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SYSTEMS AND REAGENTS FOR TISSUE CLEARING

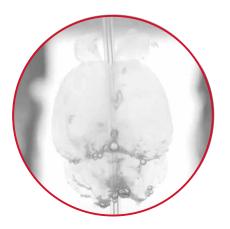




SECTIONING IS TIME. WE CUT THE TIME.

Tissues are inherently three dimensional in nature, which makes imaging intact tissues a necessity for a more complete study into the relationship between structure and function and the system-level study of cellular mechanisms. Tissue clearing has become an important step for imaging tissues in 3D at single-cell resolution.

The X-CLARITY™ is a collection of systems and ready-to-use reagents to standardize, simplify, and accelerate each step of the tissueclearing process. X-CLARITY™ is based on the CLARITY (Clear Lipid-exchanged Acrylamide-hybridized Rigid Imaging / Immunostaining /in situ-hybridization-compatible Tissue hYdrogel) method. With CLARITY, preserved tissues are embedded in a hydrogel matrix and lipids are actively extracted through electrophoresis to create a stable and optically transparent tissue-hydrogel hybrid that is chemically accessible for multiple rounds of antibody labeling and imaging.



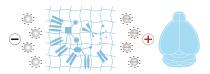
ACCELERATE YOUR 3D IMAGING WORKFLOW

STEP 1
Tissue-hydrogel
Hybridization



A fixed tissue sample is incubated in the X-CLARITY™ Hydrogel Solution Kit to allow hydrogel monomers to diffuse uniformly throughout the sample. Once the solution has permeated the tissue, the sample is placed in the X-CLARITY™ Polymerization System. Monomers polymerize in the anaerobic environment, linking the biomolecules to a hydrogel network, which preserves molecular information and structural integrity.

STEP 2Tissue Clearing



Once the tissue-hydrogel hybrid has been formed, the hybrid is cleared in the X-CLARITY™ Tissue Clearing System II using the ready-to-use Electrophoretic Tissue Clearing Solution. Lipids are extracted actively through electrophoresis or passively, leaving behind a stable and transparent tissue-hydrogel hybrid that is chemically accessible for molecular phenotyping.

STEP 3
Antibody Labeling



The transparent sample is then labled with antibodies using the DeepLabel™ Antibody Staining Kit, which enhances antibody penetration into clarified tissues. Optimized for clarified tissue samples, DeepLabel™ can be used with tissues cleared with various clearing methods such as passive or active CLARITY, iDISCO, 3DISCO, uDISCO, Visikol, or CUBIC.

STEP 4



Prior to imaging, the tissue-hydrogel hybrid is placed in X-CLARITY™ Mounting Solution, a refractive index matching solution (RIMS), to homogenize the environment within the tissue to the solution. This reduces light scatter, which in turn increases optical transparency and consequently increases image quality and imaging depth.

X-CLARITY™ Hydrogel Solution Kit C1310X-1 kit

The X-CLARITY™ Hydrogel Solution Kit is a pre-tested hydrogel solution for uniform and consistent tissue-hydrogel hybridization. The kit is composed of X-CLARITY™ Hydrogel Solution and X-CLARITY™ Polymerization Initiator .



X-CLARITY™ Hydrogel Solution

C13103 – 1 x 1 L

The X-CLARITY™ Hydrogel Solution is a readyto-use acrylamide-based solution used to create polyacrylamide. X-CLARITY™ Hydrogel Solution contains no bis-acrylamide or paraformaldehyde.



X-CLARITY™ Polymerization Initiator

C13104 – 1 x 2.5 g

The X-CLARITY™ Polymerization Initiator is a thermal free radical initiator that releases free radicals when heated in solution to initiate the polymerization of hydrogel monomers.

X-CLARITY™ Polymerization System [2000]

The X-CLARITY™ Polymerization System is a standalone, automated system developed to simplify tissue-hydrogel hybridization, a crucial step for optimal tissue clearing. Multiple samples can be placed in multi-well plates or conical tubes for rapid and efficient high-throughput sample processing. Users can control polymerization by adjusting vacuum strength, temperature, and a timer through a simple touchscreen interface.



- High throughput (up to 768 samples/run)
- **⊘** Compatible with various vessels
- Fully automated vacuum and temperature control
- Rapid and consistent polymerization

Comes with your choice of two heat blocks



X-CLARITY™ Heat Block for 6 x 50 mL tubes

C20002

92 x 134 x 75 mm / 1.9 kg



X-CLARITY™ Heat Block for flat-bottom plates

C20003

92 x 134 x 78 mm / 2.5 kg



Touchscreen interface. The simple touchscreen interfaces gives users precise control over vacuum pressure, temperature, and polymerization time.



Compatible with multiwell plates and conical tubes. Users can select the combination of heat blocks to use with the system.

X-CLARITY™ Polymerization System Specifications

Temperature Range Temperature Accuracy Vacuum Range Power Consumption Applicable Power

Dimensions (W \times D \times H)

Weight

5"TFT LCD RT - 60°C

±0.3°C -90 - 0 kPa 312 W

AC 100-240 V. 50/60 Hz Exterior: 332 x 430 x 222 mm Interior: 307 x 137 x 140 mm

28 kg

X-CLARITY™ Tissue Clearing System II cssssi

The X-CLARITY™ Tissue Clearing System II is an all-in-one, easy-to-use solution for electrophoretic tissue clearing. Its unique design accelerates the removal of lipids from tissues while preserving the structural integrity of the sample.

Users can set tissue clearing conditions through a simple and intuitive touchscreen interface. In ETC (electrophoretic tissue clearing) mode, platinum-plated electrodes generate an electric field to accelerate the removal of lipids from tissues in a highly efficient manner. A built-in temperature control system actively cools and heats buffer to maintain consistent buffer temperatures during clearing. Buffer is constantly circulated to ensure consistent buffering capacity, temperature control, and elimination of tissue clearing byproducts. This advanced system ensures efficient, rapid, and consistent tissue clearing.





Electrophoretic Tissue Clearing Solution

C13001 – 12 x 1 L

Electrophoretic Tissue Clearing Solution is a premixed SDS-based buffer optimized for use with the system.

✓ Precise temperature control

- Active buffer cooling and heating capacity
- Sensitive and accurate temperature sensor

⊘ Compatible with multiple tissue types and sizes

- Electrophoretic and passive clearing
- Holders of various sizes available

Ø Uniform electric field

- Platinum-plated electrodes
- Constant current and constant voltage modes

✓ User-friendly setup

- Simple touchscreen interface
- Ready-to-use clearing solution



Compatible sample holders





X-CLARITY™ ETC Chamber Specifications

Instrument type
External dimensions (W x D x H)
Internal dimensions (W x D x H)
Weight

Electrophoretic chamber 176 mm x 128 mm x 154 mm 57 mm x 30 mm x 93 mm 2.8 kg

X-CLARITY™ ETC Control Tower Specifications

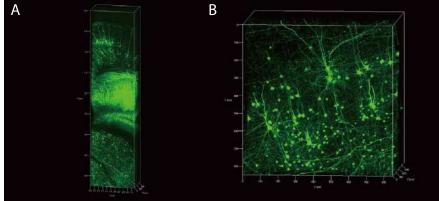
User interface
Power supply modes
Current & voltage range
Temperature range
Pump speed range
Electrical requirements
Power consumption
Dimensions (W x D x H)
Weight

Instrument type

Power supply, temperature control, and buffer circulation device 5 inch TFT LCD touchscreen Constant current or constant voltage 0.2-1.5 A, 5-70 V 30-60° C 50-200 rpm AC 100-240 V, 50/60 Hz 500 W (including the X-CLARITY** ETC Chamber) 205 mm x 430 mm x 370 mm 20 kg

Rapid, consistent, and reproducible clearing for thick tissues

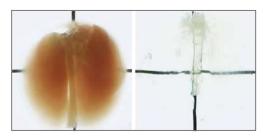
Long-term preservation of endogenous fluorescence proteins



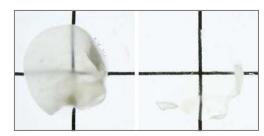
Long-term preservation of the Thy1-YFP signal in tissues cleared with the X-CLARITY™ systems and reagents. (A) Thy1-YFP signal immediately after clearing. (B) Thy1-YFP signal one month after clearing.

Thy1-YFP mouse brain slices cleared with the X-CLARITY™ systems and reagents. Thy1-YFP (green), Anti-Collagen IV (red), TO-PRO-3 (blue).

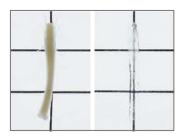
Verified with multiple tissue types



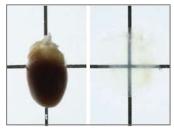
Mouse lungs and trachea cleared with the X-CLARITY $\!\!^{\mathrm{\tiny M}}$



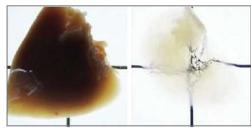
Mouse outer ear cleared with the X-CLARITY $^{\!\scriptscriptstyle{\text{\tiny M}}}$



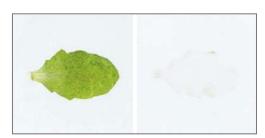
Mouse spinal cord cleared with the X-CLARITY $^{\!\scriptscriptstyle\mathsf{TM}}$



Mouse heart cleared with the X-CLARITY $^{\!\scriptscriptstyle\mathsf{TM}}$



Mouse liver cleared with the X-CLARITY $^{\!\scriptscriptstyle\mathsf{TM}}$



Arabidopsis thaliana Cleared with the X-CLARITY™

DeepLabel™ Antibody Staining Kit 33001-1 kit

DeepLabel™ Antibody Staining Kit is a set of non-toxic, ready-to-use reagents optimized for use with clarified tissues for effective antibody penetration and site-specific binding. With DeepLabel™, macromolecular probes rapidly and efficiently penetrate thick, protein-dense tissues at lower antibody concentrations. DeepLabel™ facilitates homogenous antibody staining with 2.6 X greater signal-to-background than conventional staining methods. DeepLabel™ is compatible with virtually all antibodies and all tissue clearing methods including CLARITY, PACT, iDISCO, and CUBIC.



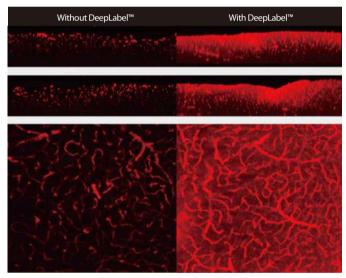
 DeepLabel™ Solution A
 C33002 - 1 x 25 mL

 DeepLabel™ Solution B
 C33003 - 2 x 25 mL

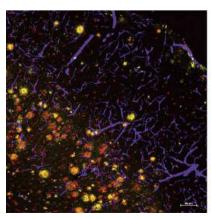
 DeepLabel™ Washing Buffer
 C33004 - 1 x 250 mL

 X-CLARITY™ Mounting Solution
 C13101 - 1 x 25 mL

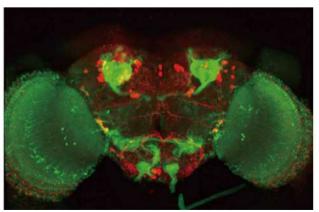
- Momogenous antibody distribution
- **⊘** Site-specific labeling
- ✓ 2.6 X greater signal-to-background
- ✓ Vibrant imaging at subcellular resolution
- **⊘** Simple protocol with ready-to-use reagents
- ✓ Compatible with multiple clearing methods



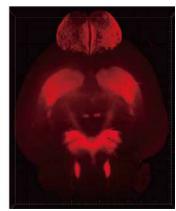
 ${\sf DeepLabel^{\sf M}}\ enhances\ anti-Collagen\ {\sf IV}\ penetration\ into\ clarified\ mouse\ brain\ tissues.$



Adult mouse brain stained using DeepLabel™ with anti-β-Amyloid (yellow), anti-lba1 (red), and anti-SMA (purple).



Whole adult *Drosophila* brain stained using DeepLabel™ with anti-GFP (green) and anti-TH (red).



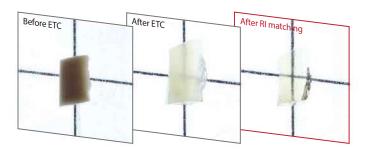
Whole adult mouse brain stained using DeepLabel $^{\mathbb{M}}$ with anti-TH (red).

X-CLARITY™ Mounting Solution C13101 – 1 x 25 mL C13102 – 10 x 25 mL C13107 – 20 x 25 mL

X-CLARITY™ Mounting Solution is a refractive index matching solution (RIMS). The RI of the solution is 1.460 at 25°C and is stable over a wide temperature range. X-CLARITY™ Mounting Solution minimizes photobleaching and preserves fluorescence signals, making it an ideal solution for mounting clarified and labeled tissue samples.



- Minimizes photobleaching
- ✓ Preserves fluorescence signals



PFA-fixed human cerebral cortex sample cleared with the X-CLARITY $^{\text{TM}}$ systems and reagents Courtesy of Dr. Hyung Jin Choi, Seoul National University College of Medicine

X-CLARITY™ cleared tissues are compatible for imaging with the following imaging systems:

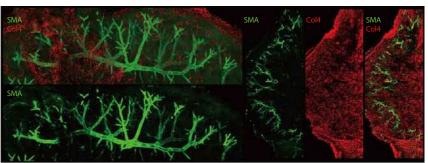


- Zeiss Lightsheet Z1
- OpenSPIM
- Lavision Ultramicroscope
- Confocal microscope (inverted)
- Confocal microscope (upright)

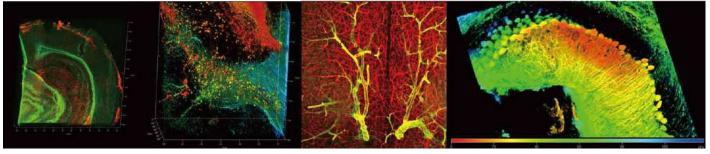
Refractive Index (value±0.001)	Temperature
1.461 1.460 1.459 1.458 1.457 1.455 1.454 1.453	20°C 25°C 30°C 35°C 40°C 45°C 50°C 55°C 60°C



Courtesy of Michal Shoshkes-Carmel, PhD and Andrea Stout, PhD, University of Pennsylvania



Mouse kidney cleared with the X-CLARITY**



What our customers are saying



"The [X-CLARITY] has actually been fundamental to the Data Brain Project because without being able to clear tissues ... we wouldn't be able to extract that structural information that is so important to understand the function of mammalian brains. It's been a huge advantage and a great leap forward in imaging."

Arti Ahluwalia, PhD Director, Centro E. Piaggio

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Cambridge, MA, USA

"We purchased the X-CLARITY for our facility and have been very satisfied with its easy-to-use design and consistent results. The X-CLARITY Tissue Clearing System has now allowed many more researchers to enter the field of tissue clearing."

Doug Richardson, PhD Director, Harvard Center for Biological Imaging



Paris, France

"The X-CLARITY allows us to perform rapid, efficient, and standardized clearing of mouse and human brain tissues. Access to the X-CLARITY technology will undoubtedly help the ICM research teams to better understand the 3D organization of protein assemblies and organelles in tissues."

Annick Prigent and Benoît Delatour, PhD
Operational Manager and Scientific Manager,
ICM Histomics

San Diego, CA, USA

"The X-CLARITY system is a delight to use and significantly lowers the barrier to tissue clearing imaging applications. With better clearing, and faster turnaround time, we were able to generate many more samples and images than would otherwise have been possible. Highly recommended!"

Uri Manor, PhD Director, Salk Institute for Biological Studies Biophotonics Core

Ul vor

Lyon, France

"Thanks to X-CLARITY, we now have access to the three dimensions of the whole heart organ."

Gabriel Bidaux, PhD Principal Investigator, INSERM

Lausanne Switzerlan

"The X-CLARITY saves a lot of time. We were able to image an entire set of brain and spinal cords in a relatively short amount of time. This allowed us to clearly see the benefit of the approaches we're having on spinal cord networks following an injury. It's a great device to save time for your research."

Quentin Barraud, PhD Lab Manager & Scientific Coordinator, EPFL

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