

Supplemental Information S4: Mechanistically Coupled PK (MCPK) model to describe enzyme induction and occupancy dependent DDI of dabrafenib metabolism

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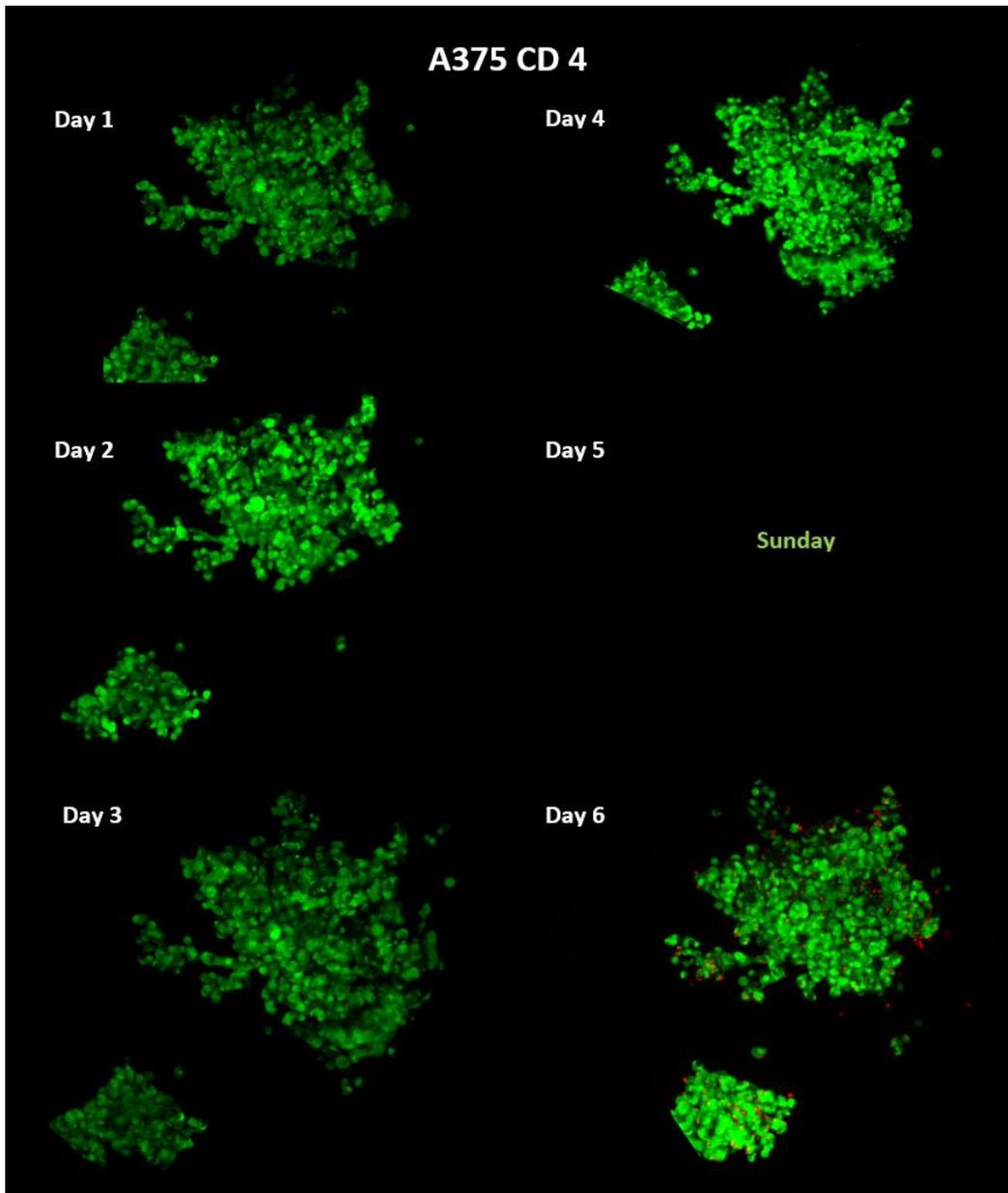
Spheroid Image Data of A375 Spheroid Growth

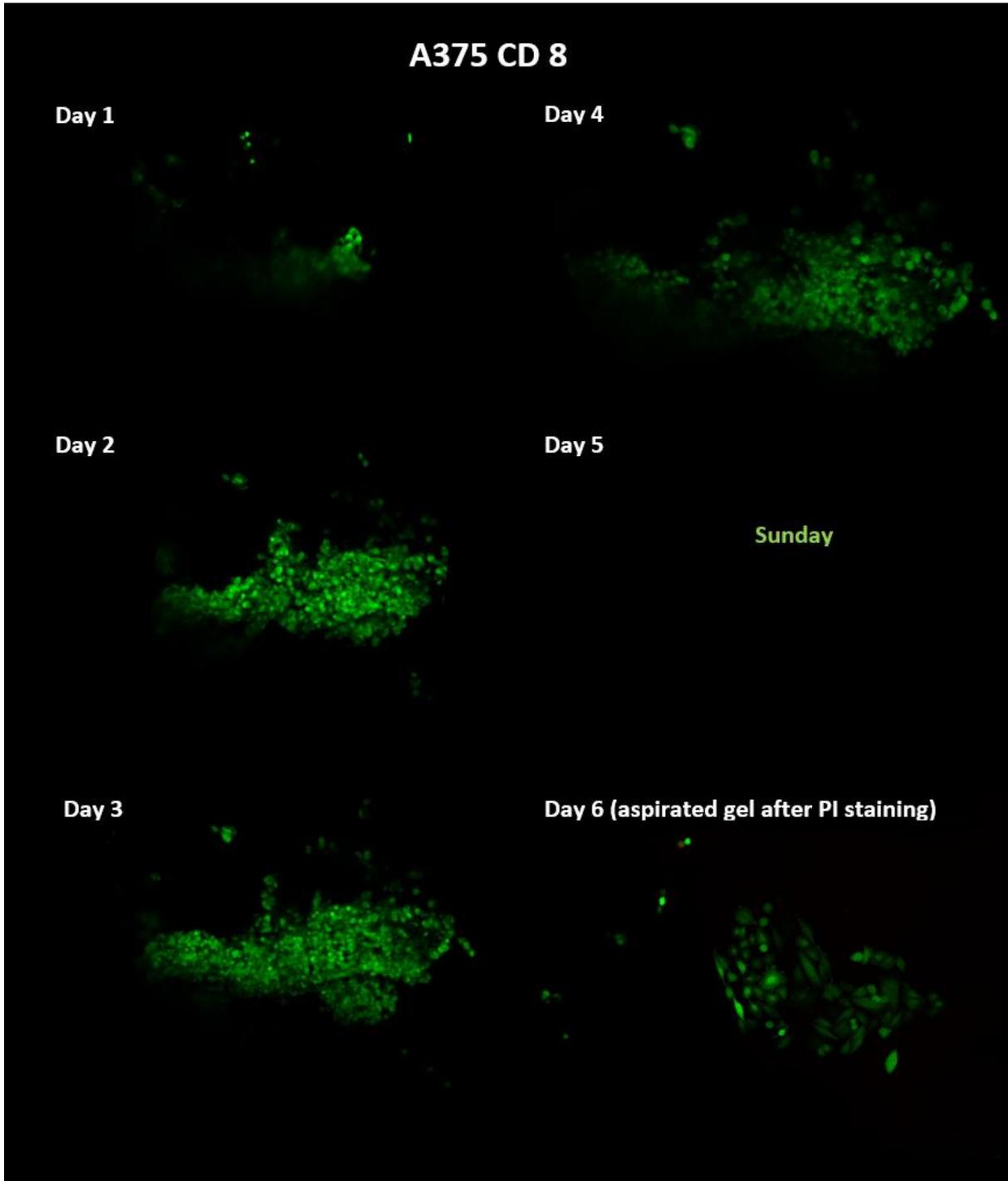
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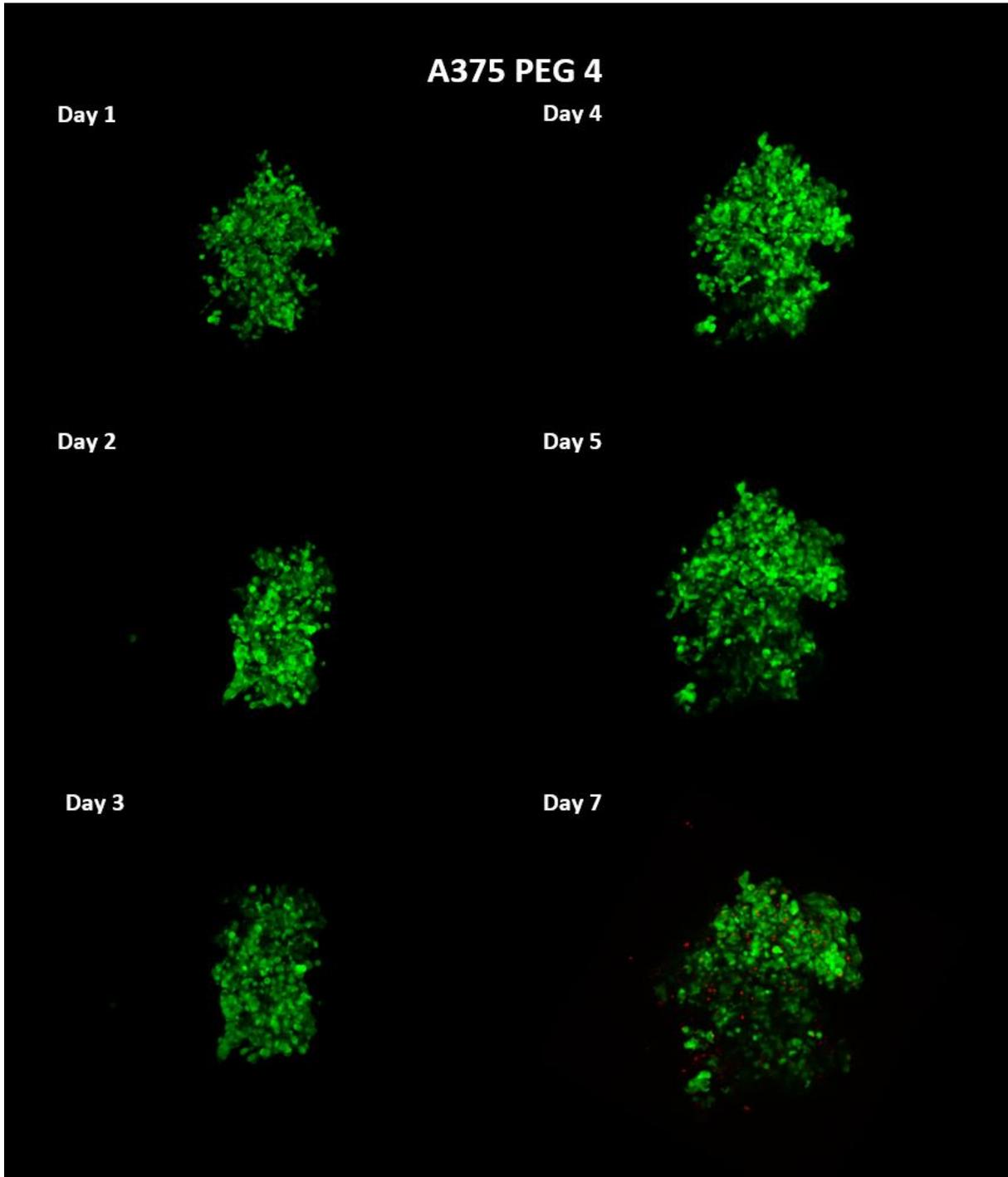
Legend

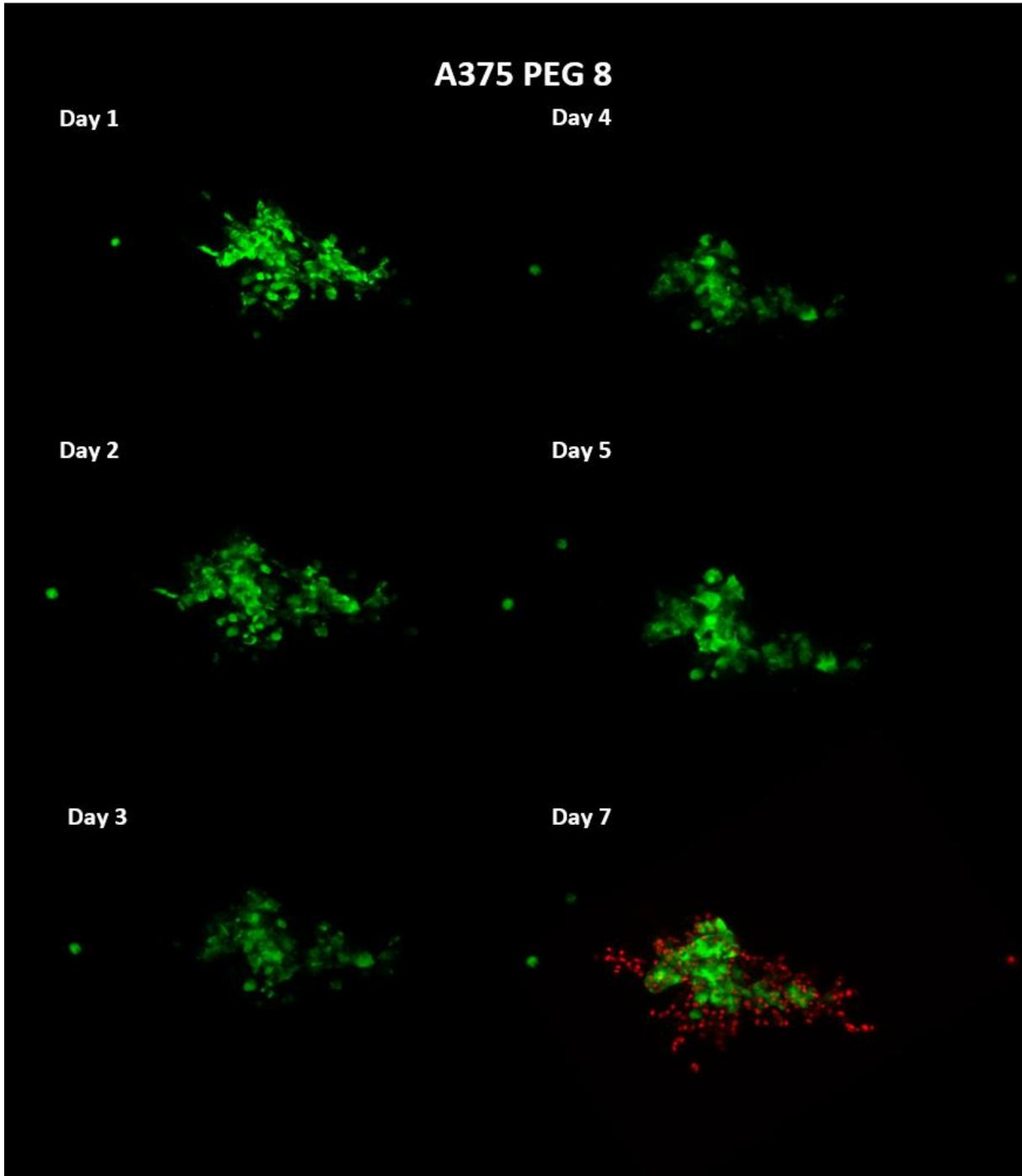
- CD: Gel with cell degradable linker
- PEG: Gel with non-degradable polyethylene glycol linker
- R1-3: Biological Replicate 1 to 3.
- A375: Metastatic Melanoma Cell Line with **green** fluorescent protein
- PI: Propidium iodide as a **red** cell death marker at last day

A375– First Iteration



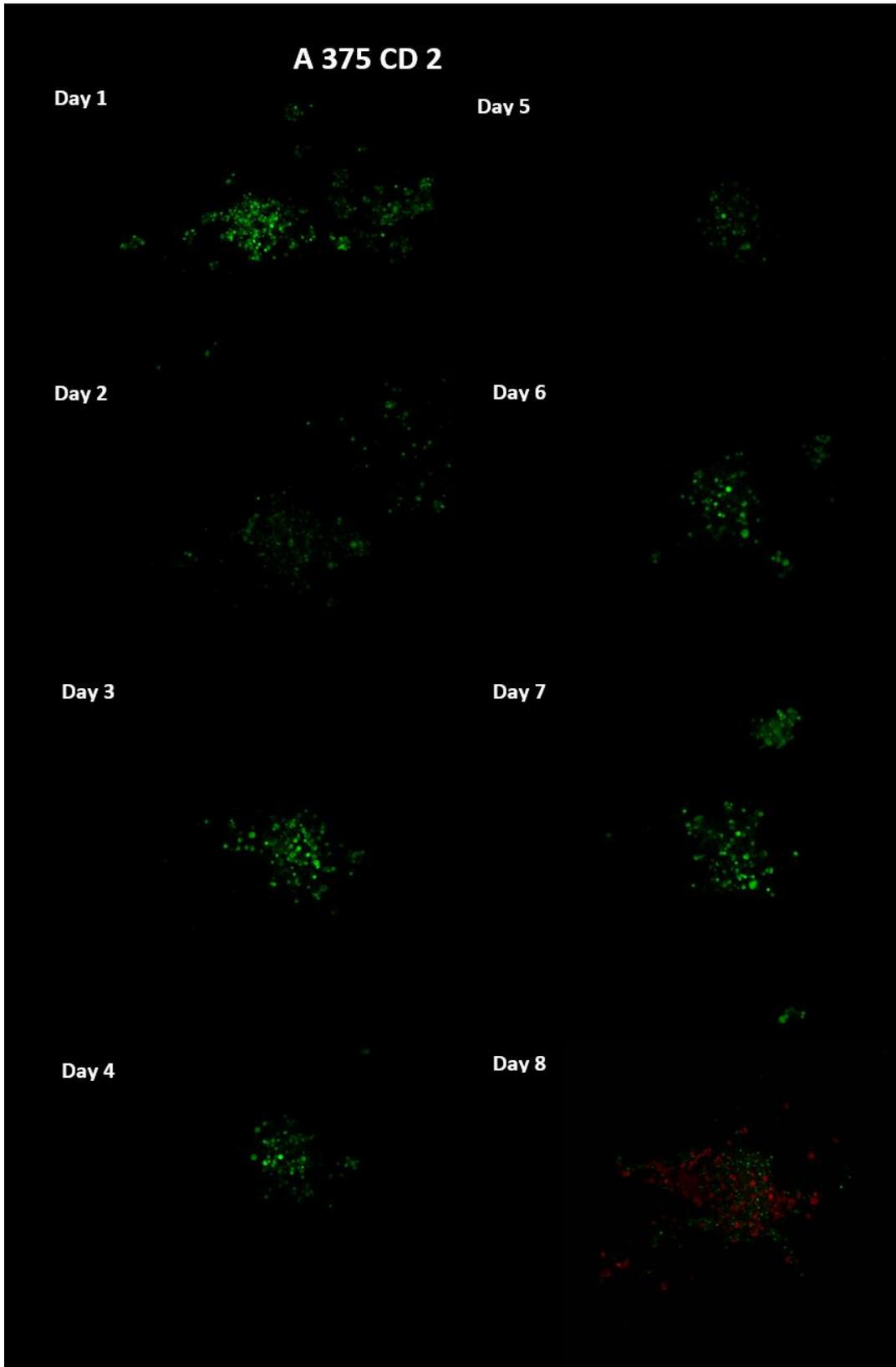


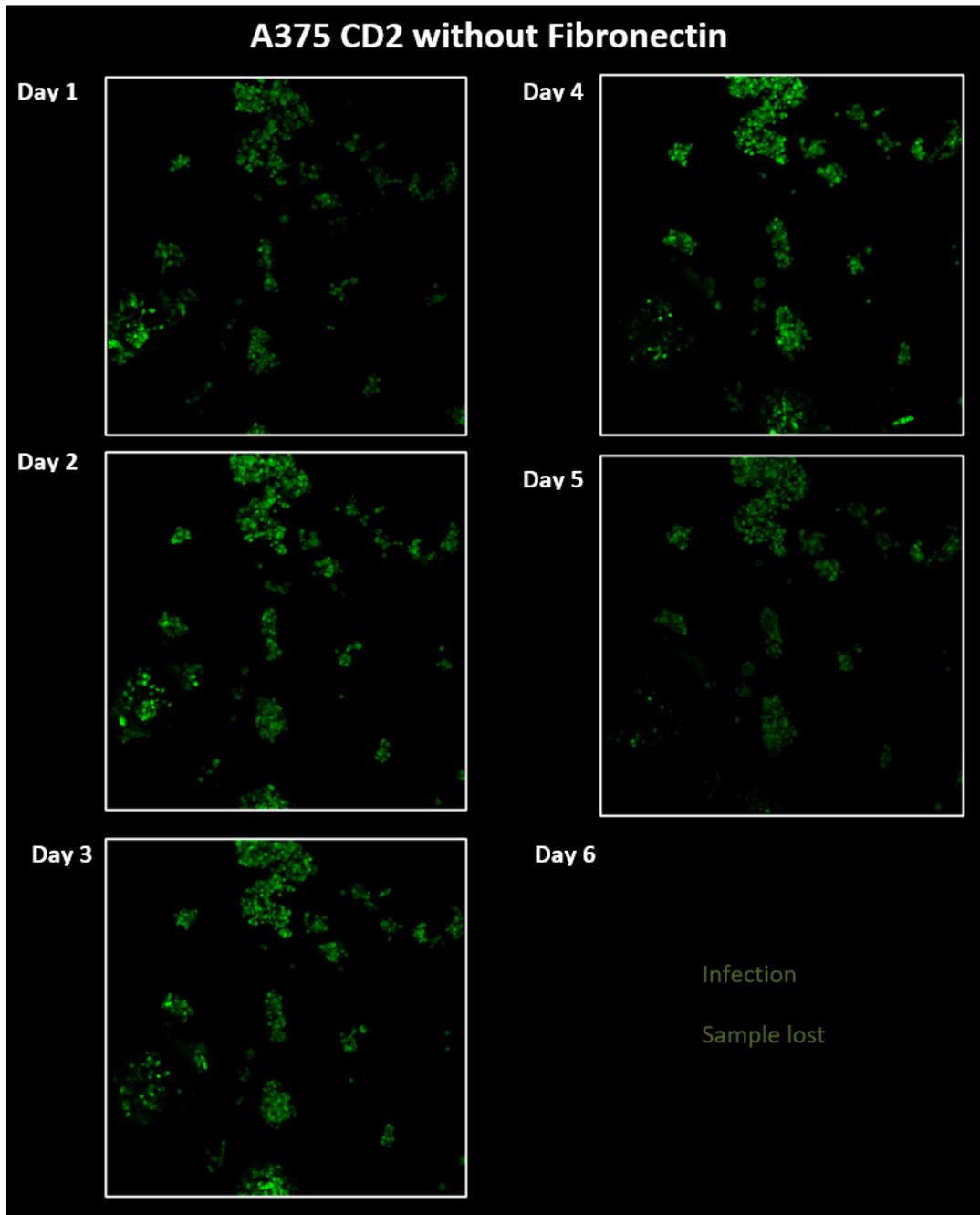


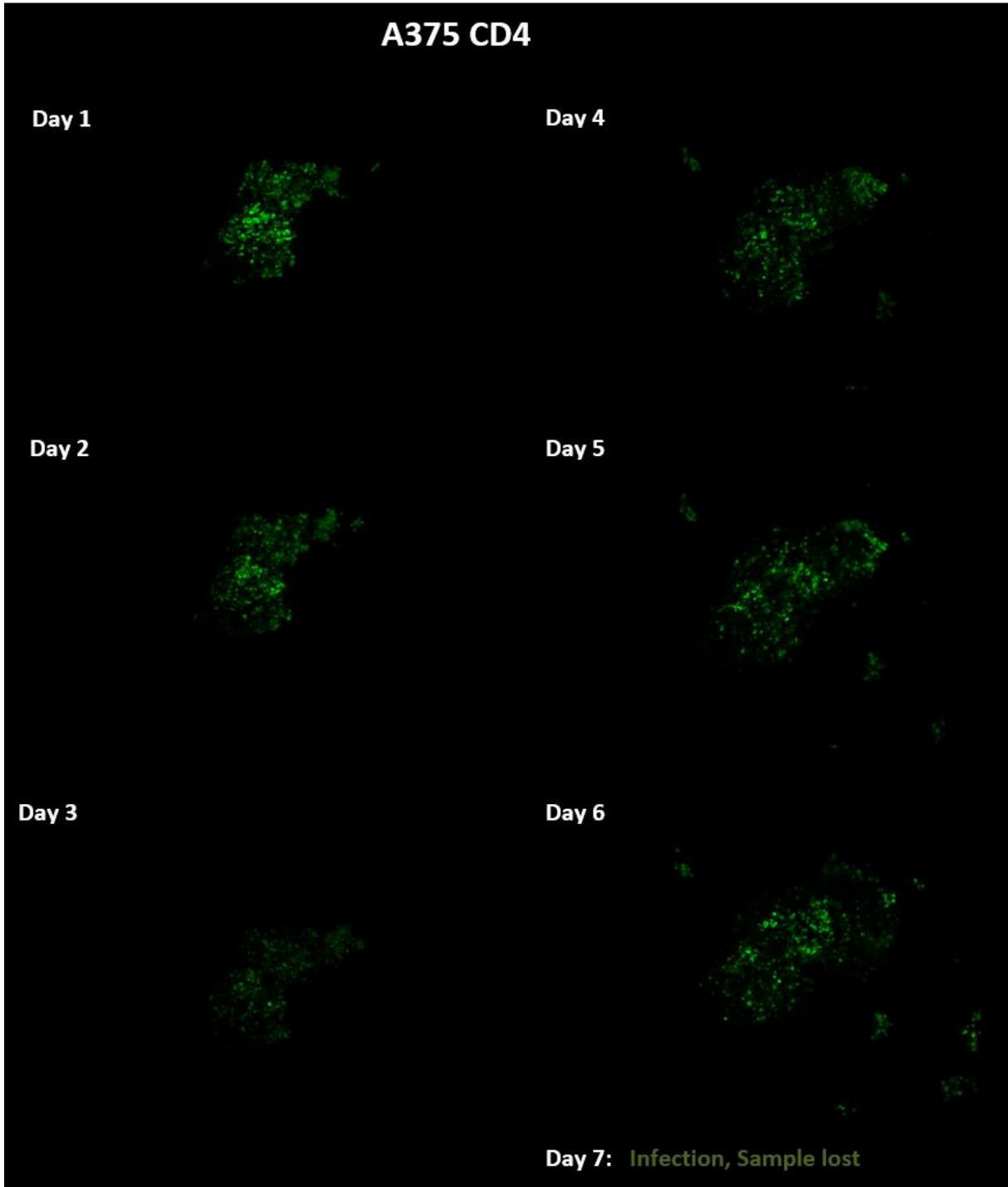


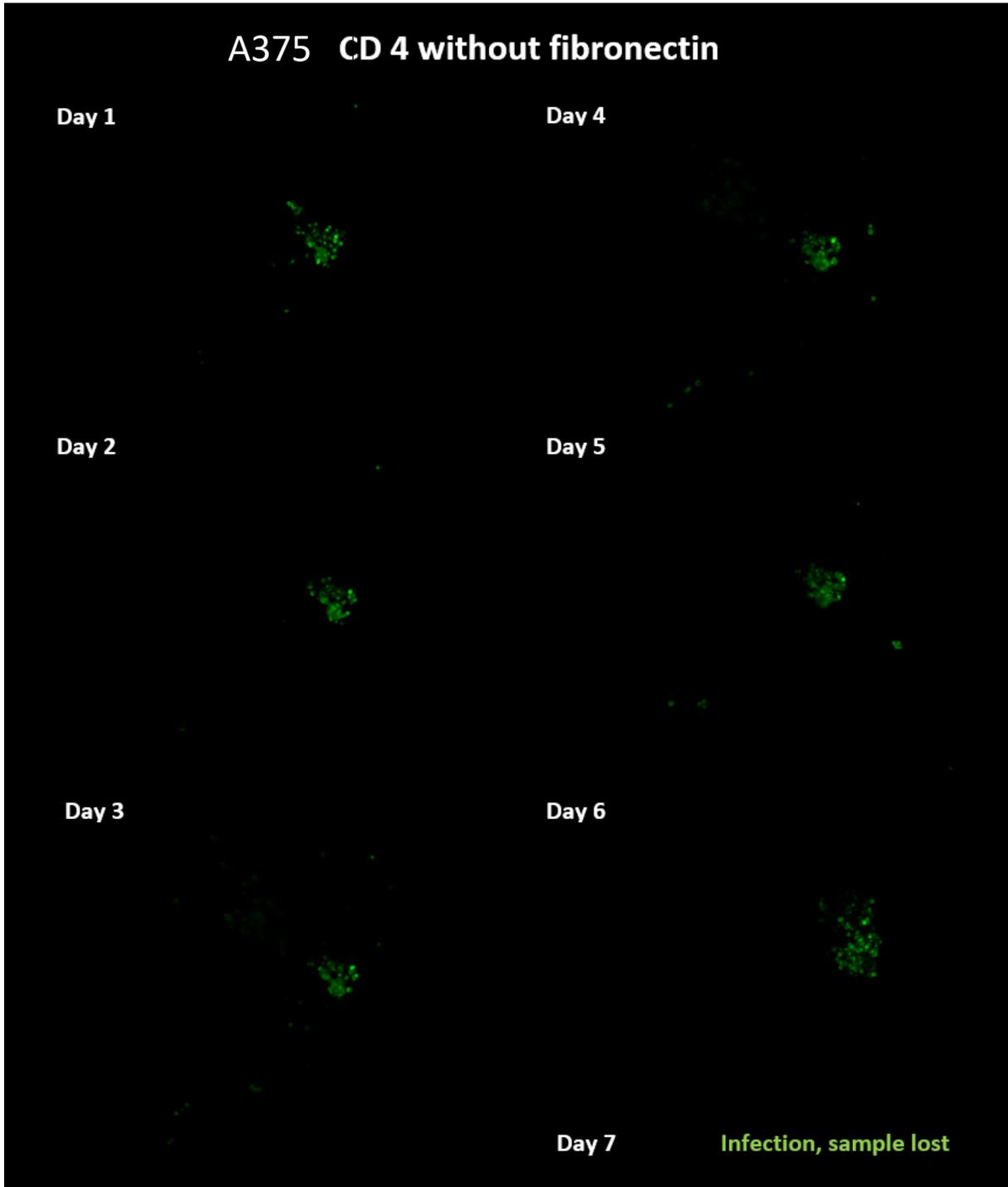
A375– Second Iteration (with or without Fibronectin)

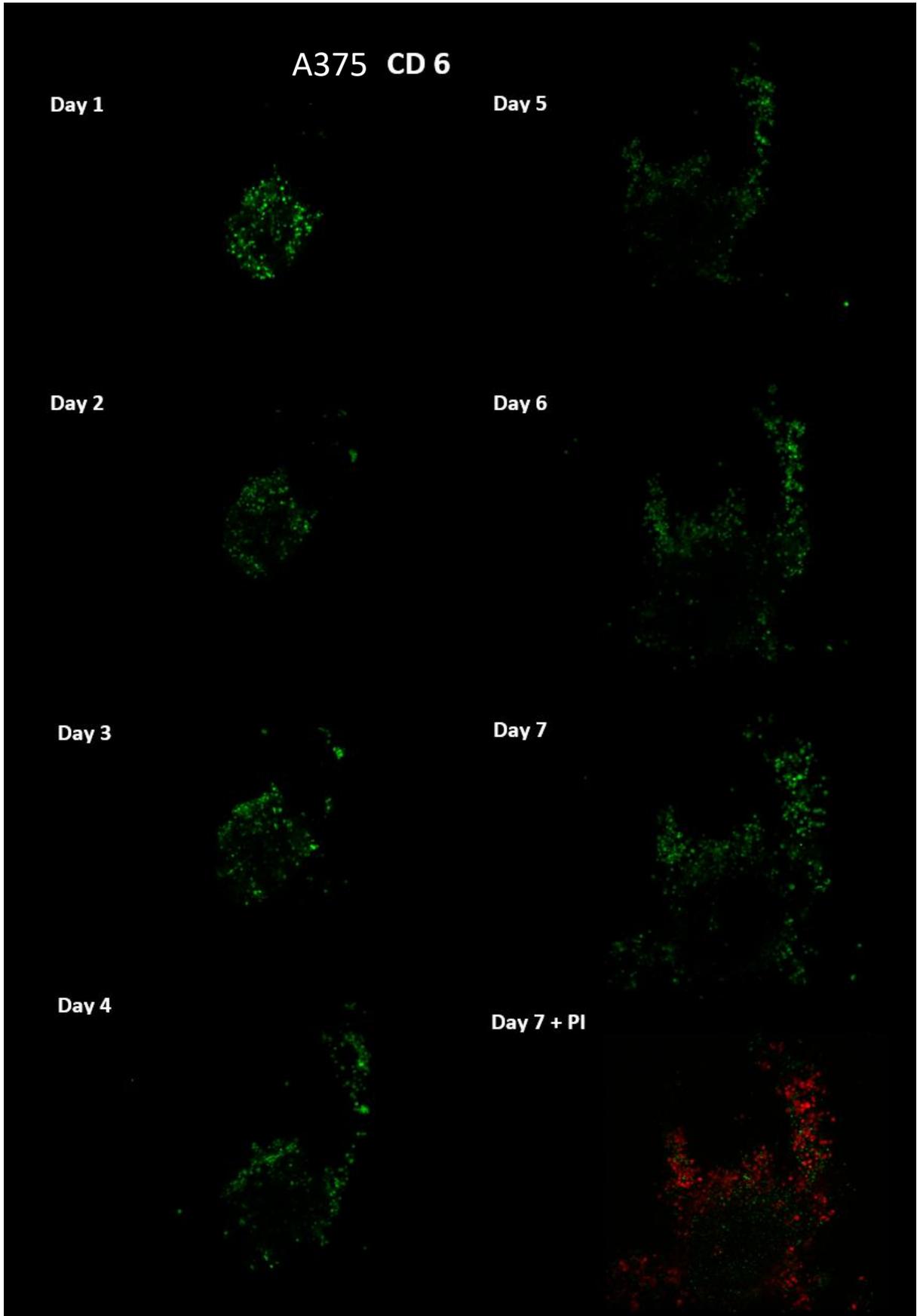
2	2 without Fibronectin	4	4 without Fibronectin	6	8	Ingredients	PEG Link	CD-Link
14.8	16.6	10.7	12.5	5.7	0.7	Water	539	539
2.4	2.4	2.5	2.5	2.5	2.5	10xCB PH 7.2	580	485
2	2	4	4	6	8	Thiol-reactive polymer	544	544
1.8	0	1.8	0	1.8	1.8	Fibronectin	SLBP0436V	SLBP0436V
6	6	5	5	5	5	Cell suspension		
3	3	6	6	9	12	CD or PEG Link	565	553
30	30	30	30	30	30			

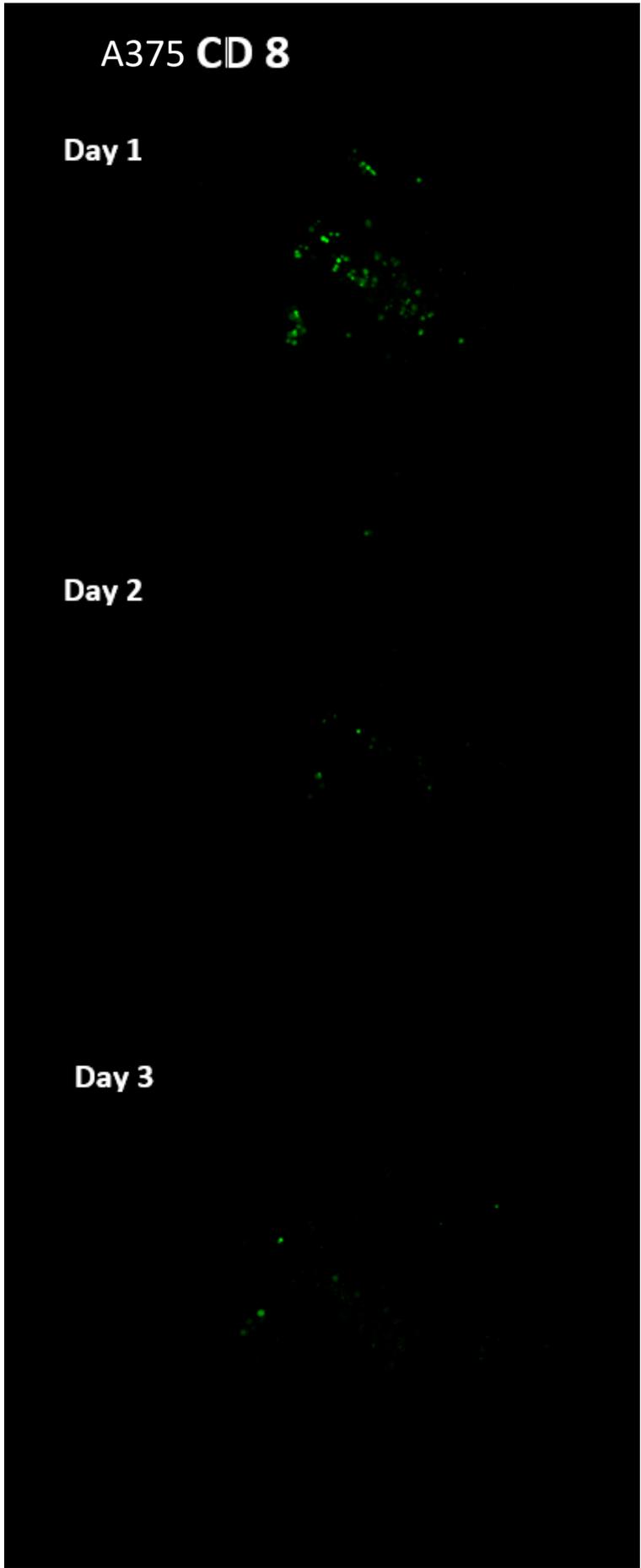


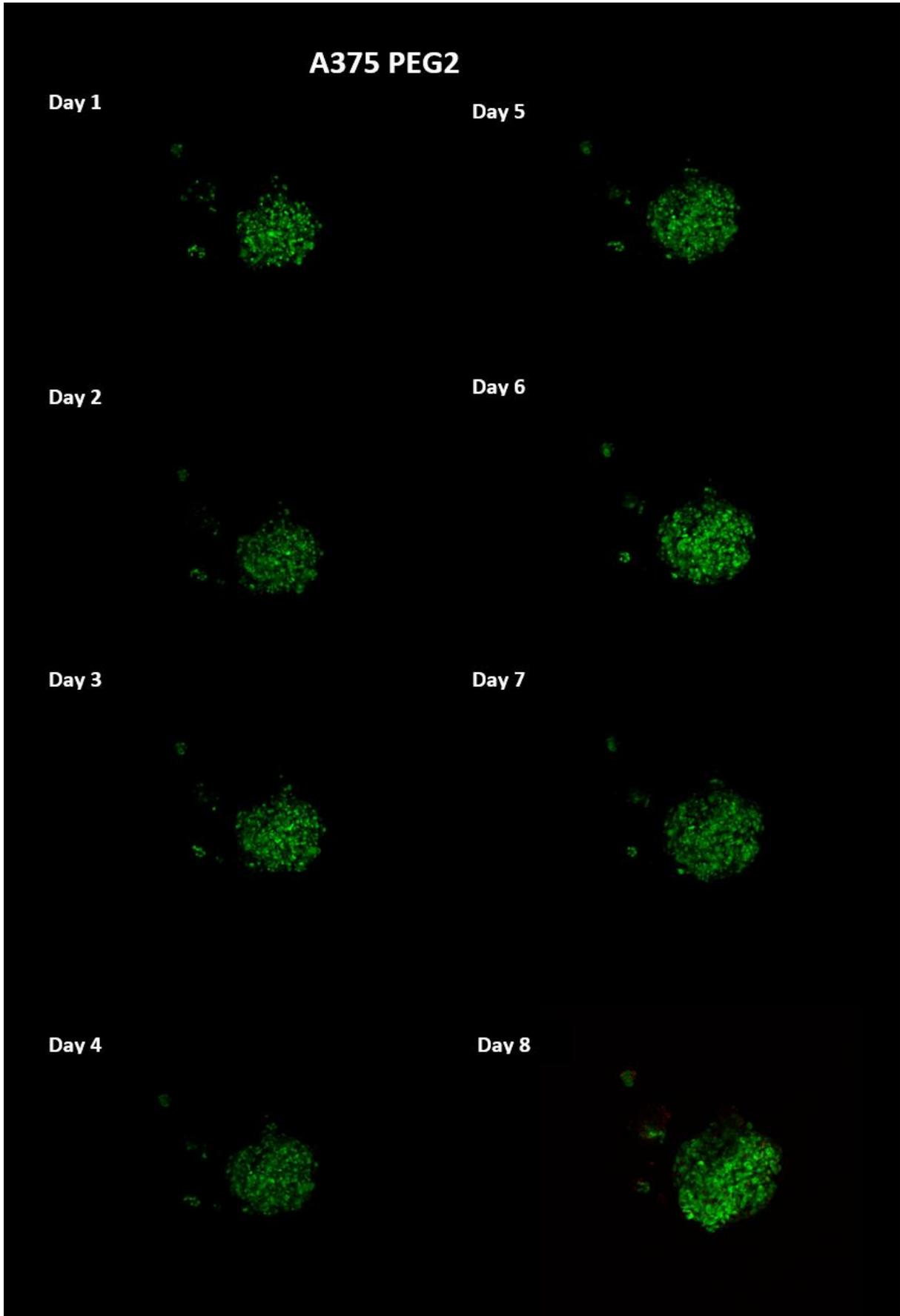


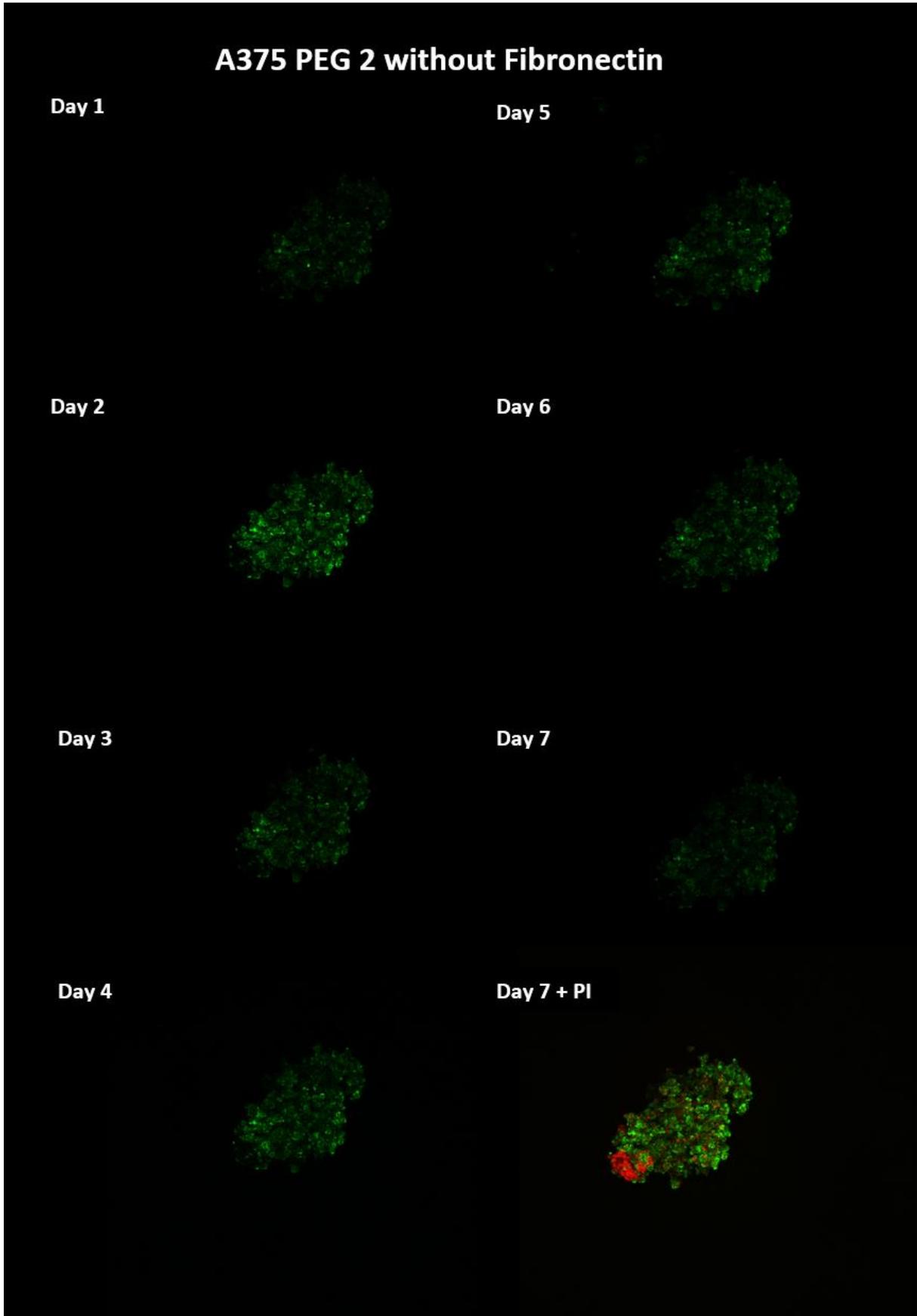


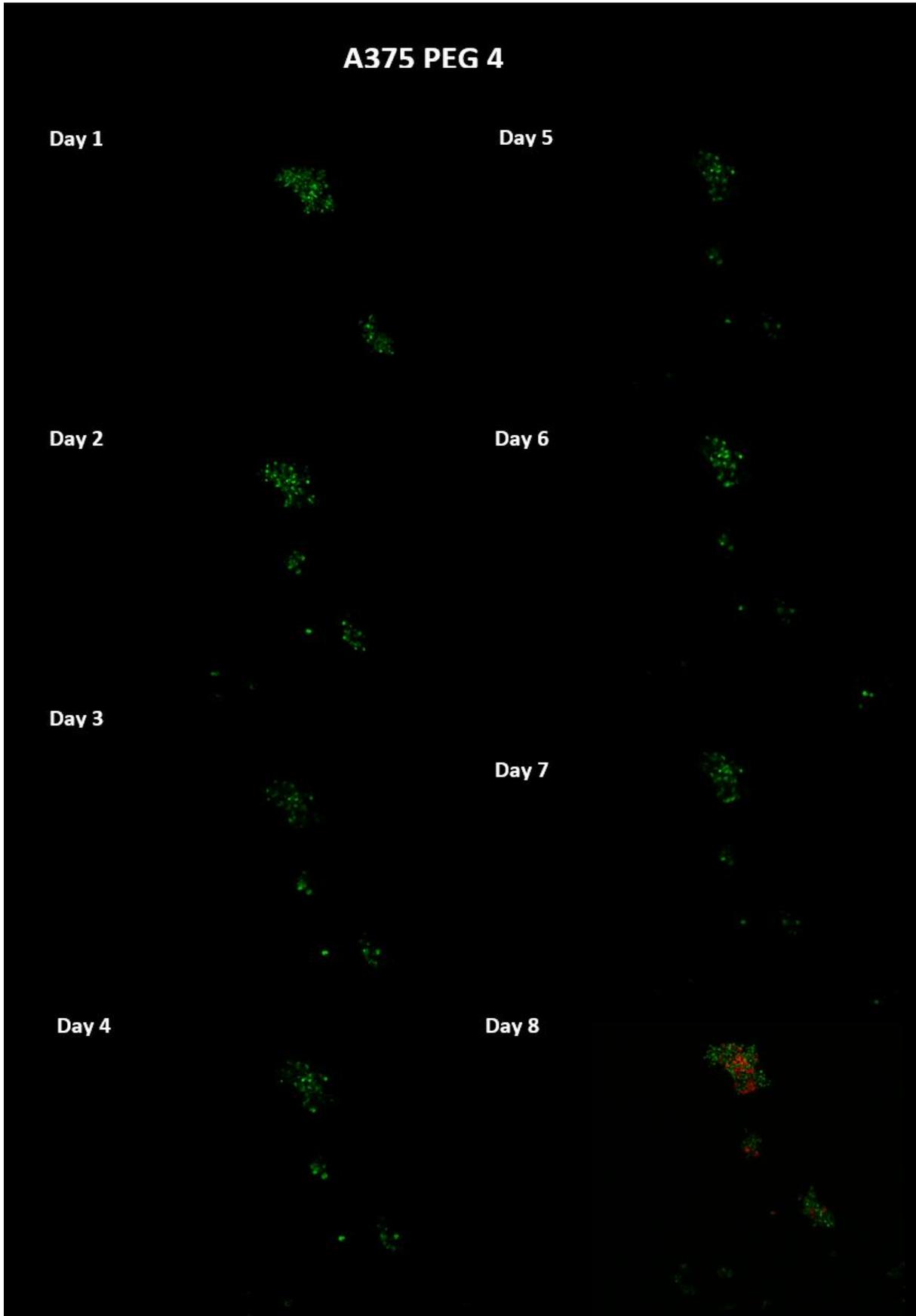


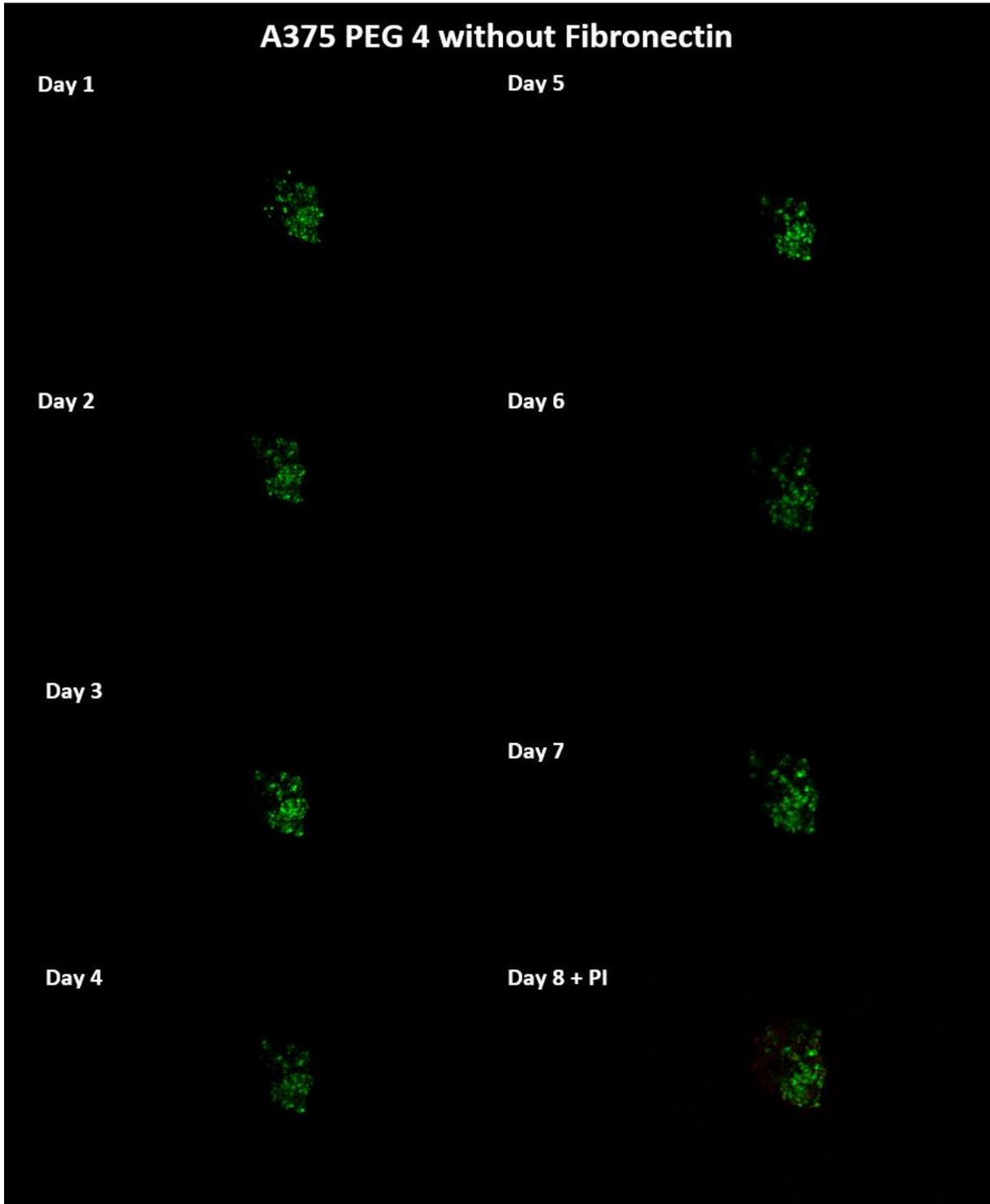


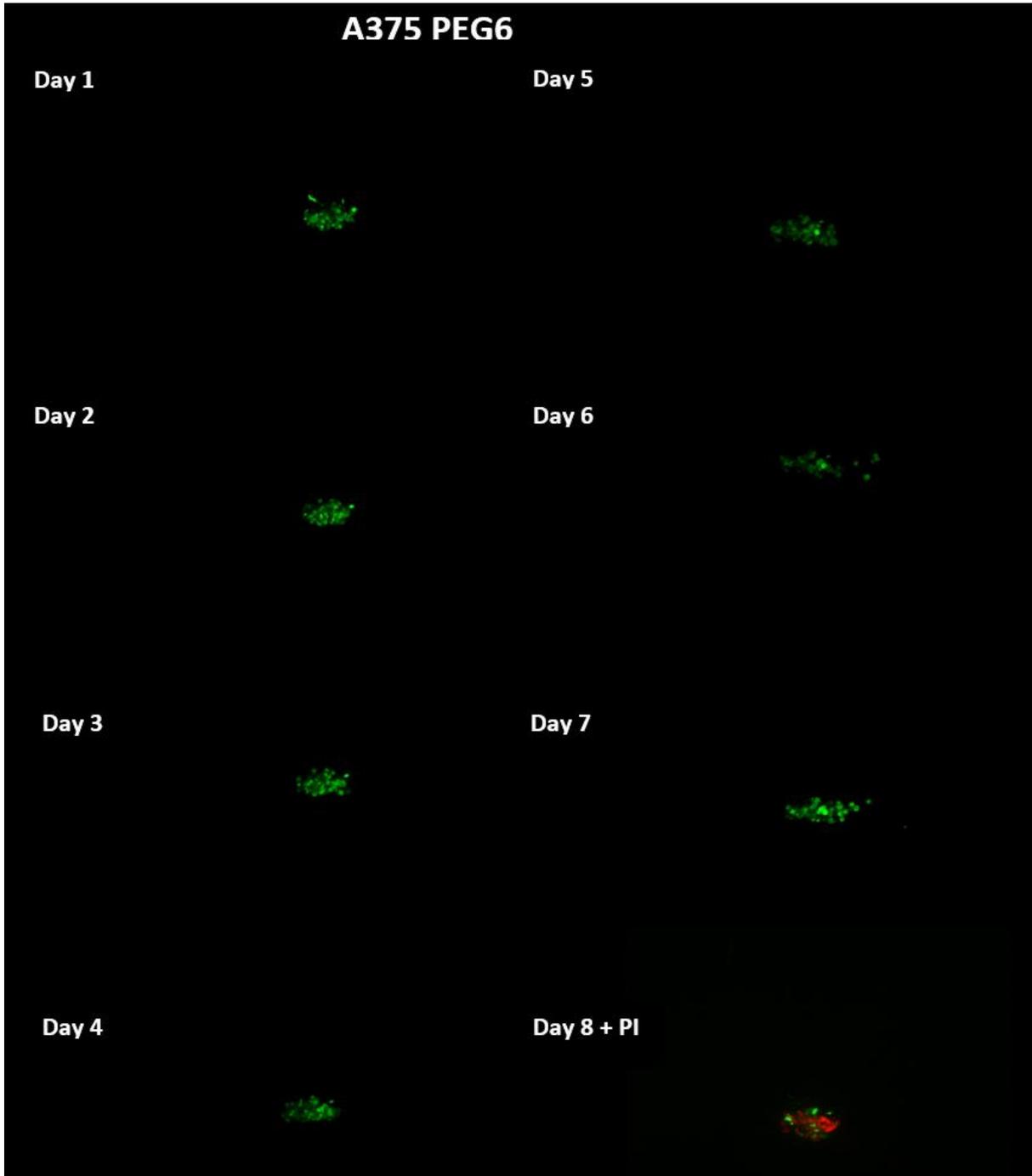


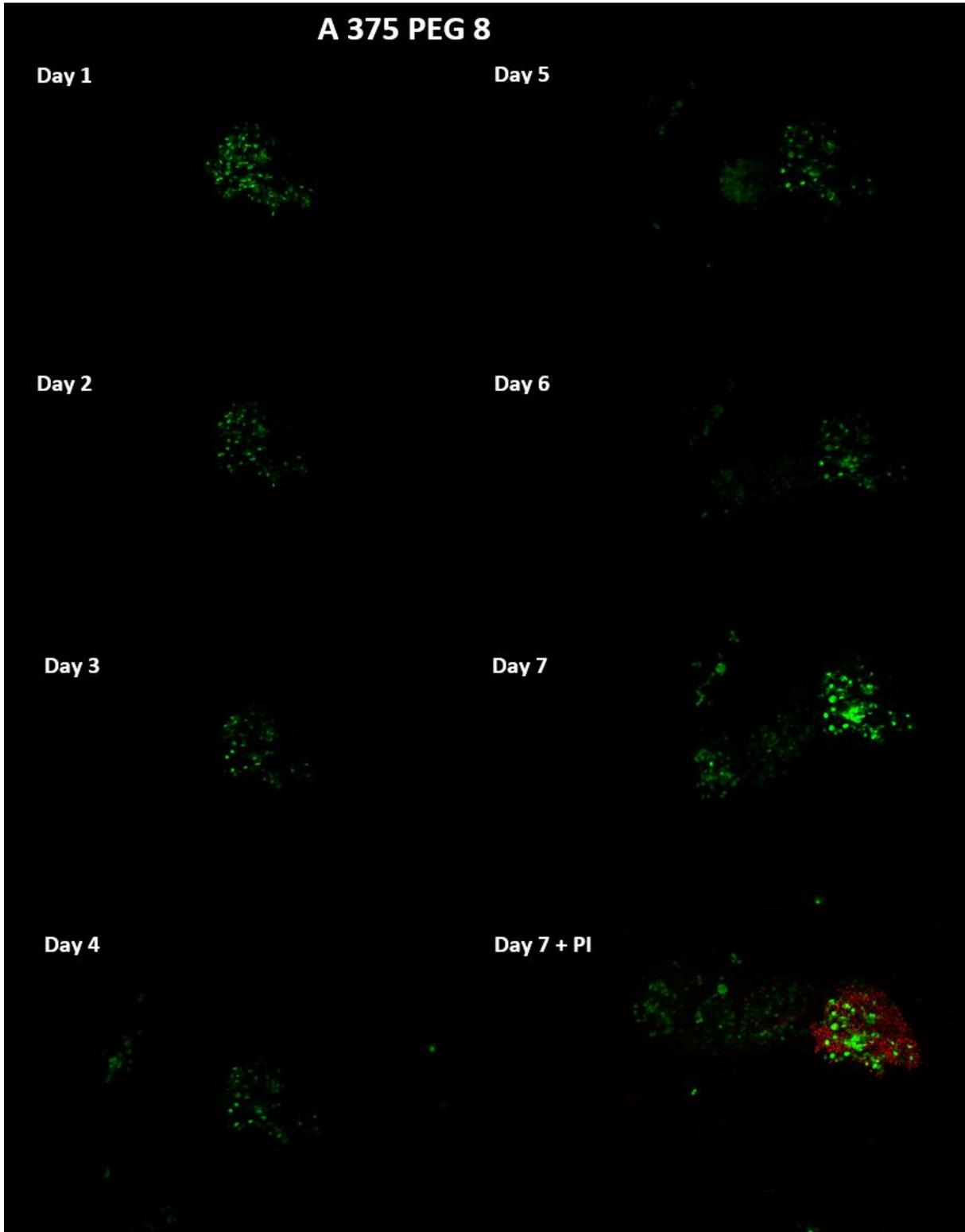












A375– Third Iteration

