

Impact of feeding strategy after pancreatoduodenectomy on delayed gastric emptying and hospital stay: nationwide study

Tessa E. Hendriks^{1,2,3,4,†} (D), Bo T. M. Strijbos^{5,†}, Michiel F. G. Francken^{1,2,†}, Mahsoem Ali^{1,2} (D), J. Annelie Suurmeijer^{1,2} (D), Marcel G. W. Dijkgraaf^{6,7}, Jana S. Hopstaken⁵ (D), Kees van Laarhoven⁵, Quintus Molenaar⁸, Vincent E. de Meijer⁹ (D), Erwin van der Harst¹⁰, Marcel den Dulk^{11,12} (D), Werner Draaisma¹³, Vincent Nieuwenhuijs¹⁴, Michael F. Gerhards¹⁵, Mike S. L. Liem¹⁶, George van der Schelling¹⁷, Eric Manusama¹⁸, Ignace de Hingh¹⁹ (D), Hjalmar van Santvoort⁷, Bas Groot Koerkamp²⁰ (D), Olivier R. Busch^{1,2} (D), Bert A. Bonsing³, Martijn W. J. Stommel^{5,‡} (D), Marc G. Besselink^{1,2,*‡} (D) and Dutch Pancreatic Cancer Group (DPCG) and Dutch Institute for Clinical Auditing (DICA)

¹Department of Surgery, Amsterdam UMC, University of Amsterdam, Amsterdam, the Netherlands

²Cancer Center Amsterdam, Amsterdam, the Netherlands

³Department of Surgery, Leiden University Medical Center, Leiden, the Netherlands

⁴Dutch Institute for Clinical Auditing, Leiden, the Netherlands

⁵Department of Surgery, Radboud University Medical Center, Nijmegen, the Netherlands

⁶Department of Epidemiology and Data Science, Amsterdam UMC, University of Amsterdam, Amsterdam, the Netherlands

⁷Amsterdam Public Health, Methodology, Amsterdam, the Netherlands

⁸Department of Surgery, Regional Academic Cancer Center Utrecht, University Medical Center Utrecht and St Antonius Hospital Nieuwegein, Utrecht, the Netherlands

⁹Department of Surgery, University of Groningen and University Medical Center Groningen, Groningen, the Netherlands

¹⁰Department of Surgery, Maasstad Hospital, Rotterdam, the Netherlands

¹¹Department of Surgery, Maastricht University Medical Center, Maastricht, the Netherlands

¹²NUTRIM-School of Nutrition and Translational Research in Metabolism, Maastricht University, Maastricht, the Netherlands

¹³Department of Surgery, Jeroen Bosch Hospital, 's-Hertogenbosch, the Netherlands

*Correspondence to: Marc G. Besselink, Department of Surgery, Amsterdam University Medical Center, University of Amsterdam, Cancer Center Amsterdam, De Boelelaan 1117 (ZH-7F), Amsterdam 1081 HV, the Netherlands (e-mail: m.g.besselink@amsterdamUMC.nl)

[†]Joint first authors [‡]Joint last authors

Members of the Dutch Pancreatic Cancer Group (DPCG) and Dutch Institute for Clinical Auditing (DICA) collaborators are co-authors of this study and are listed under the heading Collaborators.

Abstract

Background: Delayed gastric emptying is a major contributor to prolonged hospital stay following pancreatoduodenectomy. Although enhanced recovery after surgery guidelines recommend unrestricted feeding after pancreatoduodenectomy, nationwide studies evaluating the impact of different feeding strategies after surgery on delayed gastric emptying and length of hospital stay are limited. This study aimed to identify the use and impact of different feeding strategies after pancreatoduodenectomy on delayed gastric emptying and length of hospital stay.

Methods: This nationwide cohort study included consecutive patients after pancreatoduodenectomy from the Dutch Pancreatic Cancer Audit (2021–2023). Primary endpoints were delayed gastric emptying grade B/C and length of hospital stay. Feeding strategies were categorized based on structured interviews with representatives from 15 centres. Multilevel analysis was used to assess associations between feeding strategy, delayed gastric emptying, and length of hospital stay. Predictors of delayed gastric emptying were determined.

Results: Overall, 2354 patients undergoing pancreatoduodenectomy were included, of whom 526 (23%) developed delayed gastric emptying grade B/C. Median length of hospital stay was 13 days longer in patients with delayed gastric emptying (23 *versus* 10 days; P < 0.001). Feeding strategies were: unrestricted feeding (3 centres, 637 patients; delayed gastric emptying 18%); step-up feeding (9 centres, 1462 patients; delayed gastric emptying 24%); and artificial feeding (3 centres, 255 patients; delayed gastric emptying 25%). No association was observed between feeding strategy and delayed gastric emptying: step-up *versus* unrestricted feeding (odds ratio 1.14, 95% confidence interval 0.53 to 2.47) and artificial *versus* unrestricted feeding (odds ratio 1.76, 0.65 to 4.73). Similarly, no association was found between feeding strategy and length of hospital stay. The strongest predictor of delayed gastric emptying was pancreatic fistula after surgery (odds ratio 3.16, 2.47 to 4.05).

Conclusion: This study found no significant association between feeding strategy and incidence of delayed gastric emptying or length of hospital stay after pancreatoduodenectomy. Efforts to reduce delayed gastric emptying should focus on reducing pancreatic fistula after surgery.

© The Author(s) 2025. Published by Oxford University Press on behalf of BJS Foundation Ltd.

Received: January 22, 2025. Revised: March 21, 2025. Accepted: April 27, 2025

This is an Open Access article distributed under the terms of the Creative Commons Attribution License (https://creativecommons.org/licenses/by/4.0/), which permits unrestricted reuse, distribution, and reproduction in any medium, provided the original work is properly cited.

¹⁴Department of Surgery, Isala Clinics, Zwolle, the Netherlands

¹⁵Department of Surgery, OLVG, Amsterdam, the Netherlands

¹⁶Department of Surgery, Medisch Spectrum Twente, Enschede, the Netherlands

¹⁷Department of Surgery, Amphia Hospital, Breda, the Netherlands

¹⁹Department of Surgery, Catharina Hospital, Eindhoven, the Netherlands

Introduction

Complications following pancreatoduodenectomy (PD) remain significant^{1,2}, with delayed gastric emptying (DGE) being a major contributor to prolonged hospital stay^{3,4}. Clinically relevant DGE (grade B/C) is defined by the International Study Group of Pancreatic Surgery (ISGPS)⁵ as the need for nasogastric tube (NGT) drainage for more than 7 days, inability to tolerate solid foods, vomiting, and the need for nasoenteral or parenteral nutrition. Prolonged hospital stay is very common in patients with DGE⁶, and DGE is associated with an estimated €10 000 increase in hospital costs per patient⁷. Effective prophylactic or therapeutic strategies for DGE are currently lacking^{5,8}.

The impact of feeding strategies after surgery on the incidence of DGE and subsequent length of hospital stay (LOS) is unclear. The enhanced recovery after surgery (ERAS) guidelines⁸ recommend 'a normal diet after surgery without restrictions according to tolerance' (unrestricted diet) following PD. In contrast, many surgeons use a step-up approach for gradually reintroducing oral intake after PD to prevent vomiting and repeated NGT drainage, whereas others9-12 advise artificial feeding using nasoenteral or parenteral feeding. A 2022 systematic review¹³, including three retrospective studies and one randomized trial, suggested that an early oral feeding strategy after PD reduced LOS but not the incidence of DGE; however, within 'oral feeding', no distinction was made between unrestricted and step-up oral feeding. Consequently, it is unclear which feeding strategy should be advised after PD, concerning the incidence and severity of DGE and LOS. There is still debate as to whether patients should immediately commence an unrestricted diet, or a more stepwise approach or an artificial feeding strategy should be used.

This study aimed to assess the use and impact of feeding strategy on the incidence of DGE grade B/C and LOS after PD.

Methods

Study design

This was a nationwide retrospective analysis using data from the Dutch Pancreatic Cancer Audit (DPCA), coordinated by the Dutch Institute for Clinical Auditing and the Dutch Pancreatic Cancer Group (DPCG). The DPCA^{14,15} is a nationwide mandatory registry on pancreatic surgery in the Netherlands that has covered all pancreatic resections since 2014 with an estimated data completeness exceeding 97%. The scientific committee of the DPCG approved the study protocol¹⁶. According to Dutch law, no ethical approval or informed consent was required, as all data were registered anonymously.

This study included all consecutive patients who underwent PD for all indications and were registered in the DPCA between 1 January 2014 and 31 December 2023. Patients undergoing total pancreatectomy were excluded from the study. During this period, the incidence of DGE was determined per year. Additionally,

a structured interview took place with a representative surgeon from each DPCA centre in the Netherlands to assess the protocolized institutional feeding strategy after surgery.

The aim of this study was to evaluate the association between feeding strategy after surgery (for the years 2021–2023, as in these years a protocolized feeding strategy was present which remained unchanged in all participating hospitals) and the incidence of DGE grade B/C and LOS after PD. As secondary outcome, the incidence of DGE grade B/C and feeding strategy per centre in relation to the 3-year PD surgical volume and predictive factors for DGE was assessed. This study was reported in accordance with the STROBE statement¹⁷.

Data collection and definitions

Study baseline characteristics consisted of sex, age at the time of surgery, body mass index (BMI), American Society of Anesthesiologists (ASA) grade, and histopathological diagnosis before surgery. Treatment characteristics included neoadjuvant therapy, pylorus resection or preservation, minimally invasive or open surgery, venous or arterial resection, extended resection (in addition to the primary tumour, adjacent structures or organs were removed, such as mesocolon transversum gastric resection), and surgical drain placement. During the study period, neoadjuvant therapy for pancreatic cancer was mainly administered in randomized trials.

Data on feeding strategy and NGT placement (both during and after surgery) were not available in the DPCA. As there were no consistent policies on the use of NGT, this parameter varied per patient; therefore, it was not included in further analysis. Feeding strategy per hospital was determined through from interviews.

Primary outcomes were DGE and LOS per feeding strategy. Outcome parameters were collected during the entire hospital stay and, in the event of earlier discharge, up until 30 days after surgery.

DGE was defined according to the ISGPS⁵. Only clinically relevant DGE (grade B/C) was included. Other outcome parameters included major complications (Clavien-Dindo grade \geq III)¹⁸ and pancreatic surgery-related complications, LOS, and readmission within 30 days after discharge. Pancreatic surgery-related complications included postoperative pancreatic fistula (POPF)¹⁹, postpancreatectomy haemorrhage (PPH)²⁰, chyle leakage²¹, and bile leakage²², all grade B/C according to the ISGPS or International Study Group for Liver Surgery (ISGLS) criteria. Additionally, DGE was categorized into primary and secondary DGE. Primary DGE refers to the presence of DGE in the absence of other intra-abdominal surgical complications (for example PPH, POPF, bile leakage), and secondary DGE develops in association with intra-abdominal complications after surgery^{23,24}. Mortality was defined as in-hospital/30-day mortality (including in-hospital mortality during the entire primary admission or, in case of earlier discharge, up to 30 days).

¹⁸Department of Surgery, Medisch Centrum Leeuwarden, Leeuwarden, The Netherlands

²⁰Department of Surgery, Erasmus MC Cancer Institute, Rotterdam, the Netherlands



Fig. 1 Trend in the annual rate of DGE grade B/C after PD in the Netherlands (2014–2023)

Shaded area represents 95% confidence interval. DGE, delayed gastric emptying; PD, pancreatoduodenectomy.

Statistical analysis

Baseline patient, tumour, and treatment characteristics are presented using descriptive statistics. Continuous data are presented as median (interquartile range, i.q.r.) or mean(standard deviation), based on data distribution. Categorical variables are presented as counts and proportions. Normally distributed continuous data were compared using one-way ANOVA and non-normally distributed data using the Kruskal–Wallis test. Categorical data were analysed using the χ^2 test.

Multilevel logistic regression modelling was used to assess the association between different feeding strategies and DGE grade B/C, adjusting for prespecified confounders, to derive odds ratios (ORs) and 95% confidence intervals. Prespecified confounders included sex, age, BMI, ASA grade, Charlson Co-morbidity Index (CCI), diabetes, biliary drainage before surgery, type of surgery (pylorus resection versus pylorus-preserving), type of anastomosis (pancreatojejunostomy versus pancreatogastrostomy), vascular resection, extended resection, minimally invasive PD, pancreatic adenocarcinoma, POPF, PPH, bile leakage, and centre (as a random intercept). Centre was included in the multilevel model to adjust for differences in DGE incidence and feeding strategies between the included centres. The same regression model was used to assess which variables were most strongly predictive for DGE grade B/C. In this analysis, predictive factors for DGE were ranked by their likelihood ratio χ^2 value, with a higher value indicating that the variable is more strongly associated with DGE. This approach was used as it is invariant to the scale (continuous/categorical) of the variable, unlike ORs.

Linear mixed models were used to compare LOS between feeding strategies, while adjusting for the same confounders as in the multilevel logistic regression model, with centre as a random intercept, to estimate both the adjusted mean LOS (in days) per feeding strategy and the average (marginal) difference in LOS between feeding strategies.

Missing data were handled using multivariable imputation by chained equations (5 imputations and 10 iterations) with predictive mean matching. Results were pooled across imputed data sets using Rubin's rules.

In a sensitivity analysis, the association between different feeding strategies, DGE grade B/C, and LOS was assessed separately for primary and secondary DGE. In the sensitivity analysis, the same confounders were adjusted for as in the main analysis. Predictors of DGE grade B/C were identified.

Two-sided P < 0.050 was considered statistically significant. All analyses were undertaken in R version 4.3.2 (R Foundation for Statistical Computing, Vienna, Austria), using the lme4 and rms packages^{25,26}.

Results

Between 2014 and 2023, the DPCA included 7008 patients who underwent PD, with a mean DGE incidence of 21% (Fig. 1). The present study period (2021–2023), in which feeding strategies remained unchanged in all hospitals, included 2354 patients after PD. The mean age was 69 (i.q.r. 61–75) years, 44% of the patients were women, and 42% were diagnosed with pancreatic ductal adenocarcinoma. Most patients underwent

Table 1 Baseline characteristics of patients after PD (2021-2023)

	Overall	Unrestricted feeding $(n - 627)$	Step-up feeding $(n - 1462)$	Artificial feeding $(n - 255)$	P*
	(11 = 2554)	(n = 637)	(n = 1402)	(n = 255)	
Sex					0.120
Male	1306 (56%)	346 (54%)	803 (55%)	157 (62%)	
Female	1043 (44%)	291 (46%)	654 (45%)	98 (38%)	
Unknown	5	0	5	0	
Age (years), median (i.q.r.)	69 (61–75)	69 (61–75)	69 (61–75)	70 (63–76)	0.416
Unknown	8	0	8	0	
BMI (kg/m ²)					0.765
≤ 25	1418 (62%)	354 (61%)	902 (63%)	162 (64%)	
> 25	858 (38%)	225 (39%)	540 (37%)	93 (36%)	
Unknown	78	58	20	0	
ASA grade					0.750
I–II	1425 (62%)	382 (61%)	890 (62%)	153 (60%)	
\geq III	883 (38%)	242 (39%)	539 (38%)	102 (40%)	
Unknown	46	13	33	0	
CCI score					0.029
0–1	1325 (63%)	412 (66%)	750 (60%)	163 (65%)	
≥2	793 (37%)	210 (34%)	495 (40%)	88 (35%)	
Unknown	236	15	217	4	
Histological diagnosis					< 0.001
Pancreatic adenocarcinoma	955 (42%)	254 (40%)	586 (42%)	115 (45%)	
Cholangiocarcinoma	335 (15%)	83 (13%)	222 (16%)	30 (12%)	
Papillary cancer	291 (13%)	63 (9.9%)	188 (13%)	40 (16%)	
Duodenal cancer	142 (6.2%)	57 (9.0%)	71 (5.0%)	14 (5.5%)	
Neuroendocrine neoplasm	97 (4.2%)	31 (4.9%)	61 (4.3%)	5 (2.0%)	
IPMN, SPN, MCN	189 (8.2%)	68 (11%)	106 (7.5%)	15 (5.9%)	
Other	290 (13%)	79 (12%)	175 (12%)	36 (14%)	
Unknown	55	2	53	0	

PD, pancreatoduodenectomy; i.q.r., interquartile range; BMI, body mass index; ASA, American Society of Anesthesiologists; CCI, Charlson Co-morbidity Index; IPMN, intraductal papillary mucinous neoplasm; SPN, solid pseudopapillary neoplasm; MCN, mucinous cystic neoplasm. *Pearson's χ^2 test or Kruskal–Wallis rank-sum test.

Table 2 Operative data and treatment characteristics of patients after PD (2021-2023)

	Overall	Unrestricted	Step-up feeding	Artificial feeding	P‡
	(n = 2354)	feeding	(n = 1462)	(n = 255)	
	, , , , , , , , , , , , , , , , , , ,	(n = 637)	, , , , , , , , , , , , , , , , , , ,	. ,	
Surgical approach					< 0.001
Open	1661 (71%)	515 (81%)	988 (68%)	158 (62%)	
Robot-assisted	617 (26%)	120 (19%)	412 (29%)	85 (33%)	
Laparoscopic	56 (2.4%)	1 (0.2%)	43 (3.0%)	12 (4.7%)	
Unknown	20	1	19	0	
Vascular resection	411 (19%)	85 (15%)	284 (21%)	42 (16%)	0.006
Unknown	136	57	79	0	
Extended resection*	293 (13%)	42 (6.6%)	225 (16%)	26 (10%)	< 0.001
Unknown	15	1	14	0	
Surgery type					< 0.001
Pylorus-preserving	801 (34%)	258 (41%)	396 (27%)	147 (58%)	
Pylorus resection	1553 (66%)	379 (59%)	1066 (73%)	108 (42%)	
Unknown	0	Ö	0	Ó	
Drain after surgery	2155 (95%)	577 (100%)	1325 (92%)	253 (99%)	< 0.001
Unknown	78	59	19	0	
Neoadjuvant therapy†	419 (19%)	98 (16%)	282 (20%)	39 (16%)	0.059
Unknown	140	42	84	14	

*In addition to the primary tumour, adjacent structures or organs were removed (such as mesocolon transversum, gastric resection). \uparrow Neoadjuvant therapy only in patients with pancreatic adenocarcinoma. PD, pancreatoduodenectomy. \ddagger Pearson's χ^2 test.

open surgery (71%), followed by robot-assisted surgery (26%). The median LOS was 12 (i.q.r. 8–20) days with an in-hospital/ 30-day mortality rate of 2.7% (Tables 1–3). Grades of ISGPS-defined POPF, PPH, chyle leak, and ISGLS-defined bile leakage are reported in Table S3.

Feeding strategies

Three distinct feeding strategies were identified: unrestricted feeding (as recommended by ERAS) was used in 637 patients

(27%) across 3 centres, with an 18% incidence of DGE grade B/C; step-up feeding, which starts with fluids and gradually builds up towards normal oral intake, was used in 1462 patients (62%) across 9 centres, with a 24% incidence of DGE grade B/C; and artificial feeding, which starts with nasojejunal feeding or total parenteral nutrition, was used in 255 patients (11%) across 3 centres, with a 25% incidence of DGE grade B/C (P = 0.007, difference in incidence grade B/C DGE between 3 feeding strategies) (Table 3 and Fig. 2).

Table 3 Characteristics of patients after PD (2021-2023)

(1	Overall n = 2354)	Unrestricted feeding $(n = 637)$	Step-up feeding (n = 1462)	Artificial feeding (n = 255)	P†
DGE					0.007
Grade B/C 5	26 (23%)	115 (18%)	348 (24%)	63 (25%)	
Unknown	18	1	17	O	
LOS (days), median (i.q.r.)	12 (8–20)	12 (8–23)	11 (7–18)	13 (9–20)	< 0.001
Unknown	38	13	25	`0	
Major complications [*] 8	57 (37%)	220 (35%)	573 (40%)	64 (25%)	< 0.001
Únknown	28	6	20	2	
POPF					0.424
Grade B/C 4	.89 (22%)	142 (22%)	304 (22%)	43 (18%)	
Unknown	82	1	61	20	
Bile leakage					0.021
Grade B/C 1	44 (6.2%)	53 (8.3%)	80 (5.5%)	11 (4.3%)	
Unknown	21	2	19	0	
РРН					0.467
Grade B/C 2	21 (9.5%)	68 (11%)	130 (9.0%)	23 (9.1%)	
Unknown	24	2	21	1	
Reintervention 8	30 (35%)	208 (33%)	560 (39%)	62 (24%)	< 0.001
Unknown	14	2	12	0	
Reoperation 1	77 (7.6%)	41 (6.5%)	117 (8.1%)	19 (7.5%)	0.420
Unknown	28	3	23	2	
Readmission 4	80 (20%)	112 (18%)	311 (21%)	57 (22%)	0.499
Unknown	150	79	62	9	
Death 6	54 (2.7%)	22 (3.5%)	34 (2.3%)	8 (3.1%)	0.322
Unknown	7	0	7	0	

*Clavien–Dindo grade ≥ III¹⁹. PD, pancreatoduodenectomy; DGE, delayed gastric emptying⁵; LOS, length of hospital stay; i.q.r., interquartile range; POPF, postoperative pancreatic fistula¹⁹; PPH, postpancreatectomy haemorrhage²⁰. †Pearson's χ^2 test, Kruskal–Wallis rank-sum test or Fisher's exact test.



Fig. 2 Centre-specific rate of DGE after PD (2021–2023) and routine feeding strategy per centre in relation to 3-year surgical volume DGE, delayed gastric emptying; PD, pancreatoduodenectomy; c.i., confidence interval.

DGE and LOS

Among the 2354 included patients, 526 (23%) developed DGE grade B/C, with 275 (12%) classified as having primary DGE and 251 (11%) secondary DGE. Median LOS was 23 (i.q.r. 16–34) days in patients with DGE grade B/C and 10 (7–15) days in those without DGE (P < 0.001) (Tables 3, 4, S1, and S3).

Multilevel analysis

Compared with the unrestricted feeding strategy, there was no significant association between step-up feeding (OR 1.14, 95% c.i. 0.53 to 2.47) and artificial feeding (OR 1.76, 0.65 to 4.73) and DGE grade B/C. Similarly, there was no association between feeding strategy and LOS (*Table 5*). A post hoc sensitivity analysis including additional adjustment for pancreatic duct size and texture did not affect the results materially (*Table S5*).

Primary and secondary DGE

In patients with primary DGE, there was no association between step-up feeding (OR 1.18, 95% c.i. 0.58 to 2.40) or artificial feeding (1.78, 0.71 to 4.50) compared with unrestricted feeding for the incidence of DGE. Moreover, LOS did not differ significantly between the step-up and unrestricted feeding strategies, both for patients with primary DGE (22 *versus* 20 days; mean difference 2 (95% c.i. –6 to 9) days) and secondary DGE (32 *versus* 41 days; mean difference –9 (–17 to –1) days). Similarly, no differences in LOS were found between the artificial and unrestricted feeding strategies for primary (22 *versus* 20 days; mean difference 2 (–8 to 12) days) and secondary DGE (33 *versus* 41 days; mean difference –7 (–24 to 10) days) (*Tables* 5 and S2).

Predictors

The following predictors were identified for DGE grade B/C after PD: POPF, CCI, PPH, extended resection, and bile leakage

Table 4 LOS of patients with and without clinically relevant DGE

	Overall (n = 2336)	No DGE (n = 1810)	DGE grade B/C (n = 526)	Р*
LOS (days), median (i.q.r.)	12 (8–20)	10 (7–15)	23 (16–34)	< 0.002
Unknown	33	22	11	

DGE, delayed gastric emptying $^{\rm 5};$ LOS, length of hospital stay; i.q.r., interquartile range. *Wilcoxon rank-sum test.

(P < 0.050). Among all predictors analysed, POPF had the strongest association with DGE (OR 3.16, 95% c.i. 2.47 to 4.05) (Fig. 3).

Post hoc sensitivity analysis—enteral versus parenteral nutrition

A post hoc sensitivity analysis excluding patients receiving parenteral nutrition showed comparable effects on DGE (*Table S4*). In the unadjusted analyses, a statistically significant difference was found between unrestricted feeding (18% DGE), step-up feeding (24%), and artificial feeding (25%) (P = 0.007).

Discussion

In this nationwide study among 2354 patients undergoing PD, no association was found between feeding strategy after surgery (unrestricted, step-up, and artificial feeding) and the incidence of clinically relevant DGE (grade B/C) and LOS. Notably, despite ERAS guidelines⁸ recommending unrestricted feeding, only one-quarter of patients received this approach; despite this, no meaningful difference in DGE or LOS after PD was observed. Patients who developed DGE experienced a median LOS of 13 days longer than those without DGE. Among the predictors assessed, POPF emerged as the strongest determinant of DGE, along with CCI, PPH, extended resection, and bile leakage.

Limited multicentre studies have assessed the specific association between feeding strategy after PD and DGE, making direct comparisons challenging; however, a meta-analysis²⁷ of studies published between 2000 and 2019 evaluated various nutritional approaches, including enteral nutrition after surgery, enteral feeding before surgery, immunonutrition after surgery, and total parenteral nutrition, but did not assess unrestricted oral feeding. Additionally, a more recent single-centre retrospective cohort study²⁸ among 428 patients after PD found that an an early oral feeding strategy was associated with a lower incidence of DGE (7.4 *versus* 15%; P = 0.005) compared with nasojejunal early enteral nutrition.

The 23% incidence of DGE grade B/C in the present study is higher than the 16–19% reported in previous multicentre studies^{4,24,29,30}; however, this discrepancy may be explained by differences in DGE definitions, as not all studies utilized the ISGPS criteria, potentially leading to underestimation in previous reports. Additionally, variations in surgical technique,

Table 5 Multilevel a	analyzcic of foodin	a atratagy in all	notionte and in	thoco with prim	any and cocondany D(ידי
able 5 multilevel a	allalysis of feedill	g su alegy m an	patients and m	ulose with print	ary and secondary Do	JE
	1		1	1	1	

	Total* (n = 2354)	Primary DGE† (n = 1686)	Secondary DGE‡ (n = 668)	
	((n = 275 with DGE)	(n = 251 with DGE)	
Feeding strategy, adjusted OR				
Unrestricted feeding	1.00 (reference)	1.00 (reference)	1.00 (reference)	
Step-up feeding	1.14 (0.53, 2.47)	1.18 (0.58, 2.40)	1.32 (0.49, 3.57)	
Artificial feeding	1.76 (0.65, 4.73)	1.78 (0.71, 4.50)	1.39 (0.36, 5.41)	
LOS, adjusted MD (days)§				
Unrestricted feeding	Reference	Reference	Reference	
Step-up feeding	-2 (-5, 1)	2 (-6, 9)	-9 (-17, -1)	
Artificial feeding (days)	-2 (-6, 2)	2 (-8, 12)	-7 (-24, 10)	

Values in parentheses are 95 per cent confidence intervals. *Patients without delayed gastric empyting (DGE)⁵ were included in primary and secondary DGE groups for the multilevel analysis; the exact numbers of patients with DGE are shown. †Patients without any of the following: bile leakage (grade B/C)²³, postpancreatectomy haemorrhage (PPH) (grade B/C)²¹, POPF (grade B/C)²⁰. ‡Patients with any of the following: bile leakage (grade B/C)²¹, POPF (grade B/C)²⁰. §Patients with any of the following: bile leakage (grade B/C)²¹, POPF (grade B/C)²⁰. §Patients with any of the following: bile leakage (grade B/C)²¹, POPF (grade B/C)²⁰. §Patients with any of the following: bile leakage (grade B/C)²¹, POPF (grade B/C)²⁰. §Multilevel analysis corrected for centre (random effect), sex, age, body mass index, American Society of Anesthesiologists grade, Charlson Co-morbidity Index score, pre-existing diabetes, preoperative biliary drainage, type of surgery (pylorus resection versus pylorus-preserving), type of anastomosis (pancreatojejunostomy versus pancreatogastrostomy), vascular resection, extended resection (in addition to the primary tumour, adjacent structures or organs were removed, such as mesocolon transversum, gastric resection), minimally invasive pancreatoduodenectomy, pancreatic adenocarcinoma, POPF, PPH grade B/C, and bile leakage grade B/C. OR, odds ratio; LOS, length of hospital stay; MD, mean difference.



Fig. 3 Predictors of DGE grade B/C after PD

*In addition to the primary tumour, adjacent structures or organs were removed (such as mesocolon transversum, gastric resection). †Preoperative biliary stenting, ‡Pre-existing diabetes. §Pancreatojejunostomy versus pancreatogastrostomy. ¶Pylorus resection versus pylorus-preserving. DGF, delayed gastric emptying; PD, pancreatoduodenectomy; POPF, postoperative pancreatic fistula; PPH, postpancreatectomy haemorrhage grade B/C; CCI, Charlson Co-morbidity Index; ASA, American Society of Anesthesiologists; PDAC, pancreatic ductal adenocarcinoma; MIPD, minimally invasive pancreatoduodenectomy; BMI, body mass index.

patient population, or care protocols after surgery may have contributed to differences in incidence of DGE across studies. Previously, audit-based studies^{4,29} using the American College of Surgeons National Surgical Quality Improvement Program reported an incidence of DGE of around 15%, but did not assess feeding strategies after surgery in relation to DGE.

Several studies^{31,32} have explored predictive factors for DGE to identify potential targets for prevention. A Swedish audit-based study³⁰ involving 2503 patients after PD also reported POPF as the strongest independent predictor of DGE, which is in line with the present findings. Additionally, the authors found that pylorus-preserving PD (PPPD) and reconstruction with a pancreatogastrostomy were associated with a lower risk of DGE. A pylorus preservation rate of 20% was reported, with a DGE incidence of 19%³⁰. In comparison, the present study had a higher pylorus preservation rate of 34%, but a comparable DGE incidence of 23%. Notably, type of resection was not a significant predictor of DGE in the analysis, a finding consistent with previous meta-analysis³³ showing no clear advantage of pylorus-resecting PD over PPPD in reducing DGE or other complications.

This study represents the largest cohort regarding feeding strategy after PD in relation to DGE. Despite the identified risk factors, the underlying mechanisms of DGE and strategies for its prevention remain unclear. As the primary outcomes did not differ between the three feeding strategies, surgeons should also take secondary downsides into account. For instance, nasojejunal feeding could cause more patient discomfort compared with step-up or unrestricted feeding. Additionally, costs of artificial feeding could be higher compared with those of step-up or unrestricted diet; therefore, the present findings create a new insight into feeding strategies after surgery and could contribute to future changes in national protocols and ERAS guidelines³⁴.

Several limitations must be considered when interpreting the results of this study. First, the retrospective nature of the analysis carries inherent risks of bias and confounding, which may affect the validity of findings³⁵. To mitigate these concerns, the analysis adjusted for confounders and centre-level effects. was Additionally, interviews revealed that feeding strategies changed over time within hospitals, making it increasingly difficult to recall feeding strategies that were used in the past. To minimize recall bias, the authors restricted the analysis to the most recent 3 years (2021-2023), during which hospitals maintained a single feeding strategy after surgery; however, protocol deviations could have occurred. Second, the categorization of feeding strategies may have introduced some degree of misclassification bias, as some differences could still exist between hospitals in the same category. Third, placement of an NGT for gastric decompression following PD was not recorded in the DPCA, and potentially this could have influenced the incidence and severity of DGE. Fourth, although the results did not show a clinically relevant benefit (such as 10% fewer cases of DGE), it is possible that the strategies do slightly influence the main outcome. Fifth, the present Dutch audit data set does not capture potentially relevant clinical parameters, such as grade A complications, Clavien–Dindo I–II complications, 90-day mortality and baseline nutritional status. Although grade B/C complications are considered clinically relevant, it is possible that grade A complications may still influence recovery after surgery. Furthermore, 90-day mortality might have provided more insight into potential differences between groups, but was not available. Additionally, baseline nutritional status was not captured in the DPCA; therefore, the authors were unable to account for this, although potentially it could still have influenced DGE incidence. Sixth, the artificial feeding category included total parenteral nutrition (1 centre) and enteral feeding (2 centres). A much larger study population would be required to adequately power an

analysis capable of detecting clinically meaningful differences³⁶. The primary strength of this study lies in its large, nationwide cohort, which provides a comprehensive evaluation of feeding strategies after surgery and their impact on DGE.

DGE remains a challenging complication after PD, influenced by multiple different factors. Future studies should prospectively compare unrestricted feeding (ERAS guidelines) with alternative feeding strategies in randomized trials to provide high-level evidence for clinical guidelines. Additionally, investigating patient-centred outcomes, such as quality of life and functional recovery, in relation to different feeding strategies, could provide a more comprehensive assessment of their benefits. Integrating multimodal approaches, including nutritional support, pharmacological interventions, and enhanced recovery protocols, could further optimize outcomes for patients undergoing PD.

In conclusion, this nationwide study found no evidence to support a specific feeding strategy after PD (unrestricted feeding, step-up feeding, and artificial feeding) to reduce the incidence of DGE grade B/C and LOS. Nevertheless, these findings highlight the need for targeted interventions to reduce DGE, particularly by addressing modifiable risk factors, such as POPF.

Collaborators

J. Haver, Department of Nutrition and Dietetics, Amsterdam University Medical Center, University of Amsterdam, Amsterdam, the Netherlands. E. Steenhagen, Department of Dietetics, University Medical Center Utrecht, Utrecht, the Netherlands.

Funding

The authors have no funding to declare.

Author contributions

Tessa Hendriks (Conceptualization, Data curation, Formal analysis, Investigation, Methodology, Project administration, Resources, Software, Supervision, Validation, Visualization, Writingoriginal draft, Writing-review & editing), Bo T. M. Strijbos (Conceptualization, Data curation, Investigation, Resources, Software, Supervision, Visualization, Writing-original draft), Michiel Francken (Conceptualization, Formal analysis, Investigation, Methodology, Project administration, Resources, Software, Supervision, Validation, Writing-original draft, Writing-review & editing), Mahsoem Ali (Formal analysis, Methodology, Resources, Supervision, Validation, Visualization, Writing-review & editing), J. Annelie Suurmeijer (Conceptualization, Methodology, Supervision, Validation, Writing-review & editing), Marcel Dijkgraaf (Formal analysis, Investigation, Methodology, Supervision, Validation, Writing-review & editing), Kees van Laarhoven (Supervision, Writing-review & editing), Jana Hopstaken (Supervision), Quintus Molenaar (Data curation, Supervision, Writing-review & editing), Vincent de Meijer (Supervision, Writing-review & editing), Erwin Van der Harst (Data curation, Supervision, Writing-review & editing), Marcel den Dulk (CRediT contribution not specified), Warner Draaisma (Data curation, Supervision, Writing-review & editing), Vincent Nieuwenhuijs (Data curation, Supervision, Writing-review & editing), Michael Gerhards (Data curation, Supervision, Writingreview & editing), Mike Liem (Data curation, Supervision, Writing-review & editing), George van der Schelling (Data curation, Supervision, Writing-review & editing), E. R. Manusama

(Data curation, Supervision, Writing—review & editing), Ignace de Hingh (Data curation, Supervision, Writing—review & editing), Olivier Busch (Data curation, Supervision, Validation, Writing review & editing), Bas Grootkoerkamp (Conceptualization, Data curation, Methodology, Supervision, Validation, Writing—original draft, Writing—review & editing), Hjalmar van Santvoort (Data curation, Methodology, Supervision, Writing—review & editing), Bert Bonsing (Conceptualization, Methodology, Resources, Supervision, Validation, Visualization, Writing—original draft, Writing—review & editing), Martijn Stommel (Data curation, Methodology, Resources, Supervision, Validation, Writing—review & editing), and Marc Besselink (Conceptualization, Data curation, Methodology, Resources, Software, Supervision, Validation, Visualization, Writing—original draft, Writing—review & editing)

Disclosure

The authors declare no conflict of interest.

Supplementary material

Supplementary material is available at BJS Open online.

Data availability

Study data are available on request to the corresponding author.

References

- Sánchez-Velázquez P, Muller X, Malleo G, Park JS, Hwang HK, Napoli N et al. Benchmarks in pancreatic surgery: a novel tool for unbiased outcome comparisons. Ann Surg 2019;270:211–218
- Cameron JL, He J. Two thousand consecutive pancreaticoduodenectomies. J Am Coll Surg 2015;220:530–536
- Snyder RA, Ewing JA, Parikh AA. Delayed gastric emptying after pancreaticoduodenectomy: a study of the national surgical quality improvement program. *Pancreatology*. 2020;20:205–210
- Dominguez OH, Grigorian A, Wolf RF, Imagawa DK, Nahmias JT, Jutric Z. Delayed gastric emptying is associated with increased risk of mortality in patients undergoing pancreaticoduodenectomy for pancreatic adenocarcinoma. Updates Surg 2023;75:523–530
- Wente MN, Bassi C, Dervenis C, Fingerhut A, Gouma DJ, Izbicki JR et al. Delayed gastric emptying (DGE) after pancreatic surgery: a suggested definition by the International Study Group of Pancreatic Surgery (ISGPS). Surgery 2007;142:761–768
- Jasmijn Smits F, Verweij ME, Daamen LA, van Werkhoven C H, Goense L, Besselink MG et al. Impact of complications after pancreatoduodenectomy on mortality, organ failure, hospital stay, and readmission: analysis of a nationwide audit. Ann Surg 2022;275:E222–E228
- Francken MFG, van Roessel S, Swijnenburg RJ, Erdmann JI, Busch OR, Dijkgraaf MGW *et al.* Hospital costs of delayed gastric emptying following pancreatoduodenectomy and the financial headroom for novel prophylactic treatment strategies. HPB 2021;23:1865–1872
- Melloul E, Lassen K, Roulin D, Grass F, Perinel J, Adham M et al. Guidelines for perioperative care for pancreatoduodenectomy: Enhanced Recovery After Surgery (ERAS) recommendations 2019. World J Surg 2020;44:2056–2084

- 9. Federatie Medisch Specialisten. Richtlijn Pancreascarcinoom. https://richtlijnendatabase.nl/richtlijn/pancreascarcinoom (accessed 26 July 2024)
- Hwang SE, Jung MJ, Cho BH, Yu HC. Clinical feasibility and nutritional effects of early oral feeding after pancreaticoduodenectomy. Korean J Hepatobiliary Pancreat Surg 2014;18:84–89
- 11. Halle-Smith JM, Pathak S, Frampton A, Pandanaboyana S, Sutcliffe RP, Davidson BR *et al.* Current postoperative nutritional practice after pancreatoduodenectomy in the UK: national survey and snapshot audit. *BJS Open* 2024;**8**:zrae021
- Bassi C, Marchegiani G, Giuliani T, Di Gioia A, Andrianello S, Zingaretti CC et al. Pancreatoduodenectomy at the Verona Pancreas Institute: the evolution of indications, surgical techniques, and outcomes: a retrospective analysis of 3000 consecutive cases. Ann Surg 2022;276:1029–1038
- Halle-Smith JM, Pande R, Powell-Brett S, Pathak S, Pandanaboyana S, Smith AM et al. Early oral feeding after pancreatoduodenectomy: a systematic review and meta-analysis. HPB 2022;24:1615–1621
- van Rijssen LB, Koerkamp BG, Zwart MJ, Bonsing BA, Bosscha K, van Dam RM et al. Nationwide prospective audit of pancreatic surgery: design, accuracy, and outcomes of the Dutch Pancreatic Cancer Audit. HPB (Oxford) 2017;19:919–926
- Van Der Werf LR, Voeten SC, Van Loe CMM, Karthaus EG, Wouters MWJM, Prins HA. Data verification of nationwide clinical quality registries. BJS Open 2019;3:857–864
- 16. Strijker M, Mackay TM, Bonsing BA, Bruno MJ, Van Eijck CHJ, De Hingh IHJT et al. Establishing and coordinating a nationwide multidisciplinary study group: lessons learned by the Dutch Pancreatic Cancer Group. Ann Surg 2020;**271**:E102–E104
- von Elm E, Altman DG, Egger M, Pocock SJ, Gøtzsche PC, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement: guidelines for reporting observational studies. *Lancet* 2007;**370**: 1453–1457
- Clavien PA, Barkun J, De Oliveira ML, Vauthey JN, Dindo D, Schulick RD et al. The Clavien–Dindo classification of surgical complications: five-year experience. Ann Surg 2009;250:187–196
- Bassi C, Marchegiani G, Dervenis C, Sarr M, Abu Hilal M, Adham Met al. The 2016 update of the International Study Group (ISGPS) definition and grading of postoperative pancreatic fistula: 11 years after. Surgery 2017;161:584–591
- Wente MN, Veit JA, Bassi C, Dervenis C, Fingerhut A, Gouma DJ et al. Postpancreatectomy hemorrhage (PPH): an International Study Group of Pancreatic Surgery (ISGPS) definition. Surgery 2007;142:20–25
- Besselink MG, van Rijssen LB, Bassi C, Dervenis C, Montorsi M, Adham M et al. Definition and classification of chyle leak after pancreatic operation: a consensus statement by the International Study Group on Pancreatic Surgery. Surgery 2017;161:365–372
- 22. Koch M, Garden OJ, Padbury R, Rahbari NN, Adam R, Capussotti L et al. Bile leakage after hepatobiliary and pancreatic surgery: a definition and grading of severity by the International Study Group of Liver Surgery. Surgery 2011;**149**:680–688

- Noorani A, Rangelova E, Del Chiaro M, Lundell LR, Ansorge C. Delayed gastric emptying after pancreatic surgery: analysis of factors determinant for the short-term outcome. Front Surg 2016;3:25
- 24. Marchegiani G, Di Gioia A, Giuliani T, Lovo M, Vico E, Cereda M et al. Delayed gastric emptying after pancreatoduodenectomy: one complication, two different entities. *Surgery* 2023;**173**: 1240–1247
- 25. Harrell FE. Regression Modeling Strategies. New York: Springer New York, 2001.
- Bates D, Mächler M, Bolker BM, Walker SC. Fitting linear mixed-effects models using lme4. J Stat Softw 2015;67:1–48
- 27. Wang SY, Hung YL, Hsu CC, Hu CH, Huang RY, Sung CM et al.Optimal perioperative nutrition therapy for patients undergoing pancreaticoduodenectomy: a systematic review with a component network meta-analysis. Nutrients 2021;**13**: 4049
- Jing W, Wu S, Gao S, Shi X, Liu W, Ren Y et al. Early oral feeding versus nasojejunal early enteral nutrition in patients following pancreaticoduodenectomy: a propensity scoreweighted analysis of 428 consecutive patients. Int J Surg 2024; 110:229
- 29. Werba G, Sparks AD, Lin PP, Johnson LB, Vaziri K. The PrEDICT– DGE score as a simple preoperative screening tool identifies patients at increased risk for delayed gastric emptying after pancreaticoduodenectomy. HPB 2022;**24**:30–39
- Zdanowski AH, Wennerblom J, Rystedt J, Andersson B, Tingstedt B, Williamsson C. Predictive factors for delayed gastric emptying after pancreatoduodenectomy: a Swedish National Registry-based study. World J Surg 2023;47:3289–3297
- Robinson JR, Marincola P, Shelton J, Merchant NB, Idrees K, Parikh AA. Peri-operative risk factors for delayed gastric emptying after a pancreaticoduodenectomy. HPB (Oxford) 2015;17:495–501
- 32. Ellis RJ, Gupta AR, Hewitt DB, Merkow RP, Cohen ME, Ko CY et al.Risk factors for post-pancreaticoduodenectomy delayed gastric emptying in the absence of pancreatic fistula or intra-abdominal infection. J Surg Oncol 2019;119:925
- 33. Klaiber U, Probst P, Strobel O, Michalski CW, Dörr-Harim C, Diener MK et al. Meta-analysis of delayed gastric emptying after pylorus-preserving versus pylorus-resecting pancreatoduodenectomy. Br J Surg 2018;105:339–349
- Gerritsen A, Wennink RAW, Besselink MGH, Van Santvoort HC, Tseng DSJ, Steenhagen E et al. Early oral feeding after pancreatoduodenectomy enhances recovery without increasing morbidity. HPB 2014;16:656–664
- 35. Hendriks TE, Balduzzi A, van Dieren S, Suurmeijer JA, Salvia R, Stoop TF et al. Interobserver variability in the International Study Group for Pancreas Surgery (ISGPS)-defined complications after pancreatoduodenectomy: an international cross-sectional multicenter study. Ann Surg 2024;280:728–733
- 36. de Graaff MR, Hendriks TE, Wouters M, Nielen M, de Hingh I, Koerkamp BG et al. Assessing quality of hepato-pancreatobiliary surgery: nationwide benchmarking. Br J Surg 2024;111: znae119