



Supplementary Materials

Circulating Exosomal Integrin β3 Is Associated with Intracranial Failure and Survival in Lung Cancer Patients Receiving Cranial Irradiation for Brain Metastases: A Prospective Observational Study

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Table S1. Clinical factors associated with intracranial progression-free survival and extracranial progression-free survival.

		Median IC-PFS (month)	95% CI	p-value	Median EC-PFS (month)	95% CI	<i>p</i> -Value
Sex	Male	8.0	4.2-9.3	0.11	4.8	3.0-9.1	0.22
	Female	7.9	6.3–19.8		5.7	3.7–10.9	
Number of BM	<5	8.6	5.2-NA	0.93	5.8	2.4-NA	0.86
	5-10	7.2	4.2-NA		3.7	2.8-12.1	
	>10	7.9	5.1–13.7		5.0	3.5–9.3	
LM carcinomatosis	Present	7.4	4.1-9.3	0.93	5.0	3.1-9.1	0.12
	Absent	7.9	5.2–11.9		4.4	2.7–9.5	
Histology	ADC	8.0	7.4–11.7	0.002	5.6	4.8-9.3	0.02
	Non-ADC	3.5	2.4-NA		2.4	1.8-NA	
GPA score	0-1.0	3.6	3.2-NA	0.002	3.3	2.8-NA	0.11
	1.5-2.0	8.0	5.0-13.7		4.8	2.7-9.1	
	2.5-3.0	9.1	7.9–31.3		5.8	5.6-14.3	
	3.5-4.0	19.8	9.3-NA		Not reached	1.4-NA	

Abbreviations: IC-PFS, intracranial progression-free survival; CI, confidence interval; EC-PFS, extracranial progression-free survival; BM, brain metastases; LM, leptomeningeal; ADC, adenocarcinoma; GPA; graded prognostic assessment.

 Table S2. Univariate Cox regression analysis of factors associated with overall survival.

	HR	95% CI	<i>p</i> -Value
Age (per 1-year increase)	1.03	0.99-1.07	0.12
Sex (female vs. male)	0.62	0.33-1.16	0.14
Histology (adenocarcinoma vs. non-adenocarcinoma)	0.36	0.18-0.73	0.004
Prior SRS (yes vs. no)	0.98	0.81-1.19	0.88
Number of BM (<5, 5–10, >10)	0.99	0.70-1.40	0.94
High GPA (high vs. low)	0.26	0.14-0.48	< 0.001
Integrin β3 (per 1 ng/mL increase)	1.15	1.005-1.32	0.04
Integrin β6 (per 1 ng/mL increase)	1.06	0.87-1.29	0.56

Abbreviations: HR, hazard ratio; CI, confidence interval; vs., versus; SRS, stereotactic radiosurgery; BM, brain metastases; GPA; graded prognostic assessment.

Table S3. Univariate subdistribution hazards for factors associated with intracranial failure with death as competing risk.

	sHR	95% CI	<i>p</i> -Value
Age (per 1-year increase)	1.01	0.97-1.06	0.54
Sex (female vs. male)	1.1	0.59-2.05	0.76
Histology (adenocarcinoma vs. non-adenocarcinoma)	1.26	0.48-3.32	0.64
Prior SRS (yes vs. no)	1.04	0.99-1.08	0.12
Number of BM (<5, 5–10, >10)	1.05	0.72-1.55	0.8
High GPA (high vs. low)	1.56	0.71-3.42	0.27
Integrin β3 (per 1 ng/mL increase)	1.16	0.995-1.34	0.058
Integrin β6 (per 1 ng/mL increase)	0.995	0.79-1.25	0.96

Abbreviations: sHR, subdistribution hazard ratio; CI, confidence interval; vs., versus; SRS, stereotactic radiosurgery; BM, brain metastases; GPA; graded prognostic assessment.