

# Synchronous carcinoma breast with renal cell carcinoma: A case report

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## ABSTRACT

**Introduction:** Incidence of double malignancy (synchronous or metachronous) has increased significantly. Renal cell cancer or Breast Cancer as second primaries are reported in the literature, but synchronous carcinoma breast with renal cell carcinoma is extremely rare. We are reporting a case of carcinoma breast with synchronous renal cell carcinoma. **Case Report:** A 38-year-old female was investigated for a lump in right breast. Biopsy from breast lump was reported as infiltrating duct carcinoma. Metastatic workup revealed a large mass at lower pole of right kidney. Right MRM and right radical nephrectomy were done for synchronous carcinoma breast and renal cell

carcinoma. **Conclusion:** Genetic predisposition, tobacco, HPV, improvement in survival and long term surveillance after first primary cancer treatment, history of radiation or chemotherapy can be associated with second cancer but exact cause of second cancer is still unknown. Larger studies and research is warranted as the risk of double malignancy has been increased substantially. The plausible mechanisms behind the synchronous cancers have to be investigated.

**Keywords:** Carcinoma breast, Rare, Renal cell carcinoma, Synchronous

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## INTRODUCTION

With to availability of effective cancer therapy and improvement in survival, the incidence of double malignancy has risen [1]. Double malignancy now comprises of the sixth most common cancers and it makes 16% of all incident cancers [2]. Renal cancer is known to be associated with multiple tumors involving bladder, prostate, rectum, lung, non-Hodgkin's lymphoma,

stomach and melanoma etc. [3]. Breast cancer is also associated with increased risk of second tumor involving colon, vulva, lung, larynx, liver, uterus and thyroid [4]. All these associations can be metachronous (develop consequently) or synchronous (tumors coexist at the time of diagnosis). Synchronous breast and renal cell carcinoma (RCC) can occur rarely [3]. We hereby report a case of carcinoma breast with synchronous renal cell carcinoma.

## CASE REPORT

A 38-year-old premenopausal, multipara female with no significant family history presented with a lump in right breast since eight months duration. Sonomammography of ipsilateral breast showed a fibroadenoma of 23x18 mm at 9 o'clock position, surrounding fibroadenosis and reactive axillary lymph nodes. Excision Biopsy was done by a surgeon at a peripheral hospital and it was reported as pT 4.5x3 cm, infiltrating duct carcinoma with predominant ductal carcinoma in situ (cribriform pattern).

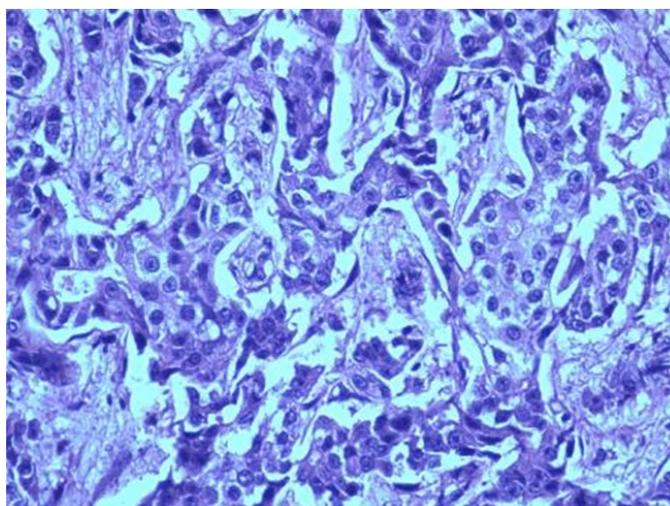
The patient was referred to our hospital for further management. Computed tomography scan of abdomen for metastatic workup showed a 45x42 mm mass in mid-lower pole of right kidney with small exophytic component and associated perilesional fat density suspected to be malignant (Figure 1). Bone Scan, CT chest and other routine investigations were within normal limits. Right modified radical mastectomy and right radical nephrectomy were done. Histopathology of breast specimen was reported as lumpectomy cavity of size 6x3.5x3 cm, no residual primary tumor, 4/26 lymph nodes positive for metastatic duct carcinoma (Figure 2), immunohistochemistry profile of axillary lymph node showed positivity for GCDFP and negativity for ER/ PR/ Her 2 neu/ CD 10/ PAX 8 and Vimentin. Gross size of the right kidney specimen was 12.5x7x3 cm with ureter 4 cm in length.

On cutting a well circumscribed tumor identified in middle region of kidney measuring 4.5x3.2x3 cm, with variegated cut surface, and Gerota's fascia adherent to the tumor at one focus measuring 3x1.2 cm. Histopathology of renal mass was reported a well-circumscribed tumor, grade I, clear cell renal cell carcinoma, Gerota's fascia uninvolved, vascular and ureteric cut end unremarkable, renal sinus and perinephric fat free of tumor (Figure 3).

Immunohistochemistry of renal mass was positive for CD 10, PAX8, Vimentin and negative for GCFDP. She was finally diagnosed as a case of synchronous carcinoma right breast (pT2pN2cM0) along with conventional renal cell carcinoma right kidney (pT1bpN0cM0, Grade I). She received adjuvant chemotherapy (FEC x 4 followed by Taxol x 4 cycles). She further received external beam radiotherapy to ipsilateral chest wall and SCF.



Figure 1: Computed tomography scan of abdomen showing a mass in mid-lower pole of right kidney with small exophytic component and associated perilesional fat density.



Figures 2: Malignant ductal cells diagnostic of Infiltrating duct carcinoma (H&E stain, x400).

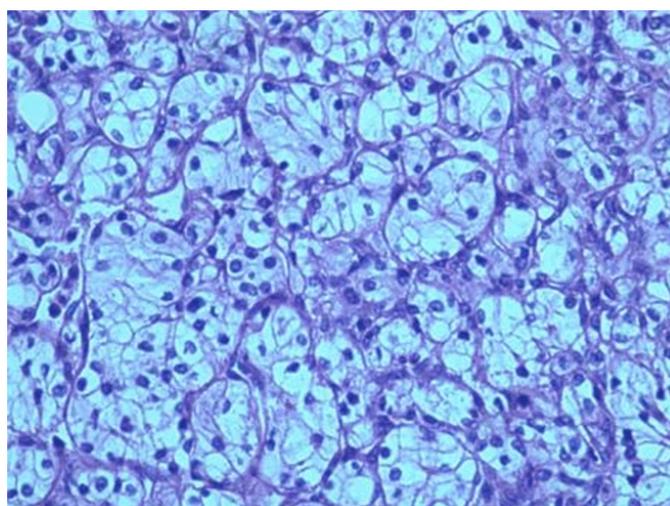


Figure 3: Slide showing clear cells diagnostic of clear cell renal cell carcinoma (H&E stain, x400).

## DISCUSSION

Renal cell carcinoma and breast cancer are associated with increased risk of double malignancy [3, 4]. Sato et al. [3] found multiple primary malignant tumors in up to 12% of patients with renal cell cancer. Rabbani et al. [5] also reported 30–40% incidence of other primary malignancies on autopsy series in patients with renal cell carcinoma. Synchronous breast and renal carcinomas are rarely reported [3].

The exact mechanism of double malignancy is not well understood. The predisposing risk factors for dual malignancy include tobacco and alcohol, genetic predisposition (Li Fraumeni and Beckwith–Wiedemann syndrome, Cowden syndrome and BRCA mutations), improved survival, history of prior radiation or chemotherapy, environmental risk factors, old age [1]. PTEN gene is associated with Cowden syndrome which has high risk of developing tumors of the thyroid, breast, kidney, endometrium [6].

Detection bias may be another important factor as metastatic workup or long-term surveillance for first malignancy increases chances of detection of second primary tumor.

Hormone-ER complex may be playing a role in development of RCC. Di Silvero et al. [7] reported a series of 17 cases of renal cell carcinoma associated with primary tumors of steroid hormone target tissue such as breast, ovary and endometrium. Liu et al. [8] in a study of differentially expressed genes (DEGs), reported that ER target genes were closely associated with renal cell carcinoma development. Tanaka et al. [9] investigated that exposing the Syrian hamsters to estrogens results in the development of kidney cancer and aneuploidy in renal cells. Henriksson et al. [10] found no difference in survival for metastatic renal cell carcinoma patients treated with immunotherapy (interleukin 2/interferon alpha) and tamoxifen arm versus tamoxifen alone arm in a multicentric randomized controlled trial. These results suggest that the role of endocrine manipulation for renal cell cancer needs to be investigated more. Epidemiological studies of survivors of atomic bomb irradiation links the radiation as potential carcinogenic but the proportion of second cancers that attributes to radiotherapy is still unknown. Radiotherapy techniques have changed significantly in last two decades. These rapidly evolving techniques (intensity modulated radiotherapy (IMRT), volumetric arc therapy (VMAT) tomotherapy, cyberknife, stereotactic radiotherapy, proton therapy) have different acute and late toxicity profiles compare to conventional radiotherapy. IMRT delivers more conformal radiation doses to the target tumor volume but it involves more number of radiation fields and exposes a larger volume of normal tissue to lower doses. IMRT and its implication as increased second cancer incidence is a matter of debate.

Methodological research is required on association of dual malignancy with late side-effects of chemotherapy/

radiotherapy, genetic modifiers, environmental risk factors.

## CONCLUSION

As the survival of cancer patients has improved due to early detection and availability of advance treatment options, the risk of second cancer or double malignancy has also increased significantly. Long-term surveillance and high index of suspicion of double malignancy by treating physician and reviewing pathologist is warranted for early detection of second tumor. Further research is required to quantify the association between second malignancy and genetic, environmental and treatment related factors like IMRT and newer chemotherapy.

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## Author Contributions

Soni Tej Prakash – Substantial contributions to conception and design, Acquisition of data, Analysis and interpretation of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published

Goel Sajal – Acquisition of data, Revising it critically for important intellectual content, Final approval of the version to be published

Sharma Lalit Mohan – Acquisition of data, Revising it critically for important intellectual content, Final approval of the version to be published

Gupta Anil Kumar – Acquisition of data, Revising it critically for important intellectual content, Final approval of the version to be published

Sharma Shantanu – Acquisition of data, Revising it critically for important intellectual content, Final approval of the version to be published

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## Guarantor

The corresponding author is the guarantor of submission.

## Conflict of Interest

Authors declare no conflict of interest.

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