Instructions for the Use of global® total® w/ HEPES

(Catalogue Numbers: LGTH-050, LGTH-100, LGTH-250)

PRECAUTIONS AND WARNINGS

1. **Caution:** Federal Law (USA) restricts this device to sale by or on the order of a physician (or properly licensed practitioner).

2. **Caution:** The user should read and understand the Instructions for Use, Precautions and Warnings, and be trained in the correct procedure before using global® total® w/ HEPES for human oocyte and embryo washing and manipulation, ICSI or embryo transfer.

3. Not to be used for injection.

4. Do not resterilize.

5. Do not use the product if:
   - the product packaging appears damaged or if the seal is broken
   - the expiry date has been exceeded
   - the product becomes discolored, cloudy, or shows evidence of particulate matter

6. This product contains human serum albumin, a derivative of human blood. The human serum albumin used in the preparation of this product has been heated at 60°C for ten hours.

   **Caution:** Standard measures to prevent infections resulting from the use of medicinal products prepared from human blood or plasma include selection of donors, screening of individual donations and plasma pools for specific markers of infection and the inclusion of effective manufacturing steps for the inactivation/removal of viruses. Despite this, when medicinal products prepared from human blood or plasma are administered, the possibility of transmitting infective agents cannot be totally excluded. This also applies to unknown or emerging viruses and other pathogens. There are no reports of virus transmissions with albumin manufactured to European Pharmacopoeia specifications by established processes. It is strongly recommended that every time that global® total® w/ HEPES is administered to a patient, the name and batch number of the product are recorded in order to maintain a link between the patient and the batch of the product.

7. global® total® w/ HEPES contains the antibiotic gentamicin sulfate. Appropriate precautions should be taken to ensure that the patient is not sensitized to this antibiotic.

8. To avoid problems with contamination, practice aseptic techniques.

9. Discard unused medium within 7 days of opening. Do not use after expiry date.

GENERAL INFORMATION

**Indications for Use**

Human oocyte and embryo washing and manipulation, fertilization by intracytoplasmic sperm injection (ICSI), embryo transfer.

**Storage and Shelf Life**

Store at 2-8°C and protected from light. Ten (10) weeks from the date of manufacture.

**Disposal Consideration**

Treat or dispose of waste material in accordance with all local state/provincial, and national requirements. Dispose with laboratory waste.

**Composition**

A HEPES-buffered medium is required when oocytes and embryos are handled outside of a CO₂ incubator.
### QUALITY CONTROL SPECIFICATIONS

<table>
<thead>
<tr>
<th>Assay (performed for each batch)</th>
<th>Specification</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Physicochemical Tests</strong></td>
<td></td>
</tr>
<tr>
<td>pH</td>
<td>7.2-7.4</td>
</tr>
<tr>
<td>Osmolality</td>
<td>260-270 mOsM</td>
</tr>
<tr>
<td><strong>Biological Tests</strong></td>
<td></td>
</tr>
<tr>
<td>Endotoxin (LAL)</td>
<td>≤ 0.5 EU/ml</td>
</tr>
<tr>
<td>Sterility Test (bacterial and fungal screen, SAL 10⁻³)</td>
<td>PASS</td>
</tr>
<tr>
<td><strong>Biological Assays</strong></td>
<td></td>
</tr>
<tr>
<td>1-cell Mouse Embryo Assay (% expanded blastocysts at 96 h of culture)</td>
<td>≥ 80%</td>
</tr>
</tbody>
</table>

### INSTRUCTIONS FOR USE

The procedures described below have been found to be effective for human oocyte and embryo washing and manipulation, ICSI or embryo transfer. Every laboratory must define and optimize its own procedures.

1. For oocyte and embryo washing and manipulation, prepare culture dishes containing 25-100 μl droplets under oil, or in larger volumes (0.5-1.0 ml) of global® total® w/ HEPES, according to general laboratory practice.

2. Before introducing the oocytes or embryos, warm the culture dishes for sufficient time to ensure temperature equilibration.

3. For washing, pass the oocytes or embryos through several droplets of global® total® w/ HEPES, according to general laboratory practice.

4. Before ICSI or other manipulation, wash the oocytes or embryos, according to general laboratory practice.

5. For ICSI, prepare 5-10 small droplets (5-10 µl) of global® total® w/ HEPES in the ICSI dish, next to the PVP droplets and cover them with oil, according to general laboratory practice. Transfer the oocytes to the respective droplets just before the ICSI procedure.

6. For transfer, wash the embryos according to general laboratory practice, and transfer to the uterus in 20-30 µl of global® total® w/ HEPES.

7. Immediately prior to transfer, rinse the transfer catheter with global® total® w/ HEPES.

### SYMBOLS

- **STERILE**
- **RX Only**
- **REF**
- **LOT**
- **Consult Instructions For Use**
- **Manufacturer**
- **Keep Away From Sunlight**
- **Temperature Limitation**
- **Authorized Representative in the European Community**
- **Use By**
- **GS1 DataMatrix Barcode**
- **Do Not Resterilize**