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# High-content screening

Recent publications

June 2016



# What is high-content screening?

High-content screening (HCS) or high-content analysis (HCA) describes a set of analytical methods using automated microscopy, multiparameter image processing, and visualization tools to extract quantitative data from cell populations. HCS typically employs fluorescence imaging of samples in a high-throughput format and reports quantitatively on parameters such as spatial distribution of targets, and individual cell and organelle morphology. Most importantly, HCS is able to correlate measurements back to individual cells.

Since the introduction of Thermo Scientific™ ArrayScan™ High Content Analysis (HCA) Readers in 1999, over 1,000 peer-reviewed cellomics publications attest to a legacy of innovation in HCA that continues today. More peer-reviewed publications reference Thermo Scientific™ high-content screening instruments than any other high-content platform.

This list includes peer-reviwed publications from 2014 to June 2016 that reference the Thermo Scientific™ CellInsight™ or ArrayScan high-content platforms. Use it to find examples and applications that will validate your strategy and inform the next steps in your research.

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### **Publications 2016**

### (year-to-date, alphabetical by first author)

Ahn SO et al. (2016) Stronger uricosuric effects of the novel selective URAT1 inhibitor UR-1102 lowered plasma urate in tufted capuchin monkeys to a greater extent than benzbromaron. *J Pharmacol Exp Ther* 357:157–166.

Baker P et al. (2016) Exopolysaccharide biosynthetic glycoside hydrolases can be utilized to disrupt and prevent *Pseudomonas aeruginosa* biofilms. *Sci Adv* 2:e1501632.

Belzile J-P et al. (2016) Trehalose, an mTOR-independent inducer of autophagy, inhibits human cytomegalovirus infection in multiple cell types. *J Virol* 90:1259–1277.

Bengtsson E et al. (2016) The leucine-rich repeat protein PRELP binds fibroblast cell-surface proteoglycans and enhances focal adhesion formation. *Biochem J* 473:1153–1164.

Bruinsma W et al. (2016) Tousled-like kinase 2 regulates recovery from a DNA damage-induced G2 arrest. *EMBO Rep* 17:659–670.

Fujisawa T et al. (2016) The ASK1-specific inhibitors K811 and K812 prolong survival in a mouse model of amyotrophic lateral sclerosis. *Hum Mol Genet* 25:245–253.

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Hoelting L et al. (2016) Stem cell-derived immature human dorsal root ganglia neurons to identify peripheral neurotoxicants. *Stem Cells Trans Med* 5:476–487.

Hopcroft PJ and Fisher DI (2016) Development of a medium-throughput targeted LCMS assay to detect endogenous cellular levels of malonyl-CoA to screen fatty acid synthase inhibitors. *J Biomol Screen* 21:111–116.

Kong SL et al. (2016) Cellular reprogramming by the conjoint action of ERa, FOXA1, and GATA3 to a ligand-inducible growth state. *Mol Syst Biol* 7:526.

Li J et al. Downregulation of SMC1A inhibits growth and increases apoptosis and chemosensitivity of colorectal cancer cells. *J Intl Med Res* 44:67–74.

Mahara S et al. (2016) HIFI-α activation underlies a functional switch in the paradoxical role of Ezh2/PRC2 in breast cancer. *Proc Natl Acad Sci* 113:E3735–E3744.

Martinez E et al. (2016) *Coxiella burnetii* effector CvpB modulates phosphoinositide metabolism for optimal vacuole development. *Proc Natl Acad Sci* 113:E3260–E3269.

Matsa E et al. (2016) Human induced pluripotent stem cells as a platform for personalized and precision cardiovascular medicine. *Physiol Rev* 96:1093–1126.

Pattabhi S et al. (2016) Targeting innate immunity for antiviral therapy through small molecule agonists of the RLR pathway. *J Virol* 90:2372–2387.

Sun J et al. (2016) Comprehensive RNAi-based screening of human and mouse TLR pathways identifies species-specific preferences in signaling protein use. *Sci Signal* 9:ra3.

Sun L et al. (2016) NHERF1 regulates actin cytoskeleton organization through modulation of  $\alpha$ -actinin-4 stability. *FASEB J* 30:578–589.

Ulferts R et al. (2016) Screening of a library of FDA-approved drugs identifies several enterovirus replication inhibitors that target viral protein 2C. *Antimicrob Agents Chemother* 60:2627–2638.

Westwell-Roper C et al. (2016) Differential activation of innate immune pathways by distinct islet amyloid polypeptide (IAPP) aggregates. *J Biol Chem* 291:8908–8917.

Young HE et al. (2016) Discovery of the elusive UDP-diacylglucosamine hydrolase in the lipid A biosynthetic pathway in *Chlamydia trachomatis*. *MBio* 7:e00090–e00116.

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Amazit L et al. (2015) Finerenone impedes aldosteronedependent nuclear import of the mineralocorticoid receptor and prevents genomic recruitment of steroid receptor coactivator-1. *J Biol Chem* 290:21876–21889.

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Bailey ML et al. (2015) Dependence of human colorectal cells lacking the FBW7 tumor suppressor on the spindle assembly checkpoint. *Genetics* 201:885–895.

Barber JA et al. (2015) Quantification of drug-induced inhibition of canalicular cholyl-L-lysyl-fluorescein excretion from hepatocytes by high content cell imaging. *Toxicol Sci* 148:48–59.

Callihan P et al. (2015) Convergent regulation of neuronal differentiation and Erk and Akt Kinases in human neural progenitor cells by lysophosphatidic acid, sphingosine 1-phosphate, and LIF: specific roles for the LPA1 receptor. *ASN Neuro* 6:pii:1759091414558416.

Callis R. et al. (2015) A screening assay cascade to identify and characterize novel selective estrogen receptor downregulators (SERDs). *J Biomol Screen* 20:748–759.

Cassar PA et al. (2015) Integrative genomics positions MKRN1 as a novel ribonucleoprotein within the embryonic stem cell gene regulatory network. *EMBO Rep* 16:1334–1357.

Chatterjee SS et al. (2015) Inhibition of  $\beta$ -catenin-TCF1 interaction delays differentiation of mouse embryonic stem cells. *J Cell Biol* 211:39–51.

Chen Y et al. (2015) PHD3 stabilizes the tight junction protein occludin and protects intestinal epithelial barrier function. *J Biol Chem* 290:20580–20589.

Chetty S et al. (2015) A Src inhibitor regulates the cell cycle of human pluripotent stem cells and improves directed differentiation. *J Cell Biol* 210:1257–1268.

Cho SW et al. (2015) CXCL16 signaling mediated macrophage effects on tumor invasion of papillary thyroid carcinoma. *Endocr Relat Cancer* 23:113–124.

Chou Y-C et al. (2015) Pharmacological induction of human fetal globin gene in hydroxyurea-resistant primary adult erythroid cells. *Mol Cell Biol* 35:2541–2553.

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Deglincerti A et al. (2015) Coco is a dual activity modulator of TGFβ signaling. *Development* 142:2678–2685.

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Dopie J et al. (2015) Genome-wide RNAi screen for nuclear actin reveals a network of cofilin regulators. *J Cell Sci* 128:2388–2400.

Franco MC et al. (2015) Nitration of Hsp90 on Tyrosine 33 regulates mitochondrial metabolism. *J Biol Chem* 290:19055–19066.

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Hanniford D et al. (2015) Identification of metastasissuppressive microRNAs in primary melanoma. *J Natl Cancer Inst* 107:pii:dju494.

Her Z et al. (2015) Loss of TLR3 aggravates CHIKV replication and pathology due to an altered virus-specific neutralizing antibody response. *EMBO Mol Med* 7:24–41.

Hirst J et al. (2015) Contributions of epsinR and gadkin to clathrin-mediated intracellular trafficking. *Mol Biol Cell* 26:3085–3103.

Hirst J et al. (2015) Loss of AP-5 results in accumulation of aberrant endolysosomes: defining a new type of lysosomal storage disease. *Hum Mol Genet* 24:4984–4996.

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Liu Z et al. (2015) A novel antibody engineering strategy for making monovalent bispecific heterodimeric IgG antibodies by electrostatic steering mechanism. *J Biol Chem* 290:7535–7562.

Mahajan S et al. (2015) VX-509 (decernotinib) is a potent and selective janus kinase 3 inhibitor that attenuates inflammation in animal models of autoimmune disease. *J Pharmacol Exp Ther* 353:405–414.

Nakayama M et al. (2015) Attenuation of the phosphatidylinositol 3-kinase/Akt signaling pathway by porphyromonas gingivalis gingipains RgpA, RgpB, and Kgp. *J Biol Chem* 290:5190–5202.

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Peppard JV et al. (2015) High-content phenotypic screening and triaging strategy to identify small molecules driving oligodendrocyte progenitor cell differentiation. *J Biomol Screen* 20:382–390.

Poussin C et al. (2015) Systems biology reveals cigarette smoke-induced concentration-dependent direct and indirect mechanisms that promote monocyte-endothelial cell adhesion. *Toxicol Sci* 147:370–385.

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Reed DM et al. (2015) An autologous endothelial cell:peripheral blood mononuclear cell assay that detects cytokine storm responses to biologics. *FASEB J* 29:2595–2602.

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Sekimoto T et al. (2015) Both high-fidelity replicative and low-fidelity Y-family polymerases are involved in DNA rereplication. *Mol Cell Biol* 35:699–715.

Senkowski W et al. (2015) Three-dimensional cell culture-based screening identifies the anthelmintic drug nitazoxanide as a candidate for treatment of colorectal cancer. *Mol Cancer Ther* 14:1504–1516.

Sharanek A et al. (2015) Cellular accumulation and toxic effects of bile acids in cyclosporine A-treated HepaRG hepatocytes. *Toxicol Sci* 147:573–587.

Silva-Miranda M et al. (2015) High-content screening technology combined with a human granuloma model as a new approach to evaluate the activities of drugs against *Mycobacterium tuberculosis*. *Antimicrob Agents Chemother* 59:693–697.

Smith JG et al. (2015) Dental pulp cell behavior in biomimetic environments. *J Dent Res* 94:1552–1559.

Stroedicke M et al. (2015) Systematic interaction network filtering identifies *CRMP1* as a novel suppressor of huntingtin misfolding and neurotoxicity. *Genome Res* 25:701–713.

Stylianou M et al. (2015) Novel high-throughput screening method for identification of fungal dimorphism blockers. *J Biomol Screen* 20:285–291.

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Vellinga TT et al. (2015) SIRT1/PGC1α-dependent increase in oxidative phosphorylation supports chemotherapy resistance of colon cancer. *Clin Cancer Res* 21:2870–2879.

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Xia H-G et al. (2015) Degradation of HK2 by chaperone-mediated autophagy promotes metabolic catastrophe and cell death. *J Cell Biol* 210:705–716.

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Xie J et al. (2015) Zinc inhibits hedgehog autoprocessing: linking zinc deficiency with hedgehog activation. *J Biol Chem* 290:11591–11600.

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Zhang B et al. (2015) Human umbilical cord mesenchymal stem cell exosomes enhance angiogenesis through the Wnt4/β-catenin pathway. *Stem Cells Trans Med* 4:513–522.

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Altman M et al. (2014) Silencing of regulator of G-protein signaling 10 (RGS10) increases phosphorylation of eukaryotic initiation factor 4E binding protein (4E-BP1), and enhances growth and survival signaling pathways in ovarian cancer. *FASEB J* 28:843.11.

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Bachour-El Azzi P et al. (2014) Impact of inflammation on chlorpromazine-induced cytotoxicity and cholestatic features in HepaRG cells. *Drug Metab Dispos* 42:1556–1566.

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Bailey TA et al. (2014) A kinase inhibitor screen reveals protein kinase C-dependent endocytic recycling of ErbB2 in breast cancer cells. *J Biol Chem* 289:30443–30458.

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Dean RA et al. (2014) Identification of a putative Tdp1 inhibitor (CD00509) by *in vitro* and cell-based assays. *J Biomol Screen* 19:1372–1382.

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Feng Y et al. (2014) Platelets direct monocyte differentiation into epithelioid-like multinucleated giant foam cells with suppressive capacity upon mycobacterial stimulation. *J Infect Dis* 210:1700–1710.

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Poe JA et al. (2014) Development and validation of a high-content bimolecular fluorescence complementation assay for small-molecule inhibitors of HIV-1 Nef dimerization. *J Biomol Screen* 19:556–565.

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Zhang M et al. (2014) Progesterone receptor membrane component 1 is a functional part of the glucagon-like peptide-1 (GLP-1) receptor complex in pancreatic  $\beta$  cells. *Mol Cell Proteomics* 13:3049–3062.

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