

# Supplementary Table S1

Criteria for study enrollment		Clinical trial	Gilbert	Gilbert	Chinot	Stupp	Westphal	Stupp	Kong	Weller	Liau	Herringer
			JCO (2013)	NEJM (2014)	NEJM (2014)	Lancet (2014)	EJC (2015)	JAMA (2017)	Oncotarget (2017)	Lancet Oncol. (2017)	JTM (2018)	Lancet (2019)
Age	18-70				§	●		●	§	●	●	
	≥ 18	●	●	●	●		●		●			
Performance status	ECOG 0-2 *ECOG 0-1 only			●	*				●			
	KPS ≥ 70 *KPS ≥ 60	*	●	●		●	●	*		●	●	
Physiology	Adequate hematologic, hepatic & renal function *Adequate coagulation	●	●	*	●	●	●	*	●	●	*	
Pathology	Newly diagnosed glioblastoma *Specify supratentorial	*	●	*	*	●	●	*	●	*	●	
Treatment	Bopsy/resection *Resection required	●	*	●	●	●	●	●	●	*	*	●
	RT/TMZ #Planned for 60Gy/TMZ *Completed RT/TMZ	*	#	#	#	#	#	*	#	*	*	#
Glucocorticoids	Stable or decreasing doses	●	●	●	●					●		●
Study specific criteria for enrollment	E.g. molecular stratification					MGMT methylated GBMs only				EGFRvIII positive GBMs only		MGMT methylated GBMs only
Disease status	Progressive disease before randomization	●	●	●	●					●	●	
Disqualifying comorbidities	E.g. active heart, lung, hepatic, renal disease, other malignancy etc.	●	●	●	●		Not described	●	●	●	●	●

Time of randomization	After completion of concurrent RT/TMZ	Randomization after 3 weeks (midway) into RT/TMZ	Before start of concurrent RT/TMZ	Before start of concurrent RT/TMZ	Before start of concurrent RT/TMZ	After completion of concurrent RT/TMZ	Before start of concurrent RT/TMZ	After completion of concurrent RT/TMZ	After completion of concurrent RT/TMZ	Before start of concurrent RT/TMZ

- Inclusion criteria for enrollment trials
- Inclusion criteria used in a minority of trials investigating specific molecular subgroup (not considered a criteria in this study)
- Exclusion criteria for enrollment in trials

§ The trials do not report an upper age limit as an inclusion criterion, however, the patient characteristics do not describe any patients above 70 years (the reported median age and ranges < 70 years), and they do not describe any data for patients > 70 years in the article nor supplementary material.

## Supplementary Table S2

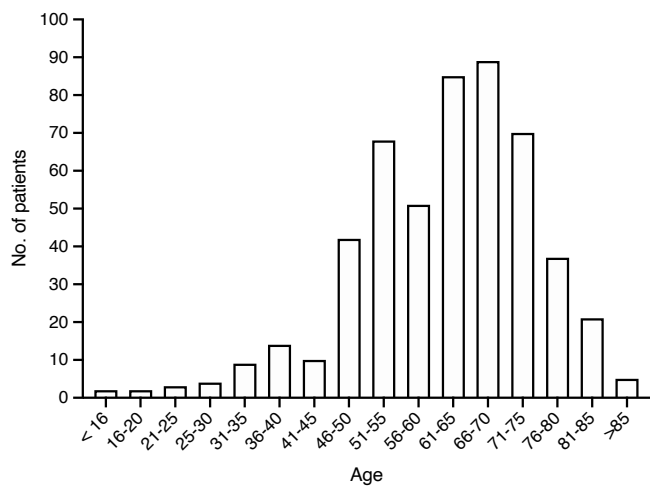
	No RT/no TMZ		RT (any dose)/no TMZ		RT (any dose) /discontinued TMZ		Completed RT/TMZ		Total (%)
	Number (%)	Median survival, months	Number (%)	Median survival, months	Number (%)	Median survival, months	Number (%)	Median survival, months	
No surgery	16 (3.1)	1.7	6 (1.1)	3.4	4 (0.8)	6.4	2 (0.4)	8.6	28 (5.5)
Stereotactic biopsy	8 (1.6)	1.6	5 (1.0)	2.7	7 (1.4)	5.5	18 (3.5)	8.5	38 (7.4)
Open biopsy	6 (1.2)	0.9	6 (1.2)	6.0	5 (1.0)	6.5	16 (3.5)	10.7	33 (6.4)
Resection	22 (4.3)	1.1	47 (9.2)	8.2	39 (7.6)	9.3	302 (59.0)	15.6	410 (80.1)
Total (%)	52 (10.2)	1.3	64 (12.5)	6.7	55 (10.7)	8.0	338 (66.0)	14.8	509* (99.4%) <sup>a</sup>

## Supplementary Table S3

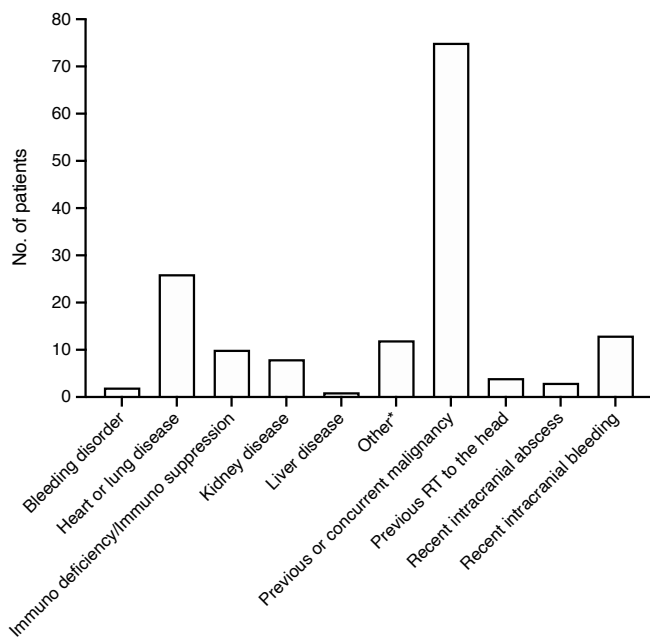
Criterion	1) Before RT/TMZ (n=excluded)	2) Midway through RT/TMZ (n=excluded)	3) At the end of RT/TMZ (n=excluded)
Age 18-70	136	136	136
Supratentorial GBM	17	17	17
Open biopsy/resection	29	29	29
No disqualifying comorbidities	72	72	72
Planned further oncological treatment	23	23	23
No progression during RT/TMZ	Not applicable	4	5
Completed RT/TMZ	Not applicable	Not applicable	12
ECOG $\leq 2$	5	6	3
Stable/no steroids	8	11	6
Adequate blood levels	0	2	17
Total excluded patients (%)	290 (57%)	300 (59%)	320 (63%)

# Supplementary Figure S1

A

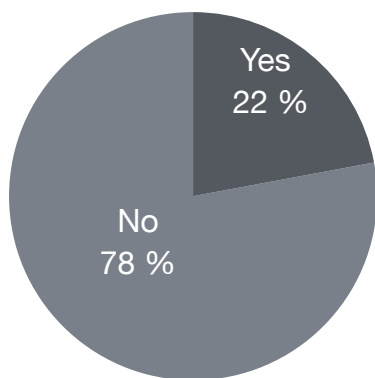


B



\*Diagnoses within the category: dementia, congenital intellectual disability, severe psychiatric disorder, downs syndrome. Also one patient that was pregnant.

C



	Number (%)	Median age (range) in years	Median survival (95% CI) in months
Polypharmacy			
Yes	113 (22.0)	69 (37-85)	10.1 (8.1-11.9)
No	399 (78.0)	61 (10-97)	12.1 (11.1-13.3)