## Long-term survival following repeat liver resections in metastatic ovarian granulosa cell tumor: Case report with review of the literature

**DOI:** 10.4103/2278-330X.208852

Dear Editor,

The natural history of Granulosa cell tumor (GCT) is characterized by long disease free interval with multiple episodes of recurrences. Hepatic metastasis occurs in 4% of cases and the treatment in this scenario is usually palliative. We report a case of a stage IIIb GCT undergoing HR twice with good outcome and hope this encourages gynecologic oncologists to consider HR in selected cases.

A 35-year-old lady underwent total abdominal hysterectomy with bilateral salpingoovariotomy and omentectomy for menorrhagia on 11/11/1997 for a stage IIIb (omental metastasis) grade 3 GCT. She refused adjuvant chemotherapy and was lost to follow-up.

She was evaluated for abdominal discomfort and detected to have ascites with pelvic and hepatic recurrence in 04/02/2003 and was managed with four cycles of bleomycin, etoposide and cisplatin, followed by debulking of pelvic disease with removal of remnant omentum and pelvic node sampling. She

had residual disease in omentum and received a postoperative abdominopelvic radiation.

She was referred to our institute for pain in the right hypochondrium with nausea, loss of weight and appetite and computed tomography (CT) abdomen showing a large hypodense lesion in segment VII of the liver. Tumor markers alfa fetoprotein, carcinoembryonic antigen and CA 125 were normal. Biopsy was consistent with metastatic GCT. In view of good performance status, long disease free interval and solitary nature of the lesion a decision for surgical resection of the lesion was taken. Intraoperatively, there was a mass of 5 cm × 5 cm arising from the diaphragm and partially infiltrating segment VII of the liver while the rest of the abdomen was free of disease. She underwent excision of diaphragmatic deposit with nonanatomical liver resection on 27/04/2007. During her postoperative stay, she developed right-sided pneumothorax which was managed conservatively. No adjuvant treatment was offered, this being a solitary lesion which was completely excised.

She developed a second hepatic recurrence in 21/01/2010 with CT abdomen showing a 27 mm  $\times$  22 mm lesion in segment VIII of liver. In view of the previous hepatic resection, it was decided in the multidisciplinary tumor board to offer a right hepatectomy. Again, no adjuvant treatment was given.

Table 1: Patients with metastatic GCT who underwent liver resection

Authors	Time*	Time†	Symptoms	Surgery	PHRS	OS	Present
	(months)	(months)			(months)	(months)	status
Crew et al.[2]	144	120	Dyspnoea	Partial HR, wedge excision right lower lobe lung	5	276	Alive with disease
Madhuri et al.[3]	204	32	Dyspnea	Partial HR, omentectomy, excision of abdominal disease	40	276	NED
Madhuri et al.[3]	108	12	Abdominal pain	Partial HR, stripping right subdiaphragm peritoneum, cholecystectomy, right hemicolectomy, anterior resection	34	156	NED
Madhuri et al.[3]	72	0	Abdominal pain	Partial HR, stripping bilateral subdiaphragm peritoneum, omentectomy, excision pelvic disease	24	96	NED
Chua et al.[4]	144	0	NA	Partial HR, right lower lobectomy of the lung, stripping right subdiaphragm peritoneum, right adrenalectomy	95	239	NED
Chua et al. <sup>[4]</sup>	72	172	NA	Right hepatectomy, stripping right subdiaphragm, total anterior parietal and pelvic peritonectomy, stripping of the omental bursa, cholecystectomy, small bowel resection, anterior resection, right adrenalectomy	10	254	Alive with disease
Rodríguez García <i>et al.</i> <sup>[5]</sup>	72	60	Abdominal mass	Right hepatectomy	9	141	Died of other causes
Present study	63	50	Abdominal pain Asymptomatic	Partial HR, excision subdiaphragm lesion Right hepatectomy	86	200	NED

GCT=Granulosa cell tumor, NA=Not available, PHRS=Posthepatic resection survival, OS=Overall survival, HR=Hepatic resection, NED=No evidence of disease. \*Time from primary diagnosis to hepatic recurrence, †Time from hepatic recurrence to hepatic resection

In 10/01/2012, CT abdomen showed a 2.4 cm  $\times$  1.8 cm lesion in left pelvic wall. Biopsy was consistent with recurrent GCT. Initially, patient was not willing for surgery or intravenous chemotherapy and was managed with oral etoposide and later with letrozole. But due to progressive disease, excision of the lesion was done on 04/09/13. She is clinically and radiologically free of disease till date (31/07/2014).

This is probably the first reported case in the English literature of a patient with metastatic GCT undergoing liver resection twice in a span of 3 years and still doing well 4½ years later.

Our decision for surgical management of the recurrences has been based on the available literature which has shown the benefits of debulking surgeries both in the primary and recurrent scenario for GCT.<sup>[1]</sup>

Table 1 lists the patients with metastatic GCT, who underwent liver resection.

Unlike epithelial ovarian tumors, GCT are indolent tumors and slowly grow to occupy a large volume of the liver causing compression of neighboring structures. Liver resection was successfully done in two cases presenting with dyspnea. Both cases had good quality of life (QoL) post HR.

The solitary nature of the liver metastasis strengthens the argument for HR. However, the presence of extrahepatic disease need not be a contraindication for surgery if complete debulking can be achieved as shown by Madhuri *et al.* and Chua *et al.*<sup>[3,4]</sup>

In view of the hypervascularity noted in these tumors,<sup>[5]</sup> there exists a grave danger of tumor rupture causing intraabdominal bleed and even death. Hence, it can be justified to perform HR even if asymptomatic which prompted us to offer surgery the second time without any delay.

We hope the above justifications for HR in metastatic GCT will encourage more gynecologic oncologist to consider HR and work in tandem with hepatobiliary surgeons to achieve optimal cytoreduction. The selection criteria for justifying hepatic metastectomy should be formulated by a consensus

from the available literature. Repeated liver metastasis can also be managed surgically leading to an improved QoL as seen in our case. Whether this translates to an improved recurrence-free survival and OS cannot be commented at this stage due to the limited data on the subject. However, we can probably say that an aggressive surgical approach may be justified in metastatic GCT where there is limited disease with a long recurrence-free survival and good performance status as in our case.

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**How to cite this article:** Antony MA, Pavithran K, Vijaykumar DK, Sudheer OV. Long-term survival following repeat liver resections in metastatic ovarian granulosa cell tumor: Case report with review of the literature. South Asian J Cancer 2017;6:87-8.

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