## Radiation Recall Pneumonitis on FDG PET/CT Triggered by COVID-19 Vaccination

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Abstract: A 67-year-old man with metastatic lung adenocarcinoma was initially treated with whole-brain radiotherapy for intracranial metastases, followed by chemotherapy and pembrolizumab. After completing 2 years of systemic therapy, the primary right lung lesion was biopsy-proven to have residual adenocarcinoma, which was then treated with radiation (6000 cGy in 15 fractions). Follow-up serial FDG PET/CT showed radiation fibrosis. Eighteen months after radiotherapy, the patient received 2 doses of an mRNA COVID-19 vaccine. FDG PET/CT performed 4 days following his second vaccine dose showed FDG-avid multistation lymphadenopathy and radiation recall pneumonitis, likely vaccination-induced and mimicking recurrent disease. This resolved spontaneously without therapy.

Key Words: COVID-19, pneumonitis, radiation, recall, vaccination

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**FIGURE 1.** A 67-year-old man was undergoing serial FDG PET/CT scans for restaging evaluations of metastatic lung cancer. Postradiotherapy changes are seen in the right lung on FDG PET/CT performed 15 months following completion of radiotherapy, with somewhat heterogeneous mildly FDG-avid linear consolidative opacification at the treatment site (arrows).



**FIGURE 2.** FDG PET/CT performed 3 months later and 4 days after the second dose of mRNA COVID-19 vaccination shows expected postvaccination changes of FDG uptake at the injection site in the right deltoid (arrowhead) and ipsilateral FDG-avid axillary and subpectoral lymph nodes (curved arrows). However, newly FDG-avid mediastinal lymph nodes (thin arrow) were also noted, as well as new FDG-avid ground-glass and interlobular septal thickening in the right upper lobe, superior segment right lower lobe, and left upper lobes (thick arrows). The differential diagnosis for the lung parenchymal findings included recurrent disease manifesting as lymphangitis carcinomatosis; however, given the atypical pattern of spread and the well-demarcated field of involvement, a diagnosis of radiation recall triggered by COVID-19 vaccination was proposed. The patient remained asymptomatic, and further investigation with bronchoscopy was therefore deferred.

e282 | www.nuclearmed.com



**FIGURE 3.** A short-interval follow-up FDG PET/CT was performed 2 months later, while the patient was still off therapy. This showed decreasing intensity of FDG uptake in both upper lobes (arrows), improving lung parenchymal findings on CT, and resolution of FDG uptake seen previously in multistation lymph nodes and at the intramuscular injection site, confirming improving inflammatory changes. While vaccination-induced lymphadenopathy and FDG uptake in ipsilateral regional lymph nodes at the vaccine injection site in the deltoid muscle have been widely reported,<sup>1</sup> there are fewer case reports of other inflammatory responses to COVID-19 vaccination, including systemic inflammatory response syndrome<sup>2</sup> and cutaneous radiation recall.<sup>3</sup> Here, we present an illustrative case of radiation recall pneumonitis induced by COVID-19 vaccination. Radiation recall is an inflammatory response confined to the radiation port site and may be triggered by chemotherapeutic agents, including immunotherapy, but also antibiotics, antitubercular medications, nonsteroidal anti-inflammatory response can occur in any previously irradiated tissue including the lung.<sup>4</sup> Radiation recall pneumonitis in patients with lung cancer can be mistaken for progressive malignancy on imaging,<sup>6</sup> so it is important that the interpreter be aware of this entity. Steber et al<sup>7</sup> also reported a case of COVID-19 vaccination-induced radiation recall pneumonitis in a patient with lung cancer who developed rapidly progressive symptoms, and Soyfer et al<sup>3</sup> reported 2 cases of COVID-19 vaccination-induced radiation recall pneumonities in a patient with lung cancer who developed rapidly guidelines have been published to aid imaging interpreters be aware of and consider the possibility of other inflammatory responses to COVID-19 vaccination, reporting, and management,<sup>8</sup> but the guidance is less clear beyond these manifestations, and it is important that interpreters be aware of and consider the possibility of other inflammatory responses induced by the vacci