Abstract. Background: Most cases with advanced-stage epithelial ovarian malignancies will experience recurrent disease at a certain moment of their evolution, even if maximal cytoreductive surgery has been performed at the moment of initial diagnosis. However, it seems that the best therapeutic strategy, in case of relapse, remains aggressive re-resection, with complete cytoreduction being the most efficient way to improve survival. Materials and Methods: We present the case of a 55-year-old patient diagnosed with an isolated pelvic recurrence after stage IIIC surgically-treated ovarian cancer three years after primary cytoreduction. Results: Intraoperatively, the diagnosis of an isolated pelvic recurrence invading the urinary bladder was confirmed. The recurrent tumor was resected en bloc with partial cystectomy. The postoperative course was uneventful, while histopathological studies confirmed the presence of a poorly differentiated epithelial ovarian recurrent adenocarcinoma. At two year follow-up, the patient is free of any recurrent disease. Conclusion: Isolated pelvic recurrences after surgically-treated ovarian adenocarcinomas can be safely removed and might improve survival.

Ovarian cancer remains one of the most common gynecologic malignancies affecting women worldwide and is commonly diagnosed in advanced stages of the disease when disseminated lesions are already present (1). In most of these cases, although the principles of complete cytoreductive surgery are applied, relapse will occur at a certain moment. The standard treatment for these patients has widely varied in the last decades; initially it has been considered that platinum-based chemotherapy should become the standard of care for ‘platinum-sensitive’ recurrent tumors, while the benefits of secondary cytoreductive surgery have been considered uncertain (2). However, this theory was discarded by studies demonstrating that debulking surgery is able to increase the biological efficacy of chemotherapeutic agents by removing all bulky tumors with poor vascular supply (3). The benefits of secondary cytoreduction at the time of relapse seem to be more evident in cases presenting good performance status, early stage at diagnosis, no residual disease after primary cytoreduction, association of adjuvant chemotherapy, longer disease-free survival, normal cancer antigen (CA)-125 levels, no ascites and isolated recurrences (4-10). The good postoperative outcome reported by patients presenting isolated relapse is mainly explained by the limited number of visceral resections required to obtain complete cytoreduction, transforming these cases in perfect candidates for secondary cytoreduction (11, 12). Unfortunately, it is well-known that this eventuality is a rare one as most cases present multiple recurrences necessitating an important visceral sacrifice both in the lower and upper abdomen (13-15). The aim of this work is to present the case of a 55-year-old patient diagnosed with an atypical, isolated pelvic recurrence invading the urinary bladder removed by performing a partial cystectomy three years after initial diagnosis of ovarian cancer.

Case Report

A 55-year-old patient with a medical history of stage IIIC ovarian cancer presented in our service for pelvic pain and macroscopic hematuria. The patient had been submitted to cytoreductive surgery three years before; total hysterectomy with bilateral adnexectomy, pelvic, parietal bilateral peritoneectomy, omentectomy, pelvic and para-aortic lymph node dissection. At that moment, an R0 resection was achieved, while the histopathological examination revealed...
the presence of a medium to poorly differentiated epithelial serous ovarian adenocarcinoma. Postoperatively, the patient was submitted to six cycles of adjuvant platinum- and taxane-based chemotherapy, while the follow-up revealed no recurrent disease for the next two years and half. The imagistic studies performed at the current moment revealed the presence of an isolated tumor located on the posterior wall of the urinary bladder, in close connection with the sigmoid loop. The patient was submitted to cystoscopy, which confirmed the lesion that was biopsied. The histopathological examination confirmed the ovarian metastatic origin and, thus, the patient was re-submitted to surgery; intraoperatively, the presence of the isolated recurrence in the urinary bladder wall, adherent the sigmoid loop, was confirmed (Figure 1). The recurrent tumor was resected en bloc with partial cystectomy, while the digestive structures were successfully dissected and preserved. The parietal defect of the urinary bladder was closed throughout a two layer closure of 2-0 running, delayed absorbable, non-locking stich (Figures 2-7); the transurethral Foley catheter was retrieved in the 21th postoperative day, after performing a computed tomography in order to confirm the intact repair of the urinary bladder. Since the postoperative course was uneventful, the patient was discharged in the fifth postoperative day. Histopathological examination confirmed the ovarian origin of the tumor, which was successfully removed with negative resection margins. At two years of follow-up, the patient is free of any local or distant recurrent disease.

Discussion

Cases presenting isolated recurrences with ovarian cancer origin are uncommon since most cases are exhibiting disseminated lesions at the time of diagnosis. Fortunately, this has been widely demonstrated to be a favorable prognostic factor, being associated with high rates of long-term survival whenever complete cytoreductive surgery is performed (4, 16, 17). Moreover, among patients presenting localized relapsed, cases with lymph node recurrence seem to have the best overall prognosis (17). One of the largest studies conducted on the theme of the influence on survival of the anatomic site of relapse comes from Petrillo et al. demonstrating that cases with isolated lymph node recurrences experience a significantly higher overall survival versus cases with isolated peritoneal lesions or versus cases with isolated parenchymatous lesions, with median progression-free survival being 63 versus 41 versus 24 months (11). In the meantime, the presence of peritoneal carcinomatosis has been reported in up to 70% of patients with recurrent disease but failed to be a negative prognostic factor on the overall survival if complete debulking surgery...
to no residual disease is feasible (18). The benefits of complete cytoreductive surgery at the time of relapse have been best revealed by Bristow et al.’s meta-analysis, published in 2009, including 2,019 patients from 40 studies. The authors created a regression model demonstrating that the median overall survival time increased from 18 months to 48 months if the proportion of cases submitted to complete cytoreductive surgery increased from 0 to 100% (with each 10% increase of the proportion of cases submitted to complete cytoreduction at the moment of the first relapse being associated with a 3-month increase of the median survival of the cohort) (19).

In order to determine the patients to potentially benefit most after secondary cytoreduction, Minaguchi et al. conducted a study on 80 cases submitted to secondary cytoreduction for ovarian cancer relapse; the authors studied the influence of four different favorable prognostic factors on the long-term outcome: treatment-free interval longer than 12 months, absence of distant metastases, presence of isolated lesions and 0 performance status. The authors demonstrated that cases achieving three or all of the four factors are the best candidates for secondary cytoreductive surgery (20). According to Minaguchi et al.’s theory, our patient was the perfect candidate for secondary cytoreductive surgery, with all four criteria being fulfilled.

Another particularity of our case is the location of the relapse in the urinary bladder requiring a partial cystectomy. The presence of an infiltrating tumor at this level necessitated partial urinary bladder resection with 1 cm resection macroscopic-free margin; however, this resection did not interfere with the functional outcome of the patient, with more than 30% of the urinary bladder capacity being preserved.

The feasibility of urinary tract resections, as part of debulking surgery for advanced stage ovarian cancer, was initially demonstrated by the study of Malviya et al.; the study included 22 patients submitted to surgery between 1983 and 1987. The most commonly performed urinary tract resections included partial cystectomy (in seven cases) followed by ureteroneocystostomy (in five cases), partial cystectomies with ureteroneocystostomy (in four cases) and uretero-ureteral anastomosis (in other four cases). The authors reported nine major and two minor complications that occurred in seven cases, with one patient being dead at six weeks postoperatively. When it comes to the long-term outcomes, Malviya et al. reported a mean survival period of 15.2 months for patients in whom complete cytoreduction had been achieved. The authors concluded that urinary tract resections are justified if complete debulking surgery is achievable with acceptable rates of postoperative morbidity (21).
However, literature data regarding urinary tract resections as part of debulking surgery for advanced stage or relapsed ovarian cancer are scarce; most often these resections are presented as collateral surgical procedures for pelvic tumors with ovarian origin and rectosigmoid involvement (22-24).

In Park et al.’s study, the authors included 60 patients submitted to multiple visceral resections for advanced stage or relapsed ovarian cancer; among these cases, 14 patients were introduced in the current study at the moment of secondary cytoreductive surgery, with six of them necessitating urinary tract resections. The most commonly performed urinary tract resections at the time of secondary cytoreductive surgery included ureteral resection with uretero-ureteral anastomosis and partial cystectomy and did not associate any specific urinary complication (23). Another study that reported the association of urinary tract resections as part of debulking surgery for pelvic recurrences after surgically treated ovarian cancer comes from Bristow and colleagues encompassing 56 patients submitted to surgery at Kelly Gynecologic Service, Baltimore, MD, USA, and at the Division of Gynecologic Oncology of the European Institute of Oncology, Milan, Italy. Urinary tract resections consisted of segmental ureteral resections and partial cystectomies and were performed in eight cases. In all cases, the continuity of the urinary tract was re-established by ureteroneocystostomy and psoas hitch with or without Boari flap and did not associate any urinary leaks (24).

Conclusion

Isolated pelvic recurrences after advanced-stage, surgically-treated ovarian cancer are rare eventualities and associate a good long-term outcome in terms of disease-free and overall survival if cytoreductive surgery is performed. The particularity of our case consists of the limited pelvic visceral invasion of the recurrent tumor, imposing only a partial cystectomy and no digestive visceral sacrifice. In the meantime, the tumor location out of the urinary bladder...
trigone and the absence of any sign of ureteral ostium invasion pointed to a limited cystectomy with no need of ureteral resection or reimplantation.

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