# World Journal of Gastrointestinal Surgery

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## World Journal of Gastrointestinal Surgery

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#### **ABOUT COVER**

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The primary aim of World Journal of Gastrointestinal Surgery (WJGS, World J Gastrointest Surg) is to provide scholars and readers from various fields of gastrointestinal surgery with a platform to publish high-quality basic and clinical research articles and communicate their research findings online.

WJGS mainly publishes articles reporting research results and findings obtained in the field of gastrointestinal surgery and covering a wide range of topics including biliary tract surgical procedures, biliopancreatic diversion, colectomy, esophagectomy, esophagostomy, pancreas transplantation, and pancreatectomy, etc.

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**Observational Study** 

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ORIGINAL ARTICLE

## Application of omental interposition to reduce pancreatic fistula and related complications in pancreaticoduodenectomy: A propensity score-matched study

Yang Li, Yun Liang, Yao Deng, Zhi-Wei Cai, Ming-Jian Ma, Long-Xiang Wang, Meng Liu, Hong-Wei Wang, Chong-Yi Jiang

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#### Abstract

#### BACKGROUND

The life-threatening complications following pancreatoduodenectomy (PD), intraabdominal hemorrhage, and postoperative infection, are associated with leaks from the anastomosis of pancreaticoduodenectomy. Although several methods have attempted to reduce the postoperative pancreatic fistula (POPF) rate after PD, few have been considered effective. The safety and short-term clinical benefits of omental interposition remain controversial.

#### AIM

To investigate the safety and feasibility of omental interposition to reduce the POPF rate and related complications in pancreaticoduodenectomy.

#### **METHODS**

In total, 196 consecutive patients underwent PD performed by the same surgical team. The patients were divided into two groups: An omental interposition group (127, 64.8%) and a non-omental interposition group (69, 35.2%). Propensity scorematched (PSM) analyses were performed to compare the severe complication rates and mortality between the two groups.

#### RESULTS

Following PSM, the clinically relevant POPF (CR-POPF, 10.1% vs 24.6%; P = 0.025) and delayed postpancreatectomy hemorrhage (1.4% vs 11.6%; P = 0.016) rates were significantly lower in the omental interposition group. The omental inter-



position technique was associated with a shorter time to resume food intake (7 d vs 8 d; P = 0.048) and shorter hospitalization period (16 d vs 21 d; P = 0.031). Multivariate analyses showed that a high body mass index, nonapplication of omental interposition, and a main pancreatic duct diameter < 3 mm were independent risk factors for CR-POPF.

#### **CONCLUSION**

The application of omental interposition is an effective and safe approach to reduce the CR-POPF rate and related complications after PD.

Key Words: Pancreaticoduodenectomy; Pancreatic fistula; Complication; Omental interposition

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Core Tip: Postoperative pancreatic fistula (POPF) is a life-threatening complication after pancreaticoduodenectomy. Multiple methods have been described in the literature to prevent POPF; however, few trials have demonstrated that a certain method can achieve good clinical outcomes. In this study, we proved that the application of omental interposition can reduce the incidence of clinically relevant POPF, which is associated with a trend towards accelerated recovery.

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#### INTRODUCTION

Pancreaticoduodenectomy is the gold standard for benign or malignant tumors in the periampullary region. Despite advances in surgical techniques and perioperative care, the postoperative morbidity rate remains high (20-50%), even in high-volume comprehensive hospitals[1-3]. Postoperative pancreatic fistula (POPF) is a life-threatening complication because of its interrelationship with delayed postpancreatectomy hemorrhage (PPH) and postoperative intraabdominal infection[4]. POPF is responsible for erosion of the gastroduodenal artery stump (GDAS), skeletonized hepatic artery (HA), or other adjacent abdominal vessels due to activated pancreatic enzymes.

During the last quarter of the 20<sup>th</sup> century, multiple methods have been described in the literature to prevent POPF and subsequent complications, including the usage of somatostatin or octreotide, introduction of pancreatic duct stenting, creation of various anastomosis techniques (e.g., duct-tomucosa, pancreatogastrostomy, invagination), use of polyethylene glycolic acid mesh to reinforce around the pancreatojejunostomy (PJ) site, and use of fibrin glue over the PJ site[5-9]. However, few trials have demonstrated that a certain method will reinforce the PJ site in PD with favorable clinical outcomes.

Currently, the greater omentum has been widely used to reinforce anastomoses and compensate for tissue defects in the fields of thoracic, urinary, and general surgery [10-12]. Recently, some centers have shown that fixing the omental interposition behind the anastomotic site of the PJ to protect the GDAS and nearby HA from erosive pancreatic juices is the most promising approach to reduce the incidence of severe complications<sup>[13,14]</sup>, but they did not have control group data.

Our study investigated whether the application of the omental interposition could effectively reduce the incidence of POPF and its related complications after pancreaticoduodenectomy.

#### MATERIALS AND METHODS

#### Patients

Between January 2015 and December 2019, 196 consecutive patients underwent pancreaticoduodenectomy performed by the same surgical team at our institution. The first 69 consecutive patients did not use omental interpositions, and the remaining 127 used omental interpositions. According to whether the omental interposition was applied, the patients were divided into two groups: the omental group (79 males, 48 females; mean age: 64.8 years) and the non-omental interposition group (44 males, 25 females; mean age: 62.1 years). Propensity score matching (PSM) was used to minimize bias from the



nonrandomized treatment assignments. We summarized the data on the general clinical characteristics, short-term surgical outcomes, and recovery. Moreover, the laboratory data on the drain fluid amylase obtained on the first postoperative day (DFA1) were pooled. All data were prospectively collected in our electronic media database. This study was approved by the ethics review committee of Huadong Hospital Affiliated to Fudan University (2019K087; Shanghai, China).

#### Surgical technique

At our institution, PD was accomplished with a standard approach. After the head of the pancreas had been removed, intestinal reconstruction was achieved with a modified version of the method described by Child. A reconstruction PJ was performed (by duct-to-mucosa, end-to-side reconstruction) and a pancreatic drainage tube was placed. (1) Insert the pancreatic juice drainage tube into 3-5 cm and use 4-0 polydioxanone suture to insert the needle from the ventral side of the pancreatic duct, penetrate the anterior and posterior walls of the pancreatic juice drainage tube, and suture from the back of the pancreatic duct to fix the drainage tube; (2) Place the pancreatic juice drainage tube into the distal end of the jejunal loop, and purse suture of the jejunal incision; and (3) Use 3-0 prolene to suture of seromuscular layer of pancreas and jejunum. Hepaticojejunostomy (HJ) was performed with continuous barbed sutures or interrupted sutures. Gastrojejunostomy (GJ) was performed with interrupted 3-0 polypropylene monofilament sutures.

In the omental interposition group, following complete anastomosis, we routinely placed a pedicled omental interposition in front of the adjacent vessels (HA, PV, and GDAS) and behind the anastomosis where the pancreas stump was fixed to the jejunum<sup>[15]</sup>. The omental interposition was fixed to the hepatic portal and hepatogastric ligament with several sutures to prevent postoperative mobilization (Figure 1). Generally, the upper boundary of the omental interposition was the level of the hepatogastric ligament, the left boundary was the level of the pancreatic body, and the right boundary was the right margin of the inferior vena cava, so that the omental interposition could separate skeletonized vessels from a possible anastomotic leakage. Then two double catheterization cannulas (PJ tube and HJ tube) were placed at the left anterior of the PJ anastomosis site and right posterior of the HJ anastomosis site, respectively. The blood flow of the omental interposition was reconfirmed before the abdominal cavity was closed. The application of the omental interposition in PD is shown in Figure 2.

In the non-omental interposition group, we simply placed the two drainage tubes at the aforementioned positions after completing the anastomosis. After the operation, the amylase concentration from the drainage fluid was measured daily. If the drain fluid amylase obtained on DFA1 exceeded 2000 U/L, abdominal irrigation was used to dilute the concentration of pancreatic juice around the anastomosis as soon as possible. Approximately 3000 mL normal saline was irrigated every day, with a flow rate of 200 mL/h. The flow of irrigation was modulated frequently according to the character of the secretion. The suction pressure was set with low-pressure suction between 20 and 30 cm water. Once the amylase level of the dilution fluid was lower than 30 U/L, the use of abdominal irrigation was stopped. The drainage tubes were removed until the amylase concentration was less than three times the upper limit of the normal serum level. All patients underwent routine postoperative computed tomography (CT) examinations before the drain tubes were removed to assess the presence of potential complications and peritoneal effusion.

#### Definitions

POPF was defined and graded according to the modified definition by the International Study Group of Pancreatic Fistula (ISGPF)[16]. Clinically relevant POPF (CR-POPF) was considered grades B and C. Delayed gastric emptying (DGE) and PPH were defined and classified by the International Study Group for Pancreatic Surgery[17,18]. Intra-abdominal infections were diagnosed according to the definition proposed by the Surgical Infection Society and the Infectious Diseases Society of America[19].

#### Statistical analysis

All statistical analyses were conducted using SPSS 23.0. The  $\chi^2$  test or Fisher's exact test was used for categorical variables, whereas the Student's t-test or Wilcoxon rank-sum test (whether the variables were normally distributed) were used for continuous variables. P < 0.05 was considered statistically significant. After matching, each patient who received an omental interposition was matched to a patient in the non-omental interposition group by using nearest-neighbor matching in a 1:1 ratio. A PSM analysis was used to reduce the impact of the treatment selection bias when estimating the omental interposition values using original observational indicators. Multivariable logistic regression was performed with adjustments for the propensity scores using the associated covariates.

#### RESULTS

#### Analyses of all unmatched patients

The demographic and clinically related variables of all patients including age, sex, body mass index





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Figure 1 Omental interposition was placed in front of the adjacent vessels and behind the anastomosis where the pancreas stump was fixed to the jejunum. 1: Liver; 2: Portal vein; 3: Hepatic artery; 4: Common bile duct; 5: Hepaticojejunostomy; 6: Gastrojejunostomy; 7: Celiac artery; 8: Pancreaticojejunostomy site; 9: Gastrojejunostomy. site; 10: Omental interposition; 11: Transverse colon.

(BMI), American Society of Anesthesiologists score, serum albumin content, main pancreatic duct size and pathology were similar between the two groups (P > 0.05). However, patients in the omental interposition group had a higher median serum bilirubin than those in the non-omental interposition group (96.5 [17.9-107.0] *vs* 20.5 [9.6-148.5]; P = 0.015). Laparoscopic pancreaticoduodenectomy (LPD) was more frequently performed in the omental interposition group than in the non-omental interposition group (69, 54.3% *vs* 19, 27.5%; P < 0.001). The details are shown in Table 1.

Regarding postoperative complications, a comparison revealed that the rates of CR-POPF (13, 10.2% *vs* 17, 24.6%; *P* = 0.028), biliary fistula (BF, 2,1.6% *vs* 5, 7.2%; *P* = 0.041), delayed PPH associated with POPF (1, 0.8% *vs* 8, 11.6%; *P* = 0.002), and postoperative transfusion (18,14.2% *vs* 20, 29.0%; *P* = 0.012) were significantly lower in the omental interposition group than in the non-omental interposition group. The rates of other surgery-related complications, including DGE, intra-abdominal abscess, and reoperation, did not significantly differ between the two groups. Regarding mortality, there was no significant difference between the two groups (2, 1.6% *vs* 5, 7.2%; *P* = 0.101). However, the CR-POPF-related mortality in the omental interposition group was significantly lower than the mortality in the non-omental interposition group (1, 0.8% *vs* 5, 7.2%; *P* = 0.021). The details on the deaths that occurred are shown in Table 2 and Figure 3. Fewer complications in the omental group may be related to passing the laparoscopic learning curve. However, among the 108 cases of OPD, 58 cases applied the omental interposition group had lower incidence of complications (6, 10.3% *vs* 9, 18%; *P* = 0.008) and lower mortality rate (0, 0% *vs* 4, 8%; *P* = 0.007).

When comparing relevant data on the enhanced recovery after surgery between the two groups, the HJ and PJ drainage tubes were removed earlier in the omental interposition group than in the non-omental interposition group (both P < 0.05). The omental interposition group of patients had significantly shorter postoperative durations of restarting their diet and shorter length of hospital stay than the non-omental interposition group patients (both P < 0.01). Based on the laboratory test results, the DFA1 around the HJ in the omental interposition group was dramatically lower than that in the non-omental interposition group (300.0 [74.3-893.0] *vs* 599.8 [171.1-2064.7]; P = 0.002). In the omental interposition group, the drain amylase values from the tube around the HJ were lower than those around the PJ (300.0 [74.3-893.0] *vs* 546.8 [76.4-3094.0]; P < 0.001). However, the difference disappeared in the non-omental interposition group. The details are shown in Table 2 and Figure 4A.

#### Analyses of all matched patients

To reduce the impact of selection bias and the role of the procedure (LPD and OPD), PSM was performed using nine selected baseline characteristics. After PSM, the patient demographic and clinically related characteristics, including preoperative serum bilirubin and operation methods, were similar between the two groups. The rates of CR-POPF (7, 10.1% *vs* 17, 24.6%; *P* = 0.025), delayed PPH associated with POPF (1, 1.4% *vs* 8, 11.6%; *P* = 0.016) and postoperative transfusion (9, 13.0% *vs* 20, 29.0%; *P* = 0.022) remained significantly lower in the omental interposition group than in the non-



Table 1 Comparisons of patients' characteristics between the two groups							
	Before PSM			After PSM			
	Omental interposition group (127)	Non-omental interposition group (69)	P value	Omental interposition group (69)	Non-omental interposition group (69)	P value	
Male/female	79/48	44/25	0.919	46/23	44/25	0.721	
Age (yr)	$64.8\pm10.5$	$62.1 \pm 9.9$	0.083	$64.2 \pm 9.5$	62.1 ± 9.9	0.210	
BMI (mean $\pm$ SD, kg/m <sup>2</sup> )	21.9 ± 3.0	$22.0 \pm 2.8$	0.844	21.9 ± 3.2	22.0 ± 2.9	0.933	
ASA score, $n$ (%)			0.126			0.168	
Ι	65 (51.2)	42 (60.9)		34 (49.3)	42 (60.9)		
П	60 (47.2)	24 (34.8)		34 (49.3)	24 (34.8)		
III	2 (1.6)	3 (4.3)		1 (1.4)	3 (4.3)		
Serum ALB [ $n$ (%), g/L]			0.152			1.00	
< 35	13 (10.2)	12 (17.4)		12 (17.4)	12 (17.4)		
≥ 35	114 (89.8)	57 (82.6)		57 (82.6)	57 (82.6)		
Serum bilirubin (µmol/L)	96.5 (17.9-107.0)	20.5 (9.6-148.5)	0.015	29.8 (12.4-153.7)	20.5 (9.6-148.5)	0.753	
Main pancreatic duct size $[n (\%), mm]$			0.080			0.173	
< 3	57 (44.9)	40 (58.0)		32 (46.4)	40 (58.0)		
≥3	70 (55.1)	29 (42.0)		37 (53.6)	29 (42.0)		
Operation method, <i>n</i> (%)			0.005			0.708	
LPD	69 (54.3)	19 (27.5)		21 (30.4)	19 (27.5)		
OPD	58 (45.7)	50 (72.5)		48 (69.6)	50 (72.5)		
Pathology, n (%)			0.009			0.151	
PDAC	53 (41.7)	25 (36.2)		36 (52.2)	25 (36.2)		
Bile duct cancer	10 (7.9)	13 (18.8)		4 (5.8)	13 (18.8)		
Ampulla of Vater cancer	18 (14.2)	15 (21.7)		10 (14.5)	15 (21.7)		
Duodenal cancer	11 (8.7)	2 (2.9)		2 (2.9)	2 (2.9)		
Other carcinoma	19 (15.0)	2 (2.9)		3 (4.3)	2 (2.9)		
Benign tumor	16 (12.6)	12 (17.4)		14 (20.3)	12 (17.4)		

ALB: Albumin; ASA: American Society of Anesthesiologist score; BMI: Body mass index LPD: Laparoscopic pancreaticoduodenectomy; OPD: Open pancreaticoduodenectomy.

omental interposition group after PSM. The operation time in the omental interposition group was slightly longer both before ( $388.3 \pm 68.8 vs 365.2 \pm 75.0$ ) and after ( $392.6 \pm 74.1 vs 365.2 \pm 75.0$ ) the match, which may be related to the selection, cutting, and fixing of the omental interposition. Moreover, the omental interposition group of patients had a significantly shorter postoperative duration to restart their diet (7 [5-8] vs 8 [6-15]; P = 0.048) and shorter hospital stays (16 [12-24] vs 21 [13-32]; P = 0.031] than the non-omental interposition group of patients. The non-omental interposition group had greater mortality related to POPF than the omental interposition group (5 [7.2%] vs 1 [1.4%]), but there was no significant difference, which may be related to the small number of cases. The details are shown in Table 2.

Following PSM, the omental interposition group had dramatically lower DFA1 around the HJ than the non-omental interposition group (200.0 [58-610.6] *vs* 599.8 [171.1-2064.7] P = 0.003). In the omental interposition group, the DFA1 around the HJ was lower than the DFA1 around the PJ (200.0 [58-610.6] *vs* 325.0 [75.3-2869], P < 0.001). The details on DFA1 are shown in Table 2 and Figure 4B.

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Table 2 Comparisons of the postoperative outcomes between the two groups							
	Before PSM			After PSM			
	Omental interposition group (127)	Non-omental interposition group (69)	P value	Omental interposition group (69)	Non-omental interposition group (69)	P value	
CR-POPF	13 (10.2%)	17 (24.6%)	0.028	7 (10.1%)	17 (24.6%)	0.025	
Operation time (mean ± SD, min)	388.3 ± 68.8	365.2 ± 75.0	0.031	392.6 ± 74.1	365.2 ± 75.0	0.033	
BF, n (%)	2 (1.6)	5 (7.2)	0.041	1 (1.4)	5 (7.2)	0.208	
DGE, n (%)	4 (3.1)	6 (8.7)	0.178	1 (1.4)	6 (8.7)	0.115	
PPH, n (%)	1 (0.8)	8 (11.6)	0.002	1 (1.4)	8 (11.6)	0.016	
Intra-abdominal abscess, <i>n</i> (%)	15 (11.8)	12 (17.4)	0.286	8 (11.6)	12 (17.4)	0.333	
Reoperation, n (%)	3 (2.4)	6 (8.7)	0.096	2 (2.9)	6 (8.7)	0.274	
Mortality in 30 d, $n$ (%)	2 (1.6)	5 (7.2)	0.101	2 (2.9)	5 (7.2)	0.438	
Mortality related to POPF, <i>n</i> (%)	1 (0.8)	5 (7.2)	0.038	1 (1.4)	5 (7.2)	0.210	
DFA1 around the HJ site (U/L)	300.0 (74.3-893.0)	599.8 (171.1-2064.7)	0.002	200.0 (57.5-659.8)	599.8 (171.1-2064.7)	0.003	
DFA1 around the PJ site (U/L)	546.8 (76.4-3094.0)	350.0 (50.0-2577.4)	0.255	325.0 (69.5-2972.5)	350.0 (50.0-2577.4)	0.951	
Duration until removal of the tube around the HJ site (d)	7 (5-9)	9 (7-14)	0.000	8 (6-11)	9 (7-14)	0.115	
Duration until removing the tube around the PJ site (d)	7 (6-11)	10 (7-15)	0.004	8 (6-12)	10 (7-15)	0.100	
Required blood transfusions, <i>n</i> (%)	18 (14.2)	20 (29.0)	0.012	9 (13.0)	20 (29.0)	0.022	
Length of hospital stay (d)	15 (11-22)	21 (13-32)	0.004	16 (12-24)	21 (13-32)	0.031	
Duration until restarting diet (d)	6 (5-8)	8 (6-15)	0.001	7 (5-8)	8 (6-15)	0.048	

BF: Biliary fistula; CR-POPF: Clinically relevant postoperative pancreatic fistula; DFA1: Drain fluid amylase obtained on the first postoperative day; DGE: Delayed gastric emptying; HJ: Hepaticojejunostomy; PJ: Pancreaticojejunostomy; PPH: Postpancreatectomy hemorrhage.

#### Factors associated with CR-POPF after PD

Table 3 shows the univariate and multivariate analyses of the PSM data to evaluate the risk factors associated with CR-POPF after PD. Male sex,  $BMI \ge 23 \text{ kg/m}^2$ , nonapplication of omental interposition, DFA1 around HJ  $\ge$  1000 U/L, and main pancreatic duct size < 3 mm were significantly associated with the development of CR-POPF after PD. Multivariate logistic regression analyses showed that a high BMI (odds ratio [OR] = 6.094, 95% confidence interval [CI]: 2.021-18.374; P = 0.001), nonapplication of omental interposition (OR = 3.145, 95% CI: 1.040-9.509; P = 0.042), and main pancreatic duct diameter < 3 mm (OR = 5.663, 95% CI: 1.456-22.033; P = 0.012) were independent factors that were significantly associated with the development of CR-POPF after PD.

#### DISCUSSION

To date, POPF remains the most fatal complication after PD. Pancreatic fistula, especially clinically related postoperative fistula, is the most common cause of delayed PPH and intra-abdominal infections after PD[1-4]. Leaked activated pancreatic juice is highly corrosive. Once the drainage tubes fail to effectively work, pancreatic juice accumulates in the potential cavity gap around the anastomosis. This condition may erode the vulnerable anastomosis and adjacent vascular wall. Various efforts[5-8] have been tested for their ability to reduce the incidence of CR-POPF after PD, such as improved anastomosis and the use of somatostatin. However, few randomized control trials have significantly prevented CR-POPF.

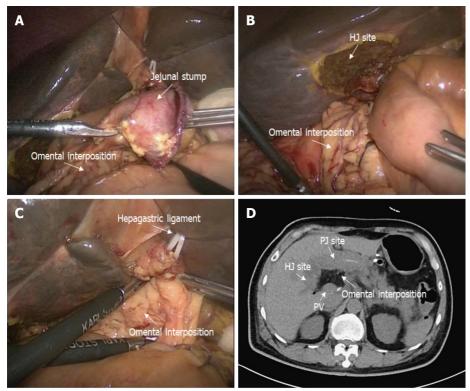


## Table 3 The univariate and multivariate analyses of the propensity score-matched data to evaluate the risk factors associated with clinically relevant postoperative pancreatic fistula after pancreaticoduodenectomy

		No CR-POPF (114)	P value	Multivariate analysis			
	CR-POPF (24)			OR	95%CI	P value	
Age (mean ± SD, yr)	$63.5 \pm 7.9$	$63.1 \pm 10.1$	0.829				
Sex, n (%)			0.040				
Male	20 (83.3%)	70 (61.4%)		2.436	0.692-8.574	0.165	
Female	4 (16.7%)	44 (38.6%)		Reference			
Operation method, <i>n</i> (%)			0.143				
LPD	4 (16.7%)	36 (31.6%)					
OPD	20 (83.3%)	78 (68.4%)					
BMI (kg/m <sup>2</sup> )			0.000				
≥23	18 (75.0%)	33 (28.9%)		6.094	2.021-18.374	0.001	
< 23	6 (25.0%)	81 (71.1%)		Reference			
Serum bilirubin (µmol/L)	96.6 (16.1-180.4)	67 (13.8-111.2)	0.185				
Serum ALB (g/L)			0.843				
≥ 35	21 (87.5%)	98 (86.0%)					
< 35	3 (12.5%)	16 (14.0%)					
ASA score, n (%)			0.122				
Grade I	11 (45.8%)	66 (57.9%)					
Grade II	11 (45.8%)	47 (41.2%)					
Grade III	2 (8.3%)	1 (0.9%)					
Pathology, n (%)			0.196				
Malignancy	23 (95.8%)	96 (84.2%)					
Benign	1 (4.2%)	18 (15.8%)					
Omental interposition, $n$ (%)			0.025				
Yes	7 (29.2%)	62 (54.4%)		Reference			
No	17 (70.8%)	52 (45.6%)		3.145	1.040-9.509	0.042	
Operating time (mean ± SD, min)	387.1±82.5	377.7±71.2	0.609				
HJ DFA1 (U/L)			0.010				
≥ 1000	13 (54.2%)	31 (27.2%)		1.000	1.000-1.000	0.834	
< 1000	11 (45.8%)	83 (72.8%)		Reference			
PJ DFA1 (U/L)			0.115				
≥1000	13 (54.2%)	42 (36.8%)					
< 1000	11 (45.8%)	72 (63.2%)					
Main pancreatic duct size [ <i>n</i> (%), mm]			0.000				
≥3	3 (12.5%)	64 (56.1%)		Reference			
< 3	21 (87.5%)	50 (43.9%)		5.663	1.456-22.033	0.012	

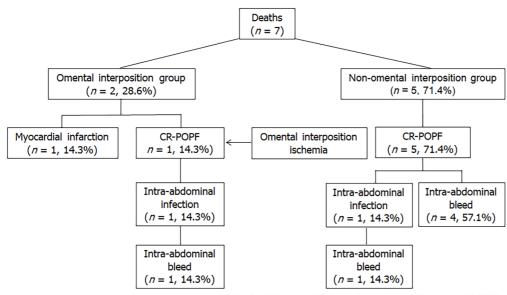
ALB: Albumin; ASA: American Society of Anesthesiologist score; BMI: Body mass index; DFA1: Drain fluid amylase obtained on the first postoperative day; HJ: Hepaticojejunostomy; LPD: Laparoscopic pancreaticoduodenectomy; OPD: Open pancreaticoduodenectomy; PJ: Pancreaticojejunostomy.

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Figure 2 Application of the omental interposition in pancreaticoduodenectomy. A: The pedicled omental interposition was placed in front of the adjacent vessels (hepatic artery, portal vein, and gastroduodenal artery stump) and behind the pancreaticojejunostomy site; B: The right boundary of the omental interposition was the right margin of the inferior vena cava; C: The upper boundary of the omental interposition was the hepatogastric ligament; the omental interposition was fixed to the hepatic portal and hepatogastric ligament with several sutures to prevent postoperative mobilization; D: Postoperative computed tomography images. The omental interposition elevated the hepaticojejunostomy site and filled the potential cavity.

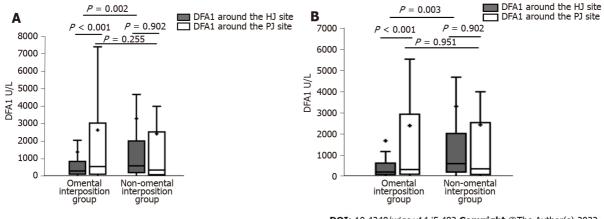


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#### Figure 3 Causes of death in the two groups.

Since pancreatic fistulas are almost inevitable after PD, it is necessary to improve the surgical techniques and accelerate the healing process of fistulas to strive for "harmless" pancreatic fistulas. Experimental results have shown that the greater omentum can resist corrosion, provide anti-infection properties, absorb the peritoneal effusion, regenerate blood vessels and repair tissue defects. Thus, we hypothesized that the omental interposition could seal the posterior wall of the PJ anastomosis, fill the potential cavity to avoid effusion at the surgical site, cover the skeletonized vessels to avoid erosion and





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Figure 4 Differences between the two groups. A: Drain fluid amylase obtained on the first postoperative day (DFA1) before propensity score-matching (PSM); B: DFA1 after PSM.

accelerate the regeneration of blood vessels to improve the blood supply of the anastomosis. The study shows that the incidence of CR-POPF and delayed PPH were lower in the omental interposition group than in the non-omental interposition group. As a result of the reduced complications, the average duration to restart diet and the length of hospital stay were shorter in the omental interposition group. Previous studies on OPD have reached similar conclusions. Maeda[14] covered the major splanchnic arteries and the PV with an omental flap in 100 patients. Although the author concluded that the incidence of POPF (20%) was not significantly different from that in other articles, he did not rule out biochemical fistulas based on the modified definition by the ISGPF. Matsuda *et al*[20] emphasized the preventive effect of omental flaps in PD against postoperative pseudoaneurysm formation. Shah *et al*[21] wrapped the omental flap around the PJ site in 101 patients and showed that it could reduce the incidence of POPF (4.0% *vs* 17.4%), PPH (0% *vs* 6.5%), BF (1.0% *vs* 13.0%), and DGE (4.0% *vs* 17.4%) compared to those in the non-omental interposition group.

In addition to the physiological function of the omental interposition, our method could elevate the height of the anastomosis and fill the potential cavity due to the physical characteristics. Because the omental interposition can elevate the position of the HJ anastomosis (Figure 2D), the erosive pancreatic fluid will flow to the left instead of remaining around the skeletonized vessels in the right upper quadrant of the abdomen. The difference in DFA1 between HJ and PJ sites confirm these physical characteristics in the omental interposition group. This finding also confirms that the application of the omental interposition, by preventing leakage from the anastomosis, reduces the incidence of delayed PPH. Because of the effective control of serious complications, the omental interposition group had their drainage tubes removed earlier, required fewer postoperative transfusions, restarted their diet earlier and had a shorter hospital stay than the non-omental interposition group. These findings are highly consistent with the aforementioned studies showing the efficacy of the omental interposition in PD.

PSM of nine baseline characteristics was performed to reduce selection bias and potential confounding factors between the two groups. After matching, the incidences of CR-POPF and delayed PPH remain significantly lower in the omental interposition group. Similarly, the difference in median DFA1 values between HJ and PJ sites in the omental interposition group remained observable. However, in the non-omental interposition group, the DFA1 around the PJ site was significantly higher than the DFA1 around the HJ site. Due to the physical characteristics of the omental interposition, the corrosive pancreatic juice would flow to the left upper quadrant of the abdomen because of gravity. Obviously, these details matter tremendously.

Previous studies[22-24] have reported that the risk factors for POPF include a high BMI, soft pancreatic texture, and small pancreatic duct size. In our study, univariate and multivariate analyses revealed that a high BMI, nonapplication of omental interposition, and main pancreatic duct diameter < 3 mm were independent factors significantly associated with the development of CR-POPF after PD. The developed statistical model had a c-index of 0.848. These findings were partially consistent with previous POPF risk scores.

Only one patient in the omental group died of delayed PPH caused by ischemic infection due to poor blood supply of the omental interposition, which resulted in delayed hemorrhage. This was the eighth case in which we applied the omental interposition with insufficient emphasis on ensuring good blood supply to the omental interposition. Since then, we detached the gastrocolic ligament along the gastric wall to ensure good blood supply to the omental interposition.

This study had several limitations, including its design as a single-center, retrospective observational study. However, all clinically related data were prospectively collected, and all operations were performed by the same surgical group with the same surgical technology. Thus, the majority of the



potential confounding factors were controlled.

#### CONCLUSION

In conclusion, we believe that the application of the omental interposition is technically simple and may help prevent CR-POPF and the associated complications following PD.

#### ARTICLE HIGHLIGHTS

#### Research background

Postoperative pancreatic fistula (POPF) is a life-threatening complication after pancreaticoduodenectomy (PD).

#### Research motivation

Several methods have attempted to reduce the POPF after PD, few have been considered effective. The safety and short-term clinical benefits of omental interposition remain controversial.

#### Research objectives

To investigate the safety and feasibility of omental interposition to reduce the POPF rate and related complications in PD.

#### Research methods

In total, 196 consecutive patients underwent PD performed by the same surgical team, the patients were divided into two groups: an omental interposition group (127, 64.8%) and a non-omental interposition group (69, 35.2%). Propensity score-matched analyses were performed to compare the severe complication rates and mortality between the two groups.

#### Research results

The clinically relevant POPF (CR-POPF; 10.1% vs 24.6%; P = 0.025) and delayed postpancreatectomy hemorrhage (1.4% vs 11.6%; P = 0.016) rates were significantly lower in the omental interposition group. The omental interposition technique was associated with a shorter time to resume food intake (7 vs 8 d; P = 0.048) and a shorter hospitalization period (16 vs 21 d; P = 0.031).

#### Research conclusions

The application of the omental interposition is an effective and safe approach to reduce the CR-POPF rate and related complications after PD.

#### Research perspectives

Prospective studies are needed on the role of omental interposition in reducing CR-POPF.

#### FOOTNOTES

Author contributions: Li Y, Liang Y, and Deng Y contributed equally to this manuscript; Jiang CY participated in the conception and design of this study; Deng Y, Cai ZW, Ma MJ, Wang LX, Liu M, and Wang HW participated in the data collection; Li Y participated in the data collection, analysis, and drafting of the article; Liang Y participated in the design of the study and data analyses; All authors have read and approved the final manuscript.

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