# Is radiofrequency ablation more effective than stereotactic ablative radiotherapy in patients with early stage medically inoperable non-small cell lung cancer?

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#### **Abstract**

A best evidence topic in thoracic surgery was written according to a structured protocol. The question addressed was 'is radiofrequency ablation more effective than stereotactic ablative radiotherapy in patients with early stage medically inoperable non-small cell lung cancer?' Altogether, over 219 papers were found, of which 16 represented the best evidence to answer the clinical question. The authors, journal, date and country of publication, patient group studied, study type, relevant outcomes and results of these papers are tabulated. Radiofrequency ablation (RFA) and stereotactic ablative radiotherapy (SABR) offer a clear survival benefit compared with conventional radiotherapy in the treatment of early stage non-small cell lung cancer (NSCLC) in medically inoperable patients. Overall survival at 1 year (68.2-95% vs. 81-85.7%) and 3 years (36-87.5% vs. 42.7-56%) was similar between patients treated with RFA and SABR. However, 5-year survival was higher in SABR (47%) than RFA (20.1-27%). Local progression rates were lower in patients treated with SABR (3.5-14.5% vs. 23.7-43%). Both treatments were associated with complications. Pneumothorax (19.1-63%) was the most common complication following RFA. Fatigue (31-32.6%), pneumonitis (2.1-12.5%) and chest wall pain (3.1-12%) were common following SABR. Although tumours ≤5 cm in size can be effectively treated with RFA, results are better for tumours ≤3 cm. One study documented increased recurrence rates with larger tumours and advanced disease stage following RFA. Another study found increasing age, tumour size, previous systemic chemotherapy, previous external beam radiotherapy and emphysema increased the risk of toxicity following SABR and suggested that risk factors should be used to stratify patients. RFA can be performed in one session, whereas SABR is more effective if larger doses of radiation are given over two to three fractions. RFA is not recommended for centrally based tumours. Patients with small apical tumours, posteriorly positioned tumours, peripheral tumours and tumours close to the scapula where it may be difficult to position an active electrode are more optimally treated with SABR. Treatment for early stage inoperable NSCLC should be tailored to individual patients, and under certain circumstances, a combined approach may be beneficial.

Keywords: Radiofrequency ablation • Stereotactic radiotherapy • Stereotactic ablative radiotherapy • Non-small cell lung cancer • Survival

#### INTRODUCTION

A best evidence topic was constructed according to a structured protocol. This is fully described in the ICVTS [1].

# **THREE-PART QUESTION**

In [patients with early stage medically inoperable non-small cell lung cancer], is [radio-frequency ablation] superior to [stereotactic ablative radiotherapy] treatment?

# **CLINICAL SCENARIO**

You are at a conference hearing about the effectiveness of radiofrequency ablation (RFA) and stereotactic ablative radiotherapy (SABR) in patients with early stage medically inoperable nonsmall cell lung cancer (NSCLC). You have an 85-year old patient who has been diagnosed with Stage IA NSCLC and is not fit for surgery due to his extensive comorbidities. You decide to do a literature search.

## **SEARCH STRATEGY**

An English language literature review was performed on MEDLINE 1948 to July 2011 using the Ovid interface: [catheter ablation/OR radiofrequency ablation.mp.OR radiosurgery/OR stereotactic radiotherapy.mp. OR stereotactic ablative radiotherapy.mp. OR stereotactic lrradiation.mp. OR radiation therapy.mp. OR stereotactic Irradiation.mp. OR radiation therapy.mp] AND [non small cell lung carcinoma.mp. OR Carcinoma, Non-Small-Cell Lung]

Table 1: Best-evide	nce papers			
Author, date, journal and country, study type (level of evidence)	Patient group	Outcomes	Key results	Comments, study weaknesses
Huang <i>et al.</i> , (2011), Eur J Cardiothorac Surg, China [2]	Three hundred and twenty-nine patients with 436 lung tumours treated with RFA from 1999 to	Median progression-free period	21.6 months	RFA is a safe and well-tolerated procedure with confirmed efficacy in the treatment of malignant lung
Retrospective study	2006 Primary	Local progression  30-day mortality	78/329 (23.7%) 0.6%	tumours  Tumours >4 cm have a significantly
	(n = 237)	Overall survival:		increased risk of local progression
	Metastatic (n = 92)	1, 2 and 5 years  Complications:	68.2, 35.5, 20.1%	Treatment-related complications were counted if within 30 days after RFA treatment
	Inclusion criteria: (1) Aged 18-80 (2) Poor pulmonary function with FEV1 < 1 I, FEV1% <50%; MVV <50% and/or high cardiac risk (3) Refusal of surgery	Pneumothorax Haemoptysis Haemothorax Pneumonia Pericardial tamponade	63/329 (19.1%) 14/329 (4.2%) 10/329 (3.0%) 15/329 (4.5%) 3/329 (0.9%)	deathen
Beland et al., (2010), Radiology, USA [3]	Seventy-nine patients with 79 tumours underwent RFA for primary NSCLC from 1998 to	Residual tumour or recurrence	34/79 (43%)	RFA is a promising treatment option for primary lung cancer in non-surgical patients, however disease recurrence is
Retrospective study	2008	Recurrence pattern: Local	38%	common, occurring in 43% of patients
	Stage IA n = 35 (78%)	Intrapulmonary Nodal	18% 18%	At 2 years, local recurrence was the most common at 28%, suggesting that
	Stage IB n = 7 (16%)	Mixed Distant metastases	6% 21%	more aggressive initial RF ablation and adjuvant radiation may offer improvement in outcomes
	Stage IIIB n = 3 (7%)		23 months	Potential for understaging of disease as patients did not undergo mediastinoscopy
	Adjuvant external beam radiation n = 19 (24%)			No standardized post-treatment follow-up imaging protocol
	Concomitant brachytherapy $n = 9 (11\%)$			
	Mean follow-up 17 months			
de Baère <i>et al.</i> , (2006), Radiology, France [4]	Sixty patients with 97 treatable lung tumours underwent 74 RFA using CT guidance	Rate of incomplete local treatment per tumour at 18 months:	7%	RFA has high local success rates of complete ablation and curative treatment in inoperable primary and
Prospective study	Patients who received systematic chemotherapy during follow-up	Tumours <2 cm Tumours >2 cm	5% 13% ( <i>P</i> = 0.66)	metastatic lung tumours and is well tolerated
	period n = 22	Survival at 18 months:	, ,	Ideal follow-up imaging to determine early treatment failure remains to be
	Primary n = 9 (15%)	Overall Lung disease free	71% 34%	improved, probably including functional imaging and CT
	Metastatic n = 51 (85%)	Complications: Pneumothorax Alveolar haemorrhage	40/74 (54%) 8/74 (11%)	Small population  Twenty-two patients received systemic
		Pleural effusion: Immediately after treatment	7/74 (9%)	chemotherapy for distant metastases during follow-up, hence difficult to evaluate the effect of such therapy on the rate of incomplete local treatment
		24-48 h after treatment	45/74 (60%)	and rate of meomplete local deadlient
		FEV <sub>1</sub> (L): Before treatment After treatment	0.62-3.65 0.72-3.65 ( <i>P</i> = 0.65)	
			. 5.55)	Continue

Table 1: (Continued)				
Author, date, journal and country, study type (level of evidence)	Patient group	Outcomes	Key results	Comments, study weaknesses
		VC (L): Before treatment After treatment Post-procedure haemoptysis	0.80-8.0 0.83-7.98 ( <i>P</i> = 0.93) 7/74 (10%)	
Lencioni <i>et al.</i> , (2008), The Lancet, Italy [5] Multicentre prospective trial	One hundred and six patients with 183 biopsy-confirmed lung tumours <3.5 cm underwent RFA NSCLC (n = 33)	Technical success  Major complications: Pneumothorax Pleural effusion	99%  n = 27  n = 4	Percutaneous CT-guided RFA yields high proportions of sustained complete ablation in patients with primary or secondary lung tumours, and is associated with acceptable morbidity
	Median follow-up 24 months	Complete tumour response lasting at least 1 year  Overall survival (NSCLC): 1 year	70%	Heterogeneous patient population  Mean follow-up not long enough to detect late tumour recurrences
		2 years  Cancer-specific survival (NSCLC): 1 year 2 years	92% 73%	PET scans not routinely used for the assessment of response
		Stage I NSCLC: 2-year overall survival 2-year cancer specific survival	75% 92%	
Hiraki <i>et al.</i> , (2007), J Thorac Cardiovasc Surg, Japan [6] Retrospective study	Twenty patients with Stage I NSCLC underwent RFA Median follow-up 21.8 months	Local progression  Local control rate: 1 year 2 years 3 years	7/20 (35%) 72% 63% 63%	Treatment of Stage I NSCLC with one or more sessions of RFA offers promising outcomes in relation to survival. However, local progression rates are relatively high
		Mean survival  Overall survival:  1 year	42 months 90% 84%	Short follow-up period Small study population
		2 years 3 years  Cancer-specific survival: 1 year 2 years 3 years  Complications: Pneumothorax	74% 100% 93% 83%	
Pennathur <i>et al.</i> , (2007), J Thorac Cardiovasc Surg, USA [7]	Nineteen patients with Stage I NSCLC underwent RFA under CT-guidance	Pleural effusion  Local progression  Initial complete response	13/20 (57%) 4/20 (17%) 8/19 (42%) 2/19 (10.5%)	RFA appears to be safe in high-risk patients with stage I NSCLC, with reasonable results in terms of survival in high-risk patients who are not fit for
Retrospective study	Stage IA ( <i>n</i> = 11)	Partial response	10/19 (53%)	surgical intervention
				Continue

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Author, date, journal and country, study type level of evidence)	Patient group	Outcomes	Key results	Comments, study weaknesses
		Overall survival:		- "
	Stage IB ( <i>n</i> = 8)	1 year 2 years Complications:	95% 68%	Small sample size
	Mean follow-up 29 months	Pneumothorax	63%	
Simon <i>et al.</i> , (2007), Radiology, USA [8]	One hundred and fifty-three patients with 189 tumours	(≤3 cm) Local tumour progression-free rates:		Lung RFA appears to be a safe treatment for Stage I NSCLC and is
	underwent 183 RFA sessions	1 year	83%	linked with promising long-term
Retrospective study	Primany	2 years 3 years	64% 57%	survival and local tumour progressior outcomes
	Primary (n = 116)	4 years	47%	outcomes
	( 110)	5 years	47%	Biopsies were not routinely performe
	Metastatic (n = 73)	(>3 cm) Local tumour		during follow-up
	(11 - 7 3)	progression-free rates:		A proportion of patients treated
	Stage I NSCLC	1 year	45%	concomitantly with systemic
	(n = 75)	2 years	25%	chemotherapy and/or external beam
		3 years	25%	radiation therapy
	Median follow-up 20.5 months	4 years 5 years	25% 25%	
		Difference between the survival rates:	(P < 0.002)	
		(≤3 cm) and (>3 cm)		
		Overall survival rates: (NSCLC)		
		1 year	78%	
		2 years	57%	
		3 years	36%	
		4 years 5 years	27% 27%	
		Complications:		
		Pneumothorax Chest tube insertion	28.4% 9.8%	
Kashima <i>et al.</i> , (2011),	Four hundred and twenty	Complications:		Lung RFA is a relatively safe procedur
Am J Roentgenol, Japan	patients with 1403 lung tumours	Death	0.4%	but it can be fatal in few cases. Know
9]	underwent 1000 RFA sessions	Aseptic pleuritis Pneumonia	2.3% 1.8%	risk factors such as age, tumour size, platelet count, previous systemic
Retrospective study		Lung abscess	1.6%	chemotherapy, previous external bea
retrospective study		Bleeding requiring transfusion	1.6%	radiotherapy, emphysema should be
		Pneumothorax requiring	1.6%	used to stratify patients
		pleural sclerosis	0.4%	
		Bronchopleural fistula Brachial nerve injury	0.4%	
		Tumour seeding	0.1%	
		Diaphragm injury	0.1%	
Zemlyak <i>et al.</i> , (2010), J Am Coll Surg, USA [10]	Sixty-four patients with Stage I NSCLC	Overall 3-year survival: SLR	87.1%	SLR, RFA and PCT are reasonable alternatives to lobectomy for patient
Retrospective study	SLR (n = 25)	RFA PCT	87.5% 77%	who are poor candidates for major surgery. Survival at 3 years is
	RFA (n = 12)		(P > 0.05)	comparable after sublobar resections and ablative therapies. Ablative
	PCT (n = 27)	3-year cancer-specific survival:		therapies appear to be a reasonable
	101 (11-21)	SLR	90.6%	alternative in high-risk patients not fi for surgery
		RFA	87.5%	54.84.7
		PCT	90.2% ( <i>P</i> > 0.05)	Small sample
			10 > 11 (15)	

Author, date, journal and country, study type (level of evidence)	Patient group	Outcomes	Key results	Comments, study weaknesses
		3-year cancer-free survival: SLR RFA PCT	60.8% 50% 45.6% ( <i>P</i> > 0.05)	Selection bias SBRT not included in comparison
Lagerwaard <i>et al.</i> , (2008), Int J Radiat Oncol Biol Phys, The Netherlands [11]	Two hundred and six patients with Stage I NSCLC underwent SRT	Median overall survival: 1-year survival rate 2-year survival rate	34 months 81% 64%	SRT is well tolerated in patients with extensive comorbidity presenting with inoperable Stage I NSCLC with high local control rates and minimal toxicity
Retrospective study	Medically inoperable $(n = 167)$ Refused surgery $(n = 39)$	Local recurrence: T1 T2	3.5% 2 /129 (1.6%) 5/90 (5.6%) (P = 0.13)	Median follow-up was only 12 months Most recurrences occur within 2 years following treatment
	Inclusion criteria: (1) Tumour <6 cm (2) Confirmed malignancy	Overall recurrence DFS:	21% (43 patients)	Only 88 patients attended for follow-u at 1 year
	(cytohistologic or CT) (3) Absence of metastases on PET scan	1 year 2 years Toxicity:	83% 68%	Only a minority of the patients had pathologic confirmation of malignancy
	Prescription dose 60 Gy in three, five or eight fractions	Fatigue Chest wall pain Nausea Dyspnoea Cough Pneumonitis Rib fractures Chronic thoracic pain	31% 12% 9% 6% 6% 3% 2% 1%	
Haasbeek <i>et al.</i> , (2010), Cancer, The Netherlands [12] Retrospective study	One hundred and ninety-three patients ≥75 years with 203 tumours treated using SRT  T1 (n = 118)	Survival rate: 1 year 3 years Median overall	85.7% 45.1% 32.5 months	SRT achieved high local control rates with minimal toxicity in patients aged ≥75 years, suggesting it should be considered as a curative alternative in the treatment of NSCLC
netrospective study	T2 (n = 85)	DFS: 1 year 3 years	89.2%	Median follow-up of 12.6 months not long enough
	80% medically inoperable 20% declined surgery Median follow-up 12.6 months	Complications: Fatigue Nausea Cough Dyspnoea Chest wall pain Rib fracture Grade ≥3 radiation pneumonitis Chronic chest wall pain Acute toxicity	72.6% 32.6% 4.1% 5.7% 5.2% 3.1% 1.6% 2.1%	
Le <i>et al.</i> , (2006), J Thorac Oncol, USA [13] Retrospective study	Thirty-two patients with inoperable lung tumours treated with single-fraction SRT  NSCLC (n = 20)	Local progression: (NSCLC) >20 Gy <20 Gy	9% 54% (P = 0.03)	Single-fraction SRT is feasible for selected patients with lung tumours and higher doses (>20 Gy) are associated with improved local contro For those who have had prior thoracion radiotherapy, doses ≥25 Gy may be to
	Metastases (n = 12)	CR rate: >20 Gy <20 Gy	57% 10% ( <i>P</i> = 0.21)	toxic Short follow-up period
	Tumour diameter 20-62 mm	1-year overall survival:	(. U.Z.)	Variation in treatment techniques (breath-holding vs. tracking)

Author, date, journal and country, study type (level of evidence)	Patient group	Outcomes	Key results	Comments, study weaknesses
	15 Gy (n = 9) 20 Gy (n = 1) 25 Gy (n = 20) 30 Gy (n = 2) Median follow-up 18 months	Toxicity: Fatigue Pneumothorax Grade 2-3 pneumonitis Pleural effusion	10/32 6/32 (19%) 4/32 (12.5%) 1/32	
Onishi H <i>et al.</i> , (2004), Cancer, Japan [14] Retrospective study	Two hundred and forty-five underwent hypofractionated high-dose STI between 1995 and 2003	Local progression overall: BED ≥100 Gy BED 100 Gy	14.5% 8.1% 26.4% ( <i>P</i> < 0.05)	Hypofractionated high-dose STI is a feasible and effective curative treatmen of patients with Stage I NSCLC
retrospective study	BED ≥100 Gy (n = 173) BED 100 Gy (n = 72)	Local disease recurrence: BED ≥100 Gy BED 100 Gy	13.5% 8.1% 26.4% (P < 0.01)	Local control and survival rates better in patients treated with BED ≥100 Gy than for BED 100 Gy Treatment parameters heterogeneous
	Stage IA n = 155 Stage IB	3-year survival rate: BED ≥100 Gy BED 100 Gy	88.4% 69.4% ( <i>P</i> 0.05)	,
	n = 9 Tumour diameter 7-58 mm (median, 28 mm)	Overall survival rates: 3 years 5 years	56% 47%	
	CT chest usually obtained 3-monthly for first year and repeated every 4-6 months thereafter	Local tumour response: CR (completely disappeared/ replaced by fibrotic tissue)	57/245 (23.3%)	
	Tumour response evaluated using previously published National Cancer Institute (NCI) criteria	PR (≥30% reduction in the maximum cross-sectional diameter)	151/245 (61.6%)	
		Overall response rate BED ≥100 Gy BED 100 Gy	84.8% 84.5% 83.3%	
		Toxicity: Radiation-induced pulmonary complications	17/245 (6.9%)	
Timmerman <i>et al.</i> , (2010) JAMA, USA [15]  Multicentre prospective study	Fifty-five patients with biopsy-proven peripheral T1-T2N0M0 NSCLC (measuring 5 cm in diameter) underwent SBRT  T1 (n = 44) T2 (n = 11)	Median overall survival  Overall 3-year survival  Disseminated recurrence at	48.1 months 55.8% 22.1%	SBRT is an effective treatment in patients with inoperable NSCLC, withigh rates of local tumour control armoderate treatment-related morbid Rarely used invasive pathological staging and histological confirmation recurrence, lowering accuracy
		3 years  3-year primary tumour control rate	97.6%	
	Prescription dose 18 Gy per fraction ×3 fractions (54 Gy total)	Local-regional control rate	87.2%	
	Median follow-up 34.4 months	DFS Adverse events: Grade 3 Grade 4	48.3% 7/55 (12.7%) 2/55 (3.6%)	
Fakiris <i>et al.</i> , (2009), Radiat Oncol Biol Phys, USA [16]	Seventy patients with biopsy confirmed NSCLC underwent SBRT	Local control at 3 years Local recurrence Nodal recurrence Distant recurrence	88.1% 4/70 (5.7%) 6/70 (8.6%) 9/70 (12.9%)	Treatment with SBRT results in high rates of local control in medically inoperable patients with Stage I NSCLO

Author, date, journal and country, study type (level of evidence)	Patient group	Outcomes	Key results	Comments, study weaknesses
Retrospective study	T1 (n = 34) T2 (n = 36)	Median survival	32.4 months	
,	,	3-year overall survival	42.7 months	
	Prescription dose:			
	60-66 Gy in three fractions	Median survival:		
	M I: 6 II 502 II	T1 tumours	38.7 months	
	Median follow-up 50.2 months	T2 tumours	24.5 months $(P = 0.194)$	
			(1 - 0.154)	
Timmerman <i>et al.</i> , (2006), J Clin Oncol [17]	Seventy patients underwent SBRT for NSCLC	2-year overall survival	54.7%	Local recurrence and toxicity occur late after this treatment; however, this
Prospective study	Treatment dose:	3-month major response rate	60%	regimen should not be used for patients with tumours near the central
	60 to 66 Gy total in three fractions during 1-2 weeks	Local control		airways due to excessive toxicity
	fractions during 1-2 weeks	2 years:	95%	Short follow-up period
	Median follow-up 17.5 months	_ /		- Processing of Parisa
	•	Deaths due to:		This trial did not define a limit on the
		Cancer	n = 5	period of observation of toxicity relate
		Treatment	n = 6	to therapy-hospitalizations/deaths
		Comorbid illness	n = 17	occurring more than a year after
		Median overall survival	32.6 months	therapy might not have been attribute to SABR
		Toxicity:		
		Grade 1-2	n = 58	
		Grade 3-4	n = 8	
		Grade 5 (death)	n = 6	

NSCLC: non-small-cell lung cancer; RFA: radiofrequency ablation; FEV<sub>1</sub>: forced expiratory volume in 1 s; VC: vital capacity; DFS: disease-free survival; MVV: maximum voluntary ventilation; SLR: sublobar resection; PCT: percutaneous cryoablation therapy; SRT: stereotactic radiotherapy; SBRT: stereotactic body radiation therapy; STI: stereotactic irradiation; BED: biologic effective dose; CR: complete resolution; PR: partial resolution; MTD: maximum tolerated dose.

AND [disease-free survival/ OR survival/ OR survival.mp. OR toxicity.mp. OR control rate.mp].

## **SEARCH OUTCOME**

The search returned 219 papers. In addition, the references of relevant papers were searched. Sixteen papers provided the best evidence to answer the question. These are tabulated in Table 1.

# **RESULTS**

The effectiveness of RFA and SABR in the treatment of inoperable NSCLC is well documented. Huang  $et\ al.\ [2]$  reported a median progression-free-interval of 21.6 months following RFA. Overall survival at 1, 2 and 5 years was 68.2, 35.3 and 20.1%, respectively, and 23.7% patients developed local progression during follow-up. There was no significant difference in outcome for tumours <3 cm, while there was a significant difference in the risk of local progression in tumours >4 cm (P = 0.01). In another study [3], 57% of primary lung tumours treated with RFA had no recurrence. The local recurrence rate was 38%, with increasing tumour size (P = 0.02) and disease stage (P = 0.007) significantly increasing its likelihood.

de Baère *et al.* [4] documented an 18-month survival rate of 71%, and a trend towards better efficacy for tumours <2 cm in diameter (P = 0.066). The respiratory function was not adversely affected when measured within 2 months of RFA treatment (P = 0.51); however, the long-term effects are unknown and the FDA have received reports of patient deaths associated with lung tumour ablation using RFA.

Lencioni *et al.* [5] achieved a technical success rate of 99% in performing RFA. 12.5% of patients with NSCLC showed incomplete response or progression of disease. The overall 2-year survival of patients with NSCLC was 48%. Hiraki *et al.* [6] observed a high local progression rate (35%) within a median of 9.0 months after the first session.

Pennathur et al. [7] documented a local progression in 42%, and the median time to progression was 27 months. The overall 2-year survival rate was 49% for primary lung cancers. Simon et al. [8] documented a significant difference in survival between patients with large (>3 cm) and small ( $\leq$ 3 cm) tumours (P < 0.002). Local recurrence was most common, suggesting that more aggressive RFA and adjuvant radiation may improve outcomes [9, 10]. Kashima et al. [9] concluded that puncture number (P < 0.02) and previous systemic chemotherapy (P < 0.05) were significant risk factors for aseptic pleuritis. Increasing age (P < 0.02) and previous external beam radiotherapy (P < 0.001) were significant risk factors for pneumonia, as were emphysema (P < 0.02) for lung

abscess and pneumothorax requiring pleural sclerosis (P < 0.02), and serum platelet count (P < 0.002) and tumour size (P < 0.02) for bleeding. Zemlyak *et al.* [10] reported comparable survival rates following sublobar resections (87.1%) and ablative therapies (87.5%).

SABR, a non-invasive technique, precisely delivers very high radiation doses in a short period of time. It is well tolerated in patients with extensive comorbidity with high local control rates and minimal toxicity, and results have been so promising that there are ongoing trials comparing SABR with surgery in operable patients. Lagerwaard *et al.* [11] observed local recurrences in only 3.5% of patients, which is much less than previously reported when using conventional radiotherapy in Stage I NSCLC. Haasbeek *et al.* [12] found no significant difference in overall survival between the older and younger patient cohorts (P = 0.18) following SABR; however, disease-free survival was slightly better in older patients (P = 0.04). Le *et al.* [13] reported an association between prior thoracic radiotherapy or chemotherapy and treatment-related toxicity.

On comparing outcomes between patients treated with biologic effective doses (BED) of  $\geq$ 100 Gy and <100 Gy, Onishi *et al.* [14] found improved local control and survival rates with BED  $\geq$ 100 Gy. They reported the most benefit in those with medically operable tumours, treated with BED  $\geq$ 100 Gy.

Timmerman *et al.* [15] reported a 3-year local control rate of 97.6%, and an overall 3-year survival rate of 55.8%. Fakiris *et al.* [16] reported lower rates of toxicity after SABR in patients with peripheral tumours. However, there was no significant difference in survival between patients with peripheral and central tumours (*P* = 0.69). The 3-year local control (88.1%) was comparable to that following lobectomy. Another study [17] reported mainly grade 1-2 toxicities (83%), consisting of fatigue, musculoskeletal discomfort and radiation pneumonitis. Most cases resolved within 3–4 months of SABR. The authors concluded that patients with perihilar/central tumours had an 11-fold increased risk of experiencing severe toxicity compared with more peripheral locations. Tumours close to the left hemidiaphragm may also be very dangerous to treat with SABR due to their proximity to the stomach.

#### **CLINICAL BOTTOM LINE**

SABR is associated with higher 5-year survival rates compared with RFA and conventional radical radiotherapy (40–47% vs. 20.1–27 vs. 19%) [18] and local control rates up to 80–90% [19] are two to three times greater than conventional fractionated radiotherapy. This modality has a favourable toxicity profile in peripheral tumours measuring ≤5 cm. RFA can be performed in one session, whereas SABR is more effective if larger doses of radiation are split over two to three fractions. RFA is more difficult in central tumours but is being increasingly performed with increased operator experience and confidence. Both treatment modalities are associated with side-effects, and risk factors should be used to stratify patients. Overlapping ablations in the same sitting also improve the outcomes for larger tumours. In certain circumstances, a combined approach may be beneficial.

Conflict of interest: none declared.

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