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Complete Disappearance of Numerous Hepatic and Adrenal Metastases after One Dose of Pembrolizumab in a Patient with Metastatic Malignant Melanoma

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Abstract Although pembrolizumab is the mainstay treatment in metastatic malignant melanoma and have shown a durable response. Heterogeneous or no response has been observed in 30-40% of the cases of metastatic malignant melanoma. Metastatic lung lesions are sensitive to checkpoint inhibitor therapy whereas liver and adrenal metastases show a variable response. We are reporting a case of malignant melanoma with numerous liver and adrenal metastases showing a significant and durable response to just a single dose of pembrolizumab.

Keywords: metastatic malignant melanoma, pembrolizumab, adrenal metastasis, hepatic metastasis

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Letter

Respected Editor,

Pembrolizumab is one of the immune checkpoint inhibitors (ICI) and is a monoclonal antibody against programmed cell death protein -1 (PD-1). It was approved by FDA in 2014 based on the results of phase 1b KEYNOTE -001 clinical trial [1]. Although, rate of fatal toxicities by ICI is 0.3 - 1.3% of the treated patients [2], currently, ICI especially the pembrolizumab is the mainstay treatment of stage IIIC or IV metastatic malignant melanoma due to considerably improved overall survival (OS) and disease related outcomes. Although, pembrolizumab have shown durable response malignant melanoma, studies hetrogeneous response or no response at all in about 30% - 40% of the cases [3,4]. A higher response to pembrolizumab have been shown for metastatic lesions to the lungs, whereas a variable response have been shown in case of liver and adrenal metastases in malignant melanoma [5,6,7]. Here we are reporting a complete dissappearance of numerous hepatic and adrenal metastases after a single dose of pembrolizumab in malignant melanoma patient.

This is an 87 year old man with a history of marginal zone non-hodgkin lympoma (NHL) of the thyroid, prostate cancer (status-post prostatectomy 17 years ago), remote history of cutaneous melanoma on his back and

face. Ten years later, the patient presented with vague abdominal pain. CT scan of the abdomen revealed multiple liver lesions. MRI of the abdomen suggested a metastatic disease. Given history of NHL, and elevated LDH, initially it was thought to be the recurrence of his lymphoma but the liver biopsy was consistent with metastatic melanoma (BRAF/c-kit/NRAS negative, MLH1 mutation positive, TP53 mutation positive). MRI of the brain was negative for intracranial metastasis. A PET/CT scan showed disffuse liver and adrenal metastases [Figure 1A]. Patient was stated on pembrolizumab. The patient developed thrombocytopenia (60x10³) and loss of appetitie after first cycle of pembrolizumab. As a result, cycle 2 was held. Patient continued to have thrombocytopenia (albiet improvement in overall platelet count upto $90x10^3$), and we continued to hold pembrolizumab therapy, as a CT scan 6 weeks after first dose of pembrolizumab showed partial response in liver and adrenal lesions. Lactate dehydrogenase (LDH) decreased from 1293 unit/L to 262 unit/L as well. Given partial response (PR) on CT scan, improving LDH, liver function tests (LFT's) and persistent but stable thrombocytopenia, while patient was doing clinically better, we opted for survillance. A survillance PET/CT scan done 3 months after the only cycle of pembrolizumab, showed no obvious disease activity in adrenal gland and liver [Figure 1B]. At the time of this report, the patient is one year since the only cycle of pembrolizumab and continued to have stable disease after initial significant response.

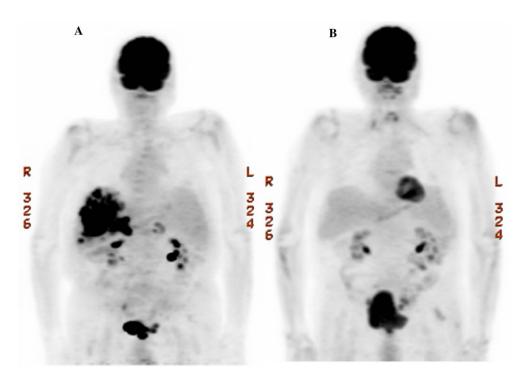


Figure 1. A= Pre-treatment PET/CT showing metastatic lesions in liver and adrenal gland, B= Disappearance of metastatic lesions after one cycle of pembrolizumab

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Conflict of Interest

None declared by both authors

Source of Funding

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Declaration of Patient Consent

We have obtained consent from the patient. The patients understand that their names and initials will not

be published, and due efforts will be made to conceal their identity.

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