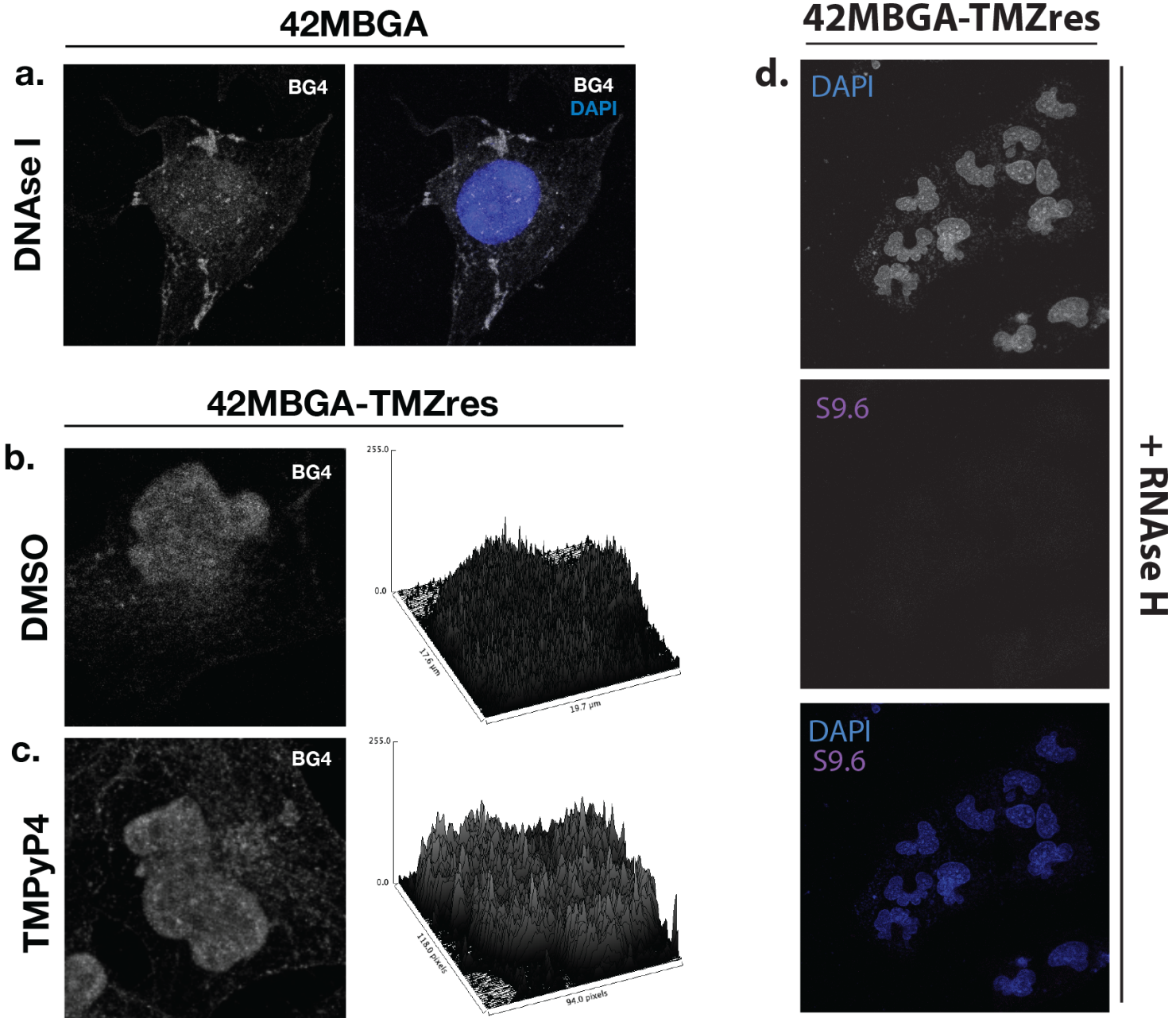


Supplementary Figure 1

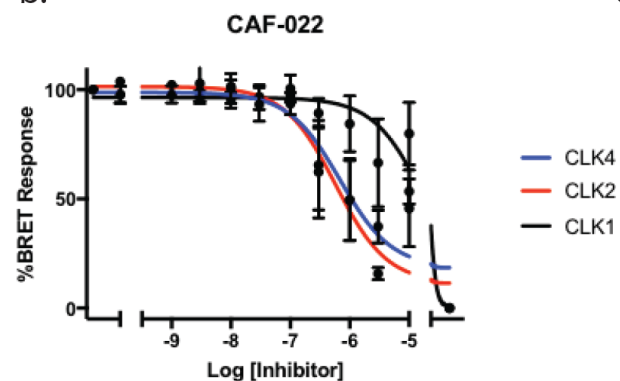


Supplementary Figure 2

a.

Kinase	Nanosyn %I		SGC DSF $\Delta T_m / ^\circ\text{C}$	DiscoverX		RBC IC ₅₀
	100 nM	1.0 μM		PoC @1 μM	K _D	
CLK1				2.90%	26 nM	10 nM
CLK2	81%	96%		8.30%	4.9 nM	0.73 nM
CLK3	4%	22%	0.4 $^\circ\text{C}$	89%	nt	nt
CLK4			5.8 $^\circ\text{C}$	27%	140 nM	8.8 nM
DYRK1A	8%	22%	1.8 $^\circ\text{C}$	73%	nt	nt
DYRK1B	5%	39%		81%	nt	nt
DYRK2	1%	6%	2.1 $^\circ\text{C}$	100%	nt	nt
DYRK3					nt	> 10 μM
DYRK4					nt	> 10 μM
HIPK1	20%	62%		20%	3.6 μM	> 10 μM
HIPK2				27%	530 nM	nt
HIPK3				74%	nt	nt
HIPK4	4%	24%		76%	nt	nt
SRPK1	-1%	-1%	0.4 $^\circ\text{C}$	100%	nt	nt
SRPK2			0.1 $^\circ\text{C}$	100%	nt	nt
MSSK1	0%	0%		100%	nt	nt
PRP4				100%		
STK16			6.5 $^\circ\text{C}$	6.90%	210 nM	100 nM
ERK8				4.40%	35 nM	120 nM
NEK6	0%	2%		11%	84 nM	> 10 μM
NEK7	3%	2%		19%	140 nM	> 10 μM

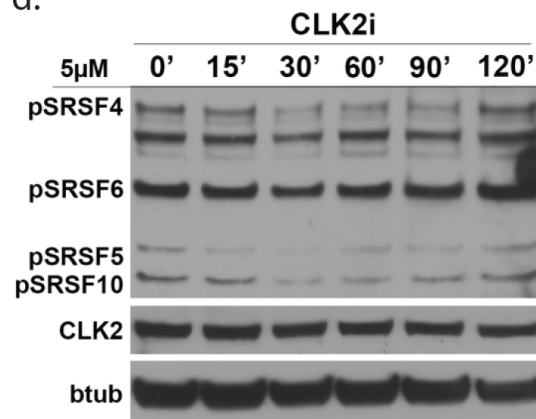
b.



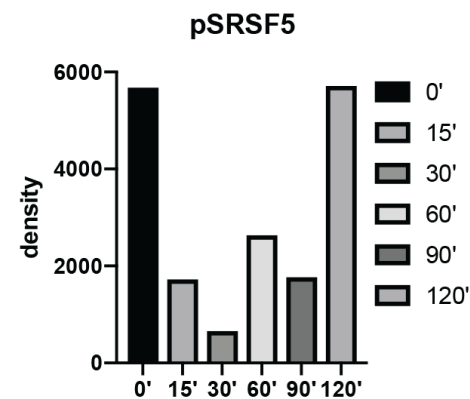
c.

NanoBRET IC ₅₀ s
CLK1 IC ₅₀ = >10,000 nM
CLK2 IC₅₀ = 580 nM
CLK4 IC ₅₀ = 690 nM

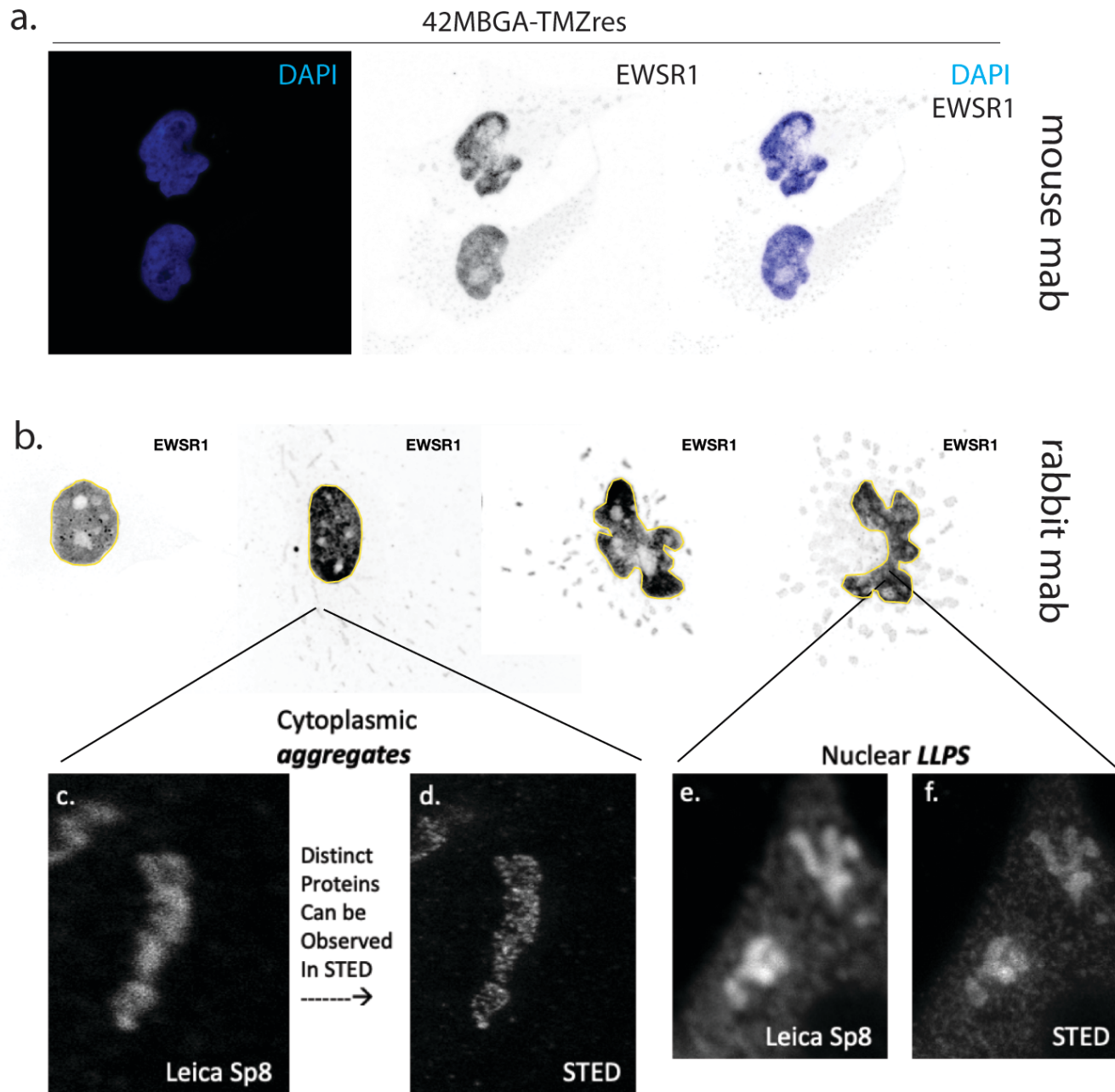
d.



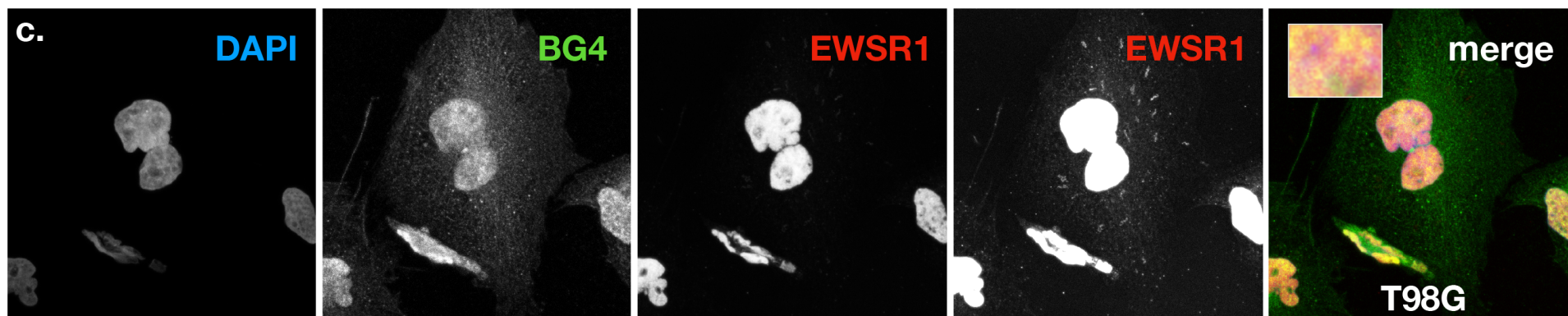
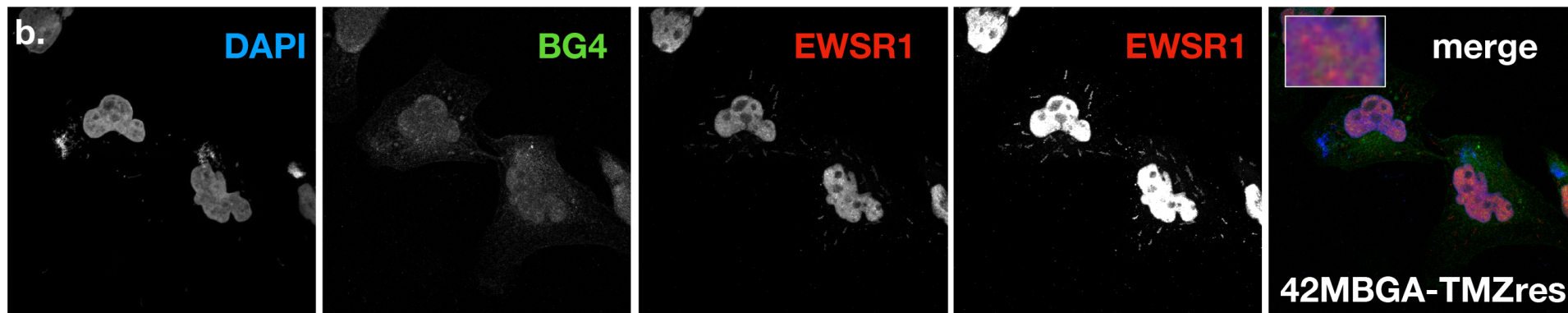
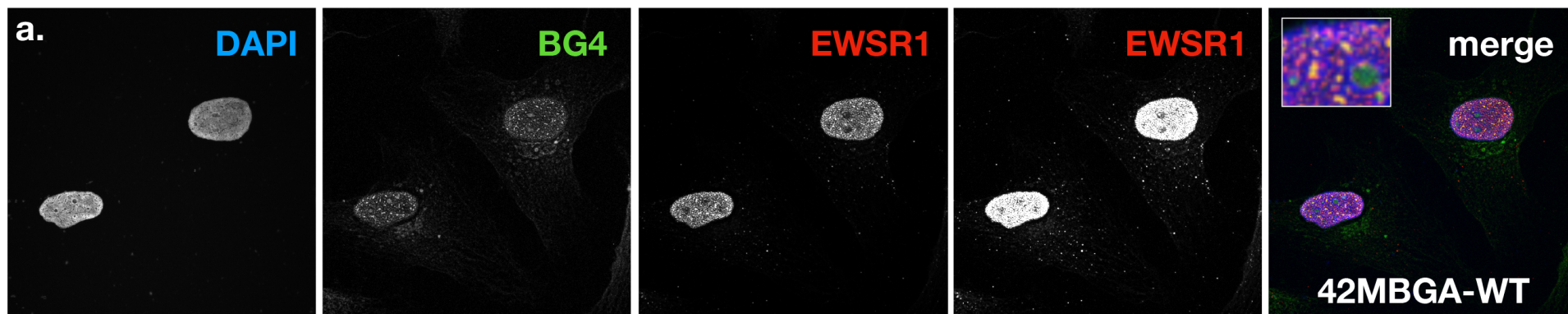
e.



Supplementary Figure 3

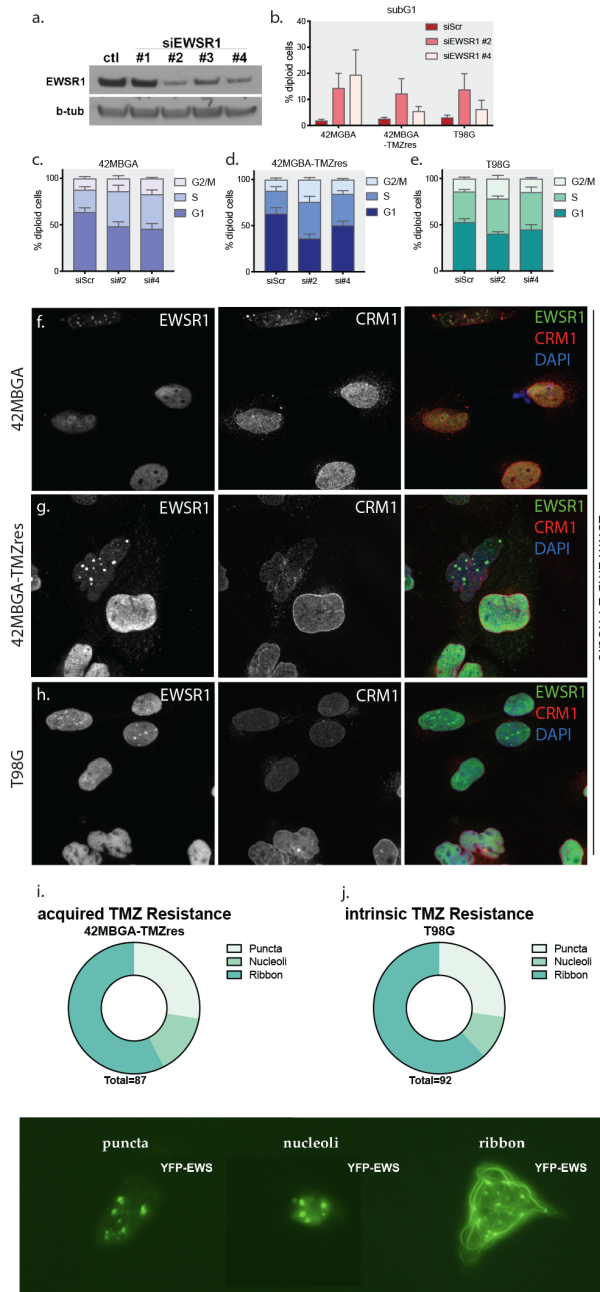


Supplementary Figure 4



enhanced*

Supplementary Figure 5



Supplementary Figure 6

