Supplementary Material

S1. Patient selection

Standard demographic data, history of chemotherapy (type, targeted therapy, duration, time between end of chemotherapy and surgery), and data on colorectal metastatic cancer were documented. Biochemical variables including liver parameters, platelets and hemoglobin were recorded before and after chemotherapy, and after surgery. The aspartate aminotransferase to platelet ratio index (APRI) was calculated before liver resection.

S2. Surgical methods

Surgery characteristics described included the extent of liver resection, combination with intraoperative radiofrequency ablation (IRFA), association with extra hepatic procedures [1]. Postoperative complications were recorded according to Dindo-Clavien system, based on medical and paramedical reports [2]. Grade 3 or higher was considered as a major complication. International Study Group of Liver Surgery (ISGLS) definition for hepatic failure, describing three grades (A to C) based on biological and clinical signs, including transitory hepatic failure (grade A), was used [3]. Postoperative mortality was defined by 90-day mortality [4].

References

[1] Evrard S, Poston G, Kissmeyer-Nielsen P, Diallo A, Desolneux G, Brouste V, et al. Combined ablation and resection (CARe) as an effective parenchymal sparing treatment for extensive colorectal liver metastases. PLoS One 2014;9(12):e114404.

[2] Dindo D, Demartines N, Clavien PA. Classification of surgical complications: a new proposal with evaluation in a cohort of 6336 patients and results of a survey. Ann Surg 2004;240(2):205-13.

[3] Rahbari NN, Garden OJ, Padbury R, Brooke-Smith M, Crawford M, Adam R, et al. Posthepatectomy liver failure: a definition and grading by the International Study Group of Liver Surgery (ISGLS). Surgery 2011;149(5):713-24.

[4] Mayo SC, Dhore AD, Nathan H, Edil BH, Hirose K, Anders RA, et al. Refining the definition of perioperative mortality following hepatectomy using death within 90 days as the standard criterion. HPB (Oxford) 2011;13(7):473-82.

Supplementary Table S1: Post-operative complications according to Dindo-Clavien system and details of liver-related complications according to ISGLS score

	Number of		Number
Martal.	Number of		Number
Variable	patients	%	(min – max)
Post-operative complications			
Yes	78	53.1	
No	69	46.9	
Major complications*	29	19.7	
Liver related complications			
	21	21.1	
Yes	31	21.1	
No	116	78.9	
Hemorrhage	4	27	
		2.7	
Biliary complications	13	8.8	
Liver failure	15	10.2	
Grade 1	13	8.8	
Grade 2	2	1.4	
Grade 3	0	0	
Hemorrhage + liver failure	1	0.7	
Average number of blood transfusion			1 (0 – 13)
Median duration of hospitalization, days			10 (4 - 47)

*Major complications: grade ≥ 3

Supplementary Table S2. Literature review of studies on the impact of chemotherapy on non-tumoural liver and its consequence on post-operative outcomes. The results concerning the impact on chemotherapy on post-operative outcomes are not presented in this table.

Reference	Hepatic resection (N)	Patients with NCT, (N)	Type of pre- operative chemotherapy (NCT)	Histologic lesions in non tumoral liver	Incidence (%)	Significate risk factor, at multivariate analysis	Post-operative (PO) impact of the histologic lesions
Aloia (2006)	92	75	5FU or FOLFOX	Sinusoidal dilation Peliosis NRH Steatosis	23 after NCT 31 after NCT 3 after NCT 13 after NCT	NCT NCT NCT	More transfusion More transfusion
Vauthey (2006)	406	248	5FU or FOLFOX or FOLFIRI	Grade 2 or 3 sinusoidal dilation Steatosis > 30% Steatohepatitis	5.4 8.9 8.4	Oxaliplatin Irinotecan	None None PO mortality, liver failure
Pawlik (2007)	212	153	5FU and/or FOLFOX and/or FOLFIRI	CALI score Grade 3 sinusoidal dilation Steatosis>30% Steatohepatitis	4.6 18.3 after NCT 1.9	NCT, diabetes, BMI>30 NCT Irinotecan, diabetes, BMI>30	None None None
Mehta (2007)	173	130	5FU and/or FOLIRI Or FOLFOX	Grade 2 sinusoidal dilation	52.8	Oxaliplatin	None
Brouquet (2009)	146	146	5FU and/or FOLFOX and/or FOLFIRI	CALI score Grade 2 or 3 sinusoidal dilation Steatosis>30% Steatohepatitis	51 34.2 21.9 10.3	BMI>27, diabetes, interval NCT- Surgery<4 weeks Oxaliplatin BMI>27 BMI>27	PO complications None None None
Rubbia-Brandt (2010)	385	274	FOLFOX and/or FOLFOXIRI	Grade 2 or 3 sinusoidal dilation Peliosis Steatosis Steatohepatitis	54.3 after NCT 10.6 after NCT 39.8 after	Oxaliplatin Oxaliplatin	Not evaluated

					NCT		
					0		
Ryan (2010)	334	132	5FU and/or	Grade 2 or 3 sinusoidal	5.9		None
			FOLFOX and/or	dilation	15.7	Oxaliplatin	None
			FOLFIRI	CVI score≥3*	9.9	BMI>30	None
				Steatosis>33%	2.4	BMI>30	None
				Steatohepatitis			
Pilgrim (2012)	232	133	NM*	Sinusoidal dilation	19	NCT (high grade	None
				Steatosis	18	lesions)	PO morbidity, respiratory
						Diabetes, BMI>30	complications and wound
				Steatohepatitis	4		infection
						Metabolic syndrome	None
Wolf (2013)	384	250	FOLFOX and/or	Sinusoidal dilation	10		None
			FOLFIRI	Steatosis	35	BMI>25, diabetes,	None
				Steatohepatitis	4	irinotecan	None
Vigano (2015)	478	478	FOLFOX and/or	NRH	18.2	Oxaliplatin	PO liver failure
			FOLFIRI	Moderate or severe SOS	38.7		None
				Steatohepatitis	10		None
Martins (2016)	140	70	5FU and/or	Peliosis	25.7 after	NCT	PO morbidity, liver failure
			FOLFOX and/or	Moderate or severe SOS	NCT	NCT	PO morbidity, liver failure
			FOLFIRI	Steatosis	21.4 after	Diabetes	Reduced PO morbidity
				Steatohepatitis	NCT		Reduced PO morbidity
					59.3		

NCT: neoadjuvant chemotherapy

NRH: nodular regenerative hyperplasia

NM: not mentioned

CVI score: combined vascular injury score