

Dear Editors and Reviewers,

Thank you very much for your comments concerning our manuscript. We have carefully considered all suggestions and questions, and modified the manuscript to address all points raised by the reviewers. We also responded point by point to the reviewers' comments as listed below.

Hope these will make it more acceptable for publication.

Yours sincerely,

Jian Wang, MD, PhD

The title must be informative, specific, and brief (Title should be no more than 10~12 words/60 bytes. Please revise it). Words should be chosen carefully for retrieval purposes. All nonfunctional words should be deleted, such as 'the', 'studies on', 'observations of', and 'roles of', *etc.*

Answer: Thank you very much for the comments. We have revised the title according to the reviewer's suggestion (P1, L1-2).

(1) Case characteristics

Please summarize main symptoms in one sentence.

Answer: A 50-year-old female with recurrent major duodenal papilla cancer (MDPC) following pancreaticoduodenectomy.

(2) Clinical diagnosis

Please summarize main clinical findings in one sentence.

Answer: Postoperatively recurrent MDPC with widespread lymph node metastases at the bilateral cervix, mediastinum, abdominal cavity, and retroperitoneal area.

(3) Differential diagnosis

Please summarize thoughts and methods for differential diagnosis in one sentence.

Answer: Periapillary cancers (pancreas, duodenum, distal common bile duct, and ampulla of Vater).

(4) Laboratory diagnosis

Please summarize laboratory testing methods and major findings in one sentence.

Answer: Serum carbohydrate antigen 19-9 was 1046 U/mL (normal range, < 34 U/mL).

(5) Imaging diagnosis

Please summarize imaging methods and major findings in one sentence.

Answer: PET with 18-fluorodeoxyglucose/CT showed hypermetabolic foci at the bilateral cervix, mediastinum, abdominal cavity, and retroperitoneal sites with mean standard uptake values of 2.8, 2.4, 1.8 and 1.8 respectively.

(6) Pathological diagnosis

Please summarize pathological methods and major findings in one sentence.

Answer: Metastatic adenocarcinoma was confirmed by ultrasonographic-guided fine-needle biopsy of the left neck.

(7) Treatment

Please summarize treatments and drugs used in one sentence.

Answer: Intravenous oxaliplatin plus oral capecitabine regimen concurrent

with image-guided radiation therapy.

(8) Related reports

Please provide other contents related to the case report to help readers better understand the present case.

Answer: To the best of our knowledge, this is the first case to demonstrate the role of chemoradiotherapy and the corresponding survival improvement in extensively recurrent MDPC.

(9) Term explanation

Please explain uncommon terms present in the case report.

Answer: MDPC is typically described as periampullary carcinoma because most tumors arise in the periampullary region.

(10) Experiences and lessons

Please summarize experiences and lessons learnt from the case in one sentence.

Answer: A regimen of intravenous oxaliplatin plus oral capecitabine concomitant with image-guided radiation therapy is a potential treatment for MDPC patients with widespread lymph node recurrence after radical pancreaticoduodenectomy.

(11) Peer- review

Please summarize the strengths and weaknesses of the article based on the reviewers' comments so that readers can obtain objective knowledge from the article.

Answer: The authors have described a rare case of recurrent MDPC with widespread lymph node metastases. The report highlights the clinical characteristics of this tumor and provides insights into the therapeutic implications.