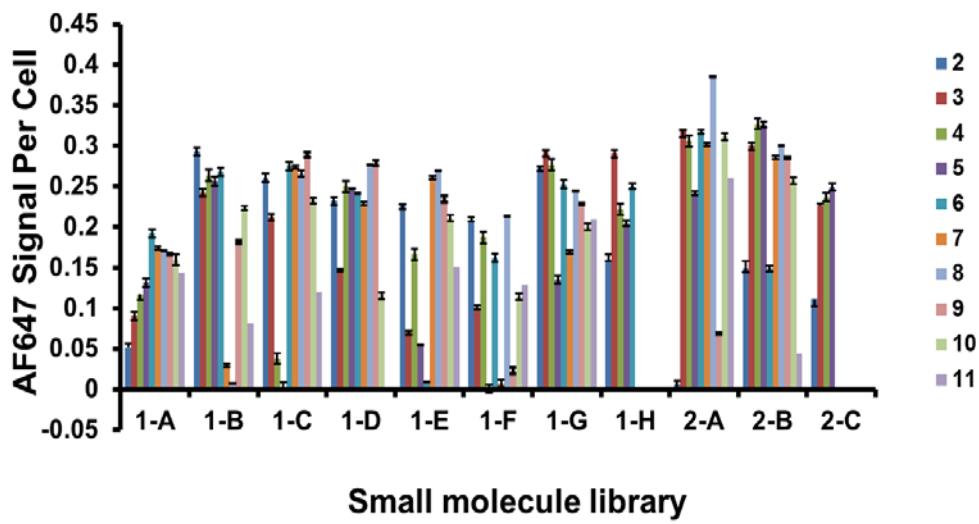


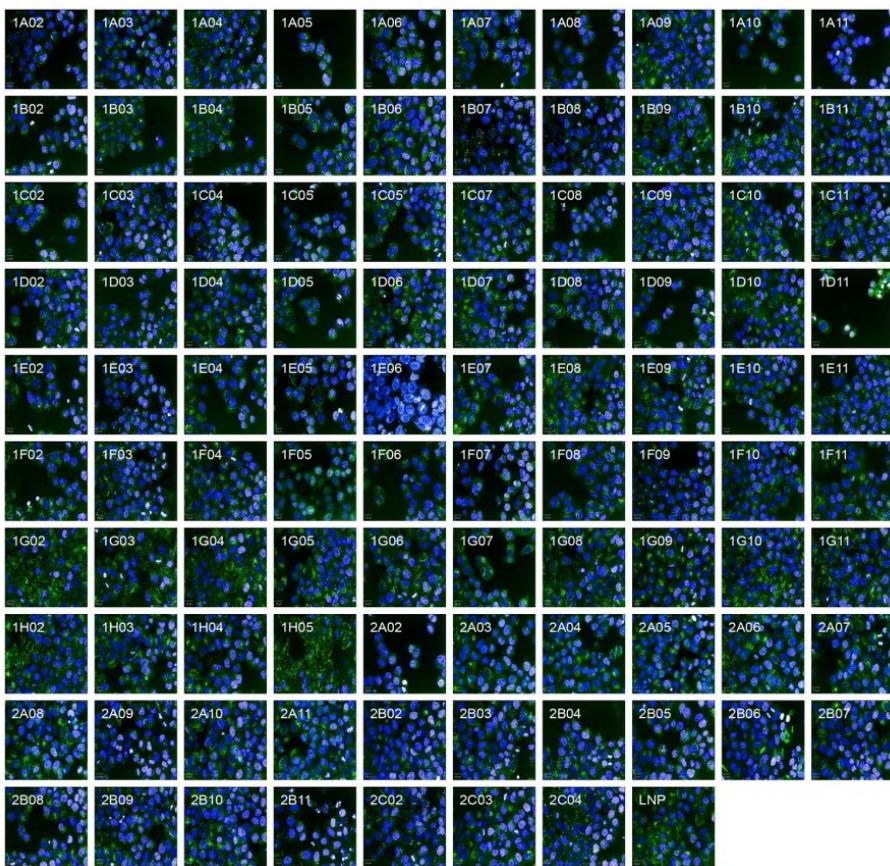
| Small Molecules | Mwt | Sample ID | Plate num | INDUCER | CAS Datab | Enzo ID | Stock | Cell signaling |
|------------------------|--------|-----------|-----------|--------------------|-------------|--------------|-------|---|
| Rapamycin | 913.6 | 1-A02 | 1 | Autophagy inducers | 53123-88-9 | BML-A275 | 10mM | mTOR inhibitor |
| Timosaponin A-III | 740.4 | 1-A03 | 1 | Autophagy inducers | 41059-79-4 | BML-AP504 | 10mM | mTOR inhib/ER stress |
| NVP-BEZ235 | 469.2 | 1-A04 | 1 | Autophagy inducers | 915019-65-7 | AC1502 | 10mM | Dual PI3K/mTOR Inhibitor |
| PI-103 | 348.1 | 1-A05 | 1 | Autophagy inducers | 371935-74-5 | ALX-270-460 | 10mM | Dual PI3K/mTOR inhib |
| AS605240 | 257.0 | 1-A06 | 1 | Autophagy inducers | 648450-29-7 | 462 BML- | 10mM | PI3Kg inhibitor |
| Lithium Chloride | 42.0 | 1-A08 | 1 | Autophagy inducers | 7447-41-8 | AC1503 | 10mM | IMPAse inhibitor/GSK-3b inhib |
| L-690,330 | 298.0 | 1-A07 | 1 | Autophagy inducers | 142523-38-4 | ALX-270-078 | 10mM | IMPAse inhibitor |
| Carbamazepine | 236.1 | 1-A09 | 1 | Autophagy inducers | 298-46-4 | 084 ALX-550- | 10mM | inhibits IP3 via inhibition of inositol synth |
| Sodium Valproate | 166.1 | 1-A10 | 1 | Autophagy inducers | 1069-66-5 | 304 ALX-550- | 10mM | inhibits IP3 via inhibition of inositol synth |
| Verapamil-HCl | 490.3 | 1-A11 | 1 | Autophagy inducers | 152-11-4 | 306 ALX-550- | 10mM | Ca2+ Blocker |
| Loperamide-HCl | 512.2 | 1-B02 | 1 | Autophagy inducers | 34552-83-5 | 253 BML- | 10mM | Ca2+ Blocker |
| Amiodarone-HCl | 681.0 | 1-B03 | 1 | Autophagy inducers | 1951-25-3 | AC105 | 10mM | Ca2+ Blocker/mTORC1 inhib. |
| Nimodipine | 418.2 | 1-B04 | 1 | Autophagy inducers | 66085-59-4 | 277 ALX-550- | 10mM | Ca2+ Blocker |
| Nitrendipine | 360.1 | 1-B05 | 1 | Autophagy inducers | 39562-70-4 | 278 | 10mM | Ca2+ Blocker |
| Niguldipine | 645.3 | 1-B06 | 1 | Autophagy inducers | 102993-22-6 | BML-CA216 | 10mM | Ca2+ Blocker(T-type) |
| Penitrem A | 637.2 | 1-B07 | 1 | Autophagy inducers | 12627-35-9 | BML-KC157 | 10mM | Ca2+/K+ channel blocker |
| Ionomycin | 678.5 | 1-B08 | 1 | Autophagy inducers | 56092-81-0 | 006 ALX-350- | 10mM | Ca2+ ionophore |
| Rotenone | 394.1 | 1-B09 | 1 | Autophagy inducers | 83-79-4 | 360 BML- | 10mM | mETC-complex1-inhib |
| TTFA | 222.0 | 1-B10 | 1 | Autophagy inducers | 326-91-0 | AC1504 | 10mM | mETC-complex2-inhib |
| Fluspirilene | 475.2 | 1-B11 | 1 | Autophagy inducers | 1841-19-6 | BML-AC116 | 10mM | Dopamine antagonist |
| Trifluoperazine-2HCl | 479.1 | 1-C02 | 1 | Autophagy inducers | 117-89-5 | 310 BML- | 10mM | Dopamine antagonist |
| Sorafenib tosylate | 636.1 | 1-C03 | 1 | Autophagy inducers | 475207-59-1 | AC1514 | 10mM | PDGFR3/FLT3/ER Stress |
| Niclosamide | 326.0 | 1-C04 | 1 | Autophagy inducers | 50-65-7 | AC1506 | 10mM | mTORC1 inhib |
| Rottlerin | 516.2 | 1-C05 | 1 | Autophagy inducers | 82-08-6 | 075 ALX-550- | 10mM | mTORC1 inhib |
| Caffeine | 194.1 | 1-C06 | 1 | Autophagy inducers | 58-08-2 | 322 ALX-270- | 10mM | mTor pathway inhibition |
| Metformin-HCl | 165.1 | 1-C07 | 1 | Autophagy inducers | 1115-70-4 | 432 ALX-550- | 10mM | mTORC1 inhib RAG/GTPase |
| Clonidine-HCl | 265.0 | 1-C08 | 1 | Autophagy inducers | 4205-90-7 | 087 | 10mM | I1R agonist |
| Rilmenidine | 296.1 | 1-C09 | 1 | Autophagy inducers | 54187-04-1 | BML-AC154 | 10mM | I1R agonist, |
| 2',5'-Dideoxyadenosine | 235.1 | 1-C10 | 1 | Autophagy inducers | 6698-26-6 | CN110 | 10mM | adenylyl cyclase inhib. |
| Suramin-6Na | 1427.9 | 1-C11 | 1 | Autophagy inducers | 129-46-4 | 022 | 10mM | Gas inhibitor |
| Pimozide | 461.2 | 1-D02 | 1 | Autophagy inducers | 2062-78-4 | BML-AC109 | 10mM | D1,D2,D3 adrenergic blocker |
| STF-62247 | 267.1 | 1-D03 | 1 | Autophagy inducers | 315702-99-5 | BML-AP501 | 10mM | trans golgi network |
| Spermidine | 145.2 | 1-D04 | 1 | Autophagy inducers | 124-20-9 | AC1515 | 10mM | inhibits HATs |
| FK-866 | 391.2 | 1-D05 | 1 | Autophagy inducers | 658084-64-1 | 501 ALX-550- | 10mM | inhibits NAD |
| Tamoxifen citrate | 563.3 | 1-D06 | 1 | Autophagy inducers | 54965-24-1 | 095 | 10mM | inhibits Ceramide |
| Minoxidil | 209.1 | 1-D07 | 1 | Autophagy inducers | 38304-91-5 | BML-KC125 | 10mM | K+ channel opener |
| Imiquimod | 240.1 | 1-D08 | 1 | Autophagy inducers | 99011-02-6 | ALX-420-035 | 10mM | TLR7 |
| Imatinib mesylate | 589.2 | 1-D09 | 1 | Autophagy inducers | 220127-57-1 | 492 ALX-270- | 10mM | Tyrosine kinase inhib. |
| AG112 | 236.1 | 1-D10 | 1 | Autophagy inducers | 144978-82-5 | 133 | 10mM | Tyrosine kinase inhib. |
| SU11652 | 414.2 | 1-D11 | 1 | Autophagy inducers | 326914-10-7 | BML-E1408 | 10mM | Tyrosine kinase inhib. |
| SB202190 | 331.1 | 1-E02 | 1 | Autophagy inducers | 152121-30-7 | BML-E1294 | 10mM | p38 MAPK inhibitor |
| Brefeldin A | 280.2 | 1-E03 | 1 | Autophagy inducers | 20350-15-6 | BML-G405 | 10mM | ER stress |
| Tunicamycin | 702.3 | 1-E04 | 1 | Autophagy inducers | 11089-65-9 | BML-CC104 | 10mM | ER Stress |
| Thapsigargin | 650.3 | 1-E05 | 1 | Autophagy inducers | 67526-95-8 | BML-PE180 | 10mM | ER Stress/SERCA inhib. |
| Calcimycin | 523.3 | 1-E06 | 1 | Autophagy inducers | 52665-69-7 | BML-CA100 | 10mM | ER stress |
| Capsaicin | 305.2 | 1-E07 | 1 | Autophagy inducers | 404-86-4 | BML-E125 | 10mM | ER stress/p38,ERK inhib |
| Dihydrocapsaicin | 307.2 | 1-E08 | 1 | Autophagy inducers | 19408-84-5 | 052 BML- | 10mM | ROS accumulation |
| Glucosamine HCl | 215.1 | 1-E09 | 1 | Autophagy inducers | 66-84-2 | AC1513 | 10mM | mTOR |

| | | | | | ALX-280- | | | |
|---------------------------------|-------|-----------|-----------|--------------------|-------------|-----------|-------|--|
| | | | | | 001 | | | |
| | | | | | BML- | | | |
| DTT | 154.0 | 1-E10 | 1 | Autophagy inducers | 3483-12-3 | | 10mM | ER stress/lysosome degradInhb |
| Deoxycholate-Na | 414.3 | 1-E11 | 1 | Autophagy inducers | 302-95-4 | AC1508 | 10mM | Beclin1- ER stress/DNA damage |
| HA 14-1 | 408.0 | 1-F02 | 1 | Autophagy inducers | 65673-63-4 | ALX-430- | 10mM | Bcl-2 inhib. |
| Licochalcone A | 338.2 | 1-F03 | 1 | Autophagy inducers | 58749-22-7 | 124 | 10mM | Bcl-2/mTOR inhib. |
| Curcumin | 368.1 | 1-F04 | 1 | Autophagy inducers | 458-37-7 | ALX-350- | 10mM | Akt/mTOR/S6K path inhib |
| Plumbagin | 188.0 | 1-F05 | 1 | Autophagy inducers | 481-42-5 | BML-AP505 | 10mM | Akt/mTOR inhib. |
| 6-Gingerol | 294.2 | 1-F06 | 1 | Autophagy inducers | 23513-14-6 | BML-CA422 | 10mM | ROS induction |
| Akt Inhibitor X-HCl | 380.1 | 1-F07 | 1 | Autophagy inducers | 925681-41-C | AC1509 | 10mM | Akt Inhibitor |
| PMSF | 174.0 | 1-F08 | 1 | Autophagy inducers | 329-98-6 | ALX-270- | 10mM | Protease Inhibitor |
| MG132 | 475.3 | 1-F09 | 1 | Autophagy inducers | 133407-82-6 | BML-PI102 | 10mM | Proteasome inhibitor |
| ALLN | 383.3 | 1-F10 | 1 | Autophagy inducers | 110044-82-1 | BML-P120 | 10mM | Proteasome inhibitor |
| 7-Ketcholesterol | 400.3 | 1-F11 | 1 | Autophagy inducers | 566-28-9 | BML-EI308 | 10mM | Mitochondrial stress |
| 17-AAG | 585.3 | 1-G02 | 1 | Autophagy inducers | 75747-14-7 | AC1511 | 10mM | Hsp90 inhibitor |
| Geldanamycin | 560.3 | 1-G03 | 1 | Autophagy inducers | 30562-34-6 | BML-EI280 | 10mM | Hsp90 inhibitor |
| C1 | 242.1 | 1-G04 | 1 | Autophagy inducers | 143413-73-4 | BML-AP500 | 10mM | ROS/non-Beclin1 dependent |
| Z36 | 393.2 | 1-G05 | 1 | Autophagy inducers | N/A | BML-ST200 | 10mM | Bcl-XL inhibitor |
| Rockout | 194.1 | 1-G06 | 1 | Autophagy inducers | 7272-84-6 | ALX-270- | 10mM | Rho kinase Inhibitor |
| Go6850 | 412.2 | 1-G07 | 1 | Autophagy inducers | 133052-90-1 | BML-EI246 | 10mM | PKC Inhibitor |
| 2-Deoxyglucose | 164.1 | 1-G08 | 1 | Autophagy inducers | 154-17-6 | AC1512 | 10mM | Blocks glycolysis |
| Etoposide | 588.2 | 1-G09 | 1 | Autophagy inducers | 33419-42-0 | BML-GR307 | 10mM | Topoisomerase II Inhibitor |
| SMER28 | 263.0 | 1-G10 | 1 | Autophagy inducers | 307538-42-7 | BML-EI397 | 10mM | Increases autophagosome clearance |
| Trehalose | 342.1 | 1-G11 | 1 | Autophagy inducers | 99-20-7 | AC1507 | 10mM | Increases Beclin-1 and autophagins |
| C ₂ -dihydroceramide | 343.3 | 1-H02 | 1 | Autophagy inducers | 13031-64-6 | BML-SL101 | 10mM | Induces autophagosome formation |
| Temozolamide | 194.1 | 1-H03 | 1 | Autophagy inducers | 85622-93-1 | ALX-420- | 10mM | Localizes LC3 to autophagosomes |
| Resveratrol | 228.1 | 1-H04 | 1 | Autophagy inducers | 501-36-0 | BML-FR104 | 10mM | Inhibits S6 kinase |
| Staurosporine | 466.2 | 1-H05 | 1 | Autophagy inducers | 62996-74-1 | BML-EI156 | 10mM | Apoptosis control |
| Small Molecules | Mwt | Sample ID | Plate num | INHIBITOR | CAS Datab | Enzo ID | Stock | Cell Signaling |
| | | | | | | | | |
| Bafilomycin A1 | | | | | 88899-55-2 | CM110 | 1mM | V-ATPase inhib |
| 3-Methyladenine | | | | | 5142-23-4 | BML-AP502 | 10mM | PI3K inhib |
| LY294002 | | | | | 154447-36-6 | BML-ST420 | 10mM | PI3K inhib |
| Wortmannin | | | | | 19545-26-7 | BML-ST415 | 10mM | PI3K inhib |
| SP600125 | | | | | 129-56-6 | BML-EI305 | 10mM | JNK inhib. |
| Chloroquine | | | | | 54-05-7 | BML-DL353 | 10mM | Lysosomes (decreases pH) |
| Hydroxychloroquine | | | | | 118-42-3 | AC1500 | 10mM | Lysosomes (decreases pH) |
| Norcloimipramine-HCl | | | | | 303-48-0 | AC1501 | 10mM | Lysosomes (decreases pH) |
| (±)Bay K8644 | | | | | 71145-03-4 | BML-CA410 | 10mM | L-type Ca2+ agonist |
| Forskolin | | | | | 66428-89-5 | CN100 | 10mM | adenylyl cyclase activator |
| Dibutyryl cAMP-Na | | | | | 16980-89-5 | CN125 | 10mM | cAMP agonist |
| Rolipram | | | | | 61413-54-5 | PD177 | 10mM | PDE4 Inhibitor(decrease cAMP) |
| 8-CPT-cAMP-Na | | | | | 93882-12-3 | CN130 | 10mM | Epac activator |
| EHNA-HCl | | | | | 51350-19-7 | PD129 | 10mM | Dynein inhibitor |
| Tolazamide | | | | | 1156-19-0 | ALX-550- | 10mM | ATP-K+ channel antagonist |
| Quinine HCl-2H ₂ O | | | | | 6119-47-7 | AC1516 | 10mM | ATP-K+ channel antagonist |
| AICAR | | | | | 2627-69-2 | BML-EI330 | 10mM | Class III PI3K/Beclin1 binding |
| PD-98059 | | | | | 167869-21-8 | BML-EI360 | 10mM | ERK pathway activator |
| Anisomycin | | | | | 22862-76-6 | BML-ST102 | 10mM | p38 MAPK activator |
| Cycloheximide | | | | | 66-81-9 | ALX-380- | 10mM | protein synthesis(CMA) |
| Pifithrin-μ | | | | | 64984-31-2 | BML-AP503 | 10mM | HSP70 inhibitor |
| Nocodazole | | | | | 31430-18-9 | BML-T101 | 10mM | Inhibits autophagosome/lysosome fusion; Microtubule depolymerization |
| LNP Alone | | | | | 2-C05 | | | |



Small molecule library

Supplemental Information-Excel : Quantitative analysis of siRNA uptake in presence of small molecules: HeLa cells were exposed to LNPs (siAF647) in presence of small molecules for 3 hrs, cells were washed and uptake was quantified with the use of Acapella software. Experiments were conducted in triplicate with error measured as Standard error Mean (S.E.M)



Supplemental Information-Excel : A representative image from different treatment groups (20 fields) are shown for the treatment groups mentioned above.