



### Introduction

Three decades ago, postoperative starvation was common practice after most types of digestive surgical procedures. In particular, gastric decompression was performed until resolution of postoperative ileus [1]. This dogma was challenged with new evidence on the healing process of intestinal anastomoses, with increased collagen deposition and strength through early feeding [2, 3]. Further, a beneficial effect on wound healing was noticed [4]. A thorough first meta-analysis provided interesting new data suggesting a reduction in infectious complications, anastomotic leak rates, wound infection, and length of hospital stay, however, with an increased risk of vomiting among early fed patients [5]. Early enteral nutrition was part of the first published enhanced recovery series by Kehlet et al. of eight patients undergoing colonic resections [6]. Early feeding was combined with epidural analgesia, mobilization, and minimally invasive surgery to provide a “stress-free” surgical experience. Further studies confirmed these results, with early postoperative resumption of normal diet as an indispensable component of all early multimodal pathways [7–10].

The concept of early enteral resumption of nutrition has to be considered as part of a more global strategy, which aims to face increased metabolic demands and catabolism during surgery [11]. A comprehensive nutritional strategy needs to be launched preoperatively. Early screening for malnutrition and nutritional conditioning are mandatory (preoperative optimi-

zation). Omission of preoperative fasting—allowing a normal meal the evening before surgery and free liquids and carbohydrate loading until 2 hours prior to surgery—further contributes to decrease surgical stress. This approach allows keeping glucose levels stable by minimizing insulin resistance [12]. Early resumption of nutrition combined with stringent perioperative fluid management and early mobilization are thus a logical continuation of events. Noteworthy, several studies demonstrated a decline in postoperative nutritional status despite preoperative treatment in low- and high-risk patients, emphasizing the importance of early resumption of diet and timely launch of nutritional support if needed [13–15].

This chapter addresses the question why early enteral nutrition should be standard of care by reviewing available evidence according to type of surgery. Further, type of nutrition and criteria for nutritional supplementation in the postoperative period including enteral (tube feeding) and parenteral nutrition are reviewed.

### Safety of Early Resumption of Diet

Oral nutrition including clear liquids can be initiated safely and immediately after surgery. This implies retrieval of nasogastric tubes by the end of the procedure, which has repeatedly been shown to be safe regardless of the type of surgery and even protective against pharyngeal and respiratory adverse events [16, 17]. A meta-analysis of randomized controlled trials of 2009 yielding 1173 patients did not find any drawback of early enteral nutrition [18]. Instead, a trend toward decreased postoperative medical and surgical complications and length of stay was observed. Even though the mechanism was not clear, early enteral nutrition within 24 hours was also associated with decreased mortality. The authors concluded that keeping patients “nil by mouth” is without any benefit and patients should be allowed to drink upon full recovery from anesthesia. Noteworthy, early postoperative feeding was also associated with increased vomiting [18]. A more recent randomized trial found a low residual

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diet to be more efficient compared to clear fluids in preventing nausea and promoting return of bowel function after colorectal surgery [19]. However, further ileus-preventing mechanisms within an enhanced recovery pathway helped to face these drawbacks of earlier experience [20].

## Evidence in Surgical Subspecialties

### Colorectal Surgery

The best evidence in favor of early resumption of enteral nutrition is available for patients undergoing colorectal surgery [21]. A systematic review of 14 randomized controlled trials described early enteral feeding after elective procedures, with 12 studies reporting almost exclusively or exclusively on patients undergoing lower gastrointestinal (GI) surgery. Seven studies reported adequately on the randomization process, whereas in the remaining studies the method of randomization was either unclear or not stated at all. Studies were heterogeneous regarding inclusion criteria, feeding policies, and reported outcomes. Most outcomes failed to reach statistical significance, but mortality and length of stay were decreased in the early feeding group. A further meta-analysis of 15 studies described a significant reduction of postoperative complications in the early feeding group, with no negative impact on anastomotic dehiscence or resumption of bowel function [22]. Individual randomized trials concluded that there was no reason to withhold early oral intake, since it was well tolerated without increasing rates of postoperative ileus, providing adequate ileus-preventing measures [23, 24]. The most recent meta-analysis providing data on 7 studies and 587 patients undergoing exclusively colorectal resections confirmed these results [25]. Hospital stay and total postoperative complications were decreased, while no significant impact on anastomotic dehiscence, pneumonia, or rate of nasogastric tube reinsertion was noticed.

Also less compelling than for colorectal surgery, the concept of early enteral nutrition embedded in an enhanced recovery pathway applies also for other types of surgery [26, 27].

### Upper Gastrointestinal Surgery

A landmark randomized trial by Lassen et al. including 447 patients demonstrated feasibility of normal food at will after major upper GI surgery [26]. In particular, functional recovery, major complications, and length of stay were decreased in the group, which tolerated normal food at will from the first day after surgery, as compared to the “nil by mouth” and

tube feeding groups. A recent meta-analysis showed further improved cellular immunity and decreased postoperative complications in gastric cancer patients undergoing major resections [28]. The meta-analysis of Willcutts et al. came to similar conclusions [29]: Early oral feeding was associated with shorter hospital stay, while no increase in relevant complications was observed. For esophageal cancer patients undergoing esophagectomy, improved nutritional parameters at the eighth day were observed in the early oral nutrition groups, and pulmonary complications and anastomotic leaks were decreased compared to patients receiving parenteral nutrition. Further studies on esophagectomy patients confirmed safety and feasibility of early enteral nutrition, by emphasizing in particular a restorative effect on intestinal barrier function postoperatively [30]. Early oral intake as part of standardized care pathways has also been recommended and endorsed by several societies after bariatric surgery [31–33]. As a common conclusion of most studies on upper GI surgery, early feeding is feasible and safe. However, more evidence particularly in the field of esophageal surgery is warranted.

### Pancreatic Surgery

In particular after pancreaticoduodenectomy, the evidence is ambiguous. Malnutrition is preponderant among patients with pancreatic cancer, and morbidity rates of up to 40% after major pancreatic surgery, including specific complications such as delayed gastric emptying (DGE), request thorough identification and timely support of patients at nutritional risk [27, 34, 35]. Early normal diet according to tolerance is safe and feasible, according to several randomized trials and systematic reviews [26, 36–38], even in the presence of delayed gastric emptying or pancreatic fistulae [27, 39]. Hence, early normal diet at will and according to tolerance should be encouraged [40]. A combined approach of early enteral nutrition with parenteral nutrition might have to be considered in some patients unable to cover their needs by the enteral route alone [41]. In this latter study, patients with a combined nutritional strategy presented with lower infectious complications, reduced rate of gastric emptying, and improved liver function compared to the comparative group receiving solely parenteral nutrition. However, a recent randomized study showed an increased postoperative complication rate including pancreatic fistulae and discouraged early enteral nutrition through a nasojejunal tube. Hence, an individual approach based on patients' nutritional status, disease presentation, and expected postoperative course should guide postoperative support strategies when normal diet at will is not sufficient.

## Hepatic Surgery

In the multicenter trial of Lassen et al., 66 patients underwent liver surgery, with the aforementioned beneficial outcomes in the early nutrition group confirming its safety after major hepatic resections [26]. A randomized controlled trial by Hendry et al., combining early oral administration of nutritional supplements with administration of laxatives, accelerated bowel recovery, however, without shortening hospital stay [42]. These results of accelerated functional recovery in early fed patients were confirmed by a meta-analysis, which further demonstrated decreased infection rates and improved immune competence and concluded early enteral nutrition to be safe after liver resection [43].

## Nutritional Supplementation Strategies

As discussed above, free diet should be aimed for starting from the first postoperative day. The amount of oral initial intake should be tailored to individual tolerance, since resumption of a normal everyday diet by the second postoperative day may not be an achievable goal for every patient [22, 44, 45]. Hence, energy needs might not be covered by free diet alone, since oral intake was shown to rarely exceed 1200–1500 kcal per day [46]. Oral nutritional supplements (ONS), in particular immunonutrition, may thus need to be considered to cover additional metabolic needs. According to recent European Society for Clinical Nutrition and Metabolism (ESPEN) guidelines [47], perioperative nutritional supplementation should be initiated if it is anticipated that patients are:

- Unable to eat for more than 5 days after surgery
- Unable to maintain above 50% of recommended intake for more than 7 days

Enteral nutritional support needs further strong consideration in patients at severe nutritional risk, which has been defined as follows by the ESPEN working group (2006):

- Weight loss >10–15% within 6 months
- Body mass index <18.5 kg/m<sup>2</sup>
- Nutritional Risk Score (NRS 2002) >5
- Hypoalbuminemia (<30 g/L) with no evidence of hepatic or renal dysfunction

All parameters reflect undernutrition and disease-associated catabolism [48–50].

In all patients fulfilling the aforementioned criteria, nutritional therapy should be started independently of the type of surgery, and the enteral route should always be preferred (Fig. 22.1) [47]. Early tube feeding with standard whole pro-

tein formulas, either through a nasojunal tube or a catheter jejunostomy for long duration, has to be considered within 24 hours of surgery in patients undergoing head and neck surgery or severely traumatized or brain injured patients [51, 52]. Several historical and more recent large-scale randomized controlled studies confirmed the superiority of the enteral route in preventing infectious complications, length of stay, and costs across all types of surgery [13, 48, 53–55]. Regarding the postoperative situation, the European and American guidelines [47, 56] recommend initiating postoperative nutritional supplementation within 24 hours. This is even more important considering that postoperative nutritional status deteriorates *despite* nutritional supplementation [15].

## Postoperative Immunonutrition

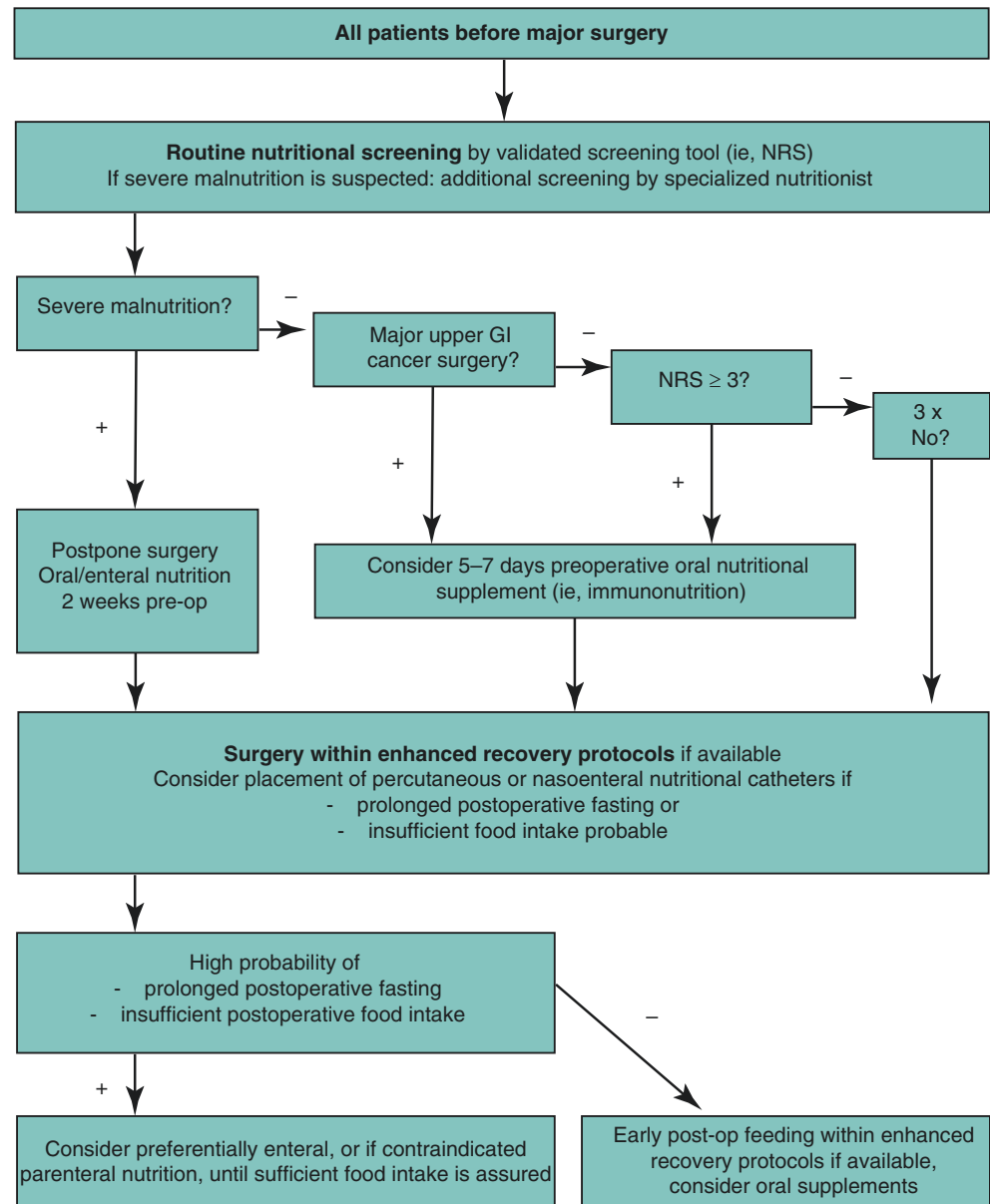
The evidence regarding immune-enhanced nutrition (arginine, glutamine, omega-3 fatty acids, and nucleotides) is somewhat ambiguous [47, 57]. While a beneficial effect on postoperative outcome was repeatedly shown in patients undergoing major cancer surgery, studies differed considerably regarding regimens, control groups, and outcomes, and a recent study revealed potential industry bias [58]. Further, the optimal timing could not be defined beyond doubt [59]. As a general rule, preoperative supplementation for 5–7 days should be considered in patients at nutritional risk according to standard definitions or screening tools, i.e., Nutritional Risk Score (NRS) or Malnutrition Universal Screening Tool (MUST) [50, 60]. However, more recent evidence also supports the administration of postoperative immunonutrition [46, 59, 61, 62]. While a randomized controlled trial of Klek et al. failed to demonstrate any clear advantage of routine postoperative immunonutrition [63], two recent studies by Moya et al. showed a significant decrease of medical and surgical infectious complications [64, 65]. Because of its cost-efficiency compared to parenteral administration, enteral immunonutrition was endorsed by recent ESPEN recommendations based on the principle of no harm [47] and has to be strongly considered in malnourished patients undergoing cancer surgery [66].

## Parenteral Nutritional Supplementation

The following contraindications to enteral nutritional support may warrant the use of parenteral support strategies [47]:

- Ileus
- Severe shock
- Intestinal ischemia
- High-output fistula
- Severe intestinal hemorrhage

**Fig. 22.1** Suggested treatment algorithm for nutritional support of surgical patients. NRS Nutritional Risk Score, GI gastrointestinal



Chen et al. presented a meta-analysis of randomized controlled trials to confirm safety and efficacy of parenteral nutrition [67]. Interestingly, an effect on leukotriene synthesis in patients with fish-oil-supplemented parenteral nutrition was observed. These findings were confirmed more recently in severely ill intensive care unit patients, especially regarding a modulated postoperative immune response [68–70].

As a common conclusion, postoperative parenteral nutrition should only be considered in patients who cannot be adequately fed enterally or who present the aforementioned contraindications [47].

## Conclusion

There is overwhelming evidence to support early resumption of a normal enteral diet, which should be the standard of care after most types of surgery. Specific criteria upon nutritional screening should guide clinicians in deciding whether nutritional support is warranted, especially in malnourished and cancer patients. The enteral route should always be the first choice; however, parenteral nutrition might be indicated in some circumstances when enteral supplementation is not feasible or sufficient.

## References

- Catchpole BN. Smooth muscle and the surgeon. *Aust N Z J Surg.* 1989;59(3):199–208.
- Uden P, Blomquist P, Jiborn H, Zederfeldt B. Impact of long-term relative bowel rest on conditions for colonic surgery. *Am J Surg.* 1988;156(5):381–5.
- Irvin TT, Hunt TK. Effect of malnutrition on colonic healing. *Ann Surg.* 1974;180(5):765–72.
- Schroeder D, Gillanders L, Mahr K, Hill GL. Effects of immediate postoperative enteral nutrition on body composition, muscle function, and wound healing. *JPEN J Parenter Enteral Nutr.* 1991;15(4):376–83.
- Lewis SJ, Egger M, Sylvester PA, Thomas S. Early enteral feeding versus “nil by mouth” after gastrointestinal surgery: systematic review and meta-analysis of controlled trials. *BMJ.* 2001;323(7316):773–6.
- Bardram L, Funch-Jensen P, Jensen P, Crawford ME, Kehlet H. Recovery after laparoscopic colonic surgery with epidural analgesia, and early oral nutrition and mobilisation. *Lancet.* 1995;345(8952):763–4.
- Muller S, Zalunardo MP, Hubner M, Clavien PA, Demartines N, Zurich Fast Track Study G. A fast-track program reduces complications and length of hospital stay after open colonic surgery. *Gastroenterology.* 2009;136(3):842–7.
- Jakobsen DH, Sonne E, Andreasen J, Kehlet H. Convalescence after colonic surgery with fast-track vs conventional care. *Color Dis.* 2006;8(8):683–7.
- Wind J, Polle SW, Fung Kon Jin PH, Dejong CH, von Meyenfeldt MF, Ubbink DT, et al. Systematic review of enhanced recovery programmes in colonic surgery. *Br J Surg.* 2006;93(7):800–9.
- Henriksen MG, Jensen MB, Hansen HV, Jespersen TW, Hessov I. Enforced mobilization, early oral feeding, and balanced analgesia improve convalescence after colorectal surgery. *Nutrition.* 2002;18(2):147–52.
- Kehlet H. The surgical stress response: should it be prevented? *Can J Surg.* 1991;34(6):565–7.
- Soop M, Carlson GL, Hopkinson J, Clarke S, Thorell A, Nygren J, et al. Randomized clinical trial of the effects of immediate enteral nutrition on metabolic responses to major colorectal surgery in an enhanced recovery protocol. *Br J Surg.* 2004;91(9):1138–45.
- Beattie AH, Prach AT, Baxter JP, Pennington CR. A randomised controlled trial evaluating the use of enteral nutritional supplements postoperatively in malnourished surgical patients. *Gut.* 2000;46(6):813–8.
- Garth AK, Newsome CM, Simmance N, Crowe TC. Nutritional status, nutrition practices and post-operative complications in patients with gastrointestinal cancer. *J Hum Nutr Diet.* 2010;23(4):393–401.
- Grass F, Benoit M, Coti Bertrand P, Sola J, Schafer M, Demartines N, et al. Nutritional status deteriorates postoperatively despite pre-operative nutritional support. *Ann Nutr Metab.* 2016;68(4):291–7.
- Weijs TJ, Kumagai K, Berkelmans GH, Nieuwenhuijzen GA, Nilsson M, Luyer MD. Nasogastric decompression following esophagectomy: a systematic literature review and meta-analysis. *Dis Esophagus.* 2017;30(3):1–8.
- Rao W, Zhang X, Zhang J, Yan R, Hu Z, Wang Q. The role of nasogastric tube in decompression after elective colon and rectum surgery: a meta-analysis. *Int J Color Dis.* 2011;26(4):423–9.
- Lewis SJ, Andersen HK, Thomas S. Early enteral nutrition within 24 h of intestinal surgery versus later commencement of feeding: a systematic review and meta-analysis. *J Gastrointest Surg.* 2009;13(3):569–75.
- Lau C, Phillips E, Bresee C, Fleshner P. Early use of low residue diet is superior to clear liquid diet after elective colorectal surgery: a randomized controlled trial. *Ann Surg.* 2014;260(4):641–7; discussion 7–9.
- Martos-Benitez FD, Gutierrez-Noyola A, Soto-Garcia A, Gonzalez-Martinez I, Betancourt-Plaza I. Program of gastrointestinal rehabilitation and early postoperative enteral nutrition: a prospective study. *Updat Surg.* 2018;70(1):105–12.
- Andersen HK, Lewis SJ, Thomas S. Early enteral nutrition within 24h of colorectal surgery versus later commencement of feeding for postoperative complications. *Cochrane Database Syst Rev.* 2006;18(4):CD004080.
- Osland E, Yunus RM, Khan S, Memon MA. Early versus traditional postoperative feeding in patients undergoing resectional gastrointestinal surgery: a meta-analysis. *JPEN J Parenter Enteral Nutr.* 2011;35(4):473–87.
- Han-Geurts IJ, Hop WC, Kok NF, Lim A, Brouwer KJ, Jeekel J. Randomized clinical trial of the impact of early enteral feeding on postoperative ileus and recovery. *Br J Surg.* 2007;94(5):555–61.
- Feo CV, Romanini B, Sortini D, Ragazzi R, Zamboni P, Pansini GC, et al. Early oral feeding after colorectal resection: a randomized controlled study. *ANZ J Surg.* 2004;74(5):298–301.
- Zhuang CL, Ye XZ, Zhang CJ, Dong QT, Chen BC, Yu Z. Early versus traditional postoperative oral feeding in patients undergoing elective colorectal surgery: a meta-analysis of randomized clinical trials. *Dig Surg.* 2013;30(3):225–32.
- Lassen K, Kjaeve J, Fetveit T, Trano G, Sigurdsson HK, Horn A, et al. Allowing normal food at will after major upper gastrointestinal surgery does not increase morbidity: a randomized multicenter trial. *Ann Surg.* 2008;247(5):721–9.
- Buscemi S, Damiano G, Palumbo VD, Spinelli G, Ficarella S, Lo Monte G, et al. Enteral nutrition in pancreaticoduodenectomy: a literature review. *Nutrients.* 2015;7(5):3154–65.
- Cheng Y, Zhang J, Zhang L, Wu J, Zhan Z. Enteral immunonutrition versus enteral nutrition for gastric cancer patients undergoing a total gastrectomy: a systematic review and meta-analysis. *BMC Gastroenterol.* 2018;18(1):11.
- Willcutts KF, Chung MC, Erenberg CL, Finn KL, Schirmer BD, Byham-Gray LD. Early oral feeding as compared with traditional timing of oral feeding after upper gastrointestinal surgery: a systematic review and meta-analysis. *Ann Surg.* 2016;264(1):54–63.
- Xiao-Bo Y, Qiang L, Xiong Q, Zheng R, Jian Z, Jian-Hua Z, et al. Efficacy of early postoperative enteral nutrition in supporting patients after esophagectomy. *Minerva Chir.* 2014;69(1):37–46.
- Allied Health Sciences Section Ad Hoc Nutrition C, Aills L, Blankenship J, Buffington C, Furtado M, Parrott J. ASMBS allied health nutritional guidelines for the surgical weight loss patient. *Surg Obes Relat Dis.* 2008;4(5 Suppl):S73–108.
- Torres AJ, Rubio MA. The Endocrine Society’s Clinical Practice Guideline on endocrine and nutritional management of the post-bariatric surgery patient: commentary from a European Perspective. *Eur J Endocrinol.* 2011;165(2):171–6.
- Mechanick JI, Youdim A, Jones DB, Garvey WT, Hurley DL, McMahan MM, et al. Clinical practice guidelines for the perioperative nutritional, metabolic, and nonsurgical support of the bariatric surgery patient—2013 update: cosponsored by American Association of Clinical Endocrinologists, The Obesity Society, and American Society for Metabolic & Bariatric Surgery. *Obesity (Silver Spring).* 2013;21(Suppl 1):S1–27.
- Bozzetti F, Mariani L. Perioperative nutritional support of patients undergoing pancreatic surgery in the age of ERAS. *Nutrition.* [Review]. 2014;30(11–12):1267–71.
- Akizuki E, Kimura Y, Nobuoka T, Imamura M, Nagayama M, Sonoda T, et al. Reconsideration of postoperative oral intake tolerance after pancreaticoduodenectomy: prospective consecutive analysis of delayed gastric emptying according to the ISGPS definition and the amount of dietary intake. *Ann Surg.* 2009;249(6):986–94.

36. Lassen K, Revhaug A. Early oral nutrition after major upper gastrointestinal surgery: why not? *Curr Opin Clin Nutr Metab Care*. [Review]. 2006;9(5):613–7.
37. Gerritsen A, Wennink RA, Besselink MG, van Santvoort HC, Tseng DS, Steenhagen E, et al. Early oral feeding after pancreatoduodenectomy enhances recovery without increasing morbidity. *HPB*. [Observational Study Research Support, Non-U.S. Gov't]. 2014;16(7):656–64.
38. Gerritsen A, Besselink MG, Gouma DJ, Steenhagen E, Borel Rinkes IH, Molenaar IQ. Systematic review of five feeding routes after pancreatoduodenectomy. *Br J Surg*. 2013;100(5):589–98; discussion 99.
39. Fujii T, Nakao A, Murotani K, Okamura Y, Ishigure K, Hatsuno T, et al. Influence of food intake on the healing process of postoperative pancreatic fistula after pancreatoduodenectomy: a multi-institutional randomized controlled trial. *Ann Surg Oncol*. [Multicenter Study Randomized Controlled Trial]. 2015;22(12):3905–12.
40. Braga M, Capretti G, Pecorelli N, Balzano G, Dogliani C, Ariotti R, et al. A prognostic score to predict major complications after pancreaticoduodenectomy. *Ann Surg*. 2011;254(5):702–7; discussion 7–8.
41. Zhu XH, Wu YF, Qiu YD, Jiang CP, Ding YT. Effect of early enteral combined with parenteral nutrition in patients undergoing pancreaticoduodenectomy. *World J Gastroenterol*. 2013;19(35):5889–96.
42. Hendry PO, van Dam RM, Bukkems SF, McKeown DW, Parks RW, Preston T, et al. Randomized clinical trial of laxatives and oral nutritional supplements within an enhanced recovery after surgery protocol following liver resection. *Br J Surg*. 2010;97(8):1198–206.
43. Richter B, Schmandra TC, Golling M, Bechstein WO. Nutritional support after open liver resection: a systematic review. *Dig Surg*. 2006;23(3):139–45.
44. Grass F, Bertrand PC, Schafer M, Ballabeni P, Cerantola Y, Demartines N, et al. Compliance with preoperative oral nutritional supplements in patients at nutritional risk—only a question of will? *Eur J Clin Nutr*. 2015;69(4):525–9.
45. Grass F, Schafer M, Demartines N, Hubner M. Normal diet within two postoperative days—realistic or too ambitious? *Nutrients*. 2017;9(12):1336.
46. Sultan J, Griffin SM, Di Franco F, Kirby JA, Shenton BK, Seal CJ, et al. Randomized clinical trial of omega-3 fatty acid-supplemented enteral nutrition versus standard enteral nutrition in patients undergoing oesophagogastric cancer surgery. *Br J Surg*. 2012;99(3):346–55.
47. Weimann A, Braga M, Carli F, Higashiguchi T, Hubner M, Klek S, et al. ESPEN guideline: clinical nutrition in surgery. *Clin Nutr*. 2017;36(3):623–50.
48. Bozzetti F, Braga M, Gianotti L, Gavazzi C, Mariani L. Postoperative enteral versus parenteral nutrition in malnourished patients with gastrointestinal cancer: a randomised multicentre trial. *Lancet*. 2001;358(9292):1487–92.
49. Kondrup J, Allison SP, Elia M, Vellas B, Plauth M, Educational, et al. ESPEN guidelines for nutrition screening 2002. *Clin Nutr*. 2003;22(4):415–21.
50. Kondrup J, Rasmussen HH, Hamberg O, Stanga Z, Ad Hoc EWG. Nutritional risk screening (NRS 2002): a new method based on an analysis of controlled clinical trials. *Clin Nutr*. 2003;22(3):321–36.
51. Bozzetti F, Gianotti L, Braga M, Di Carlo V, Mariani L. Postoperative complications in gastrointestinal cancer patients: the joint role of the nutritional status and the nutritional support. *Clin Nutr*. 2007;26(6):698–709.
52. Perel P, Yanagawa T, Bunn F, Roberts I, Wentz R, Pierro A. Nutritional support for head-injured patients. *Cochrane Database Syst Rev*. 2006;18(4):CD001530.
53. MacFie J, Woodcock NP, Palmer MD, Walker A, Townsend S, Mitchell CJ. Oral dietary supplements in pre- and postoperative surgical patients: a prospective and randomized clinical trial. *Nutrition*. 2000;16(9):723–8.
54. Mack LA, Kaklamanos IG, Livingstone AS, Levi JU, Robinson C, Sleeman D, et al. Gastric decompression and enteral feeding through a double-lumen gastrojejunostomy tube improves outcomes after pancreaticoduodenectomy. *Ann Surg*. 2004;240(5):845–51.
55. Smedley F, Bowling T, James M, Stokes E, Goodger C, O'Connor O, et al. Randomized clinical trial of the effects of preoperative and postoperative oral nutritional supplements on clinical course and cost of care. *Br J Surg*. 2004;91(8):983–90.
56. McClave SA, Taylor BE, Martindale RG, Warren MM, Johnson DR, Braunschweig C, et al. Guidelines for the provision and assessment of Nutrition support therapy in the adult critically ill patient: Society of Critical Care Medicine (SCCM) and American Society for Parenteral and Enteral Nutrition (A.S.P.E.N.). *JPEN J Parenter Enteral Nutr*. 2016;40(2):159–211.
57. Marimuthu K, Varadhan KK, Ljungqvist O, Lobo DN. A meta-analysis of the effect of combinations of immune modulating nutrients on outcome in patients undergoing major open gastrointestinal surgery. *Ann Surg*. 2012;255(6):1060–8.
58. Probst P, Ohmann S, Klaiber U, Huttner FJ, Billeter AT, Ulrich A, et al. Meta-analysis of immunonutrition in major abdominal surgery. *Br J Surg*. 2017;104(12):1594–608.
59. Osland E, Hossain MB, Khan S, Memon MA. Effect of timing of pharmaconutrition (immunonutrition) administration on outcomes of elective surgery for gastrointestinal malignancies: a systematic review and meta-analysis. *JPEN J Parenter Enteral Nutr*. 2014;38(1):53–69.
60. Sandhu A, Mosli M, Yan B, Wu T, Gregor J, Chande N, et al. Self-screening for malnutrition risk in outpatient inflammatory bowel disease patients using the Malnutrition Universal Screening Tool (MUST). *JPEN J Parenter Enteral Nutr*. 2016;40(4):507–10.
61. Song GM, Tian X, Zhang L, Ou YX, Yi LJ, Shuai T, et al. Immunonutrition support for patients undergoing surgery for gastrointestinal malignancy: preoperative, postoperative, or perioperative? A bayesian network meta-analysis of randomized controlled trials. *Medicine (Baltimore)*. 2015;94(29):e1225.
62. Farreras N, Artigas V, Cardona D, Rius X, Trias M, Gonzalez JA. Effect of early postoperative enteral immunonutrition on wound healing in patients undergoing surgery for gastric cancer. *Clin Nutr*. 2005;24(1):55–65.
63. Klek S, Kulig J, Sierzega M, Szybinski P, Szczepanek K, Kubisz A, et al. The impact of immunostimulating nutrition on infectious complications after upper gastrointestinal surgery: a prospective, randomized, clinical trial. *Ann Surg*. 2008;248(2):212–20.
64. Moya P, Miranda E, Soriano-Irigaray L, Arroyo A, Aguilar MD, Bellon M, et al. Perioperative immunonutrition in normo-nourished patients undergoing laparoscopic colorectal resection. *Surg Endosc*. 2016;30(11):4946–53.
65. Moya P, Soriano-Irigaray L, Ramirez JM, Garcea A, Blasco O, Blanco FJ, et al. Perioperative standard oral nutrition supplements versus immunonutrition in patients undergoing colorectal resection in an enhanced recovery (ERAS) protocol: a multicenter randomized clinical trial (SONVI Study). *Medicine (Baltimore)*. 2016;95(21):e3704.
66. Braga M, Gianotti L, Nespoli L, Radaelli G, Di Carlo V. Nutritional approach in malnourished surgical patients: a prospective randomized study. *Arch Surg*. 2002;137(2):174–80.
67. Chen B, Zhou Y, Yang P, Wan HW, Wu XT. Safety and efficacy of fish oil-enriched parenteral nutrition regimen on postoperative patients undergoing major abdominal surgery: a meta-analysis

- of randomized controlled trials. *JPEN J Parenter Enteral Nutr.* 2010;34(4):387–94.
68. Pradelli L, Mayer K, Muscaritoli M, Heller AR. n-3 fatty acid-enriched parenteral nutrition regimens in elective surgical and ICU patients: a meta-analysis. *Crit Care.* 2012;16(5):R184.
69. Li NN, Zhou Y, Qin XP, Chen Y, He D, Feng JY, et al. Does intravenous fish oil benefit patients post-surgery? A meta-analysis of randomised controlled trials. *Clin Nutr.* 2014;33(2):226–39.
70. de Miranda Torrinhas RS, Santana R, Garcia T, Cury-Boaventura MF, Sales MM, Curi R, et al. Parenteral fish oil as a pharmacological agent to modulate post-operative immune response: a randomized, double-blind, and controlled clinical trial in patients with gastrointestinal cancer. *Clin Nutr.* 2013;32(4):503–10.