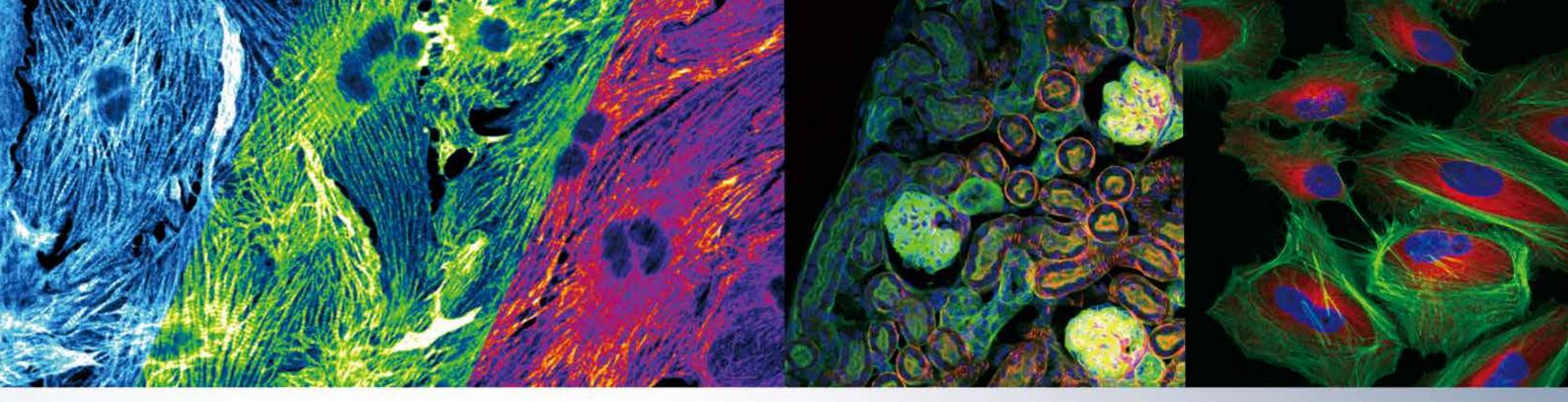




High-throughput Cytological Discovery System

Cell Voyager CV8000



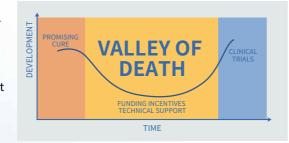


Introduction

Within the drug development market, demands on high content analysis systems for drug efficacy evaluation are increasing in accordance with the needs for cell-based assay and phenotypic screening. In order to increase screening efficiency, devices with higher speeds (higher throughput) are required.

On the other hand, in order to bridge the "valley of death" of the drug development process, the quality of screening hits must be increased. This requires the construction of more complex evaluation systems that utilize multifaceted parameters via 3D cultivation systems, live-cell imaging and higher detail image analysis.

In current drug development research, determining how to implement throughput screening and complex evaluation system screening in parallel is an important issue.



Solution

The CellVoyager CV8000 is a high-end, high content analysis system that solves this contradictory screening

Through the combination of a proprietary Yokogawa high speed confocal scanner, water immersion lens, up to four high field-of-vision cameras, a microscopic stage with cell cultivation environment, and an integrated robotic pipetter, we have realized not only high throughput, high-resolution imaging, but also phenotypic screening via a more complex evaluation

In addition, our specialized analysis software, CellPathfinder, uses deep leaning and machine learning to recognize target objects with high accuracy, supporting you from image analysis to results display using graphs.



The Yokogawa advantage

Confocal scaner unit

Live/kinetic experiment compatible

High throughput

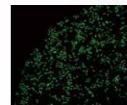
technology

Voyage to unknown worlds

-Real time confocal and label-free imaging-

Long-term live cell imaging

Stage incubator included as standard. Realization of non-stop, long-duration observation (3 days +) via humidity, temperature and CO₂ control.

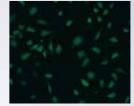


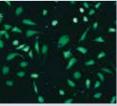
Before incubation



Kinetic assay

Drug addition during imaging is made possible by an integrated robotic pipetter with disposable tips Ideal for kinetic experiments involving the observation of high speed phenomena.

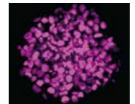




Before instillation

Organoid / Spheroid

Yokogawa's spinning disk confocal technology excels in imaging of samples with depth, such as 3D culture samples where clear and quick imaging is difficult, allowing for evaluation close to in-vivo quality.

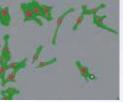




Recognition and analysis can be performed by taking bright field images from several Z positions and creating a CE bright field image using the included CellPathfinder analysis software. Analysis accuracy is further enhanced via the new Deep Learning option.

Label-free analysis





CE Bright Field

Cell Recognition image

CV8000 2

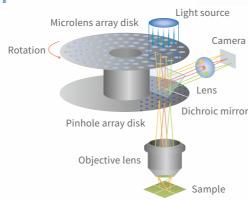
State-of-the-art technology that enables you to do what you want



Observe cells as they are

Dual spinning disk confocal system

A Yokogawa proprietary multi-scan method utilizing approximately 1,000 laser beams on the observation region and tandem disks rotating at high speed. The disks comprise a pinhole array disk with approximately 20,000 pinholes arranged in an equal pitch spiral pattern, and a microlens array disk that focuses the excitation light laser into individual pinholes. Not only does this allow high speed imaging, but it also largely prevents phototoxicity and fluorescence photobleaching.

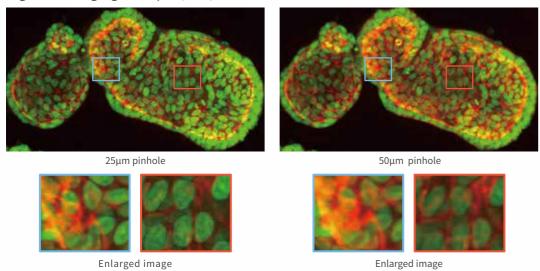


Deeper, clearer observation

Pinhole disk exchanger

Two different types of pinhole disks (25/50µm) can be used, according to the sample. For thick samples, reducing the pinhole diameter allows for higher confocality, shaper images. For dark samples, increasing the pinhole diameter allows for brighter images.

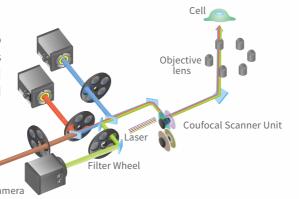
Organoid imaging example (MIP)



Higher throughput screening

Optical configuration

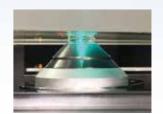
The optical system configuration can be selected according to the purpose. A single 96-well plate can be imaged in four colors in one minute by attaching four high-sensitivity wide-field sCMOS cameras. The system is also compatible with FRET and CellPainting assay.



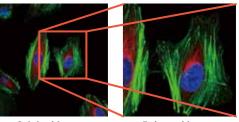
Capturing finer structures

Original water immersion lens

Water immersion lenses excel in capturing high-resolution images of cells within a liquid. The CV8000 can be equipped with a 40x or 60x water immersion objective lens. Our 40x lens is a particularly unique lens capable of highly advanced spherical aberration correction, allowing for the capture of bright high-resolution images over a full wide-field. The lens water supply is also completely automated. This equipment makes high throughput screening via water submersion lens possible.



 Cell image captured using the 60x water submersion lens



Original image

Capturing live cell movement

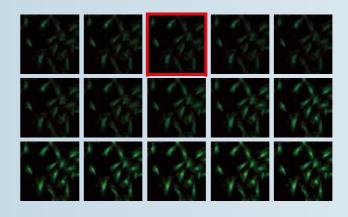
High-precision incubator and robot pipetter

The stage incubator features an airtight construction, managing humidity, temperature and CO₂ levels. The integrated robotic pipetter conducts the following process fully automatically: tip pickup → reagent collection from the reagent plate \rightarrow reagent addition to the sample plate \rightarrow tip disposal. Not only can images be rapidly obtained before and after reagent instillation, but it's also possible to add reagents to single wells multiple times, and adjust the addition speed etc., broadening the range of dynamics observation via reagent instillation.

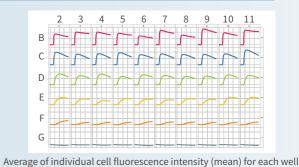


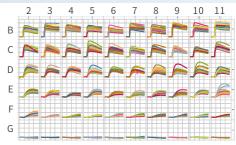
Ionomycin concentration-dependant calcium response

It's possible to conduct high speed imaging before and after ionomycin instillation, recognize individual cells from the images, and obtain timelapse data for each cell.



0.2 second interval images (2.6 seconds after imaging start to 5.4 seconds after) Ionomycin instillation conducted at 3.0 seconds after imaging start (red frame)





Fluorescence intensity (mean) for individual cells in each well

3 CV8000

A more live-cell-friendly total HCA system



Making long-duration live-cell imaging possible

Featuring a stable built-in stage incubator

HeLa cells were seeded in a 96 well plate at a density of 500 cells per well, and cultured for 24 hours. The well plate was then placed in the internal stage incubator and cell culturing was conducted for 72 hours, and the total area (hereinafter Total Area) occupied by cells was analyzed. As a result, minimal unevenness in cell multiplication was observed across the 96-wells (excluding the four corner wells) when compared to a regular CO₂ incubator.



Scan for information

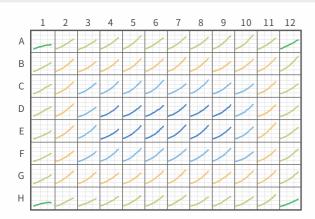
Cell multiplication comparison with regular CO₂ incubator after 72hr incubation (n=3)

	_1	2	3	4	5	6	7	8	9	10	11	12
Α	47	77	79	79	90	86	83	84	80	76	74	64
В	73	87	83	84	88	95	91	94	93	97	89	80
С	86	89	89	94	95	93	100	99	99	92	102	85
D	84	96	93	98	94	95	90	98	93	105	112	87
Ε	93	97	98	95	92	98	95	103	94	98	103	88
F	94	95	93	96	93	92	98	96	101	102	106	80
G	87	92	99	92	92	93	95	100	98	109	96	84
Н	51	77	83	80	87	84	90	87	91	91	101	64

96 well average: 90 Average of outermost 36 wells: 81 Average of 60 wells (excl. outermost): 96

The values represent the following:
CV8000 Total Area after 72hrs / Total Area at 0hrs (hereinafter
Total Area ratio) / CO2 incubator Total Area ratio x 100.
(Numbers near to 100 mean that cell multiplication was
approximately equal for the CV8000 and CO2 incubator.)
Cell multiplication near to that of the CO2 incubator was
verified, excluding the four corner wells.

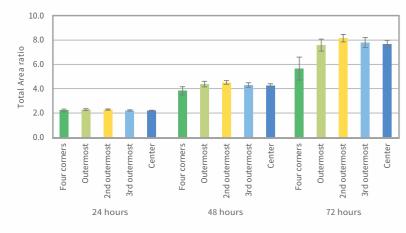
Cell multiplication curves for each well of a 96-well plate

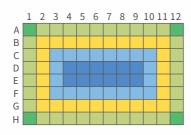


Vertical axis: Total Area Horizontal axis: Time (0-72 hours)

Cell multiplication was low in the four corner wells; however, it continued in the other wells.

Total Area ratio after cultivation start (24, 48 and 72 hours) (n=3)



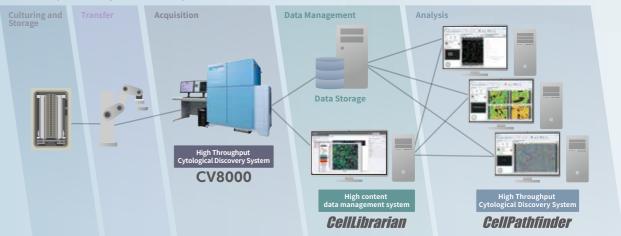


Excluding the four corner wells, even after 72 hours, there were no large differences in cell multiplication. The low variation in cell multiplication speed across the wells can be seen.

System Integration

Centralized process management, from the cultivation environment, to transfer, imaging, analysis and data management.

We offer optimum systems in response to our customers' needs.



High Content Analysis Software CellPathfinder

The software analyzes image data captured with the CV8000, creates graphs and exports various data. Beginner and expert users alike can take full advantage of the software, thanks to an abundance of templates and flexible protocol editing capability. CE bright field and machine-learning functionalities make label-free analysis possible. The new Deep Learning option has also been added, largely improving cell recognition accuracy.

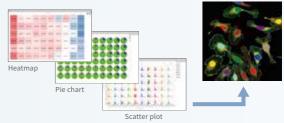
Just click the menu item for analysis

Simply follow the flow displayed at the top of the screen. The analysis menu has easy-to-understand icons. Simply click the desired menu item and the protocol will load.



Fast results for immediate verification and study Computed numeric data can be displayed in a variety of ways.

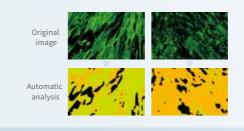
Graph plots and cell images are linked, making for easy result verification and inquiry.



Unbiased phenotype evaluation via Al

Machine-learning also provides bias-free digitization of visually-evaluated experiments.

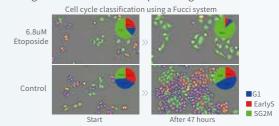
Automatic recognition is made possible simply by clicking the shape you want the software to learn.



Label-free phenotype analysis

Eliminates the time, cost and influence on cells associated with cell labelling.

Even higher precision classification is made possible through combination with deep learning.

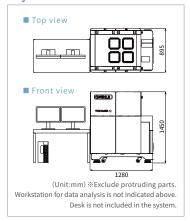


5 CV8000 CV8000 6

Specifications

Specifications							
Model	CV8000						
Sample format	Multiple well plate (6, 12, 24, 48, 96, 384, 1536 wells), Glass slide						
lmaga mada	Confocal mode: Max. 4 color simultaneous recording						
Image mode	Bright field / Phase contrast (10x, 20x for 6, 12, 24 well plates), Digital phase contrast (10x, 20x)						
Output data format	data format Image data: 16bit TIFF, PNG Numerical data: CSV, original format						
Excitation wavelength	405/445/488/561/640 nm, all solid laser, max. 5 lasers 【Option】365 nm LED						
White light illumination	LED						
Autofocus	Laser-based mode, image-based mode						
Objectives	Max. 6 lenses are available, automatically switchable						
Objectives	Dry: 2x, 4x, 10x, 20x, 40x Water immersion: 40x, 60x Phase contrast: 10x, 20x Long working distance: 20x						
Confocal unit	Microlens-enhanced wide-view dual Nipkow disk confocal scanner, 50 μm pinhole						
Comocat unit	[Option] 25 µm pinhole disk and auto pinhole disk exchanger						
Camera	sCMOS (Effective pixels: 2,000 X 2,000 pixel size: 6.5 μm) Max. 4 cameras						
Stage incubator	Temperature for inchbation: 35 - 40°C CO ₂ supply box (CO ₂ : 5 %, forced humidification)						
Robot pipetter	[Option] Disposable tip type (96 tip or 384 tip type)						
Bar code reader	[Option] 1 or 2 dimension						
Workstations	Dual-monitor work station for system control, dual-monitor work station for data analysis						
Analysis software	Granularity, Neurite, Nuclear morphology, Nuclear translocation, Plasma membrane translocation,						
(CellPathfinder)	Machine learning, Label-free analysis, 3D analysis, Deep Learning, etc.						
Operating environment	genvironment 15 - 30°C 30 - 70%RH (No condensation)						
	Measurement unit: AC100-240V, 50/60Hz, 2KVA max						
Power supply	Workstation for system control: AC100-240V, 50/60Hz, 1.3KVA max						
	Workstation for data analysis: AC100-240V, 50/60Hz, 950VA max						
Dimensions	Measurement unit: W1,280×D895×H1,450 mm						
Weight	Measurement unit: 510Kg						

Layout



Reliable after-service / Powerful technical support

We offer the best after-service program to meet your requirement and budget.

Our HCA experts will support you to obtain the best results easily.



Notice No.50, dated June 24, 2007 Yokogawa Bechic Cosporator 2-9-32 Nakacho, Musadnino-shi, Tokyo, 580-8750 Japan Manufactured KZ. Caution

Read user's manual carefully in order to use the instrument correctly and safely.

This product falls under

the category of class laser product.

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