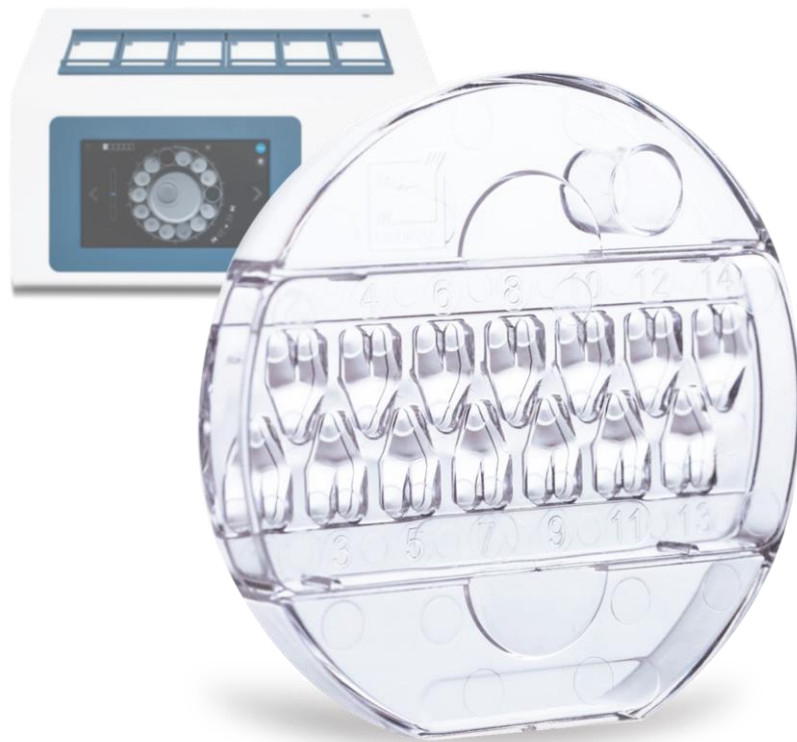




CE 0088



USER MANUAL

CultureCoin®

Rev. 3.0
Date of Revised 07.01.2020
Rx only



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Copyright Information

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The information contained in this manual and the accompanying product is copyrighted and all rights are reserved by Esco.

Esco reserves the right to make periodic minor design changes without obligation to notify any person or entity of such change.

Caution: Federal law restricts this device to sale by or on the order of a licensed healthcare practitioner.

Only to be used by a trained and qualified professional. The device is sold under exemption 21 CFR 801 Subpart D.

“Material in this manual is provided for informational purposes only. The contents and the product described in this manual (including any appendix, addendum, attachment or inclusion), are subject to change without notice. Esco makes no representations or warranties as to the accuracy of the information contained in this manual. In no event shall Esco be held liable for any damages, direct or consequential, arising out of or related to the use of this manual.”

Unpacking and Inspection

Follow standard receiving practices upon receipt of the instrument. Check the shipping carton for damage. If damage is found, stop unpacking the instrument. Notify the freight carrier and ask for an agent to be present while the instrument is unpacked. There are no special unpacking instructions, but be careful not to damage the instrument when unpacking it. Inspect the instrument for physical damage such as bent or broken parts, dents, or scratches.

Claims

Our routine method of shipment is via common carrier. Upon delivery, if physical damage is found, retain all packing materials in their original condition and contact the carrier immediately to file a claim.

If the instrument is delivered in good physical condition but does not operate within specifications, or if there are any other problems not caused by shipping damage, please contact your local sales representative or Esco Medical immediately.

Standard Terms and Conditions

Refunds & Credits

Please note only serialized products (products labelled with a distinct serial number) and accessories are eligible for partial refund and/or credit. Non-serialized parts and accessory items (cables, carrying cases, auxiliary modules, etc.) are not eligible for return or refund. In order to receive a partial refund/credit, the product must not have been damaged, and must be returned complete (meaning all manuals, cables, accessories, etc.) within 30 days of original purchase, in “as new” and resalable condition. The *Return Procedure* must be followed.

Return Procedure

Every product returned for refund/credit must be accompanied by a Return Material Authorization (RMA) number, obtained from Esco Medical Customer Service. All items being returned must be sent *prepaid* (freight, duty, brokerage and taxes) to our factory location.

Restocking Charges

Products returned within 30 days of original purchase are subject to a minimum restocking fee of 20% of the list price. Additional charges for damage and/or missing parts and accessories will be applied to all returns. Products which are not in “as new” and resalable condition, are not eligible for credit return and will be returned to the customer at their own expense.

Certification

This instrument has been thoroughly tested/inspected and found to meet Esco Medical’s manufacturing specifications when shipped from the factory. Calibration measurements and testing are traceable and done according to Esco Medicals ISO certification.

Warranty and Product Support

Esco Medical warrants this instrument to be free from defects in materials and workmanship under normal use, and service for two (2) years from the date of original purchase, provided the instrument is calibrated and maintained in accordance with this

manual. During the warranty period Esco Medical will, at our option, either repair or replace a product that proves to be defective at no charge, provided you return the product (shipping, duty, brokerage and taxes prepaid) to Esco Medical. Any and all transportation charges incurred are the responsibility of the purchaser and are not included within this warranty. This warranty extends only to the original purchaser and does not cover damage from abuse, neglect, accident or misuse, or as the result of service or modification by parties other than Esco Medical.

IN NO EVENT SHALL ESCO MEDICAL LTD. BE LIABLE FOR CONSEQUENTIAL DAMAGES. No warranty shall apply when damage is caused by any of the following:

- Power failure, surges, or spikes
- Damage in transit or when moving the instrument
- Improper power supply such as low voltage, incorrect voltage, defective wiring or inadequate fuses
- Accident, alteration, abuse or misuse of the instrument
- Fire, water damage, theft, war, riot, hostility, acts of God such as hurricanes, floods, etc.

Only serialized products (those items bearing a distinct serial number tag) and their accessory items are covered under this warranty.

PHYSICAL DAMAGE CAUSED BY MISUSE OR PHYSICAL ABUSE IS NOT COVERED UNDER THE WARRANTY. Items such as cables and non-serialized modules are not covered under this warranty.

This warranty gives you specific legal rights and you may have other rights, which vary from province to province, state to state, or country to country. This warranty is limited to repairing the instrument per Esco Medical's specifications.

When you return an instrument to Esco Medical for service, repair or calibration, we recommend shipment using the original shipping foam and container. If the original packing materials are not available, we recommend the following guide for repackaging:

- Use a double-walled carton of sufficient strength for the weight being shipped
- Use heavy paper or cardboard to protect all instrument surfaces. Use non-abrasive material around all projecting parts
- Use at least four inches of tightly packed, industrial-approved, shock-absorbent material all around the instrument

Esco Medical will not be responsible for lost shipments or instruments received in damaged condition due to improper packaging or handling. All warranty claim shipments must be made on a prepaid basis (freight, duty, brokerage, and taxes). No returns will be accepted without a Return Materials Authorization ("RMA") number. Please contact

Esco Medical to obtain an RMA number and receive help with shipping/customs documentation.

Re-calibration of instruments, which have a recommended annual calibration frequency, is not covered under the warranty.

Warranty Disclaimer

Should you choose to have your instrument serviced and/or calibrated by someone other than Esco Medical Ltd. and their representatives, please be advised that the original warranty covering your product becomes void when the tamper-resistant Quality Seal is removed or broken without proper factory authorization.

In all cases, breaking the tamper-resistant Quality Seal should be avoided at all cost, as this seal is key to your original instrument warranty. In an event where the seal must be broken to gain internal access to the instrument, you must first contact Esco Medical Ltd.

You will be required to provide us with the serial number for your instrument, as well as a valid reason for breaking the Quality Seal. You should break this seal only after you have received factory authorization. Do not break the Quality Seal before you have contacted us! Following these steps will help ensure that you will retain the original warranty on your instrument without interruption.

WARNING

Unauthorized user modifications or application beyond the published specifications may result in electrical shock hazard or improper operation. Esco Medical will not be responsible for any injury sustained due to unauthorized equipment modifications.

ESCO MEDICAL LTD. DISCLAIMS ALL OTHER WARRANTIES, EXPRESSED OR IMPLIED, INCLUDING ANY WARRANTY OF MERCHANTABILITY OR FITNESS FOR A PARTICULAR PURPOSE OR APPLICATION.

THIS PRODUCT CONTAINS NO USER-SERVICEABLE COMPONENTS.

UNAUTHORIZED REMOVAL OF THE INSTRUMENT COVER SHALL VOID THIS AND ALL OTHER EXPRESSED OR IMPLIED WARRANTIES.

Table of contents

1 How to use this manual.....	7
2 Safety warning.....	7
3 Indication for use.....	7
4 About the product.....	8
5 Safety symbols and labels	8
6 Important safety instructions and warnings	9
6.1 Pre use.....	9
6.2 During use.....	9
6.3 Post use.....	10
7 The Culture mode.....	10
8 Overview.....	10
9 Use.....	10
10 The use illustrated.....	11
11 pH measuring.....	12
12 Writing area on the CultureCoin and lid.....	13
13 User troubleshooting.....	13
14 Specifications.....	13

1 How to use this manual

The manual is designed to be read by sections, and not ideally cover to cover. This means that if the manual is read, from start to finish, there will be some repetition and overlap. We recommend the following method to go through the manual: first, familiarize yourself with the safety instructions; then proceed to the basic user functions that are needed for operating the equipment on a day to day basis; then review the alarm functions. The menu function of the user interface details information that is needed for the advanced level of users only. All parts must be read before the device is taken into use.

2 Safety warning

- Anyone working with, on or around this equipment should read this manual. Failure to read, understand and follow the instructions given in this documentation may result in damage to the unit, injury to operating personnel, and/or poor equipment performance.
- Any internal adjustment, modification or maintenance to this equipment must be undertaken by qualified service personnel.
- If the equipment must be relocated, make sure it is fixed properly on a support stand or base, and move on a flat surface. When necessary move the equipment and the support stand/base separately.
- The use of any hazardous materials in this equipment must be monitored by an industrial hygienist, safety officer or other suitably qualified individual.
- Before you proceed, you should thoroughly understand the installation procedures and take note of the environmental/electrical requirements.
- In this manual, important safety related points will be marked with the following symbols:



NOTE

Used to direct attention to a specific item.



WARNING

Use caution

- If the equipment is used in a manner not specified by this manual, the protection provided by this equipment may be impaired.

3 Indication for use

Preparing, storing and transferring human embryos. To be used only with the MIRI TL device.

4 About the product

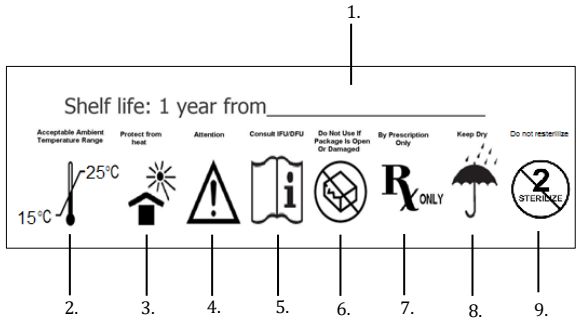
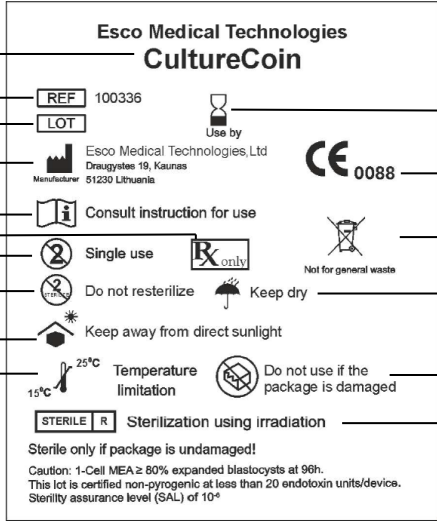
The Esco Medical CultureCoin® is a sterile, single use, culture vessel specifically developed for use in IVF. The dish holds up to 14 embryos in the centrally placed line of culture wells. The culture well features a 300µ, optically clear area where the embryo is situated. The area is optimized for microscopy. A clear circle is present around the culture area for the purpose of making it easy for an automatic focus algorithm to find the area under microscopic image acquisition. The culture wells are numbered (1-14) for easy location of each well by the user. Each culture well has two washing wells for cell manipulations, washing media or replacement media.

The lid is mounted inside the circumference of the dish covering the culture area completely – the user will only grab the dish during handling and not risk to lose the dish by holding on to the lid only – as can be the case on all traditionally designed dishes. A large well is provided outside the culture area for pH measuring without affecting the cultured samples. The dish has a flat bottom so it is ideal for heat transfer from below.

5 Safety symbols and labels

There are several user labels on the surfaces of the CultureCoin® to guide the user. User labels are shown below.

Table 5.1 Labels

Description	Image
<p>Packing box label:</p> <ol style="list-style-type: none"> 1. Shelf life is the time the unit can be in storage. If stored longer than the shelf life, the dishes must be discarded. 2. Stores temperature between +15 °C and +25 °C 3. Keep away from direct sunlight 4. Caution: Consult the accompanying documents for important safety-related information such as warnings and precautions that cannot, for a variety of reasons, be presented on the device itself 5. Consult instructions for proper use of the device 6. Do not use if the packing material is damaged 7. By prescription only 8. Keep dry 9. Do not re-use. 	 <p>The diagram shows a rectangular label with the following elements: <ul style="list-style-type: none"> 1. Shelf life: 1 year from _____ 2. Temperature range: 15°C to 25°C 3. Sun icon with 'Protect from heat' 4. Warning triangle with 'Attention' 5. Information icon with 'Consult IFU/DFU' 6. Damaged box icon with 'Do Not Use If Package Is Open Or Damaged' 7. 'Rx ONLY' icon with 'By Prescription Only' 8. Umbrella icon with 'Keep Dry' 9. 'STERILIZE' icon with 'Do not re-sterilize' </p>
<ol style="list-style-type: none"> 1. Model 2. Ref No. 3. Lot No. 4. Use – by date 5. Manufacturer’s address and country of origin 6. CE mark 7. Consult instruction for use 8. Prescription only 9. WEEE wheeled bin 10. Single use 11. Do not re-use 12. Keep dry 13. Keep away from direct sunlight 14. Temperature limitation 15. Do not use if the package is damaged 16. The method of sterilization. 	 <p>The diagram shows the 'CultureCoin' label with the following elements: <ul style="list-style-type: none"> 1. Esco Medical Technologies CultureCoin 2. REF 100336 3. LOT 4. Use by 5. Manufacturer: Esco Medical Technologies, Ltd, Draugysties 19, Kaunas, 51230 Lithuania 6. CE 0088 7. Information icon with 'Consult instruction for use' 8. Prescription only icon 9. WEEE wheeled bin icon with 'Not for general waste' 10. Single use icon 11. Do not re-sterilize icon 12. Keep dry icon 13. Sun icon with 'Keep away from direct sunlight' 14. Temperature limitation icon (15°C to 25°C) 15. Damaged box icon with 'Do not use if the package is damaged' 16. Sterilization icon: STERILE R Sterilization using irradiation <p>Additional text on the label includes: 'Sterile only if package is undamaged!', 'Caution: 1-Cell MEA ≥ 80% expanded blastocysts at 96h. This lot is certified non-pyrogenic at less than 20 endotoxin units/device. Sterility assurance level (SAL) of 10⁻⁶'.</p> </p>

6 Important safety instructions and warnings

6.1 Pre use

1. Do not use the product, if the package is damaged. Contact Esco Medical or the local representative.
2. Read the user manual completely before use.
3. Always keep these instructions at hand.

6.2 During use

1. Use only for the purpose listed under intended use.
2. The dishes are single use only. Do not attempt to clean them and reuse.

2. Only for the use with water based culture media and mineral oil.
3. Do not use this product at temperatures exceeding 40 °C.

6.3 Post use

1. Discard the dishes according to the proper procedures.

7 The Culture mode

The CultureCoin® can be used only for culturing with culture media filled into the wells and oil overlay. They cannot be used for open culture.

8 Overview

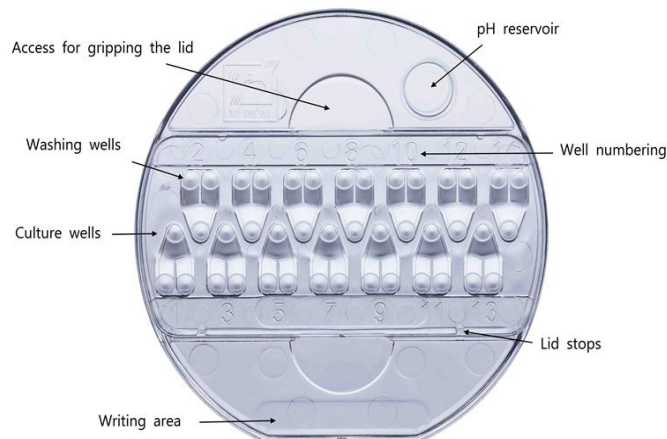


Figure 8.1 CultureCoin ®

9 Use

1. Unpack both the lid and dish in an aseptic environment.
2. Leave the dishes with the lids on for off gassing in the aseptic environment for 12 hours.
3. Warm the dishes on a heated stage or an incubator prior to filling with media.
4. Fill all the wells with app. 25 µl of pre cultured equilibrated culture media in an aseptic environment. Filling of the washing wells is optional. They can contain approximately 23 µl each.
5. Check under a microscope for any visible air bubbles and remove them with a stripper tip.
6. Fill the oil area with a confluent layer of suitable mineral or paraffin oil, so that it covers the culture media. Do not overfill.
7. Put the lid on the dish and leave it for 4 hours of equilibration in the CO₂ incubator.

8. Check for air bubbles in the culture wells after the equilibration. Remove any bubbles with a stripper tip.
9. Load the embryos in the wells. Make sure the embryos are located very precisely in the center of the culturing well and not on the side slope.
10. Optionally fill the pH reservoir and insert the silicone plug.
11. Incubate the CultureCoin® in the MIRI TL.

10 The use illustrated

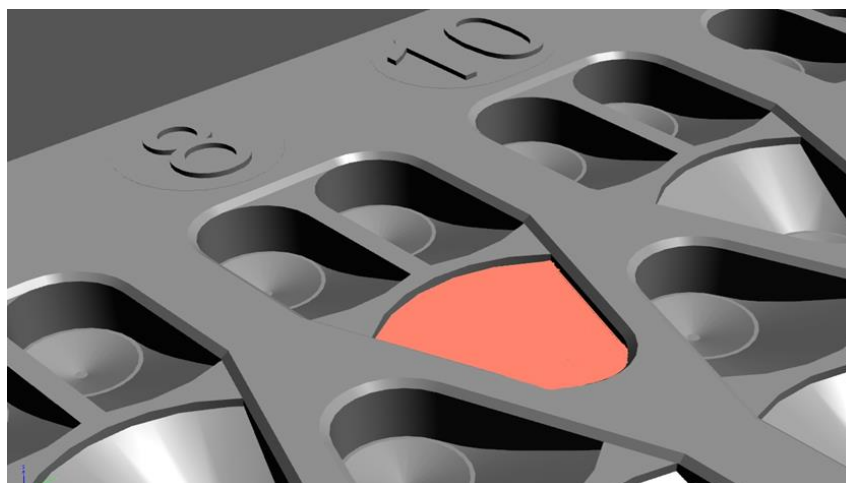


Figure 10.1 The wells

Each of the 14 wells are filled with approximately 25 μ l culture medium. The washing wells can also be filled but it is not a requirement. The embryo is placed at the bottom of the culturing area.

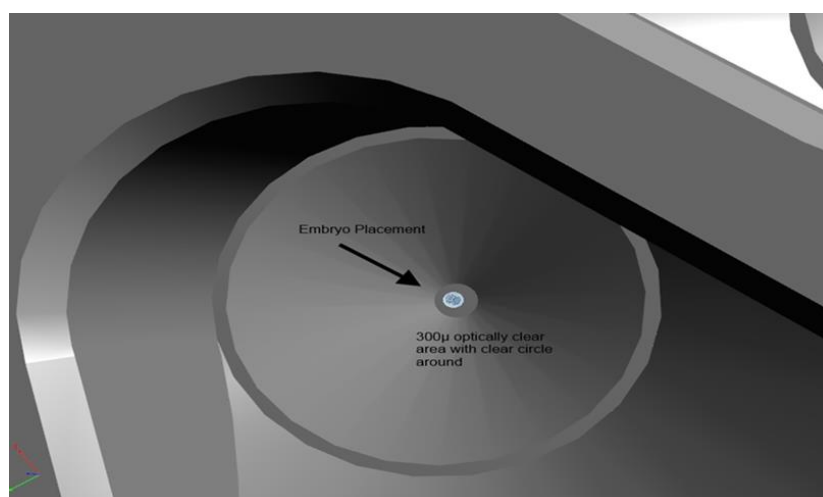


Figure 10.2 Embryo Placement in detail on the 300 μ optically clear area.

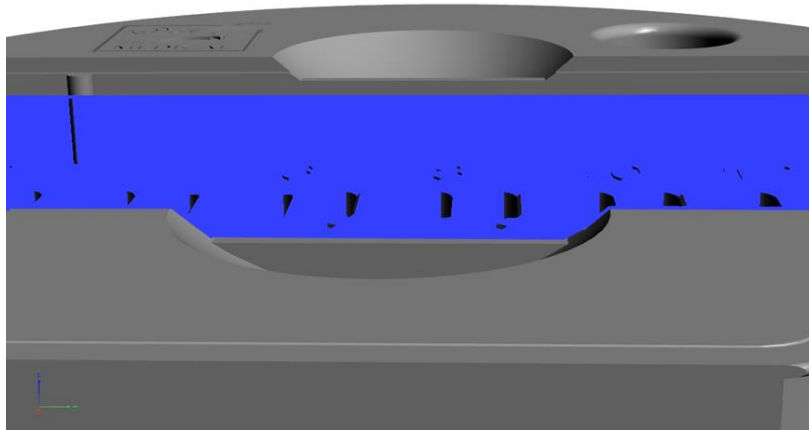


Figure 10.3 The culture wells are covered with a mineral oil layer and the lid is put on.

11 pH measuring

Validating the pH of the culture media should be a standard procedure. The CultureCoin[®] is equipped with a reservoir that is located outside the culturing area. This makes it possible to safely and conveniently make spot checks on the pH level.

Remove the silicone plug and measure with a calibrated pH probe in the reservoir. Adjust the CO₂ level if necessary.

A micro probe that can measure in a droplet can also be supplied.



Figure 11.1 Micro pH probe for 0.5 µl drop measurement

12 Writing area on the CultureCoin ® and lid



Figure 12.1 Always mark both the lid and the dish

13 User troubleshooting

Table 13.1 Heating system

Symptom	Cause	Action
Embryos drift to the side of the well.	The placement of the embryo is not done with enough accuracy.	Practice correct placement.
Air bubbles appear during culturing.	Temperature differences during loading or incorrect procedure for bubble removal.	Avoid temperature differences and follow the correct procedure. Practice correct bubble removal.
Oil overflows during handling of the dish.	Too much oil in the reservoir.	Decrease oil volume.

14 Specifications

Table 14.1 CultureCoin® specifications

Technical specifications	CultureCoin®
Overall dimensions (Diameter x Height)	Ø 71 x 10 mm
Weight empty	0.5 gram
Material	Styrolution PS 158 N
Temperature range	25 – 40 °C
CO ₂ range	1.9 – 10%
O ₂ range	20 – 4.9%
Sterilization method	Gamma beam
Lifetime	1 year
Toxicity tested	Embryo toxicity tested with thawed 1 cell mouse embryos. Expanded Blastocyst rate after 96 hours > 80 % (n=150) 0.6 bar (8.70 PSI)