ORIGINAL ARTICLE

Exposure During Pancreatic Surgery. Do We Have to Examine the Falciform and Round Ligaments for Pancreatic Cancer Metastasis?

Offir Ben-Ishay¹, Arie Ariche², Jesse Lachter⁴, Yoram Kluger^{1,3}

¹Department of General Surgery, ²HepatobiliarySurgery Service, ³Centre for Pancreatic Surgery, and ⁴Department of Gastroenterology; Rambam Health Care Campus. Haifa, Israel

ABSTRACT

Context Metastatic and locally advanced disease is unfortunately a common clinical occurrence in patients affected by pancreatic cancer. The most frequent sites of metastasis are the liver and the peritoneal lining. The falciform and round ligaments are routinely resected and sent for pathology, but no metastases have ever been reported. Objective In this study, we assess whether the falciform and round ligaments are possible sites for metastases and if routine pathology examination of these structures is justified. Design Retrospective, single institution study. Patients The charts of all patients who underwent pancreatic resection from June 2005 through January 2011 were reviewed retrospectively. Data on age, gender, type of operation performed, preoperative CA 19-9 levels, vascular and perineural invasion, tumor differentiation and pathological staging were retrieved from the hospital electronic data base. Only patients with malignant lesions of the pancreas were included in the study. Main outcome measures Presence of pancreatic metastasis in the falciform and round ligaments. Results Fifty-seven patients were included in the study. Forty patients had undergone pancreaticoduodenectomies, 10 subtotal pancreatectomies, 2 total pancreatectomies, and 1 distal pancreatectomy. Four patients had exploratory laparotomy but exploration revealed stage III or IV disease, thus the planned resection was not carried out. The falciform and round ligaments were all negative for metastasis in all patients. Discussion Metastatic progression of pancreatic cancer is poorly understood. Theoretically, metastases to the falciform and round ligaments could occur by either contiguous, hematogenic or lymphatic spread. In our study, none of the patients had metastases to the falciform and round ligaments, regardless of the pathological staging. Conclusion We concluded that routine pathological examination of the falciform and round ligaments is not justified if the surgeon elects to remove the structures for ease of exploration and retraction during surgery. This may lighten the workload and reduce costs in a busy pathology laboratory.

INTRODUCTION

The incidence of pancreatic cancer has increased during the last decade and is ranked as the 4th leading cause of death in the United States [1]. The etiology of pancreatic cancer is poorly understood [2] and overall survival is only 4% [3].

Pancreatic cancer is characterized by early systemic dissemination and extraordinary local tumor progression [4]. Seventy-five percent of the patients diagnosed with pancreatic cancer have advanced disease [5]. Metastases are often present at the time of diagnosis. Common sites for metastases are the liver and the peritoneal cavity. Improvement in the preoperative workup has reduced the number of

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Correspondence Offir Ben-Ishay

Department of General Surgery; Rambam Health Care Campus; PO Box 9602; Haifa, 531096; Israel

Phone: +972-50.206.3177; Fax: +972-4.854.2231 E-mail: o_ben-ishay@rambam.health.gov.il

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operations for non-resectable tumors in patients with tumors of the pancreatic head. A computed tomography scan is considered the gold standard in preoperative evaluation. However, its accuracy in detecting vascular involvement, liver or peritoneal metastasis is limited. Exploratory laparoscopy is an important tool in these patients' workup but some authors call for its selective use. In our institution, as a result of a study we conducted on preoperative indicators for non-resectability, laparoscopy is performed only in patients with CA 19-9 levels higher than 300 U/mL [6]. Positron emission tomography combined with computed tomography is increasingly being used for a metastatic workup of pancreatic cancer, and some authors have reported a specificity and sensitivity of 94% and 90%, respectively [7].

During pancreatic surgery we routinely dissect the falciform and round ligaments and the surrounding fat for ease of retraction and exposure [8]. All resected tissues are routinely sent for pathological examination. In a busy pancreatic surgery center, this may impose extra work on the pathology laboratory.

In this study, we assessed whether the falciform and round ligaments are a possible site for metastases and

Table 1. Patient characteristics.	
Age; years. Mean (range)	70 (48-84)
Gender: - Male	31 (54.4%)
- Female	26 (45.6%)
CA 19-9; U/mL. Mean (range)	448 (0-6,162)
Vascular invasion	3 (5.3%)
Perineural invasion	14 (24.6%)
Differentiation:	
- Well	17 (29.8%)
- Moderate	21 (36.8%)
- Poor	15 (26.3%)
- Non relevant a	4 (7.0%)
Mean lymph nodes harvested per patient	12
Positive lymph nodes:	
- Number of patients	23 (40.4%)
- Positive:total lymph node ratio	0.17
^a Patients with stage III and IV disease is	n which no complet

whether routine pathology examination of these tissues is mandatory.

PATIENTS AND METHODS

resection was performed

Electronic charts of all patients who underwent pancreatic resection in the Department of Surgery at the Rambam Health Care Campus (RHCC), Haifa, Israel from June 2005 through January 2011 were retrospectively reviewed. The data collected included age, gender, type of operation performed, preoperative CA 19-9 levels, vascular and perineural invasion, level of tumor differentiation and pathological staging.

A unilateral left subcostal incision was performed for distal pancreatectomies and a bilateral subcostal was performed for subtotal, total and pancreatico-duodenectomies. Falciform and round ligaments were routinely resected and sent to the pathology lab in all operations in which the incision crossed the midline.

Only patients with adenocarcinoma of the pancreas were included; other periampullary tumors, such as duodenal carcinoma, ampullomas, tumors of the distal common bile duct and other pancreatic malignant (neuroendocrine), premalignant (intra-pancreatic mucinous neoplasm) or benign cystic lesions were excluded from the study.

Levels of CA 19-9 are expressed in U/mL with a reference range of 0-33 U/mL; the TNM staging system was used for postoperative pathological staging.

STATISTICS

Mean, standard deviation, range and frequencies were used as descriptive statistics. The one-way analysis of variance (ANOVA) and the linear-by-linear association chi-square test were applied to analyze data by running the SPSS for Windows (version 13.0) statistical package using a personal computer.

ETHICS

Informed consent was waived due to the retrospective nature of the study and the study protocol conformed to the ethical guidelines of the "World Medical Association Declaration of Helsinki - Ethical Principles for Medical Research Involving Human Subjects" adopted by the 18th WMA General Assembly, Helsinki, Finland, June 1964, as revised in Tokyo 2004, as reflected in a priori approval by the appropriate institutional review committee.

RESULTS

One hundred and 89 patients went through pancreatic resection in the study period; only 57 patients met the inclusion criteria for the study: 40 had had pancreaticoduodenectomies, 10 subtotal pancreatectomies, 2 total pancreatectomies, and 1 patient had a distal pancreatectomy with resection of the falciform ligament. Four patients had had an exploratory laparotomy but exploration revealed non-resectable (stage III) or metastatic (stage IV) disease. Table 1 depicts patient demographics as well as pre- and post-operative variables. Twenty-three patients (40.4%) had positive lymph nodes; the mean number of lymph nodes harvested was 12 and the mean positive to total lymph node ratio was 0.17.

Table 2 depicts the pathological staging of the patients. We noted that 24 patients (42.1%) were in stage IIb. The falciform and round ligaments were all negative

The falciform and round ligaments were all negative for metastases, even when poor prognostic factors were present, such as perineural invasion (14 cases; 24.6%), vascular invasion (3 cases; 5.3%) and high CA 19-9 levels (mean CA 19-9 level was 448 U/mL; range: 0-6,162 U/mL). Table 3 depicts the pathological staging of patients correlated to age, grade of differentiation and the described prognostic factors. The falciform and round ligaments were negative for metastases even in patients with unresectable (stage III) or metastatic disease (stage IV).

DISCUSSION

Metastatic pancreatic cancer is common and 75% of patients diagnosed with pancreatic cancer have either metastatic or locally advanced disease [5, 9]. The most common sites of metastasis are the liver and the peritoneal lining but unusual sites such as the lung, muscle, skin, heart, pleura, stomach, bone, umbilicus, kidney, appendix, spermatic cord and prostate have also been described [10, 11, 12, 13, 14, 15, 16, 17, 18, 19].

The falciform ligament is a broad and thin anteroposterior peritoneal fold, which is a remnant of the

Table 2. Distribution of patients according to the Union for International Cancer Control (UICC) staging system.

TNM staging system	No. of cases				
Ia	3 (5.3%)				
Ib	21 (36.8%)				
IIa	5 (8.8%)				
IIb	24 (42.1%)				
III	2 (3.5%)				
IV	2 (3.5%)				
Total	57				

Table 3. Patient pathological staging correlated to age, differentiation and prognostic factors.

Staging	Age; mean±SD (years)	Differentiation			Prognostic factors			
		Well	Moderate	Poor	Perivascular	Perineural	Patients with positive lymph nodes	CA 19-9; mean±SD (U/mL)
Ia (No. 3)	63±7	3 (100%)	0	0	0	0	0	101±120
Ib (No. 21)	67±9	7 (33.3%)	9 (42.9%)	5 (23.8%)	0	5 (23.8%)	1 (4.8%)	654±1,358
IIa (No. 5)	75±10	0	3 (60.0%)	2 (40.0%)	1 (20.0%)	0	0	398 ± 40
IIb (No. 24)	72±8	7 (29.2%)	9 (37.5%)	8 (33.3%)	2 (8.3%)	9 (37.5%)	22 (91.7%)	352 ± 602
III (No. 2)	79±3	NR	NR	NR	NR	NR	NR	385±193
IV (No. 2)	65±6	NR	NR	NR	NR	NR	NR	303±152
P value	0.079 a		0.132 b		0.217 b	0.153 b	<0.001 b	0.885 a

NR: non relevant

ventral mesentery of the fetus. It is attached to the undersurface of the diaphragm and the posterior surface of the right rectus sheath as low down as the umbilicus. On its right margin, it extends from the notch on the anterior margin of the liver as far back as the posterior surface. The two layers of the peritoneum are closely attached together and the free edges contain the round ligament and the paraumbilical veins. The round ligament, or ligamentum teres, is a degenerative string of tissue which anatomically divides the left lobe of the liver from the medial and lateral sections. The metastatic progression of pancreatic cancer is poorly understood; falciform ligament metastasis could occur by contiguous peritoneal spread, by hematogenous spread through the arterial and venous system or by lymphatic spread with extension along the round ligament. In the English literature, there are only scattered reports of falciform ligament metastasis from stomach and renal cell carcinoma [20, 21]. To our knowledge, pancreatic cancer metastasis to the falciform ligament has never been reported. In our study, none of the patients had falciform or round ligament metastases, even when poor prognostic markers were present (e.g., perineural invasion, high preoperative levels of CA 19-9, or poor pathological staging in resectable, non-resectable or metastatic disease).

CONCLUSIONS

Based on our study, we concluded that routine falciform and round ligament pathological examination is not indicated when these are removed during surgery for pancreatic cancer. This may reduce the workload on the pathology laboratory. Additional larger scale studies are needed to confirm and determine the real incidence of falciform and round ligament metastases from pancreatic cancer.

Conflicts of interest The authors have no potential conflicts of interest

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a One-way ANOVA

^b Liner-by-linear association chi-square

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