# Early Removal of Prophylactic Drains Reduces the Risk of Intra-abdominal Infections in Patients With Pancreatic Head Resection

# Prospective Study for 104 Consecutive Patients

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**Objective:** The aim of this study was designed to determine whether the period of drain insertion influences the incidence of postoperative complications.

**Background Data:** The significance of prophylactic drains after pancreatic head resection is still controversial. No report discusses the association of the period of drain insertion and postoperative complications

**Methods:** A total of 104 consecutive patients who underwent pancreatic head resection were enrolled in this study. To assess the value of prophylactic drains, we prospectively assigned the patients into 2 groups: group I underwent resection from January 2000 to January 2002 (n = 52, drain to be removed on postoperative day 8); group II underwent resection from February 2002 to December 2004 (n = 52, drain to be removed on postoperative day 4). Postoperative complications in the 2 groups were compared.

**Results:** The rate of pancreatic fistula was significantly lower in group II (3.6%) than in group I (23%) (P=0.0038). The rate of intraabdominal infections, including intra-abdominal abscess and infected intra-abdominal collections, was significantly reduced in group II (7.7%) compared with group I (38%) (P=0.0003). Eighteen of 52 (34.6%) patients in group I had an inserted drain beyond 8 days, whereas only 2 of 52 (3.7%) patients in group II had an inserted drain beyond 4 days (P=0.0002). Cultures of drainage fluid were positive in 16 of 52 (30.8%) patients in group I, and in 2 of 52 (3.7%) patients in group II (P=0.0002). Intraoperative bleeding (>1500 mL), operative time (>420 minutes), and the period of drain insertion were significant risk factors for intra-abdominal infections (P=0.043, 0.025, 0.0003), respectively). The period of drain insertion was the only independent risk factor for intra-abdominal infections by multivariate analysis (odds ratio, 6.7).

**Conclusion:** Drain removal on postoperative day 4 was shown to be an independent factor in reducing the incidence of complications with pancreatic head resection, including intra-abdominal infections.

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Randomized clinical trials have demonstrated that prophylactic drains have not decreased the incidence of postoperative complications in elective hepatectomy, colectomy, and cholecystectomy. Tet, the significance of prophylactic drains after pancreatic head resection is still unclear. Only 1 randomized controlled trial in pancreatic surgery has suggested that routine drain insertion is unnecessary as drain insertion, compared with no drains, failed to reduce postoperative complications. However, the morbidity rate after pancreatic head resections still remains high in the range of 30% to 65%, although the mortality rate has decreased to less than 5% by recent advances in surgical techniques and perioperative management. Tels

Prophylactic drains after pancreatic head resection allow monitoring of the occurrence of intra-abdominal bleeding, as well as the detection and drainage of the pancreatic fistula. <sup>16,17</sup> Therefore, prophylactic drains usually are inserted after pancreatic head resection, even in high-volume centers of pancreatic surgery, and usually are removed around postoperative day (POD) 7. <sup>9,13,18</sup> It has been considered that surgically placed drains provide a risk of intra-abdominal infections by providing a route for ascending infections. To our knowledge, no reports in the literature have evaluated the association between the period of drain insertion and intra-abdominal infections after pancreatic head resection. We conducted this prospective study to determine whether the period of drain insertion influences the incidence of postoperative complications.

## **METHODS**

#### **Patients**

From January 2000 to December 2004, 104 consecutive patients underwent pancreatic head resection at Wakayama

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Medical University Hospital (WMUH) by 2 pancreatic surgeons (H.Y. with 25 years experience, M.T. with 20 years experience). We prospectively assigned the patients into 2 groups (by periods of time) to assess the value of prophylactic drains. In the period January 2000 to January 2002, the drains were to be removed on POD 8. Then, from February 2002 to December 2004, the drains were to be removed on POD 4 to determine whether the incidence of intra-abdominal infections could be reduced by early removal of drains, thereby inhibiting ascending infections. The protocol and study design of the present trial were approved and conducted according to the guidelines of the Ethical Committee of WMUH.

Preoperative biliary drainage was performed by percutaneous transhepatic catheter drainage (PTCD) or endoscopic nasal biliary drainage (ENBD), when the serum levels of total bilirubin were greater than 5 mg/mL, or dilatation of the intrahepatic bile duct and hepatic dysfunction (transaminase, >100 IU/mL) were detected. Patient characteristics and perioperative and postoperative parameters in the 2 groups were reviewed for the following 21 clinical variables: patient age, gender, preoperative serum level of albumin, total bilirubin and amylase, history of jaundice, history of diabetes mellitus, preoperative biliary drainage, type of resection; pancreaticoduodenectomy (PD) or pylorus-preserving pancreaticoduodenectomy (PpPD), operative time, intraoperative bleeding, red blood cell transfusion, pancreatic texture (soft or hard), presence or absence of dilatation of the main pancreatic duct, histologic diagnosis (malignant or benign), degree of lymph node dissection (D1 or D2), intended period of drain insertion (POD 4 or 8), serum amylase level on POD 1 and 4, and amylase level of drainage fluid on POD 1 and 4.

### **Perioperative Management**

Twenty patients received PD with Child reconstruction, and 84 patients received PpPD with Traverso reconstruction. Pancreatic anastomosis after PD and PpPD was performed by duct-to-mucosal, end-to-side pancreaticojejunostomy in all enrolled patients. External suture rows were performed as a single suture between the remnant pancreatic capsule, parenchyma, and jejunal seromuscular by using an interrupted sutured of 4-0 Novafil (polybutester, Tyco Healthcare Co.). Internal suture rows, duct-to-mucosa, were performed between the pancreatic ductal and jejunal mucosa by using 8 interrupted 5-0 PDS-II (polydioxanone, Johnson and Johnson Co.). A 5-French polyethylene pancreatic duct drainage tube (Sumitomo Bakelite Co.) was used in all patients. A stent placed in the pancreatic duct removed on POD 21, and the timing of stent removal was the same for both groups. No stent was used for the biliary anastomosis. Two intra-abdominal drains routinely were placed near the pancreatic and biliary anastomosis. One 10-mm Penrose drain, a silicon, multitubular flat drain, routinely was placed near the pancreaticojejunostomy, and a 10-mm silicon tube drain (Kaneka Medics Co.) was routinely placed near the cholangiojejunostomy. These drains were connected to a closed drainage system. The collection system was changed to new one daily to prevent contamination of drain system, and it was identical over the 4 years of this study. The drains were to be removed on POD 8 in group I and on POD 4 in group II, when the drainage fluid was clear, and pancreatic fistula and bacterial contamination were absent. Dilatation of the pancreatic duct was defined as a diameter greater than 3 mm and was judged intraoperatively. The amylase of serum and drainage fluid was measured on POD 1 and 4. Drainage fluid on POD 7 in group I and on POD 4 in group II was cultured in all patients.

No patient received radiotherapy preoperatively and postoperatively. All patients received prophylactic antibiotics intraoperatively and for 3 days postoperatively. Prophylactic octreotide was not administered to prevent pancreatic fistula.

### **Postoperative Complications**

Pancreatic fistula was defined as more than 50 mL drainage fluid per day with 3-fold serum amylase level on POD 4. An international study group of pancreatic surgeons (ISGPF) has proposed a consensus definition and clinical grading about postoperative pancreatic fistula, and defined as follows: grade A, called "transient fistula," it has no clinical impact; grade B, required a change in management or adjustment in the clinical pathway; grade C, a major change in clinical management or deviation from the normal clinical pathway. 19 Biliary fistula was defined as the presence of bile in drainage fluid that persisted by POD 4. Intra-abdominal abscess was defined as intra-abdominal fluid collection with positive cultures identified by ultrasonography (US) or computed tomography (CT) associated with persistent fever and elevations of white blood cells. Infected intra-abdominal fluid was defined as drainage fluid having a positive culture with clinical signs, but without detected intra-abdominal abscess. Delayed gastric emptying was defined as output from a nasogastric tube of greater than 500 mL per day that persisted beyond POD 10, or the failure to maintain oral intake by POD 14.

## Statistical Analysis

The study design to predict the number of patients necessary for statistical validity (two-sided) was based on the premise of improving intra-abdominal infection rate from 35% to 10%, with the  $\alpha$  set at 0.05 and the  $\beta$  set at 0.2, yielding a power of 80%. <sup>6,9,17</sup> We calculated that 52 patients were required in each arm of this study, for a total study population of 104 patients. Data were expressed as means  $\pm$  SD. Patient characteristics and perioperative and postoperative factors between 2 groups were compared by using  $\chi^2$  statistics, Fisher exact test, and Mann-Whitney U test. Variables with P < 0.100 were entered into a logistic regression model to determine independent risk factors of postoperative complications. The independent risk factors of the variables were expressed as odds ratios with their 95% confidence intervals. Statistical significance was defined as P < 0.05.

#### RESULTS

# Comparison of Histologic Diagnosis, Patient Characteristics, and Perioperative Status

A total of 104 patients who underwent pancreatic head resection were divided into 2 groups: 52 patients in group I (drain to be removed on POD 8) and 52 in group II (drain to be removed on POD 4).

**TABLE 1.** Characteristics of Enrolled Patients

	Group I (n = 52)	Group II (n = 52)	P
Age (yr)	67 ± 10	66 ± 10	0.6205
Gender (M/F)	31/21	26/26	0.4308
Diabetes (yes/no)	7/45	14/38	0.1428
Preoperative biliary drainage (yes/no)	19/33	15/37	0.5309
Serum albumin level (g/dL)	$3.8 \pm 0.5$	$3.7 \pm 0.5$	0.6124
Serum total bilirubin level (mg/dL)	$2.3 \pm 2.9$	$1.8 \pm 2.1$	0.3427
Serum amylase level (IU/L)	$106 \pm 107$	$148 \pm 157$	0.1087
Benign tumors/malignant tumors	15/37	9/43	0.2446
Pancreatic adenocarcinoma	19	19	
Ampullary adenocarcinoma	5	8	
Distal cholangiocarcinoma	8	9	
Duodenal adenocarcinoma	3	0	
Intraductal papillary adenoma	7	5	
Intraductal papillary carcinoma	2	7	
Pancreatic endocrine tumor	1	1	
Tumor-forming pancreatitis	4	1	
Other disease	3	2	

Group I, drain to be removed on postoperative day 8; group II, drain to be removed on postoperative day 4.

Table 1 shows the results of histologic analysis of the resected specimens, patient characteristics, and preoperative status. There was no significant difference concerning malignant (group I, n=37; group II, n=43) and benign tumors (group I, n=15; group II, n=9) between the 2 groups. There were no significant differences between the 2 groups concerning other background data.

Table 2 shows the results of perioperative status. There were no significant differences between the 2 groups concerning perioperative status except for the type of resection.

# Postoperative Changes in Amylase Level of Serum and Drainage Fluid

Table 3 shows the fluctuation of amylase levels in serum and drainage fluid. The serum amylase level on POD 4 in group I and group II was restored to the normal range.

**TABLE 2.** Comparison of Perioperative Status

	Group I (n = 52)	Group II $(n = 52)$	P
Operative time (min)	383 ± 59	407 ± 76	0.0753
Intraoperative bleeding (mL)	$1287 \pm 1374$	$1270 \pm 1220$	0.9447
Red blood cell transfusion (unit)	$3.0 \pm 3.9$	$3.4 \pm 3.7$	0.6256
Type of resection (PD/PpPD)	17/35	3/49	0.0009
Dilatation of pancreatic duct (yes/no)	27/25	27/25	0.8441
Pancreatic texture (soft/hard)	25/27	26/26	0.8445
Portal vein resection (yes/no)	4/48	5/47	0.7273
Lymph node dissection (D1/D2)	7/45	6/46	0.7668

Group I, drain to be removed on postoperative day 8; group II, drain to be removed on postoperative day 4.

TABLE 3. Amylase Levels in Serum and Drainage Fluid

	Group I (n = 52)	Group II (n = 52)	P
Serum amylase level on POD 1* (IU/L)	$429 \pm 480$	$273 \pm 257$	0.0425
Serum amylase level on POD 4 (IU/L)	$65 \pm 68$	$67 \pm 68$	0.9300
Amylase level of drainage fluid on POD 1 (IU/L)	$1982 \pm 3374$	$2025 \pm 4448$	0.9576
Amylase level of drainage fluid on POD 4 (IU/L)	$603 \pm 160$	$470 \pm 2047$	0.7908
Property of drainage fluid on POD 4 (clear/unclear)	50/2	51/1	0.5580
Property of drainage fluid on POD 7 (clear/unclear)	$44/8^{\dagger}$	$ND^{\ddagger}$	

Group I, drain to be removed on postoperative day 8; group II, drain to be removed on postoperative day 4.

The mean amylase level of drainage fluid on POD 1 was higher than on POD 4 in group I and group II (1982  $\pm$  3374 IU/L vs. 603  $\pm$  1604 IU/L; 2025  $\pm$  4448 IU/L vs. 700  $\pm$  2047 IU/L) (P = 0.0267 and .0342, respectively).

The drainage fluid on POD 4 was clear in 50 of 52 (96%) of group I and 51 of 52 (98%) of group II patients (P = 0.5580). However, the drainage fluid on POD 7 had become unclear, compared with drainage fluid on POD 4 in group I (15% vs. 4%, P = 0.0128).

# Postoperative Changes in Leukocyte Counts and C-Reactive Protein Level

Postoperative changes in leukocyte counts and C-reactive protein (CRP) level were analyzed in both groups. Leukocyte counts on POD 7 in group I were significantly higher than those in group II (P=0.011), whereas there was no difference in either group on POD 1 and 4. Leukocyte counts in group I were lower on POD 4 than on POD 1 ( $7944\pm2729/\text{mm}^3$  vs.  $10,596\pm3024/\text{mm}^3$ , P=0.0001); however, the counts in group I increased again on POD 7 ( $9325\pm3419/\text{mm}^3$ ) (P=0.013), whereas in group II leukocyte counts on POD 7 were similar to those on POD 4. On the other hand, there was no significant difference in CRP levels on POD 1, 4, and 7 between the 2 groups.

# Comparison of Postoperative Status and Complications

Postoperative complications were compared with clarify whether the period of drain insertion influenced postoperative complication rates (Table 4). The overall rate of pancreatic fistula after pancreatic head resection was 13% (14 of 104 patients); the rate of pancreatic fistula was significantly lower in group II (3.6%) than group I (23%) (P = 0.0038). There were no patients in either group with major leakage of pancreaticojejunostomy, which was defined as leakage of pancreatic juice with contamination of bile or intestinal contents. Pancreatic fistula was classified into 3

 $<sup>\</sup>label{pppp} PD\ indicates\ pancreaticoduodenectomy;\ PpPD,\ pylorus-preserving\ pancreaticoduodenectomy.$ 

<sup>\*</sup>Normal range: 15-150 IU/L.

<sup>&</sup>lt;sup>†</sup>*P* = 0.0128, compared with property of drainage fluid on POD 4 in group I. <sup>‡</sup>ND indicates not determined. Only 2 of 52 patients in group II had drain insertion on POD 7, and they had unclear drainage fluid.

**TABLE 4.** Comparison of Postoperative Complications

	Group I (n = 52)	Group II (n = 52)	P
Pancreatic fistula	12	2	0.0038
Grade A*	3	1	
Grade B	7	1	
Grade C	2	0	
Intra-abdominal infections	20	4	0.0003
Intra-abdominal abscess	10	3	0.0721
Infected intra-abdominal collection	10	1	0.0079
Intra-abdominal hemorrhage	2	0	0.4951
Gastrointestinal bleeding	0	1	0.3150
Biliary leakage	3	2	0.6467
Delayed gastric emptying	17	12	0.3818
Wound infection	0	1	0.3150
Portal vein thrombosis	1	0	0.3150
Cholangitis	2	1	0.5580
Mortality	1	0	0.3150

Group I, drain to be removed on postoperative day 8; group II, drain to be removed on postoperative day 4.

categories by ISGPF, and 12 patients of pancreatic fistula in group I were classified into grade A (n = 3), grade B (n = 7), grade C (n = 2), and 2 patients in group II into grade A (n = 1), grade B (n = 1) (Table 4). There were no significant differences between the 2 groups with regard to hemorrhage and DGE. Most importantly, the rate of intra-abdominal infections, including infected intra-abdominal collections and intra-abdominal abscess, was significantly lower in group II than group I (7.7% vs. 38%, P = 0.0003). The rate of infected intra-abdominal collections was significantly reduced in group II, compared with group I (1.9% vs. 19%, P = 0.0079).

## Cause of Long-term Drain Insertion

Eighteen of 52 (34.6%) patients in group I had inserted drains beyond 8 days, whereas only 2 of 52 (3.7%) patients in group II had inserted drains beyond 4 days (P=0.0002). Table 5 shows the details of all patients with inserted drains beyond 8 days in group I and 4 days in group II. The period of drain insertion in 18 patients was  $17 \pm 5$  days in group I (range, 12-32 days) because of 8 pancreatic fistula, 3 biliary leakage, and 7 intra-abdominal infections (6 infected intra-abdominal collections, 1 intra-abdominal abscess). Positive culture of drainage fluid was 16 of 18 (94%) cases. The period of drain insertion in 2 patients was  $15 \pm 1$  day in group II (range, 14-16 days) because of 1 pancreatic fistula and 1 biliary leakage. Cultures of drainage fluid were positive

**TABLE 5.** Cause of Long-term Drain Insertion

Case No.	Removal of Drain (POD)	Highest Amylase Level in Drainage Fluid (IU/L)	Postoperative Complications	Drainage Fluid Culture*
Group I (18/52 patients = $34.6\%^{\dagger}$ )				
1	13	860	Infected intra-abdominal collection	Streptococcus spp
2	14	286	Biliary leakage	E. faecalis
3	20	635	Intra-abdominal abscess	E. faecalis
4	21	654	Pancreatic fistula	E. cloacae
5	18	65	Infected intra-abdominal collection	Streptococcus spp
6	14	12,542	Pancreatic fistula	K. pneumoniae
7	18	43	Infected intra-abdominal collection	E. faecalis
8	14	240	Infected intra-abdominal collection	Staphylococcus spp
9	9	610	Biliary leakage	P. aeruginosa
10	32	1189	Pancreatic fistula	P. aeruginosa
11	16	65	Infected intra-abdominal collection	E. faecalis
12	21	86,769	Pancreatic fistula	E. faecalis
13	14	232	Infected intra-abdominal collection	E. faecalis
14	19	34,990	Pancreatic fistula	P. aeruginosa
15	15	21,953	Pancreatic fistula	E. faecalis
16	14	8,204	Pancreatic fistula	Negative
17	14	12,346	Biliary leakage	Negative
18	16	15,168	Pancreatic fistula	E. faecalis
Group II $(2/52 \text{ patients} = 3.7\%)$				
1	33	7977	Biliary leakage	E. faecalis
2	21	5863	Pancreatic fistula	E. cloacae

<sup>\*</sup>Drainage fluid was cultured on POD 7 in group I and on POD 4 in group II. Cultures were positive in 16 of 52 (30.8%) patients in group I and 2 of 52 (3.7%) in group II (P = .0002).

<sup>\*</sup>Pancreatic fistula was classified into 3 categories by an international study group of pancreatic surgeons (ISGPF) as follow: grade A, called "transient fistula," it has no clinical impact; grade B, required a change in management or adjustment in the clinical pathway; grade C, a major change in clinical management or deviation from the normal clinical pathway.

<sup>†</sup>Eighteen of 52 (34.6%) patients in group I had inserted drains beyond 8 days, whereas only 2 of 52 (3.7%) patients in group II had drains beyond 4 days (P = 0.0002). POD indicates postoperative day.

**TABLE 6.** Univariate Analysis of Risk Factors Influencing Intra-abdominal Infection

	Intra-abdominal Infection		
	(+) (n = 24)	(-) (n = 80)	P
Age (yr)			0.8279
≤75	18	56	
>75	6	24	
Gender			0.5290
Male	15	42	
Female	9	38	
Diabetes			0.7757
Yes	4	17	
No	20	63	
Jaundice			0.8840
Yes	14	48	
No	10	32	
Preoperative biliary drainage			0.9569
Yes	10	36	
No	14	44	
Serum albumin level (g/dL)			0.6751
≤3.8	8	21	
>3.8	16	59	
Serum total bilirubin level			0.6542
(mg/dL)			*****
≤1.5	11	43	
>1.5	13	37	
Serum amylase level (IU/L)			0.4174
≤150	17	65	
>150	7	15	
Histology			0.6506
Benign	4	19	
Malignant	20	61	
Operative time (min)			0.0253
≤420	11	58	
>420	13	22	
Intraoperative bleeding (mL)			0.0431
≤1500	15	67	
>1500	9	13	
Red blood cell transfusion (unit)			0.4839
Yes	13	50	
No	11	30	
Type of resection			0.9457
PD	4	16	
PpPD	20	64	
Dilatation of pancreatic duct			0.2480
Yes	10	45	
No	14	35	
Pancreatic texture	•		0.6446
Soft	13	38	
Hard	11	42	
Portal vein resection		.2	0.9492
Yes	2	7	0.7772
No	22	73	
1.0		13	

**TABLE 6.** (Continued)

	Intra-abdominal Infection		
	(+) (n = 24)	(-) (n = 80)	P
Lymph node dissection			0.7598
D1	3	12	
D2	21	68	
Intended period of drain insertion			0.0003
POD 4	4	48	
POD 8	20	32	
Serum amylase level on POD 1 (IU/L)			0.4428
≤150	5	25	
>150	19	55	
Serum amylase level on POD 4 (IU/L)			0.8665
≤150	21	71	
>150	3	9	

in both patients (100%). Cultures of drainage fluid were positive in 16 of 52 patients (30.8%) in group I, and in 2 of 52 patients (3.7%) in group II (P = 0.0002).

### **Risk Factors of Intra-abdominal Infections**

Univariate and multivariate analysis were used to reveal risk factors for intra-abdominal infections. Table 6 shows the results of 20 parameters univariately examined as potential risk factors for the 24 patients with intra-abdominal infections versus the 80 patients without intra-abdominal infections. Two intraoperative factors (intraoperative bleeding, >1500 mL, P=0.043; operative time, >420 minutes, P=0.025) and 1 postoperative factor (intended period of drain insertion: 4 days vs. 8 days, P=0.0003) differed significantly between these 2 groups. No significant preoperative factors, including preoperative biliary drainage, were associated with the incidences of postoperative intra-abdominal infections.

A multivariate logistic regression analysis in intraabdominal infections revealed that the intended period of drain insertion was the only independent risk factor (P = 0.0024; Table 7).

### **DISCUSSION**

The prophylactic value of drains after pancreatic head resection is still controversial. Recent reports have suggested that drain insertion after pancreatic resections failed to reduce postoperative complications, <sup>6,20</sup> whereas drains usually are

**TABLE 7.** Multivariate Analysis of Risk Factors Influencing Intra-abdominal Infection

Risk Factor	P	<b>Odds Ratio</b>	95% CI
Intended period of drain insertion*	0.0024	6.7	1.9–22.7

<sup>\*</sup>Intended period of drain insertion: 4 days versus 8 days. CI indicates confidence interval.

inserted after pancreatic head resection in most institutes. Moreover, drains placed surgically after abdominal surgery have been reported to result in increased rates of intra-abdominal infection. <sup>6,20,21</sup> However, drains after pancreatic resections are necessary for early detection of pancreatic fistula, which can lead to hemorrhage and abdominal abscess, and is associated with a high mortality rate. 21-25 No previous report has discussed the relationship between the period of drain insertion and postoperative complications. The present prospective study has been designed to clarify whether the intended period of drain insertion influenced postoperative complication rates after pancreatic head resection. Our results showed that postoperative complication rates concerning the incidence of pancreatic fistula and intra-abdominal infections were significantly lower when the prophylactic drains were to be removed on POD 4. With regard to complications after pancreas head resection, it has been suggested that high-volume centers have lower rates of morbidity and mortality. 26-28 In the period of the present study, from 2000 to 2004, more than 20 pancreatic head resections per year have been performed at WMUH, and we have performed more than 15 distal pancreatectomies per year. Thus, we did not need to take into account the hospital volume in this analysis. However, this study was not truly a randomized study, and we could not perfectly deny the possibility that these improved results were not due in part to subtle improvements in operative technique honed over the first 2 years of this study.

In this study, positive cultures of drainage fluid on POD 7 increased to 31% in group I (drains to be removed on POD 8), although there was no difference in the properties of drainage fluid on POD 4 between the 2 groups. Three of 16 bacteria (19%) detected by culture of drainage fluid in group I were indigenous bacteria of skin (2 Streptococcus spp and 1 Staphylococcus spp). It is suggested that long-term insertion of drains might be a major cause of postoperative infectious complications. Leukocyte counts on POD 7 were thought to be significantly higher than those on POD 4 in group I, which suggested the ascending infection occurred around POD 7. Indeed, despite the absence of intra-abdominal abscess, pancreatic fistula, and biliary leakage, 6 of 16 patients in group I with positive cultures of drainage fluid on POD 7 had clinical signs such as elevated leukocyte counts and fever of greater than 38.0°C. Therefore, one should consider that the onset of infection by drainage routes would appear around POD 7, and we strongly propose that drain removal at POD 4 could reduce the rates of intra-abdominal infections. However, drain removal around POD4 such as day 3 or 5 may have provided equally good results.

Pancreatic fistula, with an incidence varying between 5% to 20% in most series, has been reported as the most serious complication of morbidity after pancreatic head resection. <sup>7,8,29,30</sup> In the previous reports, many preoperative and perioperative risk factors of pancreatic fistula were evaluated, including age older than 65 years, preoperative jaundice, soft pancreatic parenchyma, small pancreatic duct, exocrine pancreatic function, longer operative time, high-volume transfusion, and intraoperative bleeding. <sup>21,29–31</sup> In this study, the incidence of pancreatic fistula was significantly higher in patients in whom

drains were removed on POD 8, compared with patients whose drains were removed on POD 4. However, pancreatic texture and size of the pancreatic duct, which generally are reported to be risk factors of pancreatic fistula, had no impact on the occurrence of postoperative complications. Because pancreaticojejunostomy has been performed by duct-to-mucosal anastomosis since 1999 in WMUH, the occurrence of postoperative pancreatic fistula is similar among patients, even in patients with a soft pancreas and a normal sized pancreatic duct.<sup>7</sup>

Ascending infections by the drain route secondarily may increase pancreatic fistula in patients with long-term drain insertion. It has been suggested that pancreatic juice should be activated by bacterial phospholipase and lipopoly-saccharide. 32–34

### CONCLUSION

Prophylactic drains placed after pancreatic head resection should be better removed on POD 4, compared with POD 8 to prevent ascending infections, and postoperative peripancreatic fluid or intra-abdominal abscess should be treated with CT- or US-guided percutaneous drainage. We are now considering to proceed the randomized controlled trial, which would have made these data much more convincing.

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