

HyClone[™] media and supplements

CDM4Retino

HyClone CDM4Retino medium is a high-performance cell culture medium. The product is developed through the HyClone Metabolic Pathway Design process (see box) to increase the process yields for the industrial manufacture of adenoviral vectors and recombinant proteins using PER.C6TM cells. This chemically defined medium contains no proteins, animal-derived components, hydrolysates, or other undefined components. CDM4Retino medium is regulatory-friendly and capable of producing superior cell yields in a variety of culture environments and applications, including bioreactor cultures. CDM4Retino is available in liquid and powder formats in userfriendly packaging (Fig 1).

Features of CDM4Retino medium include

- Chemically defined formulation free of proteins and animal-derived components
- Developed through Metabolic Pathway Design process for high adenoviral vector and recombinant protein production
- Ready-to-use liquid formulation, requires no supplementation
- Allows for direct or sequential adaptation
- Designed for large-scale culture applications, including perfusion and fed-batch strategies
- Manufactured from traceable components according to cGMP guidelines

Specifications

- Protein-free
- Does not contain phenol red
- Liquid medium contains poloxamer 188
- Store at 2°C to 8°C away from light



Fig 1. CDM4Retino medium is available as liquid or powder in pack sizes suitable for small-volume cell culture as well as large-scale bioprocessing applications.

Metabolic Pathway Design process

An optimal cell culture process is dependent of a variety of factors including the parental cell line, the genetic makeup of the specific clone, medium and feed composition, as well as process variables to maximize viable cell densities and titers while maintaining cell morphology. Our experts in medium design and development know and understand how these factors can influence the metabolic processes involved. They evaluate the culture's metabolic activities, measuring nutritional demand and waste creation to make sure the correct type and quantity of nutrients are used to minimize waste and resultant cell toxicity. Our experts use their understanding of metabolic pathways to optimize medium composition for enhanced productivity and viable cell densities. Once a medium has been optimized using this Metabolic Pathway Design process, our scientists can help you devise the most effective cell culture strategy using a combination of medium and feeds to further enrich productivity and reduce process inefficiencies.

Suggested preparation Reconstitution of CDM4Retino powder medium

- While stirring, add 13.7 g/L of CDM4Retino powder medium to cell culture-grade water (room temperature) at 90% of final preparation volume. If your water source is normally cool, it may be useful to adjust the water temperature. Using warmer room temperature water (22°C to 25°C) will improve solubilization time. Mix for 20 min until dissolved.
- 2. Add 1.0 g/L poloxamer 188 and 2.0 g/L sodium bicarbonate. Allow 20 min to mix.
- 3. Bring vessel to final volume with cell culture-grade water. Allow solution to mix for 10 to 20 min.
- 4. Check pH and osmolality. Expected values:
 - pH 7.0-7.4
 - Osmolality 290 to 340 mOsm/kg
- 5. Sterile filter into desired container using a 0.2 μm sterile filter.

Preparation notes

CDM4Retino powder medium does not contain L-glutamine. Recommended concentration: 4 mM.

General culture recommendations

- 1. Cultures should be incubated at 37°C in a 5% CO_2 environment.
- 2. The caps on culture flasks should be loosened and adequate vessel headspace should be given to provide gas exchange.
- 3. Seeding densities should be ~ 3.0×10^5 cells/mL. Cells should typically be subcultured every 3 to 5 days, as necessary.

Direct adaptation

- 1. Transfer cells grown in current medium directly into CDM4Retino medium at 5.0×10^5 cells/mL.
- 2. When viable cell density reaches 2.0 to 4.0 \times 10 6 cells/mL, subculture the cells.
- 3. Cells should be subcultured every 48 to 96 h.
- 4. If cell viability drops below 80%, proceed to sequential adaptation.

Sequential adaptation

- 1. Transfer cells grown in current medium into CDM4Retino medium at a ratio of 1:1 using a seeding density of 5.0×10^5 cells/mL.
- 2. Incubate culture until two population doublings are observed. Subculture cells by mixing equal volumes of cell suspension in conditioned medium and fresh CDM4Retino medium (1:1 ratio).
- 3. Continue to subculture the cells using this method until the previously used medium is reduced below 0.05% concentration and cell viability is > 85%.

Cryopreservation

CDM4Retino medium adapted cells can be cryopreserved in a 1:1 mixture of fresh and conditioned CDM4Retino medium supplemented with 10.0% DMSO.

Quality control testing

Quality control test specifications are listed in Table 1.

Table 1. Test specifications¹

Appearance	Clear yellow solution
Osmolality	290 to 340 mOsm/kg
рН	7.0 to 7.4
Sterility	No growth (bacteria or fungi)
Endotoxin	< 10.0 EU/mL ¹
Application	Growth promotion

¹ Refer to certificate of analysis for actual results.

Custom production

Formulations and delivery systems can be customized to your specific process requirements or optimized to maximize process yields.

Rapid Response Production (RRP)

Our RRP program manufactures up to 200 L of your custom prototype formulation within seven working days of your request. Use our RRP service to expedite the development and testing of custom buffers and process liquids for your biopharmaceutical manufacturing process.

Table 2. Supplement matrix

	Amino acids	Vitamins	Glucose	Trace elements	Growth factors	Hypoxanthine/ thymidine	ADCF* lipids	ADCF* cholesterol	Suitable for	Code number
Cell Boost 1 Supplement (R05.2)	•	•	٠						HEK293 CHO	SH30584
Cell Boost 2 Supplement (R15.4)	•		•						PER.C6™ CHO	SH30596
Cell Boost 3 Supplement (JM3.5)	•	•	•	•		٠			Hybridoma Myeloma	SH30825
Cell Boost 4 Supplement (PS307)	•	•	•	•	•		•	•	СНО	SH30857
Cell Boost 5 Supplement (CN-F)	•	٠	•	•	•	•	•	•	Hybridoma NS0 HEK293 CHO	SH30865
Cell Boost 6 Supplement (CN-T)	•	•	•	•	•	•	•	•	T-Cells Hybridoma NS0 HEK293 CHO	SH30866
LS250 supplement							•	•	NS0	SH30554
LS1000 supplement								•	NS0	SH30555

* Animal-derived component-free

Related products

Table 2 gives an overview of HyClone supplements.

HyClone Cell Boost[™] kit

Cell Boost Process Supplements (100 g each) contain samples of supplements designed to increase cell productivity in a variety of cell lines (Table 2). Each supplement is developed through the Metabolic Pathway Design process and is chemically defined and protein-free with no animal derived components.

HyClone LS250 supplement

LS250 is a chemically defined, animal-derived componentfree lipid supplement developed to stimulate cell growth and monoclonal antibody (MAb) production in NS0 cell cultures using traditional hybridoma serum-free media.

HyClone LS1000 supplement

LS1000 supplement is a chemically defined, animal-derived component-free lipid supplement developed to stimulate cell growth and MAb production in NS0 cell cultures using traditional hybridoma serum-free media.

The supplement is formulated using a proprietary complexing process for enhanced cholesterol delivery. LS1000 has been successfully tested in a variety of serum-free medium cultures, including HyClone CDM4NS0 and CDM4MAb media.

Ordering information

CDM4Retino medium is manufactured in homogenous liquid lot sizes up to 10 000 L and powder lots up to 250 000 L.

Size	Code number
1000 mL bottle	SH30520.02
5 L bag	SH30520.03
10 L bag	SH30520.04
20 L bag	SH30520.05
50 L bag	SH30520.06
100 L bag	SH30520.07
200 L bag	SH30520.08
500 L bag	SH30520.09
1 × 5 L HDPE* bottle	SH30519.01
1 × 10 L HDPE* bottle	SH30519.02
1 × 50 L HDPE* bottle	SH30519.03
1 × 100 L HDPE* bottle	SH30519.04
1 × 500 L polybag/pail	SH30519.05
1 × 1000 L polybag/pail	SH30519.06
	1000 mL bottle 5 L bag 10 L bag 20 L bag 50 L bag 100 L bag 200 L bag 200 L bag 500 L bag 100 L bag 200 L bag 100 L bag 200 L bag 1 × 5 L HDPE* bottle 1 × 10 L HDPE* bottle 1 × 50 L HDPE* bottle 1 × 100 L HDPE* bottle 1 × 100 L HDPE* bottle 1 × 500 L polybag/pail

Related products	Size	Code number
HyClone Cell Boost kit	6 × 100 g	SH30890
HyClone LS1000 cholesterol supplement	50 mL bottle 100 mL bottle	SH30554.01 SH30554.02
	500 mL bottle 1000 mL bottle	SH30554.03 SH30554.04
HyClone LS250 lipid supplement	100 mL bottle 500 mL bottle 1000 mL bottle	SH30555.01 SH30555.02 SH30555.03

* High-density polyethylene

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