

Serious Postoperative Complications Affect Early Recurrence After Cytoreductive Surgery and HIPEC for Colorectal Peritoneal Carcinomatosis

GA Simkens, TR van Oudheusden, MD Luyer, SW Nienhuijs, GA Nieuwenhuijzen, HJ Rutten, IH de Hingh

INTRODUCTION

The prognosis of patients with peritoneally metastasized colorectal cancer has improved significantly with the introduction of cytoreductive surgery followed by hyperthermic intraperitoneal chemotherapy (CRS+HIPEC). Although a macroscopically complete resection is achieved in nearly every patient, recurrence rates are high. This study aims to identify risk factors for early recurrence, thereby offering ways to reduce its occurrence.

METHODS

All patients with colorectal peritoneal carcinomatosis treated with CRS+HIPEC and a minimum follow-up of 12 months in April 2014 were analyzed. Patient data were compared between patients with or without recurrence within 12 months after CRS+HIPEC. Risk factors were determined using logistic regression analysis. Postoperative complications were graded according to the Serious Adverse Events (SAE) score, with grade 3 or higher indicating complications requiring intervention.

RESULTS

A complete macroscopic cytoreduction was achieved in 96% of all patients treated with CRS+HIPEC. Forty-six of 133 patients (35%) developed recurrence within 12 months. SAE≥3 after CRS+HIPEC was the only significant risk factor found for early recurrence (OR=2.3; p=0.046). Median survival in the early recurrence group was 19.3 months, compared to 43.2 months in the group without early recurrence (p<0.001). Patients with SAE≥3 showed a reduced survival compared to patients without such complications (22.1 vs. 31.0 months, respectively, p=0.02).

TABLE 1. Baseline Characteristics

Variable	Early Recurrence (%)	No Early Recurrence (%)	P value	
Gender male	19 (41.3)	38 (43.7)	0.79	
Age at HIPEC (years)	63.0 (24.3-77.9)	61.1 (37.7-77.1)	0.20	
Metachronous presentation	22 (47.8)	31 (35.6)	0.17	
Primary Location				
Right	21 (45.7)	26 (29.9)	0.10	
Sigmoïd	12 (26.1)	34 (39.1)	0.10	
Rectum	8 (17.4)	13 (14.9)		
Tumor differentiation				
Good	6 (13.0)	6 (6.9)		
Moderate	20 (43.5)	46 (52.9)	0.12	
Poor	5 (10.9)	19 (21.8)		
Signet cell	5 (10.9)	3 (3.4)		
T4	22 (47.8)	44 (50.6)	0.93	
N2	19 (41.3)	35 (40.2)	0.75	
Neo-adjuvant chemotherapy	7 (15.2)	14 (16.1)	0.90	
Adjuvant chemotherapy	38 (82.6)	66 (75.9)	0.37	
Type of recurrence				
Locoregional	17 (37.0)	15 (41.7)	0.58	
Systemic	15 (32.6)	8 (22.2)	0.58	
Both	14 (30.4)	13 (36.1)		

FIGURE 1. Kaplan-Meier Early Recurrence vs. No Early Recurrence

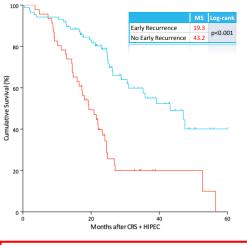


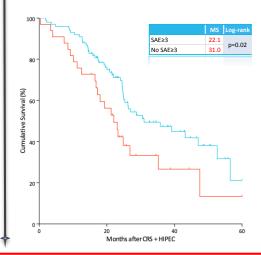
TABLE 2. Operative Outcomes

Variable	Early Recurrence (%)	No Early Recurrence (%)	P value
PCI-score	9.8 ± 5.5	8.0 ± 4.7	0.051
R-score			
1	43 (93.5)	85 (97.7)	0.34
2a	3 (6.5)	2 (2.3)	
Procedure time (minutes)	386 ± 71	367 ± 76	0.15
Blood loss (ml)	1000 (250-4000)	700 (100-6600)	0.09
SAE-grade ≥3	17 (37.0)	16 (18.4)	0.02
Intra-abdominal complication	13 (28.3)	11 (12.6)	0.03
Re-operation	8 (17.4)	11 (12.6)	0.46
Hospital stay (days)	13 (7-84)	10 (4-79)	0.003

TABLE 3. Risk Factors for Early Recurrence

Variable	Early Recurrence	No Early	P value	Odds Ratio	95% C.I.
		Recurrence			
SAE≥3	17 (37.0)	16 (18.4)	0.046	2.3	1.02-5.30
PCI-score	9.8 ± 5.5	8.0 ± 4.7	0.13	1.1	0.98-1.14

FIGURE 2. Kaplan-Meier SAE≥3 vs. No SAE≥3



CONCLUSION

Early recurrence after CRS+HIPEC is associated with a significant reduction in overall survival. This study identifies post-operative complications requiring intervention as the only significant risk factor for early recurrence, independent of the extent of peritoneal disease, highlighting the importance of minimizing the risk of post-operative complications.