

## CASE REPORT

# Extensive Paneth cell metaplasia in an ovarian Krukenberg tumor: report of an unusual case and literature review

Waleed Abdullah Alrajban<sup>1,\*</sup>, Reham Ali Khubrani<sup>1</sup>,  
Mamdouh Saad Almalki<sup>2</sup>, Ashraf Almassri<sup>3</sup>,  
and Ammar Cherkess Alrikabi<sup>1</sup>

<sup>1</sup>Department of Pathology, King Khalid University Hospital and King Saud University, Riyadh, Saudi Arabia,

<sup>2</sup>Department of Radiology, King Khalid University Hospital and King Saud University, Riyadh, Saudi Arabia,

and <sup>3</sup>Department of Pathology, Al Hammadi Hospital, Riyadh, Saudi Arabia

\*Correspondence address. Pathology Department, King Saud University and King Khalid University, 11461 Riyadh, KSA. Tel: +966-55-386-5599; E-mail: waleed.alrajban@gmail.com

## Abstract

Paneth cells are classified as secretory cells which are normally found in the cecum and ascending colon. Their presence in other parts of the gastrointestinal tract is regarded as abnormal and indicates metaplasia. Paneth cells may also be rarely found in gastrointestinal, biliary and prostatic tumors. The presence of Paneth cells in ovarian Krukenberg tumors is rare and to the best of our knowledge is restricted to metastatic appendiceal goblet cells tumors. We report a rare case of unilateral metastatic gastric carcinoma to the ovary in a 23-year-old female. This tumor showed unusual extensive Paneth cells metaplasia with classical signet ring cell morphology.

## INTRODUCTION

Krukenberg tumor accounts for 1–2% of all ovarian tumors and is characterized by the presence of metastatic adenocarcinoma cells in both ovaries. The primary source of this adenocarcinoma could be the stomach, pancreas, appendix or other gastrointestinal sites. It is an example of a selective tumor spread most commonly in the stomach ovarian axis.

Krukenberg tumor usually shows a solid morphology with numerous signet ring mucin secreting cells which are embedded in a fibrous stroma. To the best of our knowledge, Paneth cell metaplasia (PCM) has not been described in Krukenberg tumors before, except in metastatic appendiceal goblet cell

tumor [1], and for this reason our patient could represent a unique example of such an association.

## CASE REPORT

A 23-year-old female patient was referred to our institution because of a history of vague abdominal pain associated with significant distension. She reported no other symptoms. Clinical examination revealed a large lower abdominal and right sided mass which was thought to be arising from the right adnexa. Full laboratory work-up showed no significant changes. Abdominal CT scan showed a large, right sided and heterogeneous adnexal mass (Fig. 1).

Received: September 30, 2018. Accepted: November 20, 2018

Published by Oxford University Press and JSCR Publishing Ltd. All rights reserved. © The Author(s) 2018.

This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (<http://creativecommons.org/licenses/by-nc/4.0/>), which permits non-commercial re-use, distribution, and reproduction in any medium, provided the original work is properly cited. For commercial re-use, please contact [journals.permissions@oup.com](mailto:journals.permissions@oup.com)

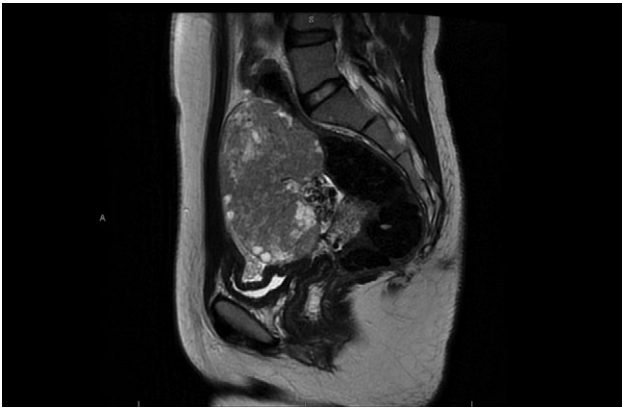


Figure 1: CT scan showing large right sided heterogeneous lesion 10 × 10 × 7 cm<sup>3</sup>.

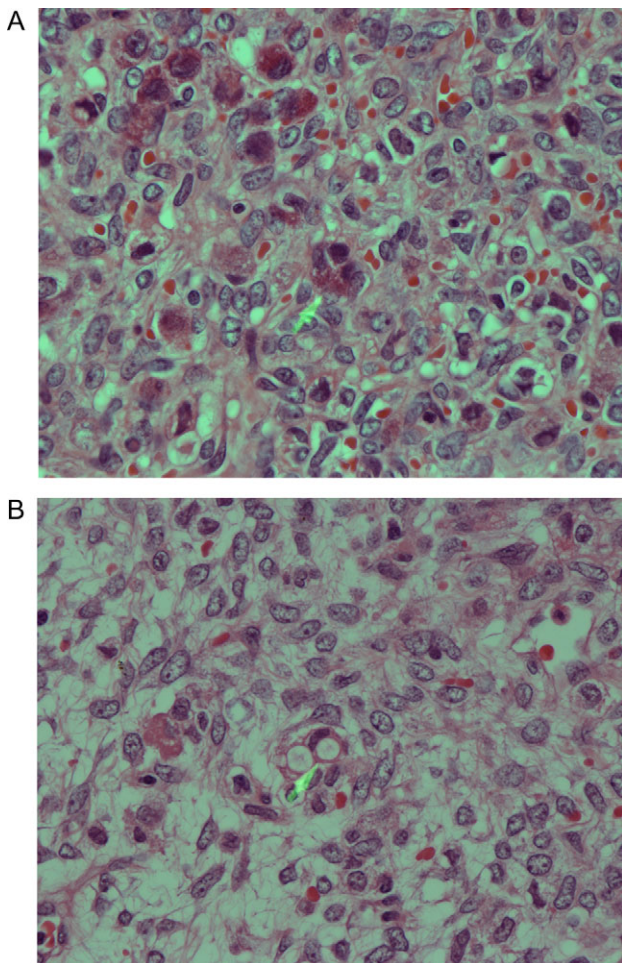


Figure 2: (A) Poorly differentiated carcinoma showing a mixture of vacuolated signet ring-cell malignant cells, some of which with eosinophilic cytoplasmic granules consistent with Paneth cells differentiation (PCD). H/E stain ×400. (B) Metastatic adenocarcinoma with signet ring cells morphology. Note also the presence of malignant cells containing orangophilic granules indicating PCD.

The mass was later on excised and sent for histopathological assessment.

The specimen received consisted of a 10 cm maximum diameter right ovarian mass with adherent fallopian tube. On

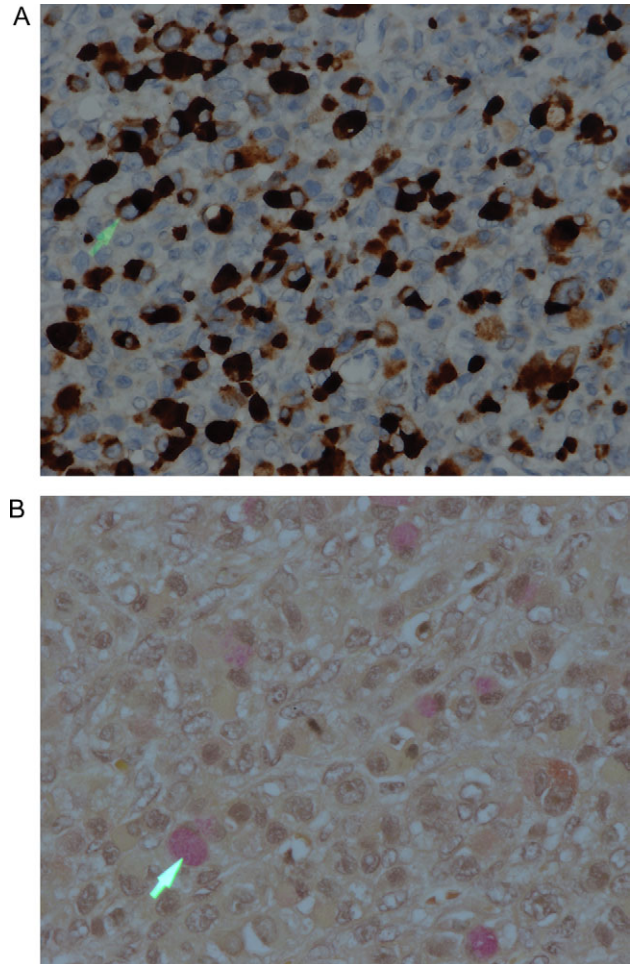


Figure 3: (A) Tumor cells showing strong positive cytoplasmic staining with the EMA immunohistochemistry. IHC stain ×200. (B) Tumor cells showing intracytoplasmic mucin, which is highlighted by the mucicarmine special stain (arrowhead). Mucicarmine stain ×400.

section the mass showed a predominantly solid cut surface with few small scattered cystic areas. Histopathological sections showed a poorly differentiated carcinoma consisting of a mixture of signet ring malignant cells with other cells showing granular and or eosinophilic cytoplasm consistent with PCM (Fig. 2A and B).

Immunohistochemical stains showed that the tumor cells were strongly positive with cytokeratin cocktail, EMA and CK7. Special stain for mucin (mucicarmine) was also positive (Fig. 3A and B).

Immunohistochemical stains for lymphoma and mesenchymal malignancies were negative. A final diagnosis of metastatic adenocarcinoma with features favouring gastric origin was rendered.

A subsequent gastroscopy and biopsies were performed, and they showed a gastric adenocarcinoma of the diffuse signet ring cell subtype.

## DISCUSSION

PCM is best defined as the presence of Paneth cells outside their normal site of origin which is the base of the Lieberkühn crypts of the small intestine. In the colon, a Paneth cells normally be found in the cecum and proximal parts of the colon. Their

presence outside these sites in the gastrointestinal tract is regarded as pathologic and indicates metaplasia [2]. PCM is described in a variety of pathologic conditions including some neoplasms [3–6] and cases of chronic inflammatory bowel disease [7, 8].

Some authorities have found that PCM affects the prognosis of certain pre-neoplastic conditions and tumors. Furthermore, it was found by some investigators that the presence of PCM in distal colorectal adenoma is inversely associated with synchronous severely dysplastic adenoma or adenocarcinoma [9]. Chen *et al.* [10] also found that the presence of PCM in cases of Barrett's esophagitis is associated with a higher risk for disease progression.

However, Kinoshita *et al.* [11] suggested that the presence of PCM may have anti-neoplastic and growth inhibiting effects and is hence associated with low risk of malignancy.

In general, the literature is still lacking regarding the significance and the prognosis of PCM. Needless to say, the presence of PCM in Krukenberg ovarian tumors is not clear and the impact of those cells on the prognosis of this advanced cancer needs further studies.

## CONFLICT OF INTEREST STATEMENT

None declared.

## REFERENCES

- Roy P. Goblet cell carcinoid tumors of the appendix: an overview. *World J Gastrointest Oncol* 2010;2:251.
- Stappenbeck T. Paneth cell development, differentiation, and function: new molecular cues. *Gastroenterology* 2009; 137:30–3.
- Mora JI. Paneth cell carcinoma of the ampulla of Vater. PubMed, NCBI [Internet]. 2004 [cited 27 June 2018]. <https://www.ncbi.nlm.nih.gov/pubmed/15270608>
- Paneth cell-rich carcinoma of the stomach: a case report. *J Pathol Transl Med* 1998; <https://www.jpatholm.org/journal/view.php?number=1746> [Internet]. [cited 27 June 2018].
- Komi N, Tamura T, Miyoshi Y, Hino M, Yada S, Kawahara H, et al. Histochemical and immunohistochemical studies on development of biliary carcinoma in forty-seven patients with choledochal cyst—special reference to intestinal metaplasia in the biliary duct. *Jpn J Surg* 1985;15:273–8.
- Park K, Chen Z, MacDonald T, Siddiqui J, Ye H, Erbersdobler A, et al. Prostate cancer with Paneth cell-like neuroendocrine differentiation has recognizable histomorphology and harbors AURKA gene amplification. *Hum Pathol* 2014;45: 2136–43.
- Elphick D. Paneth cells: their role in innate immunity and inflammatory disease. *Gut* 2005;54:1802–9.
- Simmonds N, Furman M, Karanika E, Phillips A, Bates A. Paneth cell metaplasia in newly diagnosed inflammatory bowel disease in children. *BMC Gastroenterol* 2014;14:93.
- Mahon M, Xu J, Yi X, Liu X, Gao N, Zhang L. Paneth cell in adenomas of the distal colorectum is inversely associated with synchronous advanced adenoma and carcinoma. *Sci Rep* 2016;6:26129.
- Chen W, Frankel W, Cronley K, Yu L, Zhou X, Yearsley M. Significance of Paneth cell metaplasia in Barrett esophagus. *Am J Clin Pathol* 2015;143:665–71.
- Kinoshita H, Hayakawa Y, Koike K. Metaplasia in the stomach—precursor of gastric cancer? *Int J Mol Sci* 2017; 18:2063.