Kaposi sarcoma appearing 20 year post renal transplant
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Abstract: A kidney- transplanted Saudi patient presented with a skin rash for which he was treated as fungal infection. The patient developed these lesions 20 years after transplantation. The clinical picture was that of Kaposi’s sarcoma which was confirmed by histopathology. The patient developed these lesions after he was shifted to prednisolone and mycophenolate mofetil after four years of prednisolone, azathioprine and cyclosporine immunosuppressive therapy.

Key words: Kaposi’s sarcoma, mycophenolate mofetil, cyclosporine.

Sarcoma is a cancer that develops in connective tissues such as cartilage, bone, muscle, blood vessel or fibrous tissue. In 1872 Moritz Kaposi first described “Idiopathisches multiple pigmentosarkom der Haut” which has become known as Kaposi sarcoma (KS). It was postulated that Kaposi sarcoma, especially in patients with advanced skin tumors are reactive proliferations rather than true malignancies with metastatic dissemination.

KS has several types: Classic KS which was considered a rare disease that mostly affects elderly men of Mediterranean or Jewish heritage, African KS (Endemic) including the fatal infantile KS, Transplant-Related KS (Acquired) and AIDS-Related (Epidemic) Kaposi sarcoma.

In the last 20 years most of KS cases were associated with HIV infection

It was found that KS risk is high in immunosuppressed transplant recipients, and most cases occurred in the first two years after transplantation. The risk increases steadily with recipient age, and in males.

KS is 150 – 200 times more likely to develop in transplanted patients than in general population. There is strong evidence that KS develops in recipients with pre-existing infection with human herpes virus 8(HHV-8). According to Tessari et al. only combined immunosuppressive therapy with mycophenolate mofetil + cyclosporine + prednisolone were independently associated with KS.

Case report:
A 45 –years- old man was seen in post renal transplant clinic with multiple skin lesions. The lesion were diagnosed initially and treated as fungal infection (Tinea corporis). The patient had renal transplant 20 years prior to this presentation. He was on immunosuppressive therapy. Firstly, he was on prednisolone, azathioprine and cyclosporin for four years, and then shifted to prednisolone plus mycophenolate mofetil only. He was on regular follow up by the renal unit.

Clinical evaluation of the patient revealed a large violet patch (15x10cm) over the surface of the right lower half of the abdomen (see figure 1).

Figure(1) shows a large (15x10cm) violet patch on the right lower half of the abdomen.

There were also violet multiple nodules seen over the upper part left chest over the left breast (figure 2) and over the surgical incision.
Figure(2) shows multiple nodules over the left breast.

In addition, there are two violet patches (2x3cm each) seen on the lateral side of the right upper arm and another patch on the right thigh.

**Blood tests** revealed normal white blood cells count (11000), platelet count (157) but a high ESR (45 mm/hr), a low haemoglobin (7.2 gm/dl), a high blood urea (71mg/dl) and a high creatinine (2.0mg/dl) otherwise normal chemistry.

**Sputum** was negative for AFB

**Serological tests** for both HIV and VDRL were negative.

**Chest X-ray** showed multiple infiltrations with patchy opacities bilaterally in the lower parts of both lungs suggestive of metastasis. (Figure 4)

**Discussion:**

KS is a multifocal vascular tumor that occurs most commonly in patients who have immunosuppression. It is the most common malignancy encountered in renal transplant recipients. It is seen up to 400 times more frequently in transplant patients compared to normal population. The highest percentage for post renal transplant K.S is reported from Saudi Arabia. KS was found to be the most common malignancy encountered in renal transplant recipients in Iraq with a prevalence of 2.9%. Altace et al indicated that the average latency period between transplantation and development of malignancy was 6.5 months for KS. However, Italian organ transplant recipients have an increased risk (about 100 times greater) for Kaposi sarcoma compared...
to general population especially during the first two years after transplantation. Post renal transplant KS is related more to the immunosuppressive protocol containing cyclosporine compared to azathioprine. It has been suggested that cyclosporine is a risk factor for the development of KS as it has a direct cancer promoting effect by cell-autonomous mechanisms.

On the other hand, regression of KS has been reported on replacing cyclosporine by mycophenolate mofetil. Hussein et al reported disappearance of KS after replacing cyclosporine and azathioprine by mycophenolate mofetil. However, even after the cessation of cyclosporine and azathioprine and replacing them with mycophenolate mofetil in our reported case, KS developed. This might supports earlier reports. It is not very clear whether the introduction of mycophenolate mofetil contributed to the appearance of KS in our patient or not. Some reports showed a higher (0.8%) incidence for developing KS while on mycophenolate mofetil treatment compared with only 0.01% in patients not on mycophenolate mofetil treatment.

As for the average latency period between transplantation and development of KS, some reports were as low as 6.5 months others gave up to 21 months. However it was found that most cases occurred in the first two years after transplantation. Surprisingly our patient had developed KS almost 20 years after the transplantation.

Conclusions:
It is concluded that renal transplant patients may develop KS even if shifted from cyclosporine to mycophenolate mofetil therapy. Post renal KS can occur up to twenty years after the transplant.

References: