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Technical Report

Preliminary Approach to Implementing a COVID-19 Thoracic Radiation Therapy Program

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Abstract

The value of low-dose whole thoracic radiation therapy (LD-WTRT) for SARS-CoV-2 (COVID-19) pneumonia is unknown. Should ongoing clinical trials demonstrate that LD-WTRT proves effective for COVID-19 pneumonia recovery, widespread rapid implementation will be helpful globally. Our aim was to outline a pragmatic process for safe and efficient administration of LD-WTRT to patients with COVID-19 pneumonia that could be implemented successfully in a community hospital setting based on participation in the PreVent trial of LD-WTRT.

Introduction

Radiation therapy with low doses of x-rays was historically used to treat pneumonia from the 1920s to the 1940s. Owing to the advent of penicillin and increasing concerns about radiation injury, its use for nonneoplastic disease diminished sharply after World War II. The COVID-19 pandemic presents a clinical scenario where low-dose radiation therapy may again provide a benefit for patients with certain types of pneumonia.

Recent and ongoing trials are investigating the efficacy and safety of low-dose whole thoracic radiation therapy (LD-WTRT) to treat COVID-19 pneumonia. If LD-WTRT is shown to be efficacious, radiation oncology practices will have to adapt new procedures into their workflow to efficiently ensure the safety of both cancer center patients and staff. Here we aim to outline a pragmatic process with which to safely administer LD-WTRT to patients with COVID-19 pneumonia.

The PreVent trial is a multicenter phase II clinical trial randomizing patients to best supportive care, 0.35 Gy, or 1.0 Gy of LD-WTRT. In this report, we outline the feasibility of implementing an LD-WTRT program based upon participation in the PreVent clinical trial.

Methods and Materials

Data access and patient screening

We worked with inpatient nursing operations and information technology to access patient data. We designed an auto-populated Excel spreadsheet (Microsoft Corporation,
Redmond, WA) to screen current inpatients for identifying information, admission date, confirmed COVID-19 polymerase chain reaction (PCR) positive test results, and oxygen use. The radiation oncologist screened patients for eligibility at 5 AM to ensure adequate time for chart review. Criteria for eligibility are shown in Table 1.

**Chart review**

Electronic medical record review was done for hospitalized patients deemed appropriate for LD-WTRT per discussion with the primary medical team. Information including presenting symptoms, treatment to date, whether they met protocol eligibility criteria, and diagnostic imaging was reviewed. Chest x-rays were available, but we looked for prior chest computed tomography (CT) scans to estimate field size and potential virtual simulation for treatment.

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**Table 1  Inclusion and exclusion criteria per the PreVent clinical trial**

| Inclusion criteria                                                                 | Exclusion criteria                                                                                           |
|------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------|
| 1. Laboratory-confirmed diagnosis of SARS-CoV-2 pneumonia                          | 1. Currently requiring mechanical ventilation                                                              |
| 2. Currently hospitalized with COVID-19                                             | 2. Prior thoracic RT, with the exception of the following:                                                |
| 3. Age ≥50 years                                                                   |   a. Breast or postmastectomy chest wall radiation (without regional nodal irradiation) may be included at the discretion of the site primary investigator, and |
| 4. Symptomatic fever, cough, and/or dyspnea for <9 days                             |   b. thoracic skin RT (without regional nodal irradiation) is allowed                                      |
| 5. Patient or his or her legal/authorized representatives can understand and sign the study informed consent document | 3. Known hereditary syndrome with increased sensitivity to RT, including ataxia-telangiectasia, xeroderma pigmentosum, and Nijmegen breakage syndrome |
| 6. Able to be positioned on a linear-accelerator couch for RT delivery               | 4. Known prior systemic use of the following drugs: bleomycin, carmustine, methotrexate, busulfan, cyclophosphamide, or amiodarone |
| 7. AND at least 1 of the following risk factors for significant pulmonary compromise: | 5. History of or current diagnosis of pulmonary fibrosis or an alternative pulmonary condition responsible for significant lung compromise, at the discretion of the site primary investigator |
| a. Fever >102°F during index admission                                             | 6. History of lung lobectomy or pneumonectomy                                                              |
| b. Respiratory rate of ≥26/min within 24 h of screening                            | 7. Known history of pulmonary sarcoidosis, Wegener granulomatosis, systemic lupus erythematosus, rheumatoid arthritis, systemic sclerosis, polymyositis/dermatomyositis, Sjögren syndrome, mixed connective tissue disease, Churg-Strauss syndrome, Goodpasture syndrome, or ankylosing spondylitis |
| c. SpO2 ≤95% on room air within 24 h of screening                                  | 8. Symptomatic congestive heart failure within the past 6 mo including during current hospitalization |
| d. Any patient requiring 4 L/min oxygen therapy to maintain SpO2 <93% within 24 h of screening | 9. History of recent or current malignancy receiving any cytotoxic chemotherapy or immunotherapy within the past 6 mo |
| e. Ratio of partial pressure of arterial oxygen to fraction of inspired <320      | 10. History of bone marrow transplantation                                                                |
| Patients may be enrolled in this trial while concurrently enrolled in other COVID-19 clinical trials. | 11. History of any solid organ transplant (renal, cardiac, liver, lung) requiring immunosuppressive therapy |
|                                                                                   | 12. Women who are pregnant or breast-feeding                                                              |
|                                                                                   | 13. Inability to undergo RT because of any other medical or cognitive issues                              |

**Abbreviations:** COVID-19 = coronavirus disease 2019; RT = radiation therapy; SARS-CoV-2 = severe acute respiratory syndrome corona virus 2.

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**Communication and evaluation before enrollment**

For potentially eligible patients, we contacted the hospitalist to review eligibility for consideration of LD-WTRT based upon clinical need and safety of transportation. If approved, radiation oncology consultation and screening for interest as well as eligibility for LD-WTRT was done via telehealth. The patient’s nurse logged into teleconference software with a hospital-linked smart tablet, donned personal protective equipment (PPE), and brought the tablet to the patient for virtual consultation.

A focused history was taken to assess duration of pulmonary symptoms or fever (<9 days per protocol), ability to lie still in the supine or prone position for the time necessary for positioning and radiation treatment, and patient interest in the protocol. The historical rationale, the trial, and reason
for offering LD-WTRT were reviewed in a 30 to 45 minute evaluation without physical examination.

Ineligible patients were reassured that they were receiving standard therapy currently. For patients interested in the protocol, consent was obtained. The signature page was photographed and sent securely to clinical trial staff via mobile phone. Physical examination was then performed.

Coordination for transportation

To minimize risk of infection, LD-WTRT only took place at the end of the day, after the cancer center completed treatment of scheduled radiation therapy patients on the less active of 2 linear accelerators in the department. A member of the infection prevention team traveled the route from the patient unit to radiation oncology to ensure a safe path. Security unlocked and secured access along the transport route, through a back entrance used for patients either hospitalized or coming by ambulance. Nursing was called to coordinate transport to the radiation oncology department. If the patient required high-flow oxygen, respiratory therapy provided a non-rebreather mask to the patient before transport with medical floor nursing support if needed. The service engineer was notified to prevent unscheduled work on the linear accelerator.

Treatment planning

The patient was registered in the radiation oncology electronic medical record (ARIA, Varian Medical Systems, Palo Alto, CA) and placed on a protocol-specific care path. Per protocol, CT simulation was not permitted to minimize potential departmental contamination. Diagnostic chest CT and x-ray data from the current hospitalization were used to design estimated anterior/posterior (AP/PA); kilovoltage (kV) fields. CT scan import into the Eclipse treatment planning system permitted confirmation of dose distribution. The carina was contoured to help with pretreatment localization and set-up.

The anatomic target was the entire thorax to include the lungs bilaterally with an estimated 1.5- to 2-cm margin in all directions. In the absence of clear lung imaging, the superior and lateral borders were identified as 2 cm beyond the outer aspect of the ribs. The inferior border was identified at the T12-L1 interspace. The only method of blocking allowed was via the primary LINAC jaws; no multileaf collimator leaves were used. In the event of superior-inferior dimension collimator limitations, we favored covering the lung bases and diaphragmatic recesses over coverage of the lung apices.

Patients randomized to a protocol dose of 0.35 or 1.0 Gy had treatment prescribed to midplane along the central axis without heterogeneity corrections using 6 to 18 megavoltage (MV) photons, accounting for separation at the level of the carina as identified on orthogonal KV/KV, cone beam CT, or MV imaging. Isocenter was set at the carina for localization and to permit taking a separation at the central axis.

Treatment delivery

Any equipment in the linear accelerator vault not being directly used by the patient was covered with a plastic sheet, placed inside a cabinet, or removed from the room. The treatment team consisted of “hot” and “cool” members. Two hot therapists and the physician donned PPE and entered the treatment vault to set the patient up on the LINAC couch. The patient was positioned supine with arms in a comfortable position. Prone positioning was permitted if needed for patient safety or comfort. If a diagnostic CT was used for planning, the patient was set up in the same position as the diagnostic CT. A cool therapist remained in the control room, closed the vault door, and maintained audio and visual contact with the hot team members at all times. After the hot team members doffed their PPE and exited the treatment vault, the physician and 1 therapist converted from hot to cool in the control area with the LINAC console, while the remaining hot therapist donned clean PPE in preparation for posttreatment re-entry.

Kilovolt x-ray images were then acquired. The carina was identified and the patient was aligned to bony anatomy on the AP image. Vertical depth was identified on the lateral image and shifts were made to this point. After isocenter placement, 3 MV images, central superior, inferior right lateral, and inferior left lateral, were taken to verify all field borders (Fig. 1). If these actions could not

![Diagram](image_url)

**Figure 1** To verify all field borders, 3 megavoltage images were taken: central superior, inferior right lateral, and inferior left lateral.
be accomplished from outside the treatment vault due to limitations of the linear accelerator, a hot team member would re-enter the treatment vault to perform couch movements. Once the patient’s position was confirmed by the cool therapist and physician, the patient received treatment.

After the patient was transported back to the hospital floor, the hot therapists placed a “DO NOT ENTER” sign on the LINAC vault door. All 3 therapists and the physician then doffed their PPE following recommended procedures. Environmental services and housekeeping were contacted to request room decontamination. Security was notified to close the department.

Results

Using this approach, we were able to minimize COVID-19 exposure to the members of our health care team and other patients on treatment, and successfully deliver LD-WTRT to a patient with COVID-19 pneumonia. No staff or other patients contracted COVID-19 from this process. Treatment planning time to incorporate a diagnostic CT for virtual simulation, create a digitally reconstructed radiographs; and make monitor unit calculations was 20 minutes. Patient time in the department was <40 minutes, set-up for verification on the couch was 15 minutes, and treatment delivery was 12 seconds (for a 1 Gy dose in our patient’s case).

Discussion

Here, we outline an approach for minimizing the exposure of patients undergoing radiation treatment for cancer to COVID-19 by delivering LD-WTRT at the end of the day. An alternative approach to minimize exposure between these different patient populations undergoing radiation treatment would be to deliver LD-WTRT during the day using a separate LINAC dedicated to these protocols.

LD-WTRT for COVID-19 pneumonia is investigational and there are multiple ongoing trials to evaluate its efficacy, including the PreVent trial. Treating patients with cancer on special precautions due to infection (Clos tridioides difficile, methicillin-resistant Staphylococcus aureus, etc) is done routinely, and radiation therapists already use universal precautions when needed for such situations. Using radiation to treat the infection itself, COVID-19 pneumonia specifically, however, is novel. We found implementing precautions for treatment of COVID-19 pneumonia is feasible when coordinated with the inpatient team. If studies investigating LD-WTRT for this purpose confirm its safety and efficacy, our successful process designed for a community hospital may serve as a potential model for implementation in smaller nonacademic facilities. The general process can readily be adapted to larger hospitals including academic facilities. The process described here was implemented in December 2020, which was before the Food and Drug Administration gave emergency use authorization for any COVID-19 vaccine. Thus, no staff had been vaccinated at the time of LD-WTRT delivery. The ability for cancer center patients, treating physicians, and staff to be vaccinated against COVID-19 has made this treatment protocol safer to implement.

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