# JOURNAL OF CLINICAL ONCOLOGY

# Peritoneal Colorectal Carcinomatosis Treated With Surgery and Perioperative Intraperitoneal Chemotherapy: Retrospective Analysis of 523 Patients From a Multicentric French Study

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A B S T R A C T

#### Purpose

Peritoneal carcinomatosis (PC) from colorectal cancer traditionally is considered a terminal condition. Approaches that combine cytoreductive surgery (CRS) and perioperative intraperitoneal chemotherapy (PIC) have been developed recently. The purpose of this study was to assess early and long-term survival in patients treated with that strategy.

#### **Patients and Methods**

A retrospective-cohort, multicentric study from French-speaking countries was performed. All consecutive patients with PC from colorectal cancer who were treated with CRS and PIC (with or without hyperthermia) were included. Patients with PC of appendiceal origin were excluded.

#### Results

The study included 523 patients from 23 centers in four French-speaking countries who underwent operation between 1990 and 2007. The median follow-up was 45 months. Mortality and grades 3 to 4 morbidity at 30 days were 3% and 31%, respectively. Overall median survival was 30.1 months. Five-year overall survival was 27%, and five-year disease-free survival was 10%. Complete CRS was performed in 84% of the patients, and median survival was 33 months. Positive independent prognostic factors identified in the multivariate analysis were complete CRS, PC that was limited in extent, no invaded lymph nodes, and the use of adjuvant chemotherapy. Neither the grade of disease nor the presence of liver metastases had a significant prognostic impact.

## Conclusion

This combined treatment approach against PC achieved low postoperative morbidity and mortality, and it provided good long-term survival in patients with peritoneal scores lower than 20. These results should improve in the future, because the different teams involved will gain experience. This approach, when feasible, is now considered the gold standard in the French guidelines.

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## INTRODUCTION

Peritoneal dissemination or carcinomatosis from colorectal carcinoma is a form of disease progression that affects 30% to 40% of patients.<sup>1,2</sup> Natural history studies show that peritoneal carcinomatosis (PC) is uniformly fatal, with median survival (not exceeding) attaining 6 months.<sup>3</sup> For more than a decade, a handful of centers have pursued aggressive cytoreductive surgery (CRS) combined with intraperitoneal chemotherapy (PIC), initially without and then with hyperthermia, as an

alternative approach to overcome this disease. The aim of surgery is to resect visible disease as much as possible, and the aim of PIC is to treat any residual occult disease. Results from a single center cannot be extrapolated to other centers because of the vagueness of selection criteria, of what CRS represents, and of PIC techniques. This has led to a reasonable degree of skepticism among some oncologists. This is why any multicentric study that collects a large number of patients from many centers is particularly useful. The experience and the techniques are considerably different from one center to another.

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In this paper, we present a collaborative effort of 23 Frenchspeaking centers to evaluate the efficiency of this combined approach and to identify the main prognostic factors on the basis of a population of 523 treated patients.

## **PATIENTS AND METHODS**

### **Patient Population**

This multicentric, retrospective study was conducted in French (n = 20), Belgian (n = 2), Canadian (Quebec, n = 2) and Swiss (n = 1) centers. Patients were treated between January 1990 and December 2007. Inclusion criteria were histologically confirmed PC of colorectal origin treated with CRS plus hyperthermic intraperitoneal chemotherapy (HIPEC) or early postoperative intraperitoneal chemotherapy (EPIC). Exclusion criteria were appendiceal malignancies and extra-abdominal metastases.

#### Data Forms

A standard data form was created to retrieve information on the primary colorectal tumor, on the status of the patient before the combined procedure, and on previous treatment with systemic chemotherapy. The extent of PC was assessed through intraoperative exploration by using the Peritoneal Cancer Index (PCI),<sup>4-5</sup> which has scores from 0 to 3 for each of the 13 defined areas of the abdominal cavity; therefore, the index ranges from 1 to 39. Four PCI subgroups were defined for the analysis: one to 6, 7 to 12, 13 to 19, and greater than 19. Information recorded about the combined procedure included the completeness of CRS, any simultaneous resection of the primary tumor and liver metastases, the presence or absence of lymph node metastases, the type of perioperative intraperitoneal chemotherapy (ie, HIPEC or EPIC) and its modalities, and treatment with adjuvant systemic chemotherapy. Assessment of the completeness of the cancer resection (CCR) with CRS was done by the surgeon at the end of the procedure and was classified into three categories<sup>4</sup>: CCR-0 signified no macroscopic residual cancer; CCR-1, no residual nodules greater than 2.5 mm; and CCR-2, that the diameter of the residual nodules exceeded 2.5 mm. The centers were classified arbitrarily as experienced (>7 years of practice) and as inexperienced (< 7 years of practice). Major complications (ie, grades 3 and 4 complications according to the National Cancer Institute Common Toxicity Criteria) were considered, and the reasons for reoperation were detailed.

#### Statistical Analysis

Categoric variables were described in terms of frequency and percentages. The distributions of continuous variables were described with mean, standard error, median, and first and third quartiles. Influence of patient, disease, and treatment characteristics were related to the risk of postoperative morbidity and mortality events by using univariate and multivariate logistic regression models, which were adjusted by centers.

The analysis of long-term mortality censored information after the cutoff date of December 31, 2006, because an active inquiry was performed in all centers to collect the status (ie, dead or alive) of the patients at that date. The analysis of total mortality considered the delay from the first procedure to the date of death, the date of last news, or the cutoff date, whichever came first. The analysis of recurrence or death was based on the delay from the first procedure to the date of first recurrence, the date of death, the date of last news, or the cutoff date, whichever came first. Postoperative deaths were not excluded from the survival analysis. Patients who underwent CCR-2 resections with residual tumor nodules exceeding 2.5 mm were considered to have experienced immediate relapses. When the date of recurrence was unknown in patients who died, the date of death was used instead. For this analysis, 12 patients were lost to follow-up. Kaplan-Meier survival estimates were calculated and were compared between strata with the log-rank test. Influence of baseline risk factors on the hazard of death was assessed by using a multiple proportional hazard regression model stratified by centers. Stratification was justified by a large heterogeneity of hazards between centers and by a strong potential confounding effect on other risk factors. In all multivariate analyses, age and PCI were entered as continuous variables (after their linear relationship with the risk of event was checked). CCR also was entered as continuous when justified by a linear trend across categories. An indicator variable for the HIPEC procedure (ie, yes or no) was forced in the model when an associated covariate (ie, temperature, heating duration, open/close abdomen) was significant. Risk factors with a significance degree  $\leq .10$  were retained in the final model. The SAS statistical software (version 9.1; SAS Institute, Cary, NC) was used for all analyses.

## RESULTS

Five hundred twenty-three patients who had undergone 543 procedures that combined CRS and PIC for treatment of colorectal PC between May 1990 and December 2007 constituted the study population. The 2008 annual report of the Association Française de Chirurgie was devoted to this particular approach.

Twenty-five institutions were entered into a central database. One institution recorded 152 patients, one recorded 87 patients, five recorded between 20 and 50 patients, and the others recorded fewer than 20 patients.

## **Patient Characteristics**

Patient characteristics are listed in Table 1. There were 296 female patients (56.6%) and 227 male patients (43.4%), and the median age was 54 years (mean and standard deviation,  $53 \pm 12$  years; range, 16 to 88 years). The PC arose from adenocarcinomas of the right colon (31%), transverse colon (4%), left colon (14%), sigmoid (28%), rectum (7%), and multiple locations (2%). The location of the primary was unknown in 73 patients (14%). Synchronous liver metastases were present (and were resected) in 77 patients (15%). Two hundred seventy-seven patients were treated in an experienced center, and 245 were treated in a inexperienced center. Three hundred seventy patients (70%) had been treated previously with systemic chemotherapy, which did not include new, targeted molecules.

#### Treatment

At completion of the best surgical effort at CRS, 439 patients had a CCR-0 resection (84%), 53 patients a CCR-1 resection, and 22 patients had a CCR-2 resection. Seventy-seven patients (17%) had undergone simultaneous resection of liver metastases. CRS was synchronous with the resection of the primary in 161 patients (35%). Four hundred forty-three patients (86%) had undergone HIPEC, and 84 patients (16%) had undergone EPIC. Nine patients had undergone both. All HIPEC procedures were performed intraoperatively after CRS but with many variations in exposure techniques (ie, open or closed wall), duration (30 to 90 minutes), intraperitoneal temperatures (40°C to 43°C), type of perfusate, and flow rates. Mitomycinbased regimens used mitomycin 30 to 50 mg/m<sup>2</sup>  $\pm$  cisplatin 50 to 100 mg/m<sup>2</sup> during 60 to 120 minutes at 41°C. Oxaliplatin-based regimens used oxaliplatin 360 to 460 mg/m<sup>2</sup>  $\pm$  irinotecan 200 mg/m<sup>2</sup> + intravenous fluorouracil and leucovorin during 30 minutes at 43°C. EPIC was delivered over 5 days, from day 1 to day 5 after surgery. The EPIC regimen delivered was mitomycin (10 mg/m<sup>2</sup>) on day 1 and fluorouracil (600 mg/m<sup>2</sup>) during the following four days in 0.8 to 1 L/m<sup>2</sup> of dialysis solution. Drains were clamped 23 hours of 24 hours.

Two hundred thirty-two patients (47%) had received postoperative adjuvant systemic chemotherapy when they achieved an objective response to preoperative chemotherapy (if administered) or when they exhibited poor prognostic factors (CCR-1 or -2 status, invaded lymph nodes, or liver metastases).

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Characteristic No.	%
Characteristic No.	%
Age, γears < 60 368 > 60 151	73 17
Sex Male 227 Female 296	43 57
Site of primaryRight colon162Transverse colon19Left colon71Sigmoid145Rectum36Multiple8Unknown73	31 4 14 28 7 2 14
Revealing symptom Discovered at surgery Imaging examination Abdominal pain Digestive occlusion Clinical ascites CEA elevation Worsening of general status Inguinal hernia Other	36 29 13 6 4 3 4 2 3
Yes 370 No 153	71 29
rote and rote of preoperative in clientoticity 8   PC synchronous with primary 8   Yes 161	35
No300Primary lymph node status90Positive325Negative158	65 67 33
Histologic differentiation of the primary Well 165 Moderate 195 Poor 65	39 46 15
CEA level Normal 121 Elevated 112	48 52
Calcium 19.9 level Normal 119 Elevated 80	60 60
Normal 63   Elevated 16	80 20
WHO grade     0     387       1     102     2	76 20 4
Synchronous liver metastases No 376 Yes 77	83 15
Type of intraperitoneal chemotherapy HIPEC EPIC	84 16
(continued in next column)	

Table	1. Demographic and Clinical Charac	teristics of the 523 Treated		
Patients With Colorectal PC (continued)				

	Patients		
Characteristic	No.	%	
Peritoneal cancer index			
1-6	181	37	
7-12	132	28	
13-19	96	21	
> 19	69	14	
Size of residual tumor after cytoreductive surgery, mm			
0 (CCR-0)	439	85	
< 2.5 (CCR-1)	53	10	
> 2.5 (CCR-2)	22	5	
Intraperitoneal chemotherapy regimen			
Mitomycin based	287	55	
Oxaliplatin based	235	45	

Abbreviations: PC, peritoneal carcinomatosis; CEA, carcinoembryonic antigen; IV, intravenous; HIPEC, hyperthermic intraperitoneal chemotherapy; EPIC, early postoperative intraperitoneal chemotherapy; CCR, completeness of the cancer resection.

## **Postoperative Mortality and Morbidity**

Seventeen patients (3.3%) died postoperatively. The causes of death were septic shock (n = 9), respiratory complications (n = 4), hematologic toxicity (n = 2), pulmonary embolism (n = 1), and acute renal insufficiency (n = 1).

Grades 3 to 4 complications occurred in 156 patients (31%), and 11% of all the patients underwent reoperation; 9% had a digestive fistula, 4% had a profound abscess, 6% had hemorrhage, 6% had a lung infection, and 12% had grades 3 to 4 hematologic toxicity. The median duration of hospitalization was 18 days (first-to-third quartile range, 14 to 26 days).

The logistic regression analysis of factors that significantly increased the risk of postoperative mortality and morbidity identified two primary factors—the peritoneal index (odds ratio, 1.067; 95% CI, 1.035 to 1.099; P < .001) and the center in which treatment was performed (P < .0001).

#### Survival

With a median follow-up of 45 months (first-to-third quartile range, 23 to 79 months), the overall 1-year, 3-year, and 5-year survival rates were 81%, 41%, and 27%, respectively (Fig 1; respective 95% CIs, 77% to 85%, 36% to 47%, 21% to 33%). Disease-free survival rates were 47%, 15%, and 10%, respectively (respective 95% CIs, 43% to 52%, 11% to 19%, 6% to 14%). Median survival was 30.1 months.

Among the clinical factors in the univariate analysis (Table 2), sex, age, histologic grade, and also the presence of liver metastases did not have a prognostic impact. In contrast, the extent of carcinomatosis (ie, PCI; P < .001) and invaded lymph nodes (P = .02) had significant prognostic impacts.

Among the therapeutic factors, neoadjuvant chemotherapy and use of HIPEC or EPIC did not have a statistically significant prognostic impact. In contrast, the completeness of surgery had a strong impact (P < .001), and the center where the procedure was performed (P < .001) and the use of adjuvant chemotherapy (P = .04) had significant influences on the risk of death. The impact of the PIC



Fig 1. Overall and disease-free survival rates of the 523 patients with peritoneal carcinomatosis of colorectal origin.

techniques (ie, drugs, drug concentration, temperature, duration) could not be analyzed well, because there were too many variations across centers.

A multivariate analysis (Table 3) with a Cox regression model was performed to determine which clinical or therapeutic variables were more strongly correlated with overall survival. There were four independent prognostic indicators: the PCI, which reflected the extent of peritoneal disease (Fig 2), the completeness of surgery (Fig 3), the lymph node status, and adjuvant chemotherapy.

The multivariate analysis of disease-free survival selected the following five factors: the PCI (hazard ratio [HR], 1.057; P < .0001), the completeness of surgery (HR, 0.446; P = .05), the experience of the center (HR, 2,389; P = .0003), the lymph node status (HR, 1.390; P = .03), and adjuvant chemotherapy (HR, 0.682; P = .009). Four of them were common with those of the overall survival, but the experience of the center appears in this analysis.

We specifically studied the prognostic factors in the 416 patients who had undergone complete CRS (ie, CCR-0 resection). The multivariate analysis identified five significant prognostic factors: the extent of carcinomatosis (HR, 1.054; P < .0001), the presence of liver metastases (HR, 1.623; P = .01), the experience of the center (HR, 1.841; P = .01), the lymph node status (HR, 1.44; P = .07), and adjuvant chemotherapy (HR, 0.719; P = .03). In this analysis, liver metastases had an impact on survival.

## DISCUSSION

This retrospective, multicentric study that included 523 disease occurrences shows that combined treatment of CRS and PIC yields a 5-year survival rate of 27%. This rate attained 30% when CRS was complete.<sup>6</sup> It represents the worst results that will be obtained with this combined treatment approach against PC, because it takes into account all the learning curves of the different teams. This is also the largest series of selected patients with PC from colorectal carcinoma treated with combined CRS and PIC. The study was conducted in 25 centers in the French-speaking medical community and confirms a considerable part of the results obtained in the international retrospective study of 506 patients from 28 worldwide institutions, published in 2004 by Glehen et al.<sup>7</sup> Approximately one fourth of the patients were included in these two series.

Table	2. Univariate Analysis of Overall Survival After Cytoreductive Surgery
	Plus Intraperitoneal Chemotherapy in Patients With Colorectal
	Peritoneal Carcinomatosis

	Survival Data			
Variable	Median (months)	3-Year (%)	5-Year (%)	Log-Rank <i>P</i>
Sex Male Female	30 30	39 42	26 27	.37
Preoperative systemic chemotherapy Yes No	30 30	40.5 42	27 26	.85
Performance status 0-1 2-3	33 16	45 0	30 0	.002
Synchronous resection of liver metastases Yes No	23 31	34 42	21 27	.15
Institution Best Worst	31 26	44 23	29 18	.001
Age, years ≤ 60 > 60	31 28	43 34	28 20	.07
Peritoneal cancer index 1-6 7-12 13-19 > 19	40 29 25 18	55 39 40 18 5	44 22 29 7	< .0001*
Size of remaining tumor nodules, mm 0 < 2.5 $\ge 2.5$	33 20 7	45 21 8.5	29 14 0	< .0001*
Type of intraperitoneal chemotherapy EPIC HIPEC	32 31	45 40	30 25.5	.965
Positive lymph nodes No Yes	32 24	46 36.5	30 22.5	.02*
Using oxaliplatin inside peritoneum Yes No	32 25	43 38	27 23	.02
Adjuvant systemic chemotherapy Yes No	31 27	43 40	31 23	.04*
Degree of differentiation Well Moderately Poorly	32 30 26	42 44 36	28 26 23	1.0

Abbreviations: EPIC, early postoperative intraperitoneal chemotherapy; HIPEC, hyperthermic intraperitoneal chemotherapy.

\*Variable also significant in multivariate analysis.

A comparison of these two series shows that mortality (3% and 4% for this study and the 2004 study, respectively) and morbidity (23% and 31%, respectively) are low and are no longer acceptable criticisms against this procedure. It also underlines the strong impact of the completeness of CRS on the survival rate: the 5-year survival rate was 19% in the series by Glehen et al<sup>7</sup>; 53% were CCR-0.<sup>7</sup> The 5-year survival rate was 30% in this study, and the CCR-0 in was 79%. Median survival rates were similar in the two series for

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Table 3. Multivariate Analysis of Prognostic Factors for Overall Survival of 523 Patients Treated With Cytoreductive Surgery Combined With Perioperative Intraperitoneal Chemotherapy			
	Multivariate Analysis		
Variable	Р	Hazard Ratio	95% CI
Peritoneal cancer index*	< .001	1.052	1.029 to 1.076
Completeness of surgery†	.07	1.398	0.970 to 2.014
Positive lymph nodes‡	.02	1.534	1.058 to 2.224
Adjuvant chemotherapy‡	.002	0.578	0.407 to 0.820

"For each additional point in the peritoneal cancer index, the risk of death of the relative risk increases (ie, by 5.2%).

†Completeness was divided into three categories: completeness of the cancer resection (CCR)-0, CCR-1, and CCR-2. Passing from one category to another increases the risk of death by 39%.

‡Compared in two classes (ie, yes or no).

patients who had a similar CCR-0 resection (ie, 32.4 months and 33 months, respectively); this was, of course, reassuring. Future results will be better, as confirmed by the 5-year overall survival rates recently reported by expert centers after CCR-0 : 32% for the 100 patients reported by da Silva and Sugarbaker,<sup>8</sup> 43% for the 59 patients reported by Verwaal et al,<sup>9</sup> and 48% for the 30 patients reported by Elias et al.<sup>10</sup> In 2009, the completeness of CRS must be considered fundamental because of its strong prognostic impact and the because it is a prerequisite for the use of any kind of PIC.

The main criticism medical oncologists level against this combined treatment is that it is performed exclusively in highly selected patients. Their skepticism stems from the fact that they think systemic treatment alone could yield similar results in this selected population. A recent study in patients who had resectable PC in both groups compared the results of complete CRS and HIPEC with those of systemic chemotherapy. Median survival was 60 months in the first group versus 25 months in the second group,<sup>11</sup> which underlined, at least, the positive impact of CRS.

Additional important information was obtained in this study. First, mortality (3%) and morbidity (31%) are low and are no longer acceptable reasons for not using this combined treatment. The expertise of the center has a strong prognostic impact. The procedures required for aggressive cytoreduction are lengthy, challenging, and



**Fig 2.** Prognostic impact of the extent of carcinomatosis (ie, peritoneal cancer index; P < .001) on overall survival.



Fig 3. Prognostic impact of the completeness of the surgery (P < .001) on overall survival.

morbid, and they utilize a great deal of resources. Clearly, there is a significant learning curve, and this is not a procedure that can be undertaken by the occasional operator.<sup>12,13</sup> A high volume of treated patients per center resulted in lower morbidity and mortality (data not shown) in this series, as already proven in other complex surgeries.<sup>14</sup> This impact appears to be particularly strong in this series.

Second, there is a strong correlation between the PCI (ie, extent of peritoneal disease), the completeness of CRS, and also morbidity and mortality. When the PCI is greater than 20, the 5-year survival rate is less than 10%, which proves that extensive disease becomes a relative contraindication for this combined treatment. We currently believe that a PCI greater than 20 and associated with another poor prognostic factor, such as evidence of lymph node involvement or a poor general status or progression under chemotherapy, should be considered an absolute contraindication to this treatment.

Third, when liver metastases were resected concomitantly, there was no significant prognostic impact on the entire series. However, in patients who attained a CCR-0 status, the synchronous resection of liver metastases impacted negatively on disease-free survival in the multivariate analysis of prognostic factors. Liver metastases, therefore, should be considered before deciding to perform this combined treatment, and liver metastases could be a relative contraindication if associated with a high peritoneal index.

Fourth, it was surprising to discover that there was no statistical difference in survival rates between HIPEC or EPIC. In fact, the scientific rationale for using HIPEC is more solid than that supporting the use of EPIC, <sup>15-19</sup> but it has yet to be proven in humans. In this study, postoperative morbidity also appeared to be lower (P = .03) with HIPEC than with EPIC. A randomized study comparing the two techniques in terms of tolerance, survival, quality of life, and cost would be necessary. However, before conducting such a trial, it seems more important to prove unequivocally that HIPEC or EPIC impacts on the survival rate after complete CRS and is able to cure residual occult tumor seedings. This is, in fact, the objective of the randomized trial currently ongoing in France that is comparing HIPEC with no HIPEC after complete surgical resection of PC (ie, study Prodige 7 of Fédération Francophone de Cancérologie Digestive and Fédération Nationale des Centres de Lutte Contre Ie Cancer groups).

Fifth, HIPEC techniques are miscellaneous and nonstandardized. This is a constant criticism leveled against HIPEC. In this

Information downloaded from jco.ascopubs.org and provided by at UNIV MED DNSTRY NEW JERSEY on April 6, 2014 Copyright © 2010 Americanfi@orciet@02119lit&cab@ncology. All rights reserved. multi-institutional study, although the use of oxaliplatin rather than mitomycin, and an intraperitoneal temperature greater than 42°C, seemed preferable (but more morbid) in the univariate analysis of prognostic factors for survival, the interpretation of these technical aspects appeared difficult. Clearly, more standardized and unified techniques should be used in the future.

Sixth, it is important for surgeons to realize that they play a decisive role in this treatment, which mainly is based on the peritoneal index and on the quality of the completeness of surgery. In this respect, their experience and perseverance during this long treatment will be determinative. In addition, because adjuvant systemic chemotherapy improved survival in this study, as in others,<sup>7</sup> surgeons must ensure that patients receive it.

Finally, it is important to consider the four prognostic factors for disease-free survival identified in the multivariate analysis in patients with completely resected disease, because they provide us with the keys for future improvements. These factors were the extent of PC, the presence of liver metastases (ie, a validated score incorporating the PCI and hepatic metastases is required to indicate or contraindicate complete CRS plus HIPEC.), the experience of the center (accreditation should only be delivered to a few centers), and adjuvant systemic treatment (which has improved dramatically).

In conclusion, the survival rate obtained in this multicentric study makes this combined treatment of PC the current gold standard therapy, when feasible. The analysis of prognostic factors allowed us to specify the contraindications. Future improvements in the selection of patients, in mastery of the technical process, and in the use of new molecules will rapidly increase its efficiency.

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## AUTHORS' DISCLOSURES OF POTENTIAL CONFLICTS OF INTEREST

Although all authors completed the disclosure declaration, the following author(s) indicated a financial or other interest that is relevant to the subject matter under consideration in this article. Certain relationships marked with a "U" are those for which no compensation was received; those relationships marked with a "C" were compensated. For a detailed description of the disclosure categories, or for more information about ASCO's conflict of interest policy, please refer to the Author Disclosure Declaration and the Disclosures of Potential Conflicts of Interest section in Information for Contributors.

**Employment or Leadership Position:** None **Consultant or Advisory Role:** Pierre Dubè, Roche (C), AstraZeneca (C), Novartis (C), sanofi-aventis (C) **Stock Ownership:** None **Honoraria:** None **Research Funding:** Pierre Dubè, Novartis **Expert Testimony:** None **Other Remuneration:** None

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**Conception and design:** Dominique Elias, François Gilly, Olivier Glehen **Administrative support:** Dominique Elias, Olivier Glehen **Provision of study materials or patients:** Dominique Elias, François Quenet, Jean-Marc Bereder, Baudouin Mansvelt, Gérard Lorimier, Pierre Dubè, Olivier Glehen

Collection and assembly of data: Florent Boutitie Data analysis and interpretation: Dominique Elias, Olivier Glehen Manuscript writing: Dominique Elias, Olivier Glehen Final approval of manuscript: Dominique Elias, François Gilly, Florent Boutitie, François Quenet, Jean-Marc Bereder, Baudouin Mansvelt, Gérard Lorimier, Pierre Dubè, Olivier Glehen

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## CORRECTIONS

## Author Corrections

The November 10, 2008, article by McLaughlin et al, entitled "Prevalence of Lymphedema in Women With Breast Cancer 5 Years After Sentinel Lymph Node Biopsy or Axillary Dissection: Objective Measurements" (J Clin Oncol 26:5213-5219, 2008), contained an error.

In the Results section, under Lymphedema, the *P* value in the first sentence of the third paragraph was given as P < .0001, whereas it should have been P < .93, as follows:

"For women undergoing SLNB alone, the median number of nodes excised was three (range, one to 17 nodes) for those without lymphedema and four (range, one to nine nodes) for those with lymphedema (P < .93)."

The authors apologize to the readers for the mistake.

DOI: 10.1200/JCO.2010.29.0155

The January 1, 2010, article by Elias et al, entitled "Peritoneal Colorectal Carcinomatosis Treated With Surgery and Perioperative Intraperitoneal Chemotherapy: Retrospective Analysis of 523 Patients From a Multicentric French Study" (J Clin Oncol 28:63-68, 2010), contained errors.

In the Introduction, the last paragraph should have indicated that two of the centers conducting the study did not treat patients with carcinomatosis, as follows:

"In this paper, we present a collaborative effort of 23 French-speaking centers to evaluate the efficiency of this com-

The February 1, 2010, article by Loprinzi et al, entitled "Phase III, Randomized, Double-Blind, Placebo-Controlled Evaluation of Pregabalin for Alleviating Hot Flashes, N07C1" (J Clin Oncol 28:641-647, 2010), contained an error in the spelling of the third author's name. It was originally

The February 20, 2010, article by Küpeli et al, entitled "Evaluation of Coronary Artery Disease by Computed Tomography Angiography in Patients Treated for Childhood Hodgkin's Lymphoma" (J Clin Oncol 28:1025-1030, 2010), contained errors.

In Table 1, Risk Factors for Patient 19 should have read, "Abnormal lipid profile HT, family history, **obesity**." bined approach and to identify the main prognostic factors on the basis of a population of 523 treated patients (although 25 centers conducted the study, two did not treat any patients with carcinomatosis from colorectal cancer)."

In Table 1, under Synchronous liver metastases, the number of patients without metastases was given as 376 (83%), whereas it should have been 453 (85%).

The authors apologize to the readers for the mistakes.

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given as Ernie P. Baclueva and should have been Ernie P. Balcueva.

The authors apologize to the readers for the mistake.

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In Table 3, the percentage of patients in Group I with hypertension was given as 15%, whereas it should have been **10.5%**.

The authors apologize to the readers for the mistakes.

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