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# Partial Nephrectomy as Treatment of an Atypical Metastasis from Prostate Cancer—A Case Report and Review of Literature

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#### **Abstract**

Background: Prostate cancer is the second most common type of cancer in man and the second in cancer-specific deaths in this population in the world. Most of the causes of death related to prostate cancer are due to its distant metastases, with the most common sites being: skeleton, distant lymph nodes, liver and lung. Renal metastasis is rare, and studies suggest infiltration due to arterial microembolization of the tumor. A key point in this scenario is the clinical suspicion of differential diagnoses, to offer the patient an effective therapy in such a specific case. Aim: To report a case of a patient with prostate cancer undergoing partial nephrectomy whose histopathological report revealed a metastatic lesion of that primary site. Case Presentation: 74 years old man, referred in May 2015 due to high PSA level and lumbago. PSA 323.11 ng/dl, rectal examination cT3a; biopsy was performed and histopathological study reported bilateral prostate adenocarcinoma, Gleason's score 8 (4 + 4). Patient's staging showed multiple secondary implants on skeletal scintigraphy. Tomography revealed solid exophytic lesion in the lower pole of the right kidney (4.7 × 3.6 cm); prostate without cleavage planes with seminal vesicles and pelvic node enlargement. Hormone therapy was initiated, PSA levels dropped to 9.51 ng/dl and total testosterone < 50 ng/dl. Partial nephrectomy was planned, initially by laparoscopy, but converted to laparotomy in December 2015. Procedure lasting 3 hours, minimal blood loss, no perioperative complications, discharged on the 3rd postoperative day. Histopathological report described undifferentiated malignant neoplasm, requiring immunohistochemistry that confirmed prostate adenocarcinoma. Patient remains hormone therapy, with no progression of the disease so far. Conclusion: Given the rarity of these cases, it is not possible to presume that nephrectomy enhances the survival rates. However, we observed that partial nephrectomy was a good choice for our patient, being the first case described in the literature. More reports should be available and studies with higher levels of evidence should be conducted to assist us in patient orientation and decision making.

## **Keywords**

Advanced Prostate Cancer, Kidney Metastasis, Partial Nephrectomy

#### 1. Introduction

Prostate cancer (PC) still remains the most frequent oncologic disease in the urogenital tract, the second most frequently diagnosed cancer among men and the fifth leading cause of cancer death worldwide [1].

Screening may increase chances of disease detection in early stages, however, despite all debates about this matter, PC is often discovered in advanced stages in about to 17% of patients, in which the risk of cancer-specific mortality is increased [2].

The common routes of metastatic spread from the prostate are through veins from prostate to the paravertebral or Batson's plexus, via lymphatic system, arterial, and by direct extension. Metastatic spread is usually associated with skeletal, pulmonary or hepatic masses and more locally invades the seminal vesicles, the bladder, the rectum and regional lymph nodes [3].

Literature has shown diverse sites of possible metastases, these atypical metastatic locations being the ocular region, brain, respiratory tract, testes, mammary and parotid glands, the skin and the lymphatic system [4]. Some studies reported that 15% of men with PC might be affected by atypical metastases at diagnosis [5] [6].

Metastasis to the kidney is infrequent and its management is controversial from diagnosis to treatment, as a few publications are available.

The aim of this paper is to report a case of a patient with PC undergoing partial nephrectomy whose histopathological report revealed a metastatic lesion of that primary site. To the best of our knowledge it is the first case described in literature.

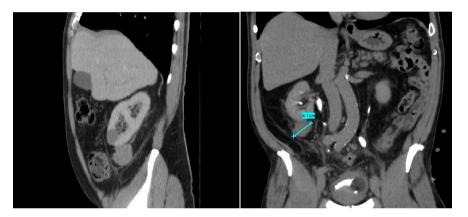
#### 2. Case Presentation

A 74-year-old male with a history of low back pain went to our hospital for clinical solution. In May 2015, present increased serum prostate specific antigen (PSA) of 323.11 ng/dl; cT3a on rectal digital examination. A prostate biopsy was performed and histopathological report was a bilateral Gleason 8 (4 + 4) adenocarcinoma. Bone scan and computed tomography (CT) was obtained. Multiple secondary bone implants was identified in bone scan. CT revealed: solid exo-

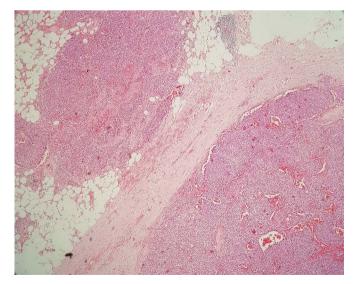
phytic lesion in the lower pole of the right kidney,  $4.1 \times 3.6$  cm (**Figure 1**), with soft tissue density, impregnated by contrast; prostate had no cleavage planes to the seminal vesicles, no retroperitoneal lymph node enlargement, but lymphadenopathy at obturator nodes bilaterally and iliac nodes on the right. Complete hormonal blockade was initiated. The first PSA after hormonal deprivation was 9.51 ng/dl and total testosterone < 50 ng/dl.

In view of all these results, partial nephrectomy was proposed and the surgery was initiated by laparoscopic approach, but converted to laparotomy access. Procedure lasted 3 hours with minimal blood loss, discharged on 3rd postoperative day.

Patient returned after 15 days to postoperative review with no pain or complications in surgery site. Histopathological report described undifferentiated malignant neoplasm, immunohistochemistry with PSA markers was found in cells, corresponding to prostate adenocarcinoma (Figure 2).



**Figure 1.** CT presenting solid exophytic lesion in the lower pole of the right kidney,  $4.1 \times 3.6$  cm with soft tissue density.



**Figure 2.** Histopathological images show undifferentiated malignant neoplasm that was identified as prostate adenocarcinoma in immunohistochemistry by positive PSA.

After 10 months of partial nephrectomy, the patient had disseminated bone pain, with difficulty to walk. This evolution of pain was in the presence of hormonal blockade, with progressive increase of PSA. New bone scan was performed, which identified an increase in disseminated osteogenic activity, mainly in the left femur, where the patient experienced more pain. X-ray of the left femur confirmed pathological fracture, which was followed up by the orthopedic team.

Chemotherapy for disease control was started. Patient still under treatment, but now with significant renal impairment and general condition, hospitalized for improvement of the health.

### 3. Discussion

Despite the great discussion within the scientific community, PC screening through PSA has led to intriguing scenarios. A Cochrane review suggests that PSA screening is associated with an increased diagnosis rate, the detection of more localized and less advanced disease (T3-4, N1, M1), but overall survival and cancer-specific survival had no benefit observed [7].

From overdiagnosis to overtreatment, the debate has never ignored the fact that patient's participation is substantial, and that all information should serve as substrate for patient's and team's decision making [8] [9].

Scosyrev and colleagues carried out a study on PC identified in elderly population; they observed frequency of M1 PC at presentation was 3% for the group aged < 75 years, 5% for the group ages 75 to 79 years, 8% for the group ages 80 to 84 years, 13% for the group ages 85 to 89 years, and 17% for the group aged 90 years. In their analyses, data indicates that, compared with younger patients (aged < 75 years), older patients were more likely to present with very advanced disease and contributed almost half (48%) of all M1 cases [2].

Gandaglia *et al.* analysed data from US Healthcare Cost and Utilization Project Nationwide Inpatient Sample between 1998 and 2010. About 74,826 patients with metastatic PC were identified. The most common metastatic sites were bone (84%), distant lymph nodes (10.6%), liver (10.2%), and thorax (9.1%). Kidney/adrenal metastasis was observed in 757 individuals. The distribution of metastatic sites in patients with a single site involved was 0.3% for those with kidney/adrenal involvement; and in those with two or more sites it becomes 4.2%.

Wu *et al.* reviewed their intra-institutional cases of metastasis to the kidney; according to the survey, 43 cases were identified; only one case had a correlation with PC [10].

As usually, challenging cases are brought to discussion rounds at scientific meetings of our service. Some differential diagnosis was discussed; a second primary tumor was possible, a tumor-to-tumor metastasis was carried out and the prostate to kidney metastasis was also considered.

In ages before PSA or CT, one journal published two articles reporting tumor-to-tumor metasis. In one paper Rabson et al. reported two cases; first a

72-year-old man which harbored a clinically silent PC with spread to an equally silent clear-cell carcinoma of the right kidney; the second one presented a 56-year-old man who also manifested spread of PC to various organs, including the right kidney that contained a renal cell carcinoma (RCC) [11]. In the other paper, Schneider described a 60-year-old man who suffered low urinary tract symptoms for six months progressed to haematuria; an enlarged and fixed prostate was observed on physical examination, cystotomy was performed and biopsy revealed adenocarcinoma; patient died 2 weeks later. Necropsy identified a left kidney mass, which histopathological study demonstrated RCC and PC in the same tissue [12].

In 1996, Inatomi *et al.* published a similar case which indicates in 1985, either before PSA screening; in this case, a 62-year-old-man was diagnosed with metastatic PC, received hormonal blockade; three years later found an image on CT suggesting RCC. It was supposed to be a second primary tumor; after follow up, death came in 1990. Necropsy was performed, demonstrated metastases in regional and para-aortic lymph nodes, lungs, left adrenal gland, pleurae, ribs, vertebrae and pelvic bone; left kidney was taken to histopathology study, observing PC in the same tissue as RCC [13].

More recently two other papers have been published about tumor-to-tumor metastasis; Horn *et al.* described a case of a kidney resected for a primary neoplasm (oncocytoma) that harbored metastases from a clinically undiagnosed PC in a 92-year-old man; Desai *et al.* reported a 72-year-old man who found a kidney mass during the staging of his newly discovered PC; after nephrectomy, histopathological found typical RCC along with extensive involvement by PC [14].

Due to the small incidence of prostatic tumor leading to renal metastasis, differential diagnosis must be thought like a second primary RCC and benign lesions such as angiomyolipoma or oncocytoma. Some groups have done biopsy before treating those masses. Kutcher *et al.* described the first pre-mortem diagnosis of prostatic metastasis to kidney; since then cases like these have been scarcely reported in literature [13]. Ibinaiye *et al.* described the second case of renal metastasis from prostatic adenocarcinoma diagnosed antemortem by percutaneous fine needle aspiration in literature [14]. Chan *et al.* reported a biochemical recurrence of an androgen-refractory metastatic PC, which staging found an enormous mass in the right kidney; biopsy was taken before hard decisions were made [15]. Chen *et al.* described a diffuse renal and retroperitoneal mass observed in CT, in which ultrasonography-guided biopsy confirmed the prostate metastatic kidney disease [16].

One of the hardest situations described came from Alaini *et al.*, who identified a prostate metastasis into a renal allograft. A 76-year-old man with a history of end-stage related to IgA nephropathy, was transplanted, and found PC two years after receiving his new organ, submitted to radiotherapy as first line treatment. Three years later, he was admitted into the hospital with pulmonary conditions and renal impairment. The tomographic findings around the allograft kidney

were essential in decision making and a biopsy was performed. It identified amyloid deposition and significant interstitial fibrosis with tubular atrophy. There was no evidence of rejection. However, up to 50% of the renal parenchyma was involved by adenocarcinoma. Diagnosis of metastatic prostate carcinoma with infiltration of the allograft, retroperitoneal, and pelvic nodes were made. Facing this diagnosis, patient opted for palliative care. Dialysis was discontinued, and he died at home [17].

Gunlusoy *et al.* and Denti *et al.* respectively reported postmortem findings of bilateral and unilateral renal metastases from PC which was initially misdiagnosed as RCC [18] [19]. Sakata *et al.* reported a 67-year-old man found to have a renal mass on CT incidentally. He had had total androgen blockade and chemotherapy for treatment of PC discovered 33 months before the incidental finding. Based on clinical features and radiologic results, it was thought to have a second malignant tumor; left nephrectomy was performed. The pathological finding of this case was renal metastasis from prostatic adenocarcinoma. Patient died 18 months post-nephrectomy [20].

The most usual clinical presentation reported was lumbar pain, but some cases are shown in different manners. Gallego *et al.* published a patient presented with acute renal failure secondary to metastasis from PC to the kidneys [21]. Gunlusoy *et al.*, reported a case which symptoms were left flank pain, hematuria and vomiting for one month [18]. Ibinaiye *et al.* presented an infected renal cyst as a result of metastatic prostatic adenocarcinoma with widespread bone metastases [14].

As shown, the incidence of renal metastasis from adenocarcinoma of the prostate is low, demonstrated in the literature only by some case reports. It is a major diagnostic challenge, due to the risk of a second malignant neoplasm. It is necessary to have the clinical suspicion of the possibility of being a metastasis, even if it is atypical.

In view of this, this article becomes necessary for the knowledge of one of the forms of presentation of the disease, aiding specialists in new future cases.

The summary of literature is found in Table 1.

#### 4. Conclusions

The data about PC is far from exhausted. Our current knowledge has not yet reached the limits of what is necessary to understand the singularities of this disease that has pushed us beyond our barriers.

Once a renal lesion is found in the context of PC, there is a possibility of facing a metastatic lesion. Even with small chances of being prostatic metastasis to the kidney, a biopsy should be considered, so the next steps will be well defined. A few options for the treatment of metastatic renal masses from primary prostate tumor are available such as nephrectomy, chemotherapy and radiotherapy, and renal artery embolization.

Given the rarity of these cases, it is not possible to presume that nephrectomy

Table 1. Summary of literature.

Year	Author	PSA	LNM	Bone Mx	Histopathological Assessment	Nephrectomy	Disclosure
1954	Rabson [11]	-	-	-	Necropsy	No	-
1955	Schneider [12]	-	Yes	Yes	Necropsy	No	-
1986	Kutcher [13]	-	Yes	No	Ultrasound-guided biopsy	No	Death in 3 months
1996	Inatomi [22]	-	No	Yes	Necropsy	No	Death in 24 months
1996	Gallego [21]	800	Yes	Yes	Laparotomic biopsy	No	Unknown
1998	Denti [19]	82.3	Yes	No	Nephrectomy	Yes	Death in 5 months
2004	Gunlusoi [18]	150	Yes	Yes	Laparotomic biopsy	No	Death in 12 months
2011	Sakata [20]	21.2	Yes	Yes	Nephrectomy	Yes	Death in 18 months
2012	Ibinaiye [14]	150	-	Yes	Ultrasound-guided biopsy	-	Patient quit treatment
2012	Horn [23]	500	-	-	Nephrectomy	Yes	-
2013	Desai [24]	19	Yes	Yes	Nephrectomy	Yes	Death in 1 month
2013	Chan [15]	200	Yes	No	CT guided biopsy	No	Death
2016	Chen [16]	36.8	Yes	No	Ultrasound-guided biopsy	No	Patient quit treatment
2017	Alaini [17]	10	Yes	No	Ultrasound-guided biopsy	No	Patient quit treatment
2017	Our case	32,311	Yes	Yes	Partial Nephrectomy	No	Patient is on follow up

PSA—Prostatic-Specific-Antigen; LNM—Lymph Nodes Metastasis; Bone Mx—Bone Metastasis.

enhances the survival rates. However, we observed that partial nephrectomy was a good choice for our patient, being the first case described in the literature.

More reports should be available and studies with higher levels of evidence should be conducted to assist us in patient orientation and decision making.

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# **List of Abbreviations**

PSA: Prostatic-Specific-Antigen CT: Computerized Tomography

PC: Prostate Cancer

RCC: Renal Cell Carcinoma