





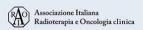




## Radioterapia nella malattia oligometastatica multifocale: limite numerico o tecnico?

#### Ciro Franzese

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#### **DICHIARAZIONE**

Relatore: Ciro Franzese

Come da nuova regolamentazione della Commissione Nazionale per la Formazione Continua del Ministero della Salute, è richiesta la trasparenza delle fonti di finanziamento e dei rapporti con soggetti portatori di interessi commerciali in campo sanitario.

- Posizione di dipendente in aziende con interessi commerciali in campo sanitario (NIENTE DA DICHIARARE)
- Consulenza ad aziende con interessi commerciali in campo sanitario (IPSEN, ASTELLAS)
- Fondi per la ricerca da aziende con interessi commerciali in campo sanitario (NIENTE DA DICHIARARE)
- Partecipazione ad Advisory Board (NIENTE DA DICHIARARE)
- Titolarità di brevetti in compartecipazione ad aziende con interessi commerciali in campo sanitario (NIENTE DA DICHIARARE)
- Partecipazioni azionarie in aziende con interessi commerciali in campo sanitario (NIENTE DA DICHIARARE)
- Altro



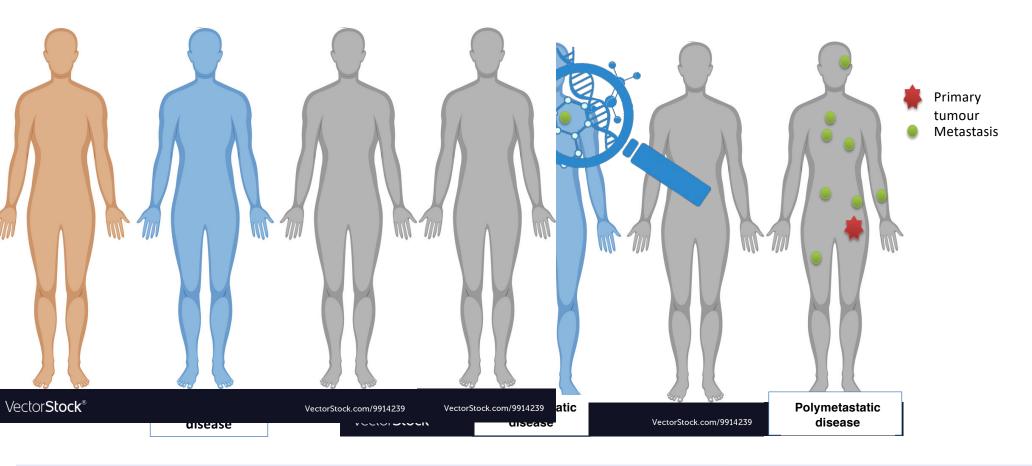




### **AIRO2022**

XXXII CONGRESSO NAZIONALE AIRO XXXIII CONGRESSO NAZIONALE AIRB XII CONGRESSO NAZIONALE AIRO GIOVAN

Radioterapia di precisione per un'oncologia innovativa e sostenibile













#### The four aces

Prognostic factor	Common definitions
Young age	Usually defined as <65 or <70, or analyzed
	as a continuous variable
Patient fitness	Karnofsky performance status ≥70
Slow-growing cancers	Metachronous presentation of oligometastases
	Longer disease-free interval between original tumor and development of metastases
Minimal disease burden	Presence of fewer metastatic sites
	Single-organ oligometastases
	Lack of extracranial disease

Palma et al. Clin Cancer Res; 21(23) December 1, 2015











#### Consensus

Defining oligometastatic disease from a radiation oncology perspective: An ESTRO-ASTRO consensus document



Yolande Lievens <sup>a,\*</sup>, Matthias Guckenberger <sup>b</sup>, Daniel Gomez <sup>c</sup>, Morten Hoyer <sup>d</sup>, Puneeth Iyengar <sup>e</sup>, Isabelle Kindts <sup>f</sup>, Alejandra Méndez Romero <sup>g</sup>, Daan Nevens <sup>h</sup>, David Palma <sup>i</sup>, Catherine Park <sup>j</sup>, Umberto Ricardi <sup>k</sup>, Marta Scorsetti <sup>1</sup>, James Yu <sup>m</sup>, Wendy A. Woodward <sup>c</sup>

While **significant heterogeneity** exists in the current OMD definitions in the literature, consensus was reached on multiple key questions.

Based on available data, OMD can to date be defined as 1–5 metastatic lesions, a controlled primary tumor being optional, but where all metastatic sites must be safely treatable.







## Stereotactic ablative body radiotherapy in patients with oligometastatic cancers: a prospective, registry-based, single-arm, observational, evaluation study

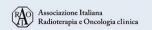
Anastasia Chalkidou, Thomas Macmillan, Mariusz T Grzeda, Janet Peacock, Jennifer Summers, Saskia Eddy, Bola Coker, Hannah Patrick, Helen Powell, Lee Berry, Gareth Webster, Peter Ostler, Peter D Dickinson, Matthew Q Hatton, Ann Henry, Stephen Keevil, Maria A Hawkins, Nick Slevin. Nicholas van As

Between 2015 and 2019, **1422 patients** were recruited from 17 hospitals in England.

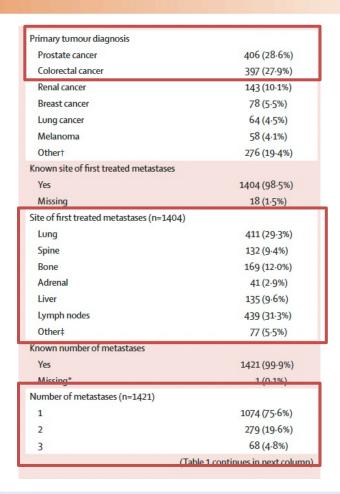
The most common primary tumour was prostate cancer (28.6% patients)

About 75% of patients treated on 1 metastasis, less than 5% on 3 metastases

Lancet Oncol 2021 Jan;22(1):98-106

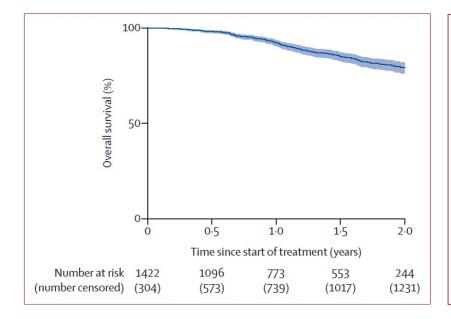












## Number of metastases had no effect on Overall survival

Number of metastases							
1	1,068 (75.1%)	144	2.453	0.113	0.751		
2	278 (19.6%)	41	2.576	0.128	0.731		
3	68 (4.8%)	6	2.428	0.078			
No of subjects used in analysis	1422						

Lancet Oncol 2021 Jan;22(1):98-106

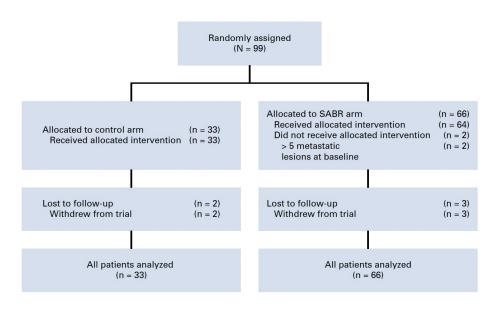


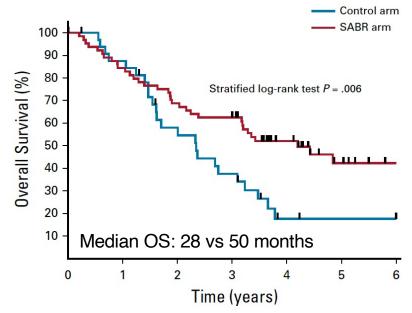






# Stereotactic Ablative Radiotherapy for the Comprehensive Treatment of Oligometastatic Cancers: Long-Term Results of the SABR-COMET Phase II Randomized Trial





Palma et al, JCO 2020









SABR was generally well tolerated, with a 29% rate of grade 2 or higher toxicity

## More than 90% of patients enrolled had 1–3 metastases

_	Arm, No. (%)		
	Control	SABR	
No. of metastases			
1	12 (36)	30 (46)	
2	13 (40)	19 (29)	
3	6 (18)	12 (18)	
4	2 (6)	2 (3)	
5	0 (0)	3 (5)	
Location of metastases (n = 191 lesions)			
Adrenal	2 (3)	7 (6)	
Bone	20 (31)	45 (35)	
Liver	3 (5)	16 (13)	
Lung	34 (53)	55 (43)	
Other <sup>a</sup>	5 (8)	4 (3)	







At present, there is no biological evidence supporting the maximal number of metastases, or the maximal lesion size, that can be treated to provide clinical benefit.

The possibility to **safely deliver** curative intent metastasisdirected radiation therapy determines the maximum number.

Lievens et al. Rad & Oncol 2020

### Are we ready to treat multiple oligometastases?



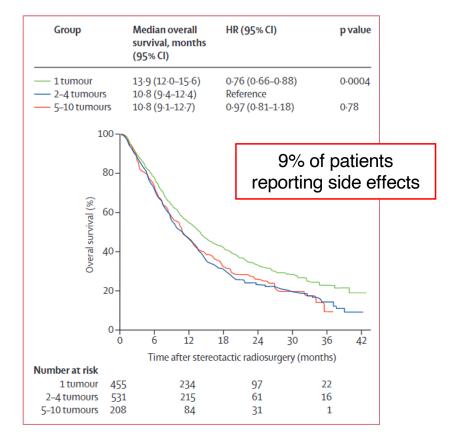




## Stereotactic radiosurgery for patients with multiple brain metastases (JLGK0901): a multi-institutional prospective observational study

Masaaki Yamamoto\*, Toru Serizawa\*, Takashi Shuto, Atsuya Akabane, Yoshinori Higuchi, Jun Kawagishi, Kazuhiro Yamanaka, Yasunori Sato, Hidefumi Jokura, Shoji Yomo, Osamu Nagano, Hiroyuki Kenai, Akihito Moriki, Satoshi Suzuki, Yoshihisa Kida, Yoshiyasu Iwai, Motohiro Hayashi, Hiroaki Onishi, Masazumi Gondo, Mitsuya Sato, Tomohide Akimitsu, Kenji Kubo, Yasuhiro Kikuchi, Toru Shibasaki, Tomoaki Goto, Masami Takanashi, Yoshimasa Mori, Kintomo Takakura, Naokatsu Saeki, Etsuo Kunieda, Hidefumi Aoyama, Suketaka Momoshima, Kazuhiro Tsuchiya

	Univariable	Univariable						
	HR (95% CI)	p value	HR (95% CI)*	p value				
Age, years (≥65 vs <65)	1-412 (1-229-1-622)	<0.0001	1-351 (1-174-1-554)	<0.0001				
Sex (male vs female)	1-427 (1-242-1-655)	<0.0001	1-377 (1-179-1-608)	<0.0001				
KPS (≤70 vs ≥80)	2.079 (1.729-2.500)	<0.0001	1.529 (1.240-1.886)	<0.0001				
Number of tumours								
2-4 vs 1	1.313 (1.131-1.525)	0.0001	1.328 (1.141-1.546)	0.0003				
5-10 vs 2-4	0.974 (0.806-1.177)	0.78	0.993 (0.819-1.204)	0.94				
Maximum diameter of largest tumour (≥1·6 cm vs <1·6 cm)	1.431 (1.249-1.638)	<0.0001	1.006 (0.771-1.314)	0.92				
Cumulative tumour volume (≥1·9 mL vs <1·9 mL)	1.503 (1.313-1.721)	<0.0001	1.172 (0.899-1.530)	0.24				
Primary tumour category								
Breast vs lung	0.743 (0.584-0.945)	0.014	0.881 (0.673-1.153)	0.36				
GI vs lung	1.750 (1.373-2.231)	<0.0001	1-407 (1-087-1-822)	0.0094				
Renal cell vs lung	1.063 (0.718-1.573)	0.76	0-964 (0-648-1-434)	0.13				
Others vs lung	1.572 (1.096-2.255)	0.021	1.333 (0.922-1.927)	0.86				
Extracerebral disease status (not controlled vs controlled)	1-385 (1-200-1-589)	<0.0001	1-272 (1-101-1-469)	0.0011				
Neurological symptoms (yes vs no)	1.779 (1.541-2.053)	<0.0001	1-334 (1-117-1-594)	0.0013				
Clinical factors were measured before stereotactic surgery. HR-hazard ratio. KPS-Karnofsky performance status, Gl-gastrointestinal. *HR adjusted for all clinical factors lister in this table.								



Lancet Oncol 2014; 15: 387-95









### Local Treatment of Unresectable Colorectal Liver

Patient and tumor

Metastases: Results of a Randomized Phase II Trial

#### 119 patients with colorectal liver metastases (< 10 mets and no extrahepatic disease) received systemic treatment alone or systemic treatment plus radiofrequency ablation +- resection

Systemic

treatment

(n = 59)

Local plus

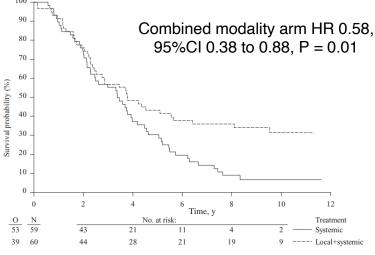
systemic treatment

(n = 60)

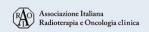
No. of liver metastases 15 (25.0) 7 (11.9) 6 (10.0) 4 (6.8) 8 (13.3) 7 (11.9) 9 (15.0) 8 (13.6) 6 (10.0) 10 (16.9) 9 (15.3) 3 (5.0)

70% 3 - 9 mets 40% 5 - 9 mets



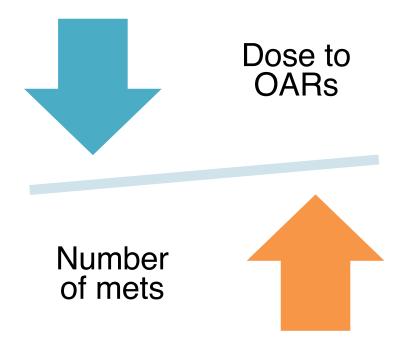


Ruers et al. JNCI J Natl Cancer Inst (2017) 109(9)









- Advanced RT techniques
- Minimum IGRT requirements
- Organ motion control
- Appropiate dose
- Established dose constraints









#### Planning Trade-offs for SABR in Patients With 4 to 10 Metastases: A Substudy of the SABR-COMET-10 Randomized Trial

Samaher Ashram, MD.\* Houda Bahig, MD, PhD.<sup>†</sup> Alsling Barry, MD.<sup>†,6</sup> Denise Blanchette, MRT(T).<sup>†</sup>
Anders Celinksi, MRT(T).\* Peter Chung, MBChB,<sup>†,6</sup> Johnson Darko, PhD.<sup>†</sup> David Donath, MD.<sup>†</sup> Robert Doucet, PhD.<sup>†</sup>
Abigail Erickson, Nat.Dip (RT).\* Meredith Giuliani, MBBS, MEd, PhD.<sup>†,6</sup> Darin Gopaul, MD.<sup>†</sup> Sort Hipwell, BA, CMD.\*
Joanna Javor, MHSc.<sup>†,6</sup> Joda Kuk, MD.<sup>†</sup> Patricia Lindsay, PhD.<sup>†,6</sup> Barbara Millman, MRT(T), BSc.\*
Michael Oliver, PhD.<sup>†</sup> Andrew Pearce, MD.<sup>†</sup> Catherine Russell, "Sashendra Senthi, MD, PhD.\*
Andrew Wamer, MSc.\* Stewart Gaede, PhD.\* and David A. Palma, MD, PhD.\*

**IJROBP 2022** 

SABR planning was **achievable without compromise** of the PTV in a large majority of patients, with **only 3%** of targets covered with a **D95 <95%** 

Compromise most commonly required for vertebral and lymph node targets

Conformality parameters such as R50, R100, and D2cm were not met in fewer than 10% of lesions

Table 3	Tumor characteristics and dosimetr	y metrics stratified b	v location of	metastases for all	patients

Characteristic	N	All lesions (n = 332)	Bone (n = 32)	Brain (n = 2)	Liver (n = 26)	Lung (n = 181)	Lymph node (n = 83)	Spine (n = 8)	P value
PTV size (cm <sup>3</sup> ), median (IQR)	296	9.0 (5.3, 24.3)	30.3 (12.9, 62.2)	3.6 (3.3, 4.0)	50.3 (16.5, 103.4)	9.1 (6.2, 18.9)	4.7 (2.7, 8.5)	14.0 (4.2, 49.7)	< .001
Dose fractionation, n (%)									
16-18 Gy in 1 fraction	332	2 (0.6)	1 (3.1)	1 (50.0)		-	-		-
20-24 Gy in 1 fraction		111 (33.4)	4 (12.5)	-	-	107 (59.1)	-	-	
24-27 Gy in 3 fractions		11 (3.3)		1 (50.0)		6 (3.3)	2 (2.4)	2 (25.0)	
25-28 Gy in 5 fractions		14 (4.2)	2 (6.3)	-	-	3 (1.7)	9 (10.8)	-	
30-33 Gy in 3 fractions		51 (15.4)	5 (15.6)	-	5 (19.2)	19 (10.5)	21 (25.3)	1 (12.5)	
30-40 Gy in 5 fractions		143 (43.1)	20 (62.5)	-	21 (80.8)	46 (25.4)	51 (61.4)	5 (62.5)	
Maximum PTV dose 0.03 cm <sup>3</sup> (%), median (IQR)	332	124.2 (117.1, 129.8)	124.3 (116.5, 129.8)	131.1 (126.5, 135.6)	118.3 (106.4, 123.4)	127.1 (122.8, 131.6)	113.0 (108.5, 123.0)	130.3 (120.9, 132.9)	< .001
Coverage D95 (%), median (IQR)	332	100.3 (100.0, 101.1)	100.3 (100.1, 100.9)	100.3 (100.0, 100.6)	100.5 (98.7, 101.7)	100.2 (100.0, 100.9)	100.5 (99.6, 101.5)	100.0 (84.6, 100.5)	.556
Coverage D95 (%), n (%)									
<95	332	11 (3.3)	0 (0)	0 (0)	1 (3.8)	1 (0.5)	7 (8.4)	2 (25.0)	< .001
≥95		321 (96.7)	32 (100)	2 (100)	25 (96.2)	180 (99.5)	76 (91.6)	6 (75.0)	
Coverage D95 (%), n (%)									
<100	332	70 (21.1)	4 (12.5)	0 (0)	7 (26.9)	33 (18.2)	23 (27.7	3 (37.5)	.210
≥100		262 (78.9)	28 (87.5)	2 (100)	19 (73.1)	148 (81.8)	60 (72.3)	5 (62.5)	
R50 status, n (%)									
Pass/acceptable	284	260 (91.6)	28 (100)	2 (100)	19 (95.0)	151 (96.2)	54 (76.1)	6 (100)	< .001
Fail		24 (8.5)	0 (0)	0 (0)	1 (5.0)	6 (3.8)	17 (23.9)	0 (0)	
R100 status, n (%)									
Pass/acceptable	293	271 (92.5)	27 (100)	2 (100)	18 (90.0)	159 (96.4)	59 (80.8)	6 (100)	.002
Fail		22 (7.5)	0 (0)	0 (0)	2 (10.0)	6 (3.6)	14 (19.2)	0 (0)	
D <sub>2cm</sub> status, n (%)									
Pass/acceptable	285								.246
		267 (93.7)	27 (96.4)	2 (100)	18 (90.0)	144 (91.1)	70 (98.6)	6 (100)	
Fail		18 (6.3)	1 (3.6)	0 (0)	2 (10.0)	14 (8.9)	1 (1.4)	0 (0)	
All organs at risk meet constraints, n (%)									
Yes	62	59 (95.2)	6 (100)	-	4 (100)	35 (100)	14 (82.4)	-	.075
No		3 (4.8)	0 (0)	-	0 (0)	0 (0)	3 (17.7)	-	









#### Single-isocenter versus multiple-isocenters for multiple lung metastases: Evaluation of lung dose



Janita E. van Timmeren <sup>a,\*</sup>, Stefanie Ehrbar <sup>a</sup>, Madalyne Chamberlain <sup>a</sup>, Michael Mayinger <sup>a</sup>, Mischa S. Hoogeman <sup>b</sup>, Nicolaus Andratschke <sup>a</sup>, Matthias Guckenberger <sup>a</sup>, Stephanie Tanadini-Lang <sup>a</sup>

**15 NSCLC patients** with **2 or 3 lesions** previously treated with SBRT was subjected to treatment planning with a **multiple-isocenter technique** and a **single-isocenter technique**.

2 margin approaches were evaluated:

- 1) identical margins for each internal target volume (ITV), assuming an average registration for all lesions in CBCT positioning verification
- 2) a smaller margin for the largest lesion, assuming an optimal registration for that lesion.

**MLD** was  $4.9 \pm 1.9$  Gy for multiple-isocenters and  $5.4 \pm 2.1$  Gy and  $5.3 \pm 2.2$  Gy for single-isocenter approach 1 and 2, respectively.

**V20Gy** was  $5.5 \pm 3.7\%$ ,  $5.5 \pm 3.2\%$  and  $5.4 \pm 3.3\%$ 

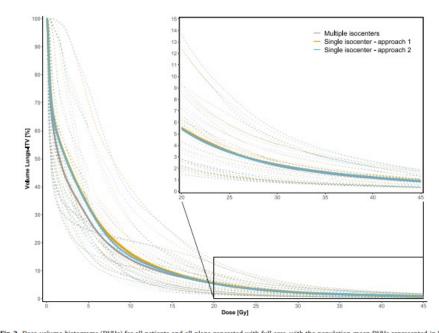


Fig. 2. Dose-volume histograms (DVHs) for all patients and all plans generated with full arcs, with the population-mean DVHs represented in bo









JAMA Oncology | Original Investigation

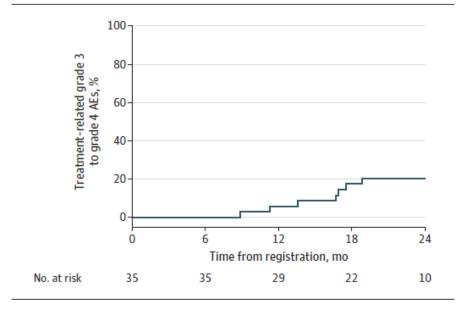
#### Evaluation of Safety of Stereotactic Body Radiotherapy for the Treatment of Patients With Multiple Metastases Findings From the NRG-BROO1 Phase 1 Trial

Steve Chmura, MD, PhD; Kathryn A. Winter, MS; Clifford Robinson, MD; Thomas M. Pisansky, MD; Virginia Borges, MD; Hania Al-Hallaq, PhD; Martha Matuszak, PhD; Sean S. Park, MD; Sun Yi, MD; Yasmin Hasan, MD; Jose Bazan, MD; Philip Wong, MD; Harold A. Yoon, MD; Janet Horton, MD; Gregory Gan, MD; Michael T. Milano, MD, PhD; Elin Ruth Sigurdson, MD; Jennifer Moughan, MS; Joseph K. Salama, MD; Julia White, MD

**42 patients enrolled**, with **3 to 4 metastases** or 2 metastases in close proximity to each other

**8 instances** of **grade 3 AEs**, most likely related to protocol therapy, occurred approximately 125 to 556 days from SBRT initiation in **7 patients** 

Figure 1. Time to Treatment-Related Grade 3 to Grade 4 Adverse Events (AEs) Occurring Greater Than 180 Days After the Start of Stereotactic Body Radiation Therapy for All Evaluable Patients









Stereotactic body radiotherapy (SBRT) for multiple pulmonary oligometastases: Analysis of number and timing of repeat SBRT as impact factors on treatment safety and efficacy



R.J. Klement <sup>a</sup>, J. Hoerner-Rieber <sup>b,c</sup>, S. Adebahr <sup>d,e,f</sup>, N. Andratschke <sup>g</sup>, O. Blanck <sup>h</sup>, J. Boda-Heggemann <sup>i</sup>, M. Duma <sup>j</sup>, M.J. Eble <sup>k</sup>, H.C. Eich <sup>l</sup>, M. Flentje <sup>m</sup>, S. Gerum <sup>n</sup>, P. Hass <sup>o</sup>, C. Henkenberens <sup>p</sup>, G. Hildebrandt <sup>q</sup>, D. Imhoff <sup>r</sup>, K.H. Kahl <sup>s</sup>, N.D. Klass <sup>t</sup>, R. Krempien <sup>u</sup>, F. Lohaus <sup>e,f,v,w</sup>, C. Petersen <sup>x</sup>, E. Schrade <sup>y</sup>, T.G. Wendt <sup>z</sup>, A. Wittig <sup>aa</sup>, M. Guckenberger <sup>g,\*</sup>



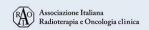
**145 patients** were treated for **multiple (2 – 6) pulmonary metastases**; 88 patients received all SBRT treatments within one month whereas 57 patients were treated with repeat SBRT

No significant association between 3-month and 6-month death rates and the number of pulmonary metastases, no grade 4 or grade 5 toxicity was observed in these patients.

Death rates of the different patient groups.

Group	3-Month death count (rate)	p-Value	6-Month death count (rate)	<i>p</i> -Value
Single SBRT	26 (6.0%)	1	62 (14.4%)	0.7221
Multiple SBRT	8 (6.3%)		16 (12.6%)	
Synchronous SBRT	3 (3.9%)	0.2729	9 (12.0%)	0.6896
Metachronous SBRT	3 (7.7%)		4 (11.1)	
Synchronous followed by metachronous SBRT	2 (11.8%)		3 (18.8%)	

Median OS was 23.5 months and **OS was not influenced by the number of SBRT treatments** or the number and timing of repeat SBRT courses





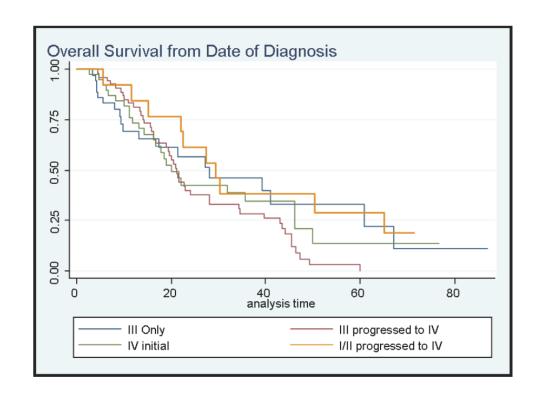


## 146 NSCLC patients (stage III and IV) treated with curative-intent radiotherapy

All stage IV NSCLC patients treated with SBRT had ≤ 8 lesions.

5-year OS was superior (p < 0.01) for those with limited metastases ( $\leq$  8 lesions) versus stage III patients who developed extensive metastases not amenable to SBRT (14% vs 0%)

No significant difference among patients with  $\leq 5$  metastases versus 6-8 (p = 0.94)



Cheruvu et al. Radiation Oncology 2011, 6:80





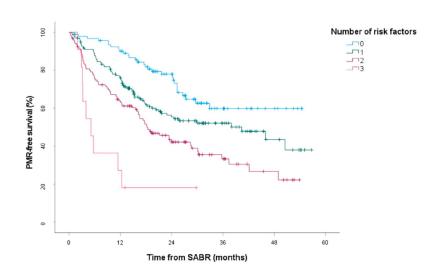




#### Predictors of Early Polymetastatic Relapse After SABR for up to 5 Oligometastases: A Secondary Analysis of the Phase II SABR-5 Trial

Sarah Baker, MD, PhD,\*\*<sup>†</sup> Benjamin Mou,\*\*<sup>‡</sup> Will Jiang, MD,\*\*<sup>†</sup> Mitchell Liu, MD, CM,\*\*<sup>§</sup> Alanah M. Bergman, PhD,<sup>§</sup> Devin Schellenberg, MD,\*\*<sup>†</sup> Abraham S. Alexander, MD,\*\*<sup>‡</sup> Hannah Carolan,\*\*<sup>§</sup> Siavash Atrchian, MD,\*\*<sup>‡</sup> Tanya Berrang, MD,\*\*<sup>‡</sup> Andrew Bang, MD,\*\*<sup>‡</sup> Nick Chng, PhD,<sup>¶</sup> Quinn Matthews, PhD,<sup>¶</sup> Scott Tyldesley, MD,\*\*<sup>§</sup> and Robert A. Olson, MD, MSc\*<sup>¶</sup>





- Worse performance status (HR = 2.01,P = .018)
- Non-prostate/breast histology (HR = 3.64, P < .001)
- Oligoprogression (HR = 3.84, P < .001)</li>

#### 3-year OS were:

- 0% Group 1
- 53% Group 1
- 77% Group 3
- 93% Group 4









## Stereotactic ablative radiotherapy (SABR) for the treatment of patients with multiple oligometastases: evaluation of safety and the impact of dose on survival.

C Franzese, V Vernier, D Franceschini, T Comito, P Navarria, E Clerici, MA Teriaca, M Massaro, L Di Cristina, B Marini, C Galdieri, P Mancosu, S Tomatis, M Scorsetti.

Humanitas Research Hospital IRCSS

Humanitas University

136 patients were treated from 2012 to 2020 on 450 oligometastases.

102 (75.0%)  $\rightarrow$  3 oligometastases 26 (19.1%)  $\rightarrow$  4 oligometastases 8 (5.9%)  $\rightarrow$  5 oligometastases

Rad & Onc, Under review





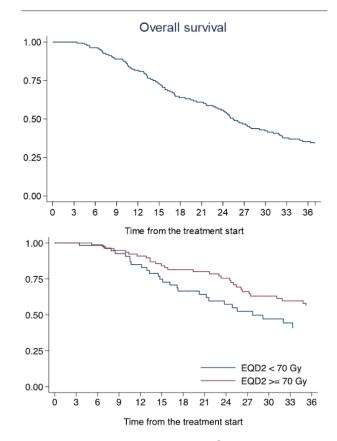


OS rates at 1 and 2 years were 81.6% and 55.7%

Performance status (p=0.022) and minimum dose EQD2 (p=0.032) were significant for OS

#### **Median OS:**

- 27.7 months with EQD2 < 70 Gy</li>
- 47.5 months with EQD2 ≥ 70 Gy



Rad & Onc, Under review







In terms of toxicity, in the acute setting we observed **26 (21.0%) patients and 11 (8.9%) patients** reporting **grade 1 and grade 2** side effect, respectively.

In the late setting, 8 (6.4%) patients reported grade 1 toxicity, and 3 (2.4%) patients reported grade 2 side effects, in the form of cough (5 instances) and dyspnea (5 instances).

No grade 3 or higher toxicity was reported both in the acute and late settings.

Rad & Onc, Under review









#### Lung multiple oligometastases – Clinical case 1

69 yo female, No comorbidities, non smoker

Jun 2018: <u>surgical resection fo rectal adenocarcinoma</u>, <u>pT3pN0</u>

Oct 2018: appearance of at least 6 liver lesions

Oct 2018 - Apr 2019 systemic therapy with FOLFOX + Panitumumab with partial response

May 2019 surgical resection of the liver lesions

Jun 2019 - Sep 2019 systemic therapy with FOLFOX + Panitumumab with bed tolerance

Nov 2020 appearance of 2 small suspicious nodules in the left lung

Mar 2021 increase of the left nodules and appearance of right lung nodule

Suggested activation of II line systemic therapy that the patient refused

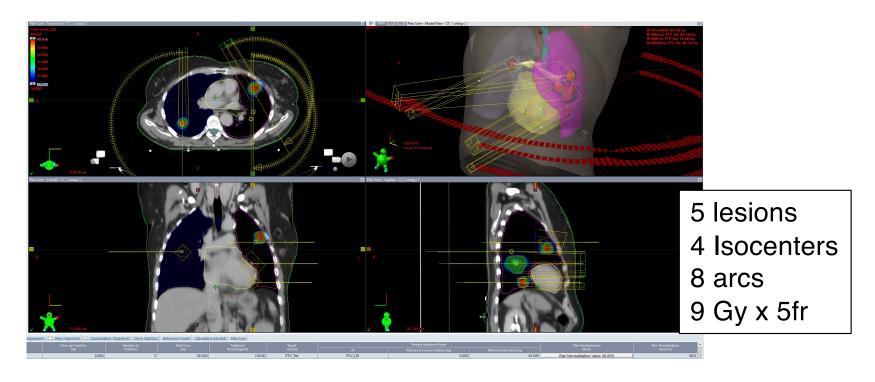








## May 2021 SBRT on 2 nodules in the upper left lung, 2 nodules in lower left lung, 1 nodule in right lung











## Liver multiple oligometastases – Clinical case 74 yo female

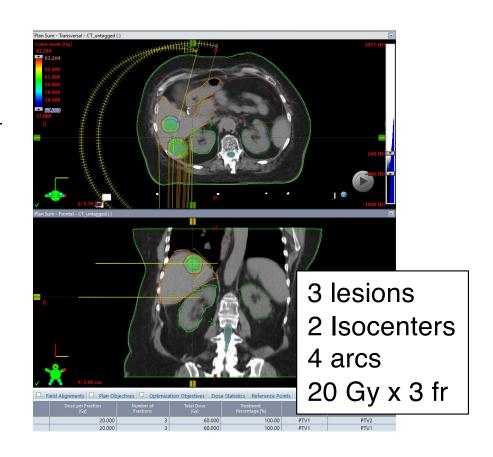
2017 Diagnosis of right breast cancer treated with <u>surgery</u>, CHT and OT

2019 Appearance of 3 liver metastases

Oct 2019 – December 2020 <u>systemic therapy</u> with scarse tolerance

Suggested surgery or RFA refused by the patient

March 2021: SBRT on the 3 liver metastases in S6, S7 and S7-8









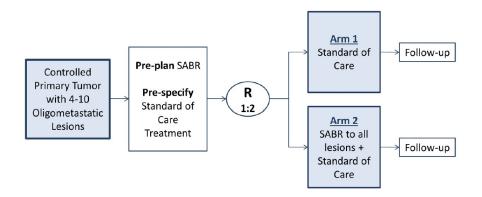


#### STUDY PROTOCOL Open Access

Stereotactic ablative radiotherapy for the comprehensive treatment of 4–10 oligometastatic tumors (SABR-COMET-10): study protocol for a randomized phase III trial



David A. Palma<sup>1</sup>, Robert Olson<sup>2</sup>, Stephen Harrow<sup>3</sup>, Rohann J. M. Correa<sup>1</sup>, Famke Schneiders<sup>4</sup>, Cornelis J. A. Haasbeek<sup>4</sup>, George B. Rodrigues<sup>1</sup>, Michael Lock<sup>1</sup>, Brian P. Yaremko<sup>1</sup>, Glenn S. Bauman<sup>1</sup>, Belal Ahmad<sup>1</sup>, Devin Schellenberg<sup>2</sup>, Mitchell Liu<sup>2</sup>, Stewart Gaede<sup>1</sup>, Joanna Laba<sup>1</sup>, Liam Mulroy<sup>5</sup>, Sashendra Senthi<sup>6</sup>, Alexander V. Louie<sup>7</sup>, Anand Swaminath<sup>8</sup>, Anthony Chalmers<sup>9</sup>, Andrew Warner<sup>1</sup>, Ben J. Slotman<sup>4</sup>, Tanja D. de Gruijl<sup>4</sup>, Alison Allan<sup>1</sup> and Suresh Senan<sup>4</sup>



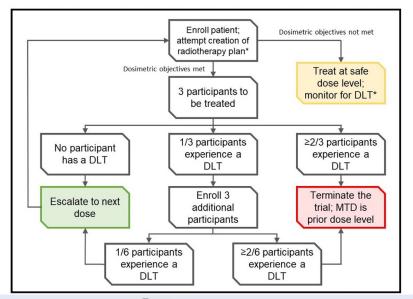
#### STUDY PROTOCOL

Open Access

Ablative radiation therapy to restrain everything safely treatable (ARREST): study protocol for a phase I trial treating polymetastatic cancer with stereotactic radiotherapy



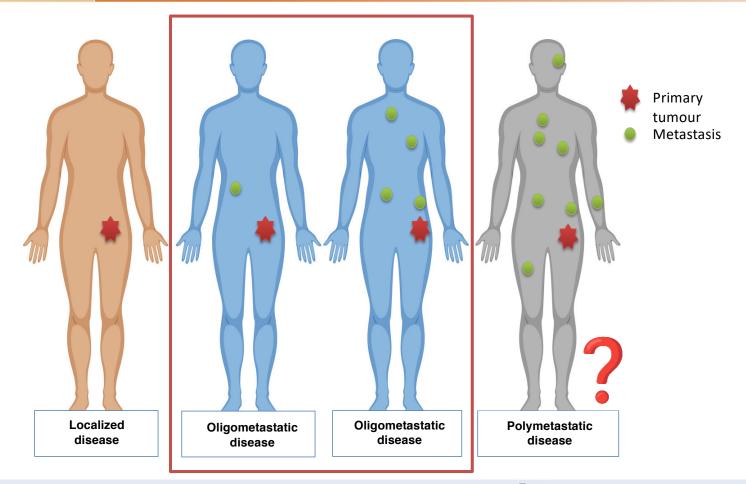
Glenn S. Bauman<sup>1\*</sup>, Mark T. Corkum<sup>1</sup>, Hatim Fakir<sup>2</sup>, Timothy K. Nguyen<sup>1</sup> and David A. Palma<sup>1</sup>

















#### **Conclusions**

- Oligometastatic setting includes a wide spectrum of disease, with a common consensus on a maximum number of 5 metastases
- Majority of published trials include patients treated with SBRT on a single lesions, few patients with 2 to 3 metastases
- Modern radiotherapy may potentially increase the number of safely treatable oligometastases with an acceptable risk of side effects
- Prospective trials are evaluating the safety and efficacy of SBRT on polymetastatic patients compared to systemic therapy alone









#### Thank you

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