

# **NEST BioFactory™ NEST**

NEST BioFactory<sup>™</sup> system is a well-designed cell culture device that utilizes the maximum culture area in a limited space, saving a large amount of space, low-cost to achieve the purpose of expanding production capacity.

NEST BioFactory<sup>TM</sup> system can be used for industrial scale production such as vaccine, monoclonal antibodies or biopharmaceuticals production, and they are suitable for adherent cell culture.

Specifications: Single layer, 2 layers, 5 layers, 10 layers and 40 layers.

Two Wide Vent Caps (with extra 16 sterile wide plug seal caps packaged separately)						
Cat.No.	Layer	Cultivation area (cm²)	Length(mm)	Width(mm)	Height (mm)	/Case
771001	1	647	335	205	44	8
771101	2	1279	335	205	61	8
771204	5	3175	335	205	112	4
771302	10	6335	335	205	197	6
771403	40	25295	335	205	712	2



### **1 Narrow Vent Cap + 1Narrow Plug Seal Cap** (with extra 8 sterile narrow vent caps & 8 sterile narrow plug seal caps packaged separately)

Cat.No.	Layer	Cultivation area (cm²)	Length(mm)	Width(mm)	Height (mm)	/Case
772001	1	647	335	205	44	8
772101	2	1279	335	205	61	8
772204	5	3175	335	205	112	4
772302	10	6335	335	205	197	6
772403	40	25295	335	205	712	2



### 1 Wide Vent Cap & 1 Narrow Plug Seal Cap

(with extra 8 sterile wide plug seal caps & 8 sterile narrowvent caps packaged separately)

Cat.No.	Layer	Cultivation area (cm²)	Length(mm)	Width(mm)	Height (mm)	/Case
773001	1	647	335	205	44	8
773101	2	1279	335	205	61	8
773204	5	3175	335	205	112	4
773302	10	6335	335	205	197	6
773403	40	25295	335	205	712	2



# **BioFactory<sup>™</sup> Versus Rolling Bottles**

# NEST

#### Advantages of BioFactory<sup>™</sup>

- Large culture area with less space occupied;
- Reduced manual operations to decrease contamination risk;
- Can be operated manually or combined with automated equipment for use;
- Can realize standardized operations to reduce intra-batch difference.

#### **Disadvantages of Rolling Bottles**

- Heavy workload of manual operations;
- High risk of contamination;
- A large amount of washing and verification processes;
- Poor controllability of the production processes with large intra-bottle difference, hard to unify the quality of different batches;
- Large space occupation, hard to rapidly expend the production scale;
- Low level of automation.

### **Comparison of Utilization Rate**

Compared Items	10-layer BioFactory™	Rolling Bottle	
Individual Culture Area	6,335cm <sup>2</sup>	1,800cm <sup>2</sup>	
Quantity	16	50	
Total Culture Area	10m <sup>2</sup>	9m²	
Occupied Space	0.216m <sup>3</sup>	0.973m <sup>3</sup>	



# Design Characteristics of BioFactory<sup>™</sup> NEST

#### Large-caliber Connection Channel

Large-caliber connection channel may quickly equilbrium the culture medium and shorten the liquid adding progress, to better increase cell viability.



One-piece Injection Molding for Mouth and Body

Ensure better sealability and better strength.





#### **Strong Packaging**

Double-layer air-exhausted packaging: meet the use requirements of pharmaceutical clients. Built-in protection layer: hard to be damaged, suitable for long-distance transportation.







#### **Unique Design of Cap**

"Petal" pattern distribution of pores, to enhance gas exchange to facilitate cell culture.



## **User Manual**

NEST

#### **Cell Culture**





Pour the prepared culture medium into the cell factory (Volume suggestion: 150 - 200 mL for each layer).

Lay the cell factory on its side to balance the liquid level.



With the liquid-adding opening upward, the culture medium will be evenly distributed to each layer after standing.



Slowly place the cell factory in a horizontal position.



Incubate in a cell incubator.

#### Precautions

- 1. Please pre-heat the cell factory and culture medium to the culture temperature: since it takes a long time for a large incubator to reach the set culture temperature, pre-heating the cell factory and culture medium to the culture temperature before starting the experiment may speed up cell attachment and significantly increase cell recovery.
- 2. Slow operating is required to avoid occurrence of air bubbles caused by sharp shaking: air bubbles may lead to flowing the medium from an upper layer to a lower layer.
- 3. Avoid spraying the alcohol onto the breathable cover, since alcohol may wet the hydrophobic membrane filter and make it impermeable and consequently affect the gas exchange or causes pressure imbalance during operations.

#### **Cell Harvest**

- 1. After the culture is completed, pour the culture medium out.
- 2. Wash the factory with the calcium- and magnesium-free phosphate buffer solution (CMF-PBS) (40-50 ml/layer) and if necessary, repeat the washing process.
- 3. Digestion: pre-heat the digestion solution (10-40 ml/layer) in advance.
- 4. Collection: centrifuge for 5 min at 1000 rpm, remove the digestion solution and collect cells.
- 5. Washing: wash the incubator with CMF-PBS or culture medium after digestion.

#### **Precautions**

- 1. Ensure that the culture surface of each layer is completely immersed in the CMF-PBS, and gently shake the cell factory forward and backward to wash off the residual culture medium.
- 2. Distribute the digestion solution evenly to each layer; gently tilt the incubator forward and backward, left and right to ensure that the digestion solution has completely covered the culture surface; gently tap the incubator to help the cells detach from the surface.
- 3. Since it is unable to clearly observe the digestion status of the cells in the middle layers of a cell factory, it is recommended to refer to the digestion status of a culture flask or a single-layer cell factory under exactly the same culture conditions. Or, use a dedicated observation platform for multiple-layer cell incubators to observe the growth status of cells in each layer.
- 4. If there are numerous cells present in the washing solution or the culture layers of the cell factory, it is necessary to wash for multiple times or adjust the procedure of cell digestion.
- 5. Even a slight deviation of the culture temperature may affect the cell harvest rate, so it is required to pay close attention as to whether or not the culture temperature is exactly the set temperature.





# **BioFactory™ Automatic Shaker NEST**

Achieve automated and high-efficiency operating of liquid adding, shaking, digesting and liquid pouring-out, etc. for BioFactory<sup>™</sup> during large-scale culturing, to reduce heavy manual workload and the errors introduced by human factors.

- The whole device is fully made of stainless steel materials, without rubber transmission parts.
- The device is fully electricity-powered; the ACFM is strictly in accordance with the requirements of EU directives; all parts comply with relevant regulations, no compressed gas source required.
- Use a complete set of imported servo motors and imported reducers, with high operating safety factors.
- Noiseless operation is achieved by using imported linear guides and ball screws.







BioFactory<sup>™</sup> Handcart









Lean Back: 0-35°



Left Rotation, Right Rotation It's available to lean back or forward while rotating.









## **BioFactory<sup>™</sup> Accessories**





744001 SPT-50 Hose Inner Dia# 3/8 Inches (9.5 mm) , Outer Dia# 5/8 Inches (15.9 mm) 50 Inches/ 15 meters 1 pcs / pack



746001 C-Flex Welding Hose Inner Dia# 3/8 Inches (9.5 mm) , Outer Dia# 5/8 Inches (15.9 mm) 50 Inches/ 15 meters 1 pcs / pack



743001 Hose Clamp Hose clamp for 12-18 mm diameter hose 10 pcs / pack



741001 Adaptor Connector Adaptor Connector to 3/8 Inches (9.5 mm)Hose 1 pcs / pack



**751001 Y Shape Connector** Y Shape CPC Connector for Inner Dia #3/8 Hose 1 pcs / pack, 5 pcs / cs



749001 T Shape Connector T Shape CPC Connector for Inner Dia #3/8 Hose 1 pcs / pack, 5 pcs / cs



 742001
 Vent Filter

 0.22 μm, 50 mm diameter
 1 pcs / pack, 5 pcs / cs

 742011
 Vent Filter

 0.22 μm, 45 mm diameter
 1 pcs / pack, 5 pcs / cs



747001 Connector CPC & PC (Inner Dia #3/8) for Hose to Hose Connecting 1 pcs / pack, 5 pcs / cs



740001Wide Mouth Seal cap740011Wide Mouth Vent cap740101Narrow Mouth Seal cap740111Narrow Mouth Vent capHDPE,1 pcs / pack, 10 pcs / cs



740201 Adaptor Cap Wide Mouth to Narrow Mouth HDPE, Spec(mm): 11.8/14.3 1 pcs / pack, 10 pcs / cs



740301 Adaptor Cap Wide Mouth to 3/8 Inches (9.5 mm)hose HDPE, Spec(mm): 7.3/11.4 1 pcs / pack, 10 pcs / cs



745001 Accessory Lite Package

Hose Clamp\*1 50 mm Vent Filter\*1 15 cm SPT-50 Hose\*1 Silicone Ring\*2 1/pk, 2/cs



745011 Accessory Basic Package Hose Clamp\*1 50 mm Vent Filter 15 cm SPT-50 Hose\*1 Adaptor Connector\*1 Silicone Ring\*2 1/pk, 2/cs



745021 Accessory Premium Package 2 different Adaptor Caps\*1 Hose Clamp\*1 Adaptor Connector\*1 50 mm Vent Filter\*1 Hose Clamp\*1 15 m SPT-50 Hose\*1

### How to use a microscope to observe the cell growth in a BioFactory™?

- 1. Use an inverted microscope to observe directly, however, it's only suitable for observing a BioFactory<sup>™</sup> with 1-10 layers. The growth status in the bottom layers 1 and 2 can be observed, but only for the cell growth uniformity. It is not suitable for a 40-layer BioFactory<sup>™</sup>. Please contact the microscope manufacturer for more details.
- 2. Observe with a microscope dedicated for BioFactory<sup>TM</sup> observation, which has a lens in two directions for side observation and bottom observation, respectively; the cell size and growth density can be calculated using a size measurement system by observing cell status with a microscope. This model of microscope is suitable for BioFactory<sup>TM</sup> with any number of layers.
- 3. Since the cell growth conditions (surface-capability relationship) of NEST BioFactory<sup>™</sup> are the same as those of NEST cell culture flasks, the cell culture flask can be used simultaneously to replace BioFactory<sup>™</sup> for observation, or use a 1-layer BioFactory<sup>™</sup> for reference.

#### How many cells may be inoculated in each type of BioFactory<sup>™</sup>?

NEST BioFactory<sup>TM</sup> are made of the same raw materials using same production processes as those of NEST cell culture flasks and the cell surface and growth status are basically the same as that of cell culture flasks. Since each kind of cells have different growth characteristics and different requirements for surface density and environment, for the number of cells that can be inoculated in each type of BioFactory<sup>TM</sup>, it cannot be simply generalized. Users are recommended to determine the number by multiplying the inoculation density per unit area in the cell culture flask with the used culture area of BioFactory<sup>TM</sup>.

#### Volumes of various working solutions used in the cell culture.

Layer	Culture Area (m <sup>2</sup> )	Volume (mL)	Washing Solution (1XPBS) mL	Digestion solution EDTA (mL)	Digestion Stop Solution (mL)
1	647	150-200	30-40	20	25
2	1279	300-400	60-80	40	50
5	3175	750-1000	150-200	100	125
10	6335	1500-2000	300-400	200	250
40	25295	6000-8000	1200-1600	800	1000

The BioFactory<sup>™</sup> allows large-scare cell culture in a compact space using its multiple-layer stacked structure. How much culture medium should be added in during use? And how much trypsin solution should be added in for cell digestion?

# The BioFactory<sup>™</sup> are divided into large-caliber ones and small-caliber ones, what are their respective characteristics?

- 1. After years of research and development, the NEST BioFactory<sup>™</sup> now have a complete product selection and clients may freely choose from BioFactory<sup>™</sup> with of two wide caps, with two narrow caps or with one wide cap and one narrow cap according to their operating and use needs.
- 2. Characteristics of two wide caps: suitable for manual operations of adding and removing liquid, can also be connected with tubing; generally used during the phase of laboratory research.
- 3. Characteristics two narrow caps: suitable for being connected with tubing to facilitate subsequent process scale-up; generally used for scale production. Characteristics of one wide cap and one narrow cap: suitable for tubing connection and manual operations; used for scale production and laboratory research. Breathable cap: the inner is covered with a 0.22um hydrophobic film, which is air penetrable and can block bacteria and water, as well as avoid gas bloating caused by adding liguid.

### High Efficiency Erlenmeyer Flask Transfer Cap

## High Efficiency Erlenmeyer Flask NEST



#### Introduction

High-efficiency, large-volume culture flasks allow cells to show strong viability with large expression amount of proteins in the culture of mammalian cells and insect cells. During the culturing process, the use rate of the shaker is significantly increased, and the survival rate and viability of cells are both dramatically elevated. NEST culture flasks also provide high repeatability, which allows highly inter-batch consistency of cell growth and yield.

#### **Feature**

- The imported medical-grade polycarbonate (PC) that meets with the requirements of ISO10993 USP<661> is chosen as the raw materials, which has high transparency, great impact resistance, oxidation resistance and can withstand a high temperature of up to 121 °C.
- The scale is clear and accurate when looked at from the outside, which facilitates the observation of the volume of culture medium.
- The vent cap is equipped with a 0.22-µm breathable film which is air-permeable and water-proof to efficiently prevent the passing of micro-organisms and contamination and to ensure gas exchange for good growth of cells or bacteria.
- The products have undergone the tightness test, drop test, high-temperature and high-voltage tests, flatness test, tensile test, endotoxin test, sterilization test, DNA/RNA enzyme-free tests and cell culture test, all the results of which meet the corresponding quality standards.
- Sterile individual package, easy to use.

Cat.No. Volume (		ume(L)    Cap Style		10000		
			Height	Bottleneck Diameter	Bottom Diameter	70a36
785101	2	Seal Cap	213.5	67	162	6
785111	2	Vent Filter Cap	213.5	67	162	6
786101	3	Seal Cap	253.5	67	162	4
786111	3	Vent Filter Cap	253.5	67	162	4
787001	5	Seal Cap	285.5	90	230	4
787011	5	Vent Filter Cap	285.5	90	230	4

# **High Efficiency Erlenmeyer Flask**



#### **Design Characteristics of High Efficiency Erlenmeyer Flask**

- 1. The lower edge of the cover is thickened and widened for better leak proofing.
- The 5L high efficiency erlenmeyer flask is designed with an detachable handle to solve the problem that liquid enters the cavity of the one-piece infected handle as residue. The handle is detachable for flexible application.
- 3. The designs of 2/3 L flasks are optimized with lengthened neck convenient for customers to grip and can avoid that the hand of the customer is too close to the flask mouth when pouring liquid or holding the flask. Meet the aseptic operating practices of pharmaceutical companies.
- The bottom area and opening size of the NEST high efficiency erlenmeyer flask are exactly the same as those of the Thomson.



#### High Efficiency Erlenmeyer Flask Versus Normal Erlenmeyer Flask

#### High Efficiency Erlenmeyer Flask Advantage

- 1. The bottom area of 3L high efficiency erlenmeyer flask is smaller than normal erlenmeyer flask. For a same shaker, more 3L high efficiency erlenmeyer flask can be placed on to which greatly reduce the usage rate of the shaker to save the client's R&D costs.
- 2. Low sheer force born by cells, The amount of foam can be efficiently controlled.



# Transfer Cap NEST

#### **Inverted Liquid Transfer Cap**

The inverted transfer cap is used along with a 2L, 3L or 5L Erlenmeyer Flask to connect a liquid inlet tubing with the required device. When liquid transfer is required, the liquid is transferred under gravity with the inverted erlenmeyer flask.

### **Multifunctional Liquid Transfer Cap**

Unlike the two-way liquid transfer cap, the multifunctional transfer cap can be directly placed in an incubator for culture after the liquid transfer is completed. It can reach a large air flux. The sampling part is composed of a sampling nozzle and a one-way valve, which can prevent the liquid from flowing backwards during the sampling process and ensure the aseptic sampling. The liquid inlet tubing is provided with a PTFE needle filter, which solves the issue of liquid remaining in the tubing during the feeding process.

### **Two-way Liquid Transfer Cap**

The two-way transfer cap is used along with a 2L, 3L or 5L shake flask to connect a liquid inlet tubing with the required device. The liquid transfer is achieved by connection of a peristaltic pump between the erlenmeyer flask and the device. Upon completion of the transfer, the transfer cap can be replaced with a vent cap for culture.

#### Feature

- Closed system reduces the risks of contamination during liquid transfer.
- The cap is molded and connected in an integrated manner, reducing the risks of leakage and media residues.
- A variety of tubing diameters are available and aseptic welding of liquid inlet tubing under normal conditions is supported.
- High-quality materials and smooth inner wall of the tubing provide an excellent transfer performance.
- Sterility level (SAL) is maintained at 10<sup>-6</sup>.
- Endotoxin-free, and no ingredients of animal origin.

#### **Application**

• It is applicable to liquid transfer and culture during mass proliferation of bacteria and suspension cells.

Cat.No.	Name	Description of Hoses	Hoses' Length (cm)	/Case
785931	Multi-Function Liquid Transfer Cap System for 2L Erlenmeyer Flasks		50 cm	4
786931	Multi-Function Liquid Transfer Cap System for 3L Erlenmeyer Flasks		50 cm	4
787931	Multi-Function Liquid Transfer Cap System for 5L Erlenmeyer Flasks		92 cm	4
785921	Bi-directional Liquid Transfer Cap System for 2L Erlenmeyer Flasks		50 cm	4
786921	Bi-directional Liquid Transfer Cap System for 3L Erlenmeyer Flasks	Thermoplastic Hose.	50 cm	4
787921	Bi-directional Liquid Transfer Cap System for 5L Erlenmeyer Flasks	Diameter: 1/8" ID, 1/4" OD	92 cm	4
785941	Inverted Liquid Transfer Cap System for 2L Erlenmeyer Flasks		50 cm	4
786941	Inverted Liquid Transfer Cap System for 3L Erlenmeyer Flasks		50 cm	4
787941	Inverted Liquid Transfer Cap System for 5L Erlenmeyer Flasks		92 cm	4







Multifunctional Liquid Transfer Cap



Two-way Liquid Transfer Cap

### Wuxi NEST Biotechnology Co.,Ltd.

- C Tel : +86-510-6878 8698 / +86-510-6878 8718
- E-mail : info@nest-wuxi.com
- Web: www.cell-nest.com

NEST

👷 Add : No. 530 Xida Road, New District, Wuxi, Jiangsu, China

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#### **NEST Scientific USA**

- **%** Tel : +1-732 381 0268
- E-mail : info@nestscientificusa.com
- Web: www.nestscientificusa.com
- 👷 Add: 1592 Hart St., Rahway, NJ 07065, USA