

# Opera High Content Screening System



A New Era of Cell Screening Sciences

# Opera

## Superior Performance, High Resolution and Ultra-High Speed

Only the combination of these key features ensures the potential for fully flexible assay development and the industrialization of the screening process. Central to high resolution are true point scanning confocal imaging and the use of water immersion objectives. Drivers for speed are simultaneous image acquisition and data processing. The most prominent areas of use are cell screening with sub-cellular resolution as well as bead based screening applications – from conventional plate formats to nanoplate scale.



### Features

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Fully automated point scanning confocal high throughput screening system prevents photo-bleaching and photo-toxicity  
Simultaneous four-color imaging and on-line analysis for multiplexed assays in high speed

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Water immersion objectives for optimized signal-to-noise

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Environmental control unit, liquid handling and robotics options enabling kinetic experiments and full screens using live cell samples

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Comprehensive and fully flexible image analysis software Acapella for user-defined applications

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### Applications

#### Toxicity and survival

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Cell viability, cell differentiation, cell proliferation, cytotoxicity, apoptosis, transporter phenomena

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#### Cell signaling and pathway screening

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Calcium flux, second messengers, ion channels, membrane potential, protein translocation and redistribution

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#### Gene expression

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Analysis of house-keeping gene and reporter gene expression, gene activity and protein regulation, and siRNA library screens

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#### Receptor activation

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Ligand binding, receptor activation and desensitization, translocation and endocytosis, recruitment of signaling molecules

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#### Phenotypic assays

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Neurite outgrowth, cell differentiation, cell adhesion and spreading

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#### Immunochemical assays

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Bead-based homogeneous antigen-antibody interaction

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**Disclaimer:** A use license to the ThermoFisher Scientific (Cellomics) HCS patent portfolio is included with each Opera™ and Acapella™ product. For the use of the assays described additional licenses may be required.

# The Opera Workflow

## Assay development

Run test plates with “high” and “low” values for the assay under investigation in micro well plates of choice:

96-well, 384-well, or 1536-well plates  
user definable

Make choice of plate reading:

one, two, three, or four colors in parallel  
10x, 20x, 40x, or 60x magnification, water immersion or air objectives  
single or multiple image fields per well  
z-sectioning  
time lapse

Read and analyze plates automatically using an existing analysis script

Optimize image analysis using the script development toolbox or generate a new analysis algorithm using the Acapella Detection Libraries

Combine modules for cell and cellular compartment detection according to cell type and staining conditions.

Create input and output parameters as required

## High Throughput Screening

Run assay in a suitable screening environment e. g. cell::explorer screening system or live cell workstation with plate handling, liquid handling, and incubator for live cell applications System with compound transfer, plate fixation and plate washing for fixed cell assays

Transfer plates to the Opera, run screen in a fully automated fashion using the pre-configured measurement and analysis parameters

Analyze images on-line

Images and data are available for display while the screen is running  
Store raw and processed data in the file based High Throughput Analysis System

Export the results automatically or batch-wise to a user defined evaluation software

Generate higher level results with the evaluation software (dose-response, QC, compound profiling, hit list generation, SAR,...)

## Virtual Re-Screening




Re-analyze data using modified or different algorithms

Optimize Z'

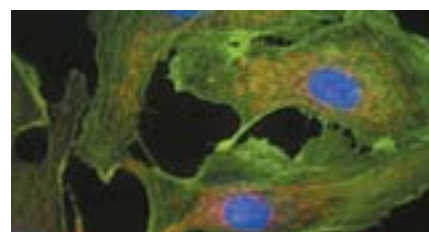
Screen for systemic artifacts

Side effects of compounds

Compound toxicity

Detection Libraries	Results (Best choice marked)
Nucleus Detection	
Cell Detection	
Compartment Detection	

Creating analysis scripts using the Acapella Detection Libraries



Three color image taken with a 60x water immersion lens



Plate handling for the Opera:  
The plate::handler™ II

# Opera

## High Content Screening System

### Optics

#### Confocal imaging unit

Point-scan confocal Nipkow-spinning disc  
Ultra fast laser-based auto focus  
Fully automated exchange of objectives, four-position objective turret  
Choice of air objectives (10x, 20x, 40x) and water immersion objectives (10x, 20x, 40x, 60x)  
Automated water immersion fluid supply  
Three parallel confocal detection channels plus optional fourth non-confocal detection channel

#### Excitation

Solid state lasers: 405 nm, 488 nm, 561 nm or 532 nm, 640 nm

### Automation

Fully automated objectives, filters, and beam splitter selection  
Computer-controlled laser power  
Scalable PC cluster for on-line data analysis  
Four-color parallel and sequential exposure image acquisition  
Data acquisition and analysis rates of > 100,000 image sets per day  
Accepts wide variety of micro titer and nano titer plates  
Integration-ready into a screening environment

### Further Options

#### UV unit

UV light source  
High pressure Xe lamp  
Additional, non-confocal detection channel including fourth camera

#### Environmental control unit

Temperature, CO<sub>2</sub>, and humidity control

#### Dispensing unit

Dispenser for µl volume reagent addition  
Software for single and multi-delay time kinetics

#### Work station extensions

Plate loading with stacker for fixed cell samples  
Live Cell Monitoring Workstation extension including plate transfer to and from an incubator

#### Fluorescence lifetime imaging (FLIM)

External pulsed laser source for confocal fluorescence lifetime imaging  
Fast gated CCD camera

### Data Handling

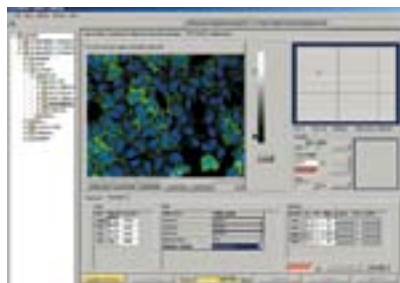
Distributed data processing system with a variable number of evaluation clients with load balancing  
Real-time image analysis  
Automatic load balancing  
Multiple field image acquisition enabling full coverage of each well  
Fully flexible choice of image acquisition: number and location fields, focus height, volume imaging  
All experiment definitions and results are stored and exported in XML files.  
Flexible export system for result to post-analysis software.  
Interface to Genedata's Screener® platform providing a seamless automated workflow from primary to secondary data analysis (pattern analysis, IC50, MOA, etc.) in high throughput.

### Image analysis

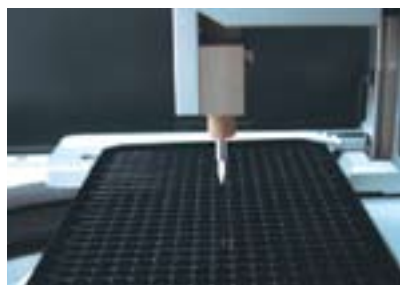
Fully flexible image analysis software, Acapella, providing a large choice of image analysis modules and standard procedures for algorithm development  
Automated flat-field correction  
Extensive image processing and analysis functions  
Cell recognition and counting  
Object recognition and identification of subcellular structures  
Classification of recognized objects based on user defined properties  
Extraction of predefined and user defined properties of objects and cell structures



Water immersion objectives on an automated turret



Microscope dialogue



Dispensing unit



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