differs internationally as a result of the differences in EN and PERT products. Consequently, methods for adding PERT to EN also differ.

The use of a fecal fat stool collection or fecal elastase may help to support the diagnosis of ongoing malabsorption in certain circumstances; however, limitations exist (see Table 2). Assessment of residual pancreatic function, signs and symptoms, in concert with clinical judgment, is needed to establish if the patient is malabsorbing. Ongoing reassessment to ensure adequate response to therapy is also important (see Table 3).

In the U.K., peptide/semi-elemental formulas are used as first line EN in patients with pancreatic disease as per European Society for Enteral and Parenteral Nutrition (E.S.P.E.N) guidelines;¹³ it is also standard practice to use PERT routinely in patients after pancreatic head resection² and



Figure 1. Enteral Nutrition and Pancreatic Enzyme Replacement Therapy Options in Pancreatic Enzyme Insufficiency^{2,4-10,17-18}

inoperable pancreatic cancer.¹⁴ In the U.S. practices vary, but at UVA, we consider standard EN as first line therapy when initiating EN and readjust as signs or symptoms of malabsorption develop or PEI is confirmed. If a patient fails on standard EN, a semi-elemental feeding will be tried, followed by a strict elemental product or initiation of PERT. Initiation of PERT is carefully considered on an individual basis as circumstances dictate. If a clinician determines PERT with EN is the best approach, they then must also decide on the best method of administration. Figure 1 provides methods that may be employed when providing PERT with EN.

In the U.S., another major hurdle when utilizing PERT with EN (or orally) is obtaining insurance coverage. Certain enzyme formulations may not be covered, or the patient may not have met their outof-pocket premiums for their particular insurance plan (see Table 4 for an overview of Medicare Part D drug coverage).

CONCLUSION

Patients with pancreatic disease are often complex and present many nutritional challenges. There are significant variations in absorptive capacity between patients with cystic fibrosis, acute and chronic pancreatitis, pancreatic resection, pancreatic cancer, and other conditions that may lead to PEI. Therefore, it is crucial that clinicians have a range of management options to optimize the nutritional care of these vulnerable patients. This is a 'how to' guide for the addition of PERT to EN, and highlights the variation in practice and patient management internationally.

PRACTICAL GASTROENTEROLOGY

Visit our website:

practicalgastro.com

References

- 1. Decher N, Berry A. Post-Whipple: A Practical Approach to Nutrition Management. Practical Gastroenterol 2012;Aug(8):30-42.
- 2. Phillips ME. Pancreatic exocrine insufficiency following pancreatic resection. Pancreatology 2015;15(5):449-455.
- Duggan, SN, Conlon KC. Pancreatogenic Type 3c Diabetes: Underestimated, Underappreciated and Poorly Managed. Practical Gastroenterol 2017;May(5):14-23.
- Yoder AJ, Parrish CR, Yeaton P. A retrospective review of the course of patients with pancreatitis discharged on jejunal feedings. Nutr Clin Pract2002 Oct;17(5):314-20.
- Krenitsky J, Makola D, Parrish CR. Parenteral Nutrition in Pancreatitis is Passe: But are we ready for gastric feeding? A practical guide to jejunal feeding: Revenge of the Cyst- Part II. Practical Gastroenterol 2007;Oct(10):54-72.
- 6. Berry AJ. Pancreatic enzyme replacement therapy during pancreatic insufficiency. Nutr Clin Pract 2014;29(3):312-321.
- Ferrie S, Graham C, Hoyle M. Pancreatic enzyme supplementation for patients receiving enteral feeds. Nutr Clin Pract 2011;26(3):349-51.
- Freedman S, Orenstein D, Black P, et al. Increased Fat Absorption from Enteral Formula Through an In-line Digestive Cartridge in Patients with Cystic Fibrosis. J Pediatr Gastroenterol Nutr 2017;65(1):97-101.
- Freedman SD. Options for addressing exocrine pancreatic insufficiency in patients receiving enteral nutrition supplementation. Am J Manag Care 2017;23:S220-S228.
- Nicolo M, Stratton K, Rooney W, et al. Pancreatic enzyme replacement therapy for enterally fed patients with cystic fibrosis. Nutr Clin Pract 2013:28(4):485-489.
- 11. NICE. Healthcare-associated infections: prevention and control in primary and community care (CG 139). 2012.
- 12. Benini L, Amodio A, Campagnola P, et al. Fecal elastase-1 is useful in the detection of steatorrhea in patients with pancreatic diseases but not after pancreatic resection. Pancreatol 2013;13(1):38-42.
- Meier, R., Ockenga, J., Pertkiewicz, M., et al. ESPEN Guidelines on Enteral Nutrition: Pancreas. Clin Nutr 2006;25(2):275-84.
- 14. NICE. Pancreatic Cancer in Adults: Diagnosis and Management (NG 85) 2018
- 15. Weiss, FU, Buddle C, Lerch MM. Specificity of a polyclonal fecal elastase ELISA for CELA3. PLoS One 2016;11(7).
- 16. Parrish CR, Krenitsky J, McCray S. Factors reducing effectiveness of enzyme therapy. Adapted with permission from University of Virginia Health System, Nutrition Support Traineeship Manual, 2016.
- 17. Actavis Inc (now Allergan). Personal correspondence with manufacturer on data for viokace stability and microbial growth when mixed in EN formula. 2014. https://www.allergan.com/assets/pdf/viokace_pi.
- Instructions for Use RELiZORB. http://relizorb.com/ docs/pdfs/Relizorb-Instructions-for-Use.pdf. Accessed on 8/29/2018.
- Costs for Medicare drug coverage. https://www.medicare. gov/part-d/costs/part-d-costs.html. Accessed on 8/29/18.