

Instructions for Use



Sperm Separation Device | ZMH3000

Important Information:

- Carefully adhere to recommended volumes for each step. Avoid over- or under-filling the device.
- Do not exceed 30 minute incubation time.
- Keep the device level during use – do not tip or rock.
- Device is single-use only and should be restricted to a single individual per device. It may not be reused.
- Do not re-use or re-sterilize device or device can be damaged and will no longer effectively separate sperm.

Note on Incubation:

Good tissue practices necessitate matching media to incubation conditions. If using a bicarbonate-buffered media, incubate in a humidified, 37°C, gassed incubator. If using a HEPES-buffered media, incubate in a humidified, non-gassed incubator. If no incubator with humidity is available, add a 35mm dish of deionized or distilled water, uncovered, to the Petri dish containing the device before placing the covered dish with the device and the 35mm dish into the 37°C incubator.

PREPARATION

1. Gather your supplies and work on a clean surface.
2. Incubate semen sample at 37°C for 20-30 minutes to allow for liquefaction.
3. Carefully open the device package without touching the device membrane.

DRAW SAMPLE

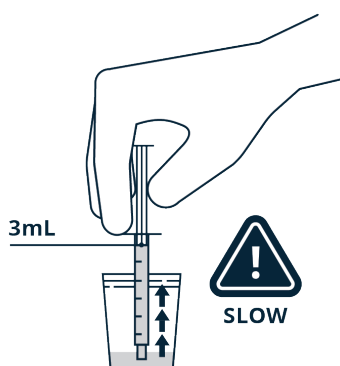


Figure 1. Draw 3mL of the sample.

4. Use a 5mL Luer-tip syringe to slowly draw a 3mL aliquot of the liquefied semen specimen. If there is insufficient volume, add sperm washing solution to give 3mL (Figure 1).

INJECT SAMPLE

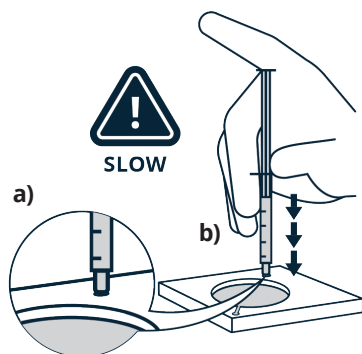


Figure 2. a) Achieve seal. b) Slowly inject sample.

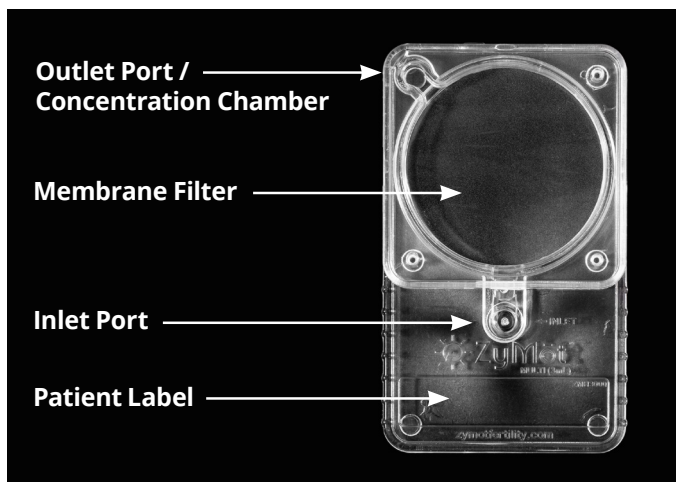
5. Holding the syringe in a vertical position, carefully insert the tip into the Inlet Port and apply gentle pressure to achieve a seal (Figure 2a). With gentle and steady pressure, inject the sample (Figure 2b). Be careful to avoid the formation of bubbles under the membrane.

Device Components:

- ZyMöt® Multi (3mL) Sperm Separation Device
- Instructions for Use

Materials/Equipment Required, But Not Supplied:

- Sperm washing solution (media): bicarbonate- or HEPES-buffered media supplemented with 2-10% protein
- 37°C incubator
- 5mL Luer-tip syringes (3) - Recommended: Norm-Ject #4050-000VZ, Henke Sass Wolf
- Sperm-safe culture tube



ADD MEDIA

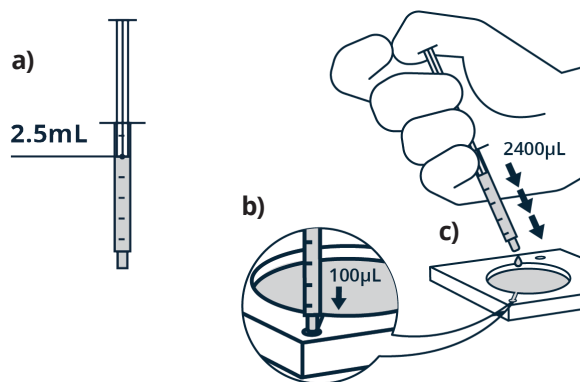


Figure 3. a) Draw 2.5mL of media. b) Prime outlet channel. c) Cover membrane surface.

6. Prepare a fresh syringe with 2.5mL of media (Figure 3a).

a) Prime the Outlet Port/Concentration Chamber by injecting a small volume of media (approximately 100µL - Figure 3b), until the media travels through the channel to the membrane.

b) Disconnect the syringe from the Outlet Port and apply the remaining media (2400µL) in the syringe to the surface of the upper membrane by dropping from approximately 2cm above the membrane (Figure 3c).

c) Completely cover the upper membrane with media, making sure media touches all the edges of the upper chamber and connects with the droplet of media that was used to prime the Outlet Port.

Note: Do not tilt the device to spread the media.

INCUBATE SAMPLE

7. Place device into a Petri dish and cover. Keep the ZyMöt device horizontal and covered at all times during the incubation. Incubate at 37°C for 30 minutes.

COLLECT SEPARATED SPERM

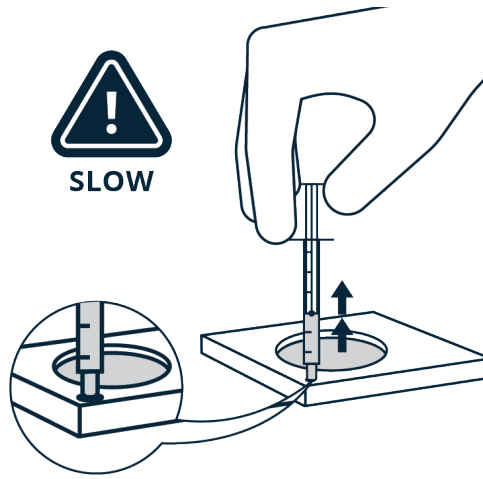


Figure 4. Slowly aspirate 1.0mL - 1.5mL.

- Insert a fresh 5mL syringe into the Outlet Port of the device. Slowly aspirate 1.0mL - 1.5mL of the sperm containing fluid (Figure 4).

Tips, Warnings and Precautions:

- Caution: Sale of this device is restricted to sale by or on the order of a physician.
- Device should be used only by properly trained operators.
- Practice universal precautions when handling human body fluids.
- Do not use if the packaging is damaged.
- Do not clean device with any solutions. Spraying or wiping with cleaning solutions can damage the device.
- Single use only — do not re-use or re-sterilize the device — it will no longer effectively separate sperm.
- Serious incidents should be reported to ZyMöt Fertility, Inc. or to the Dutch Competant Authority through our Authorized Representative MedEnvoy Global BV, Prinsessegracht 20, 2514 AP The Hague, The Netherlands, +31 70 326 2148, or vigilance@medenvoyglobal.com.

Device Description:

ZyMöt ICSI and ZyMöt Multi are sperm separation devices used to prepare motile sperm for assisted reproductive technology (ART) procedures. Both devices separate sperm based on motility. The ZyMöt ICSI and the ZyMöt Multi are sterile and single use only. The mechanism of action for both is separation of sperm based on motility within a microenvironment created by the micro channels of the ZyMöt ICSI or the micropores in the filter of the ZyMöt Multi. The primary difference between the devices is the processing volume. The ZyMöt ICSI has a processing volume of 2µL per micro channel. The ZyMöt Multi is manufactured in two (2) processing volumes, 850µL and 3mL.

The ZyMöt Multi (provided with 850µL and 3mL collection chambers) has an inlet port that communicates with the lower sample chamber. The sample chamber is separated from the upper collection chamber by a microporous filter. Untreated semen is added through the inlet port. After 30 minutes, the separated sperm are collected from the upper chamber through the outlet port/concentration chamber.

Indications for Use:

The ZyMöt Multi (3mL) Sperm Separation Device is intended for preparing motile sperm from semen for use in the treatment of infertile couples by intracytoplasmic sperm injection (ICSI), in vitro fertilization (IVF), and intrauterine insemination (IUI) procedures.

Sterilization:

The sterilization method used for the ZyMöt devices is gamma radiation, at a dose level of 25kGy to 45kGy by the VD_{max}^{25} method to meet a Sterility Assurance Level of 10^{-6} .

Storage:

Store at 15°C - 25°C (60°F - 77°F).

Disposal:

Discard the used device and materials as medical waste.

Testing Performed for Devices Used in Assisted Reproduction:

Specific testing was performed for toxicity and functional screening appropriate for products used in assisted reproduction. As required by USFDA 21 CFR 884.6160, the following Special Controls were conducted (all tests were passed): human sperm survival assay (replacing the mouse embryo assay) and endotoxin testing.

SAMPLE HANDLING AFTER COLLECTION FOR ICSI AND IUI

- Transfer the collected sample to an appropriate culture tube: a 4mL round bottom culture tube with a snap top or into the bottom of a 15mL conical tube. Tubes using HEPES-buffered media may be held on the benchtop or tightly capped in an incubator. Tubes using bicarbonate-buffered media should be stored in a CO₂ incubator with the lid loosely closed.

SAMPLE HANDLING AFTER COLLECTION FOR IVF

- Transfer the collected sample into a 15mL conical tube. Add 3mL of bicarbonate-containing media (whatever media is usually used for the final suspension of sperm for conventional insemination) to the conical tube. Mix gently. Centrifuge the conical tube for 5 minutes at 300 x g. Remove the supernatant, being careful to not disturb the lower pellet. Perform count and motility as usual and dilute if needed to achieve appropriate final insemination concentration. Store tube in a CO₂ incubator until insemination. Insemination should occur more than 1 hour, but less than 4 hours after preparation.

Endotoxin Testing Results:

Using the Limulus Amebocyte Lysate (LAL) Analysis by the Gel-Clot Method, results were <0.0729 EU per device, which meets the acceptance level of ≤20 EU per device.

Human Sperm Survival Assay Results:

Using the Human Sperm Survival Assay, results were 96.2% for ZyMöt ICSI and 97.7% for ZyMöt Multi; both results meet the acceptance level of motility ≥80% of control at 24h after exposure for 30min.

Note: The above results are from testing required prior to USFDA 510(k) clearance. These tests are conducted on each manufacturing lot of devices as part of the lot release program. A CoC can be provided upon request.

Legal manufacturer:

ZyMöt Fertility, Inc., a DxNow, Inc., subsidiary
401 Professional Drive, Suite 130
Gaithersburg, Maryland 20879-3429 USA | zymotfertility.com



















Manufactured by:

Natech Plastics, Inc.
85 Remington Blvd.
Ronkonkoma, New York 11779 USA | natechplastics.com

Licenses, Patents and Trademarks:

DxNow, ZyMöt, ZyMöt Multi and ZyMöt ICSI are trademarks of DxNow, Inc. Devices are manufactured and sold under the terms of DxNow's exclusive worldwide patent license from The Brigham & Women's Hospital, Inc., Boston, Massachusetts, USA.

Patents: US10422737B2; US11009444B2; CA2931201C; ES2915379T3; DK3071704T3; JP6524082B2; JP6850824B2; AU2014353050B2; EP3071704B1. Additional USA and other international patents pending.

 Manufacturer	 Sterilized Using Irradiation	 Storage Temperature Limits 15°C - 25°C
 Date of Manufacture	 Do Not Resterilize	 Do Not Reuse
 Expiration Date	 Do Not Use if Package is Damaged	 Consult Instructions For Use
 Lot Number	 Fragile Item, Handle Carefully	 Caution
 Catalog Number	 Keep Dry	 CE 2460
 MedEnvoy Global BV Prinses Margrietplantsoen 33 Suite 123 2595 AM, The Hague The Netherlands	 EC REP MedEnvoy Global BV Prinses Margrietplantsoen 33 Suite 123 2595 AM, The Hague The Netherlands	 MD