

## **Supplementary Online Content**

Smits FJ, van Santvoort HC, Besselink MG, et al; Dutch Pancreatic Cancer Group. Management of severe pancreatic fistula after pancreateoduodenectomy. *JAMA Surg.* Published online February 22, 2017. doi:10.1001/jamasurg.2016.5708

**eTable 1.** Crude Baseline Characteristics

**eTable 2.** Status Based on APACHE II Category

This supplementary material has been provided by the authors to give readers additional information about their work.

**eTable 1. Crude Baseline Characteristics\***

Characteristics	Catheter Drainage (n=227)	Relaparotomy (n=82)	P Value
Age – yr (median (IQR))	65 (59-72)	66 (57-71)	0.65
Male sex – no. (%)	147 (65)	62 (76)	0.07
Coexisting condition – no. (%)			
Cardiovascular disease	60 (26)	32 (39)	0.03
Pulmonary disease	14 (6)	9 (11)	0.16
Chronic renal insufficiency	4 (2)	2 (2)	0.66*
History of upper abdominal surgery	51 (23)	19 (23)	0.90
ASA class on admission (%)			0.006
I: healthy status	50 (22)	14 (17)	
II: mild systemic disease	150 (66)	46 (56)	
III: severe systemic disease	27 (12)	22 (27)	
Body-mass index (mean ±SD) †	26.4±3.6	26.7±4.5	0.47
Weight loss - no. (%)	113/222 (51)	42/77 (56)	0.58
Quantity – kg (median (IQR)) ‡	2 (0-6)	4 (0-8)	0.42
Preoperative ERCP §			0.16
Without intervention	43 (19)	8 (10)	
With stenting/papillotomy	122 (54)	45 (55)	
Preoperative cholestasis – no. (%)	165 (73)	56 (68)	0.45
Operating characteristics			
Pylorus preserving pancreatoduodenectomy	183 (81)	64 (78)	0.62
Reconstruction portal vein	7/220 (3)	3/81 (4)	0.73*
Additional organ resection	10/220 (5)	10/81 (12)	0.01
Abdominal drain	225 (99)	79 (96)	0.12*
Pathology malignant – no. (%)	210 (93)	71 (87)	0.11
Disease severity 24h before first intervention			
APACHE II (median (IQR)) #	8 (6-10)	10 (7-14)	<0.001
SIRS – no. (%)**	99 (44)	47 (58)	0.03
Organ failure – no. (%)	27 (12)	39 (48)	<0.001
Single-organ failure – no. (%)	8 (18)	26 (32)	<0.001
Multiple-organ failure – no. (%)	9/226 (4)	13 (16)	<0.001
Previous re-intervention – no. (%)	10 (4)	12 (15)	0.002

\* Normal distributed values are presented as mean ± SD, ASA denotes American Society of Anaesthesiologists; P-values denoted with \* are obtained using the Fishers exact Test

† Body-mass index, weight in kilograms divided by the square of the height in meters; out of 212 patients in drainage group and 80 patients in relaparotomy group

‡ Crude data calculated over 220 patients in drainage group and 74 patients in relaparotomy group

§ Crude data calculated over 226 patients in drainage group and 82 patients in relaparotomy group

|| Defined as jaundice, elevated bilirubin and/or need for preoperative biliary drainage

¶ Crude data calculated over 220 patients in drainage group and 78 patients in relaparotomy group

# Acute Physiology and Chronic Health Evaluation (APACHE II) scale from 0-71, higher scores indicating more severe disease

\*\* Systemic inflammatory response syndrome as defined by the American College of Chest Physicians and the Society of Critical Care Medicine

eTable 2. Status Based on APACHE II Category		APACHE II <9				APACHE II 9-12				APACHE II >12			
Outcome		Catheter Drainage (n=136)	Relaparotomy (n=33)	RR (95%CI)	P Value	Catheter drainage (n=74)	Relaparotomy (n=19)	RR (95%CI)	P Value	Catheter Drainage (n=17)	Relaparotomy (n=28)	RR (95%CI)	P Value
Death – no. (%)		14 (10)	8 (24)	0.43 (0.19-0.93)	0.044*	7 (10)	6 (32)	0.30 (0.11-0.79)	0.023*	4 (24)	16 (57)	0.41 (0.16-1.03)	0.035*
Secondary endpoints													
Major complications after first intervention for pancreatic fistula – no. (%)													
New-onset single-organ failure		27 (20)	16 (49)	0.41 (0.25-0.67)	0.001	17 (23)	6 (31)	0.73 (0.33-1.59)	0.552*	2 (12)	10 (36)	0.33 (0.08-1.33)	0.096*
New-onset multiple-organ failure		11 (8)	11 (33)	0.24 (0.12-0.51)	0.001*	12 (16)	9 (47)	0.34 (0.17-0.70)	0.011*	2 (12)	12 (43)	0.28 (0.07-1.08)	0.046*
Post pancreatectomy haemorrhage †		18 (13)	6 (18)	0.73 (0.31-1.69)	0.578*	19 (26)	2 (11)	2.44 (0.62-9.57)	0.224*	3 (18)	9 (32)	0.55 (0.17-1.75)	0.488*
Gastroenterostomy leakage †		2 (2)	2 (6)	0.24 (0.04-1.66)	0.172*	2 (3)	0 (0)	1.35 (0.06-27.03)	>0.99*	1 (6)	0 (0)	4.83 (0.20-112.35)	0.378*
Bile leakage †		10 (7)	3 (9)	0.81 (0.24-2.78)	0.720*	6 (8)	1 (5)	1.54 (0.20-12.04)	>0.99*	1 (6)	5 (18)	0.33 (0.04-2.59)	0.385*
Long term complications – no. (%)													
New-onset diabetes ‡		18/122 (15)	13/31 (42)	0.35 (0.19-0.64)	0.001	7/69 (10)	7/19 (37)	0.28 (0.11-0.69)	0.010*	3/16 (19)	7/19 (37)	0.51 (0.16-1.65)	0.285*
New-onset exocrine pancreatic insufficiency ‡		48/118 (41)	16/28 (57)	0.71 (0.48-1.05)	0.114	31/65 (47)	7/18 (39)	1.23 (0.65-2.31)	0.507	6/15 (40)	11/16 (69)	0.58 (0.29-1.17)	0.108
Interventions													
Completion pancreatectomy – no. (%)		4 (3)	10 (30)	0.10 (0.03-0.29)	<0.001*	2 (3)	5 (26)	0.10 (0.02-0.49)	0.004*	1 (6)	6 (21)	0.28 (0.04-2.09)	0.227*
No. of additional relaparotomies					<0.001				0.923				0.849
No. of patients (%)		22 (16)	16 (48)			12 (16)	3 (16)			6 (35)	11 (39)		
Median (range) per patient		0 (0-2)	0 (0-8)			0 (0-4)	0 (0-3)			0 (0-5)	0 (0-5)		
No. of additional catheter drainages				0.391					0.061				0.108
No. of patients (%)		55 (40)	16 (48)			40 (55)	7 (37)			11 (65)	11 (39)		
Median (range) per patient		0 (0-11)	0 (0-5)			1 (0-9)	0 (0-2)			1 (0-7)	0 (0-5)		
No. of interventions for pancreatic fistula				0.048					0.533				0.137
Median (range) per patient		1 (0-8)	2 (1-5)			2 (0-9)	1 (1-4)			3 (0-7)	1 (1-7)		
No. of interventions during admission				0.001					0.345				0.803
Median (range) per patient		1 (0-12)	3 (1-13)			2 (0-13)	2 (1-5)			3 (0-9)	3 (1-7)		
Hospitalization course													
New ICU admission after first intervention – no. (%)		32 (24)	26 (79)	0.29 (0.21-0.42)	<0.001	24 (33)	19 (100)	0.32 (0.23-0.45)	<0.001	11 (65)	27 (96)	0.67 (0.47-0.96)	0.008*
Length of ICU stay (median (IQR)) §,		0 (0-0)	5 (0-10)		<0.001	0 (0-2)	8 (4-12)		<0.001	2 (0-17)	9 (4-29)		0.110
Length of hospital stay (median (IQR))		26 (19-38)	54 (40-61)		<0.001	36 (25-63)	47 (31-68)			0.171	47 (30-73)	61 (51-86)	0.137
Duration of pancreatic fistula (median (IQR))   ,¶		28 (16-48)	25 (15-48)		0.750	33 (21-66)	39 (12-87)			0.989	35 (22-65)	43 (29-73)	0.381