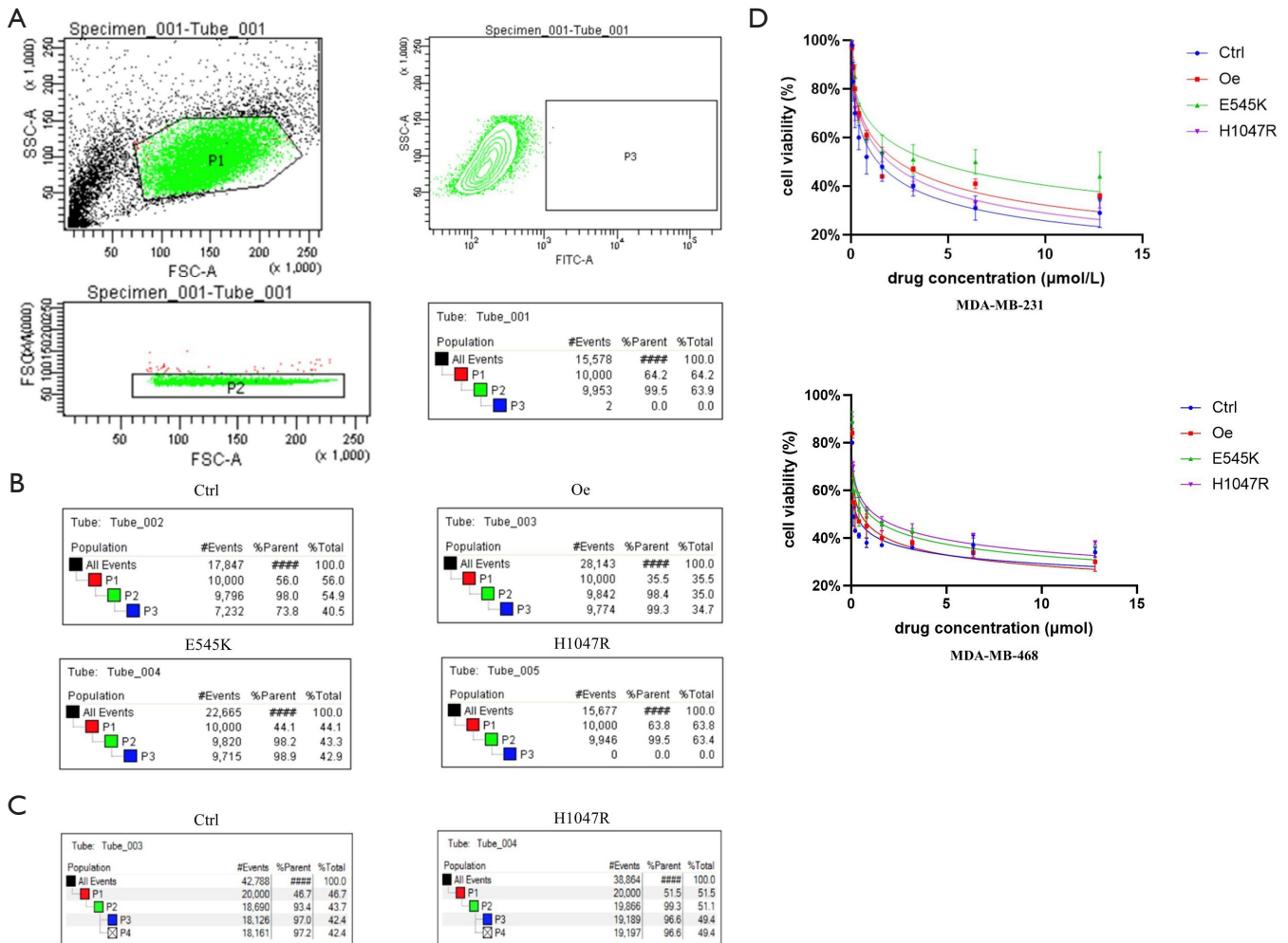


**Table S1** The sequences of the specific primers for PIK3CA<sup>Oe</sup>, PIK3CA<sup>E545K</sup>, and PIK3CA<sup>H1047R</sup> in plasmids

Primer	Sequence (5' to 3')
PIK3CA <sup>Oe</sup>	Forward: GAGGATCCCCGGGTACCGGTGCGCCACCATGCCTCCACGACCATCATCAGGTGAAC
	Reverse: TCCTTGTAGTCCATACCGTTCAATGCATGCTGTTAATTGTGTG
PIK3CA <sup>E545K</sup>	Forward: GAGGATCCCCGGGTACCGGTGCGCCACCATGCCTCCACGACCATCATCAGGTGAAC
	Reverse: TCCTTGTAGTCCATACCGTTCAATGCATGCTGTTAATTGTGTG
PIK3CA <sup>H1047R</sup>	Forward: GAGGATCCCCGGGTACCGGTGCGCCACCATGCCTCCACGACCATCATCAGGTG
	Reverse: TCCTTGTAGTCCATACCGTTCAATGCATGCTGTTAATTGTGTGGAAGATCCAATCCATTTTTGTG



**Figure S1** Isolation of single cell populations and IC<sub>50</sub> analysis for epirubicin. (A) The procedure for isolating and characterizing transfected cells (e.g., MDA-MB-231, wild-type). The P3 section represents the cells isolated. In this figure, few cells can be seen in the P3 section, since wild-type cells did not carry the GFP protein. (B) Initial isolation of transfected populations of MDA-MB-231 cells. Proportion of cells carrying GFP: 73.8% for PIK3CA<sup>ctrl</sup>, 99.3% for PIK3CA<sup>Oe</sup>, 98.9% for PIK3CA<sup>E545K</sup>, and 0% for PIK3CA<sup>H1047R</sup>. (C) Second isolation of PIK3CA<sup>ctrl</sup> and PIK3CA<sup>H1047R</sup> MDA-MB-231 cells for their low proportions in initial isolation. Proportion of cells carrying GFP: 97.2% for PIK3CA<sup>ctrl</sup> and 96.6% for PIK3CA<sup>H1047R</sup>. (D) The drug susceptibility of each cell line was determined by cellular drug resistance assay. Cell viability was measured under different concentrations of epirubicin, and each IC<sub>50</sub> value is listed in Table S2.

**Table S2** The IC<sub>50</sub> value for epirubicin in each cell group (μmol/L)

Cell lines	PIK3CA <sup>Ctrl</sup>	PIK3CA <sup>Oe</sup>	PIK3CA <sup>E545K</sup>	PIK3CA <sup>H1047R</sup>
MDA-MB-231	1.38	2.43	4.23	1.97
MDA-MB-468	0.27	0.50	0.95	1.23

**Table S3** Information of primary antibodies used in Western blot

Antibody	Dilution	Catalogue number	Manufacturer
PI3 Kinase p110α	1:1,000	4249	CST (Cell Signaling Technology)
AKT (pan)	1:2,000	2920	CST
p-AKT (Ser473)	1:2,000	4060	CST
p-AKT (Thr308)	1:1,000	13038	CST
p-mTOR (Ser2448)	1:1,000	5536	CST
p-p70 S6 Kinase (Thr389)	1:1,000	9206	CST
p-p70 S6 Kinase (Ser 371)	1:1,000	9208	CST
p-4E-BP1 (Thr37/46)	1:1,000	2855	CST
Xiap	1:1,000	14334	CST
Bcl-2	1:1,000	3498	CST
β-actin	1:200	47778	Santa Cruz
mTOR	1:200	517464	Santa Cruz
Pten	1:200	7974	Santa Cruz

**Table S4** The sequences of the specific primer for mTOR, Pten, PIK3CA, AKT, EIF4EBP1 and β-actin in RT-qPCR

Primer	Sequence (5' to 3')
mTOR	Forward: GCCGCGGAATATTAAGGAA
	Reverse: TGGTTTCTCATTCCGGCTC
Pten	Forward: ACCCACCACAGCTAGAACTT
	Reverse: GGAATAGTTACTCCCTTTTGTGTC
PIK3CA	Forward: TTACCCTTCTGCCGGAGG
	Reverse: AAGTGGATGCCCCACAGTTC
AKT	Forward: GAAGGACGGGAGCAGGC
	Reverse: CTCACGCGCTCCTCTCAG
EIF4EBP1	Forward: GGAGTGTCGGAACCTCACCTG
	Reverse: ACACGATGGCTGGTGCTTTA
β-actin	Forward: CCGTTCCGAAAGTTGCCTTTT
	Reverse: ATCATCCATGGTGAGCTGGC