

Review began 04/03/2025 Review ended 04/08/2025 Published 04/12/2025

© Copyright 2025

Tang et al. This is an open access article distributed under the terms of the Creative Commons Attribution License CC-BY 4.0., which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

DOI: 10.7759/cureus.82138

CyberKnife Versus Four-Dimensional Computed Tomography-Guided Stereotactic Body Radiation Therapy in the Treatment of Lung Cancer: A Case Report

Lingrong Tang ¹, Lei Yao ¹, Guang Li ¹

1. Department of Radiotherapy, The First Hospital of China Medical University, Shenyang, CHN

Corresponding author: Guang Li, lg13804058616@163.com

Abstract

Stereotactic body radiation therapy (SBRT) has emerged as a critical therapeutic option for treating lung and other solid tumors. Two prominent high-precision SBRT techniques currently in use are four-dimensional computed tomography-guided linear accelerator-based SBRT (4DCT-SBRT) and CyberKnife. This case presents a patient diagnosed with two distinct pulmonary lesions, each treated separately using 4DCT-SBRT and CyberKnife. By comparing target delineation, dose distribution, lesion response, and the capability to spare normal tissues, we evaluate the specific advantages of CyberKnife for particular clinical scenarios.

Categories: Radiation Oncology, Oncology

Keywords: 4-dimensional, cyberknife, lung cancer, real-time tumor tracking, stereotactic body radiation therapy

Introduction

Precision radiotherapy is a pivotal component of modern oncology, delivering high-dose radiation accurately to tumor sites in a brief treatment period while minimizing damage to adjacent normal tissues. Stereotactic body radiation therapy (SBRT) has emerged as a critical therapeutic option for treating inoperable stage I non-small cell lung cancer and other solid tumors [1,2], offering outcomes comparable to surgery, although current evidence remains conflicting regarding optimal patient selection between SBRT and surgical intervention [3,4]. Leading modalities of SBRT include 4DCT-SBRT, CyberKnife, proton SBRT, MR-guided SBRT, and helical tomotherapy [5-9]. Among these techniques, four-dimensional computed tomography-guided linear accelerator-based SBRT (4DCT-SBRT) and CyberKnife are more frequently used in clinical practice.

4DCT-SBRT utilizes four-dimensional computed tomography (4DCT) to capture respiratory-induced tumor motion, improving treatment accuracy. Techniques like respiratory gating and internal target volume (ITV) expansion effectively manage tumor movements, making this modality ideal for a broad range of solid tumors in organs such as the lungs, liver, and pancreas [2,10-12].

CyberKnife integrates a robotically controlled linear accelerator combined with Synchrony Respiratory Tracking technology. This approach tracks fiducials placed in the tumor in real-time without additional gating or ITV expansions, allowing for smaller target volumes and enhanced protection of adjacent normal tissues [13,14].

In this case report, we analyze a patient treated with 4DCT-SBRT in 2020 and subsequently with CyberKnife in 2024, highlighting these techniques' clinical distinctions and comparative advantages.

Case Presentation

A 70-year-old male patient presented in September 2020 with a chronic cough. Computed tomography (CT) imaging identified a solid lesion in the left lower lung lobe, which was subsequently confirmed as adenocarcinoma via CT-guided biopsy. The lesion tested negative for mutations in EGFR, ALK, ROS1, BRAF, MET, and RET. Following a comprehensive clinical evaluation, the patient was diagnosed with early-stage lung cancer, classified as T1bN0M0, stage IA2, according to the American Joint Committee on Cancer (AJCC) 8th edition. Due to underlying conditions including cerebral thrombosis and coronary artery disease, the patient declined surgical intervention. Consequently, treatment using 4DCT-SBRT was initiated on an Elekta Versa HD linear accelerator.

Planning involved respiratory-gated 4DCT imaging with the ITV method. The gross tumor volume (GTV) was delineated on lung window settings (window width: 1600 HU; window level: -600 HU), and the ITV encompassed the combined trajectory of the GTV across all respiratory phases. The clinical target volume (CTV) was equivalent to the GTV without additional margins, while the planning target volume (PTV) was determined by expanding the ITV by a uniform 5 mm margin. A total dose of 50 Gy was delivered in five



fractions over two weeks, each lasting approximately 10 minutes (Figure 1).

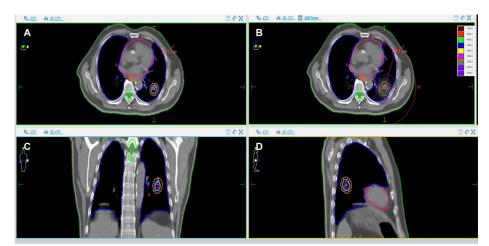


FIGURE 1: Target delineation and radiotherapy plan of 4DCT-SBRT

(A, C, D) Axial, coronal, and sagittal views showing target volume delineation. The blue line indicates ITV, and the orange line indicates PTV. (B) VMAT treatment plan.

ITV: Internal target volume; PTV: Planning target volume; VMAT: Volumetric modulated arc therapy; 4DCT-SBRT: Four-dimensional computed tomography-guided linear accelerator-based stereotactic body radiation therapy

Treatment resulted in a partial tumor response; however, mild radiation pneumonitis (Common Terminology Criteria for Adverse Events (CTCAE) version 5.0, grade 1) appeared eight months post-treatment, which gradually resolved (Figures 2A, 2B). A new enlarging lesion in the left upper lung lobe was identified during a routine follow-up at another medical facility. Enhanced CT scans demonstrated marked enhancement suggestive of malignancy (Figure 2C). Due to the lesion's proximity to the heart, biopsy was deemed high-risk and thus avoided. Whole-body imaging confirmed the absence of other malignant lymph nodes or distant lesions.



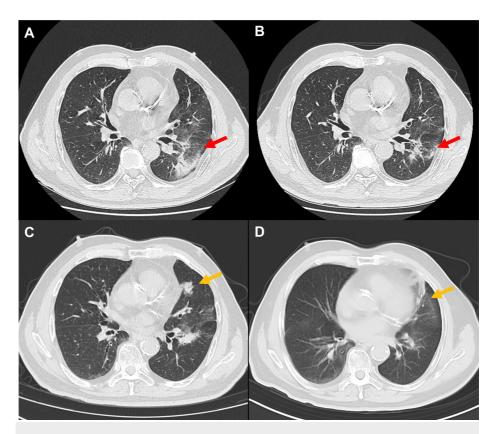


FIGURE 2: Radiographic images illustrating radiation pneumonitis and tumor response

(A) Axial CT image showing grade 1 radiation pneumonitis at eight months post-treatment (red arrow). (B) Axial CT image demonstrating gradual resolution of radiation pneumonitis at 15 months post-treatment (red arrow). (C) Axial CT scan obtained in January 2024 showed a newly detected lesion in the left upper lobe (orange arrow). (D) Axial CT image demonstrating significant tumor regression at 12 months after treatment (orange arrow).

CT: Computed tomography

In February 2024, this lesion was treated using CyberKnife SBRT on the Accuray CyberKnife M6 system. The treatment utilized Synchrony Respiratory Tracking technology with fiducial markers, eliminating the need for ITV-based planning. The simulation was performed using contrast-enhanced CT imaging with a slice thickness of 1.5 mm, acquired during end-expiratory breath-hold. The GTV was delineated on lung window settings (window width: 1600 HU; window level: -600 HU), and the PTV was defined as the GTV expanded uniformly by a 5 mm margin. The lesion was treated with a dose of 50 Gy in five fractions, which was delivered over two weeks, with each fraction lasting approximately 30 minutes (Figure 3). This treatment also achieved a partial tumor response; notably, no side effects were observed (Figure 2D).



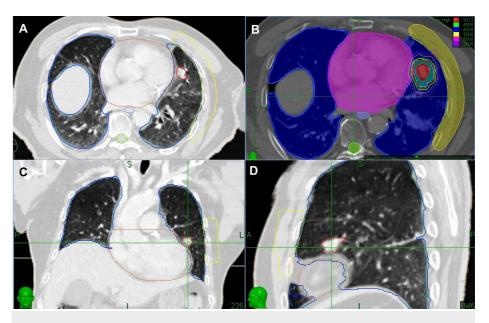


FIGURE 3: Target delineation and radiotherapy plan of CyberKnife

(A, C, D) Axial, coronal, and sagittal views showing target volume delineation. The red line indicates the GTV. (B) CyberKnife treatment plan.

GTV: Gross tumor volume

Discussion

Comparing precision and target volume

Respiratory-induced tumor movements pose challenges in lung cancer radiotherapy, typically managed either by restricting tumor mobility such as abdominal compression, deep inspiration breath-hold, active breathing control, or expanding the target volume via ITV strategies [14,15]. The former is difficult for elderly and weak patients to cooperate with, while ITV strategies may increase irradiated lung volumes, consequently elevating radiation-induced side effects.

4DCT-SBRT relies on ITV-based planning, which is derived from all phases of respiratory motion. While this approach effectively covers tumor movement, it requires larger margins around the tumor, potentially increasing the volume of normal tissue irradiated [12]. Consequently, this could elevate the risk of pneumonitis or fibrosis, particularly in cases of lesions close to critical structures, such as the heart in this case.

CyberKnife differs from 4DCT-SBRT in its approach to sparing normal tissue. The CyberKnife system employs real-time respiratory tracking technologies, such as the fiducial, X-sight lung tracking, and synchrony respiratory tracking system, allowing continuous adjustment of the radiation beam to follow tumor motion during respiration accurately [16-18]. This technology facilitates highly accurate radiotherapy without needing target volume expansions, minimizing radiation exposure to adjacent healthy tissues and critical structures [18,19].

Treatment duration and patient tolerance

Conventional SBRT was performed using 8-12 non-coplanar static beams, and treatment delivery generally took more than 20min. However, through VMAT, the delivery time was much shorter (approximately 10 minutes), ensuring better patient tolerance [20].

Studies have demonstrated that CyberKnife lung SBRT is safe and achieves progression-free survival and overall survival comparable to conventional SBRT [13,19,21,22]. The study by Diamant et al. demonstrated that CyberKnife lung SBRT achieved superior distant metastasis-free survival compared with VMAT [5]. Multiple dosimetric studies comparing CyberKnife with conventional linear accelerator-based SBRT plans have indicated that both modalities achieve clinical dosimetric goals; however, CyberKnife plans typically deliver higher monitor units (MUs) but produce lower doses at 2 cm from the target (D2cm) [23-25]. Consequently, CyberKnife treatments require longer delivery times (approximately 30 minutes per fraction) but offer superior sparing of organs at risk (OARs). Clinically, this translates into reduced radiation-induced toxicity and potentially enhanced patient quality of life.



It is important to note that CyberKnife treatment for lung SBRT necessitates continuous respiratory tracking, and circumstances such as coughing may require recalibration, thereby prolonging the treatment duration and potentially impacting patient tolerance.

Moreover, CyberKnife treatments usually require pre-treatment fiducial marker implantation to achieve precision tracking. This invasive procedure is associated with risks such as pneumothorax or bleeding, complications that do not occur with 4DCT-SBRT [17].

Clinical recommendations

CyberKnife and 4DCT-SBRT are both suitable for lung SBRT. CyberKnife is better suited to treat highly mobile tumors (e.g., lower lobe lung, liver, pancreatic tumors), leveraging its precise real-time tracking capabilities to offer superior sparing of OARs.

4DCT-SBRT is particularly suitable for pulmonary tumors with minimal movement or high fiducial marker implantation-associated risk, and for elderly or weak patients unable to stand prolonged treatment duration.

Conclusions

This case study demonstrates that both 4DCT-SBRT and CyberKnife effectively treat pulmonary tumors. CyberKnife, however, provides superior precision and improved sparing of normal tissues due to its smaller target volumes. Personalized choice of radiotherapy modality should consider patient health status, tumor characteristics, and proximity to critical structures to optimize outcomes and minimize side effects.

Additional Information

Author Contributions

All authors have reviewed the final version to be published and agreed to be accountable for all aspects of the work

Concept and design: Lingrong Tang, Guang Li

Drafting of the manuscript: Lingrong Tang

Critical review of the manuscript for important intellectual content: Lingrong Tang, Lei Yao, Guang Li

Acquisition, analysis, or interpretation of data: Lei Yao

Disclosures

Human subjects: Consent for treatment and open access publication was obtained or waived by all participants in this study. Conflicts of interest: In compliance with the ICMJE uniform disclosure form, all authors declare the following: Payment/services info: All authors have declared that no financial support was received from any organization for the submitted work. Financial relationships: All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. Other relationships: All authors have declared that there are no other relationships or activities that could appear to have influenced the submitted work.

References

- Buchberger DS, Videtic GM: Stereotactic body radiotherapy for the management of early-stage non-smallcell lung cancer: a clinical overview. JCO Oncol Pract. 2023, 19:239-49. 10.1200/OP.22.00475
- 2. Timmerman RD, Herman J, Cho LC: Emergence of stereotactic body radiation therapy and its impact on current and future clinical practice. J Clin Oncol. 2014, 32:2847-54. 10.1200/JCO.2014.55.4675
- Chang JY, Mehran RJ, Feng L, et al.: Stereotactic ablative radiotherapy for operable stage I non-small-cell lung cancer (revised STARS): long-term results of a single-arm, prospective trial with prespecified comparison to surgery. Lancet Oncol. 2021, 22:1448-57. 10.1016/S1470-2045(21)00401-0
- Chang JY, Senan S, Paul MA, et al.: Stereotactic ablative radiotherapy versus lobectomy for operable stage I non-small-cell lung cancer: a pooled analysis of two randomised trials. Lancet Oncol. 2015, 16:630-7. 10.1016/S1470-2045(15)70168-3
- Diamant A, Heng VJ, Chatterjee A, et al.: Comparing local control and distant metastasis in NSCLC patients between CyberKnife and conventional SBRT. Radiother Oncol. 2020, 144:201-8.
 10.1016/j.radonc.2020.01.017
- Figlia V, Mazzola R, Cuccia F, et al.: Hypo-fractionated stereotactic radiation therapy for lung malignancies by means of helical tomotherapy: report of feasibility by a single-center experience. Radiol Med. 2018, 123:406-14. 10.1007/s11547-018-0858-7
- Westover KD, Seco J, Adams JA, Lanuti M, Choi NC, Engelsman M, Willers H: Proton SBRT for medically inoperable stage I NSCLC. J Thorac Oncol. 2012, 7:1021-5. 10.1097/JTO.0b013e31824de0bf
- 8. Fast M, van de Schoot A, van de Lindt T, Carbaat C, van der Heide U, Sonke JJ: Tumor trailing for liver SBRT



- on the MR-Linac. Int J Radiat Oncol Biol Phys. 2019, 103:468-78. 10.1016/j.ijrobp.2018.09.011
- Brennan VS, Burleson S, Kostrzewa C, et al.: SBRT focal dose intensification using an MR-Linac adaptive planning for intermediate-risk prostate cancer: an analysis of the dosimetric impact of intra-fractional organ changes. Radiother Oncol. 2023, 179:109441. 10.1016/j.radonc.2022.109441
- Faccenda V, Panizza D, Niespolo RM, et al.: Synchronized contrast-enhanced 4DCT simulation for target volume delineation in abdominal SBRT. Cancers (Basel). 2024, 16:10.3390/cancers16234066
- Harada K, Katoh N, Suzuki R, et al.: Evaluation of the motion of lung tumors during stereotactic body radiation therapy (SBRT) with four-dimensional computed tomography (4DCT) using real-time tumortracking radiotherapy system (RTRT). Phys Med. 2016, 32:305-11. 10.1016/j.ejmp.2015.10.093
- Li Y, Ma JL, Chen X, Tang FW, Zhang XZ: 4DCT and CBCT based PTV margin in stereotactic body radiotherapy (SBRT) of non-small cell lung tumor adhered to chest wall or diaphragm. Radiat Oncol. 2016, 11:152. 10.1186/s13014-016-0724-5
- Temming S, Kocher M, Stoelben E, et al.: Risk-adapted robotic stereotactic body radiation therapy for inoperable early-stage non-small-cell lung cancer. Strahlenther Onkol. 2018, 194:91-7. 10.1007/s00066-017-1194-x
- Inoue T, Katoh N, Onimaru R, et al.: Stereotactic body radiotherapy using gated radiotherapy with real-time tumor-tracking for stage I non-small cell lung cancer. Radiat Oncol. 2013, 8:69. 10.1186/1748-717X-8-69
- Guckenberger M, Andratschke N, Alheit H, Holy R, Moustakis C, Nestle U, Sauer O: Definition of stereotactic body radiotherapy: principles and practice for the treatment of stage I non-small cell lung cancer. Strahlenther Onkol. 2014, 190:26-33. 10.1007/s00066-013-0450-y
- Collins BT, Erickson K, Reichner CA, et al.: Radical stereotactic radiosurgery with real-time tumor motion tracking in the treatment of small peripheral lung tumors. Radiat Oncol. 2007, 2:39. 10.1186/1748-717X-2-39
- Nemoto H, Saito M, Suzuki T, et al.: Evaluation of computed tomography metal artifact and CyberKnife fiducial recognition for novel size fiducial markers. J Appl Clin Med Phys. 2023, 24:e14142. 10.1002/acm2.14142
- Bibault JE, Prevost B, Dansin E, Mirabel X, Lacornerie T, Lartigau E: Image-guided robotic stereotactic radiation therapy with fiducial-free tumor tracking for lung cancer. Radiat Oncol. 2012, 7:102. 10.1186/1748-717X-7-102
- van der Voort van Zyp NC, Prévost JB, Hoogeman MS, et al.: Stereotactic radiotherapy with real-time tumor tracking for non-small cell lung cancer: clinical outcome. Radiother Oncol. 2009, 91:296-300. 10.1016/j.radonc.2009.02.011
- Verbakel WF, Senan S, Cuijpers JP, Slotman BJ, Lagerwaard FJ: Rapid delivery of stereotactic radiotherapy for peripheral lung tumors using volumetric intensity-modulated arcs. Radiother Oncol. 2009, 93:122-4. 10.1016/j.radonc.2009.05.020
- Sarihan S, Tunc SG, Irem ZK, Kahraman A, Ocakoglu G: Results of stereotactic body radiotherapy with CyberKnife-M6 for primary and metastatic lung cancer. World J Oncol. 2024, 15:711-21. 10.14740/wjon1865
- Ahn SH, Han MS, Yoon JH, Jeon SY, Kim CH, Yoo HJ, Lee JC: Treatment of stage I non-small cell lung cancer with CyberKnife, image-guided robotic stereotactic radiosurgery. Oncol Rep. 2009, 21:693-6. 10.3892/pr. 00000273
- Dai ZT, Ma L, Cao TT, Zhu L, Zhao M, Li N: Dosimetric and radiobiological comparison of treatment plan between CyberKnife and EDGE in stereotactic body radiotherapy for pancreatic cancer. Sci Rep. 2021, 11:4065. 10.1038/s41598-021-83648-5
- Yu S, Xu H, Sinclair A, Zhang X, Langner U, Mak K: Dosimetric and planning efficiency comparison for lung SBRT: CyberKnife vs VMAT vs knowledge-based VMAT. Med Dosim. 2020, 45:346-51. 10.1016/j.meddos.2020.04.004
- Song Y, Chen X, Yu X, et al.: Dosimetric comparison of multiple SBRT delivery platforms for pancreatic cancer. Eur J Med Res. 2024, 29:533. 10.1186/s40001-024-02080-3