

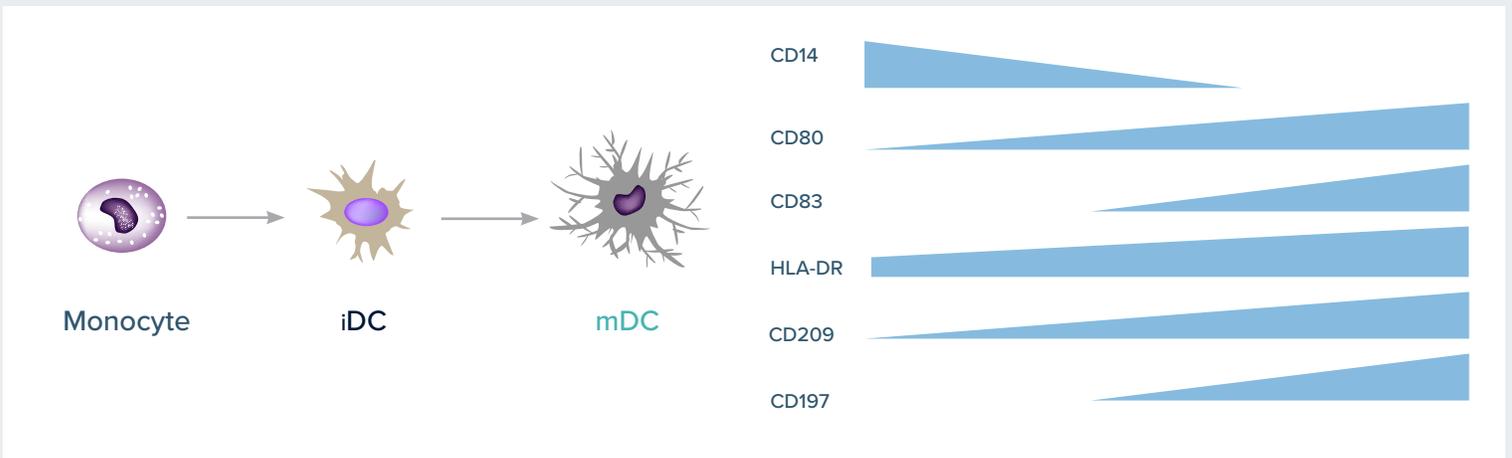
# PRIME-XV Dendritic Cell Maturation CDM

Chemically-defined, animal component-free medium for dendritic cell culture

- Optimized for differentiation of monocytes into immature dendritic cells (iDCs) and subsequent maturation of mDCs
- Achieve high yields of DCs with the desired characteristics and morphology
- Maintains potency of dendritic cells for T cell stimulation
- Ready-to-use, just supplement with desired cytokine cocktails
- Designed and manufactured to facilitate transfer from research to clinic

High yields of viable cells in chemically-defined conditions

Dendritic cells (DCs) are the most potent antigen-presenting cells capable of priming naïve T cells and are therefore essential tools in the activation of immune responses to tumors. They can be generated *in vitro* from CD14<sup>+</sup> peripheral blood or cord blood derived monocytes through a maturation process. It is essential monocyte-derived DCs (Mo-DCs) maintain their functional capacity – the launching of Ag-specific immunity – leading to T cell proliferation and polarization into effector T cells. Serum and undefined components found in animal- or human-derived raw materials can impact this functional capacity. PRIME-XV Dendritic Cell Maturation CDM provides a controlled, robust process to generate Mo-DCs in chemically-defined culture conditions thereby maintaining their capacity to induce T cell response.



# Get to Market Faster with Media Customization and Manufacturing Services

## Custom formulas and packaging

Customize our catalog products to suit your requirements for formulation or packaging.

## Media Development and Optimization

Save time and costs – let us design the medium that precisely meets your requirements with our Media Development and Optimization (MDO) program.

## Express Media Service

Typically provided within 10 working days, our rapid prototyping service offers flexible, small-scale, non-GMP media production of liquid and powder formulations. By using the same raw materials sourced and quality-controlled as for our large-scale GMP manufacturing, this approach greatly facilitates the step from research to commercialization.

## Custom cGMP manufacturing

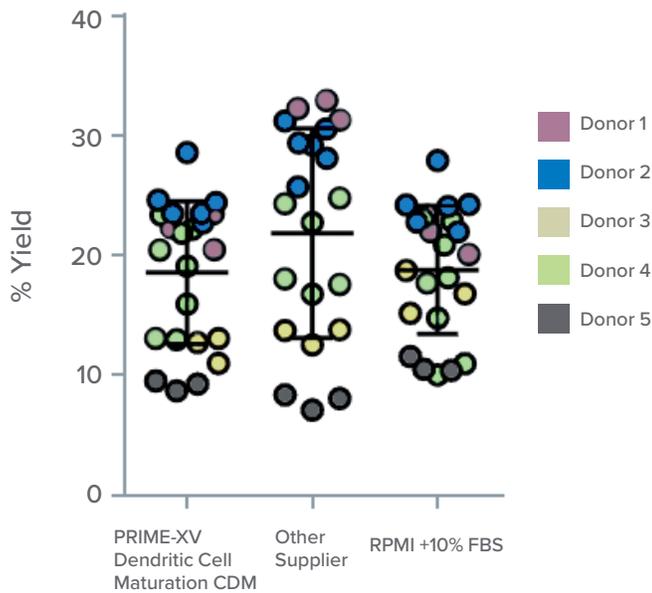
FUJIFILM Irvine Scientific offers industry-leading turnaround times for large-scale manufacturing of powder and liquid media in our state-of-the-art facilities. Comprehensive documentation, including information from Supply Chain Management through to Drug Master Files, is available to help minimize your regulatory burden.

To discuss your requirements, contact us at [getinfo@irvinesci.com](mailto:getinfo@irvinesci.com) or visit our website at [www.irvinesci.com/contact-us](http://www.irvinesci.com/contact-us)

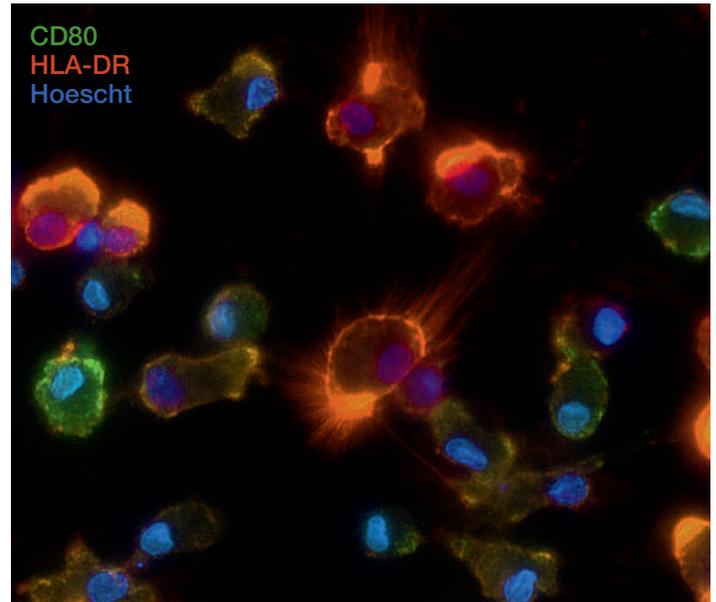
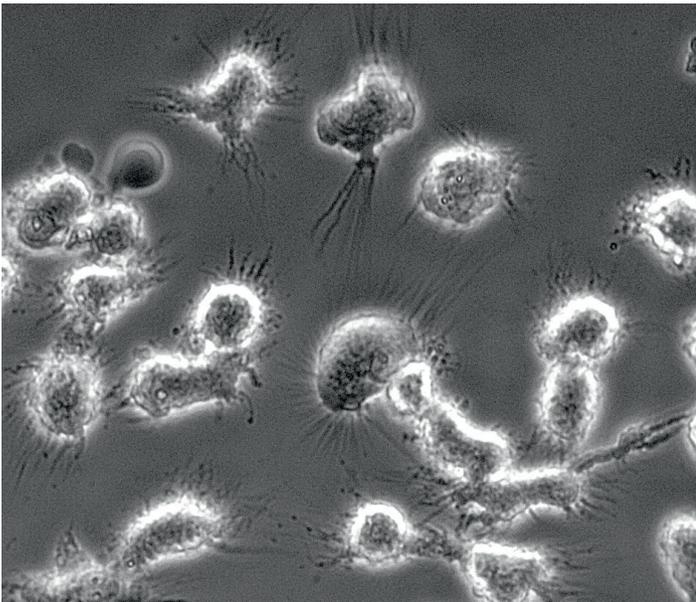
- FDA-regulated
- cGMP compliant manufacture
- 13485: 2016 certified
- Drug Master Files registered with FDA



## Active high yields of cells with desired characteristics



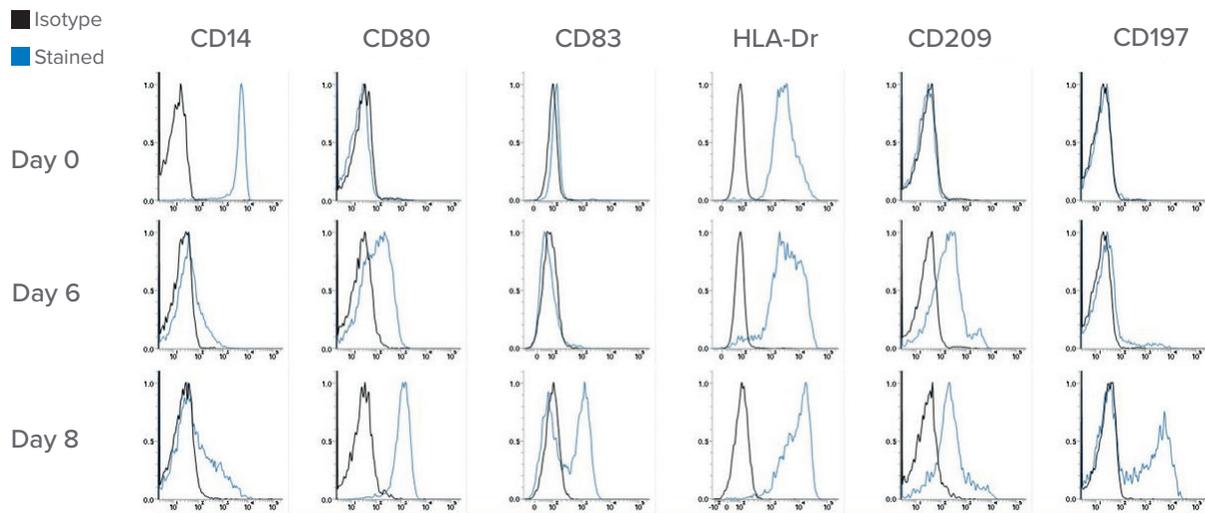
**Figure 1. Achieve high viable cell yields in chemically-defined culture conditions.** Enriched CD14<sup>+</sup> monocytes from 5 different donors were cultured in either PRIME-XV Dendritic Cell Maturation CDM, other commercially available medium, or RPMI +10% FBS for 6 days in the presence of GM-CSF and IL-4 to induce differentiation, then subsequently cultured for 2 more days in TNF- $\alpha$ , IL-1 $\beta$ , IL-6, and PGE<sub>2</sub> to induce maturation for a combined culture duration of 8 days in each respective medium. Resulting cells were analyzed by flow cytometry to measure viable cell density. Yield is shown as total viable cells as a percentage of initial CD14<sup>+</sup> monocytes plated on day 0.



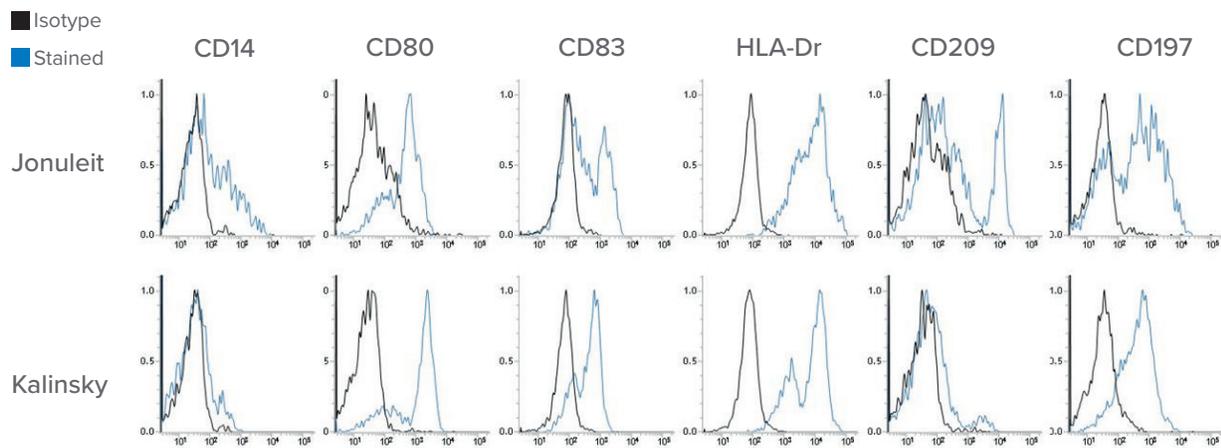
**Figure 2. Dendritic cells matured in PRIME-XV Dendritic Cell Maturation CDM demonstrate desired phenotype and morphology.** Negatively enriched CD14<sup>+</sup> monocytes were cultured in PRIME-XV Dendritic Cell Maturation CDM for 6 days in the presence of GM-CSF and IL-4 to induce differentiation, then subsequently cultured for 2 more days in TNF- $\alpha$ , IL-1 $\beta$ , IL-6, and PGE<sub>2</sub> to induce maturation for a combined culture duration of 8 days. The cells were labeled with mouse anti-human CD80 (Biolegend, Catalog # 305211), HLA-DR (Biolegend, Catalog# 307611), and Hoechst 33342 (ThermoFisher, Catalog# 62249). Immunofluorescence images were taken at 30x magnification.

# Proper marker expression maintained throughout maturation

A

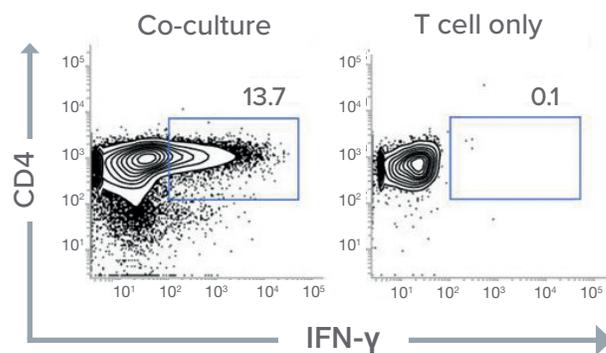
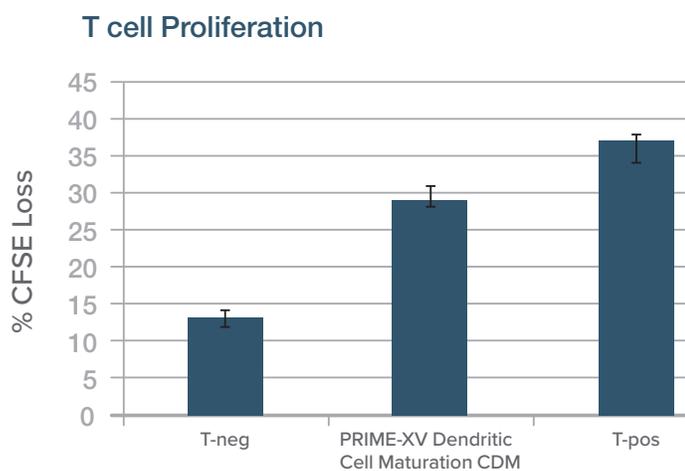
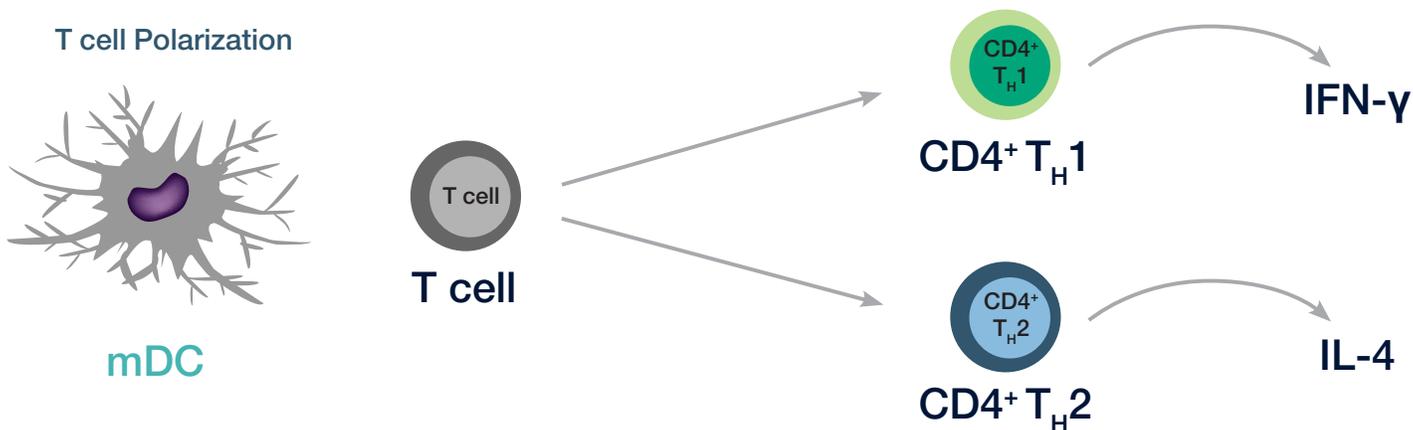


B



**Figure 3. Functionality demonstrated by surface marker expression profile.** Negatively selected CD14<sup>+</sup> monocytes were cultured in PRIME-XV Dendritic Cell Maturation CDM for 6 days in the presence of GM-CSF and IL-4 to induce differentiation, and subsequently cultured for 2 more days in either the Jonuleit cocktail (TNF- $\alpha$ , IL-1 $\beta$ , IL-6, and PGE<sub>2</sub>) or the Kalinsky cocktail (TNF- $\alpha$ , IL-1 $\beta$ , IFN- $\gamma$ , IFN- $\alpha$ , and Poly(I:C)) to induce maturation for a combined culture duration of 8 days. The cells were harvested and analyzed by flow cytometry for surface marker expression. Cells cultured with the Jonuleit cocktail were analyzed at day 0 (monocyte), day 6 (immature DC), and day 8 (mature DC) and demonstrated typical surface marker expression profiles throughout DC maturation (A). Jonuleit cocktail cultures and Kalinsky cocktail cultures were analyzed at day 8 and demonstrated typical surface marker expression profiles characteristic of maturation in each respective cocktail methodology (B).

## Maintains potency for T Cell stimulation



**Figure 4. PRIME-XV Dendritic Cell Maturation CDM generates DCs capable of inducing T cell proliferation.** Day 8 Mo-DCs cultured in PRIME-XV Dendritic Cell Maturation CDM were harvested and co-cultured with allogeneic CD4<sup>+</sup> T cells at a ratio of 1:20 (DC to T) in T cell expansion medium supplemented with IL-2 for 6 days. T cell stimulatory capacity was confirmed by loss of CFSE (n=3). CD4<sup>+</sup> T cells cultured without Mo-DC were cultured in IL-2 alone or supplemented with anti-CD3/anti-CD28 to serve as negative and positive controls, respectively.

**Figure 5. PRIME-XV Dendritic Cell Maturation CDM generates DCs capable of inducing T cell polarization.** Day 8 Mo-DCs cultured in PRIME-XV Dendritic Cell Maturation CDM were harvested and co-cultured with allogeneic CD4<sup>+</sup> T cells at a ratio of 1:10 (DC to T) in T cell expansion medium supplemented with IL-2 for 6 days. Flow cytometry performed on day 6 shows increased IFN-γ expression in co-cultured cells.

## PRIME-XV Dendritic Cell Maturation - Manufactured to facilitate transfer from research to clinic

- Chemically-defined, animal component-free formula minimizes risks from adventitious agents
- Traceability documentation provided including Certificates of Analysis, Certificates of Origin, and a Drug Master File (DMF) filed with the US FDA
- Robust raw material controls and supply chain management
- Extensive QA testing including functionality, sterility, and endotoxin
- Custom sizes and packaging available on request

## Ordering Information

Media	Catalog #	Size*	Additional Information
PRIME-XV Dendritic Cell Maturation CDM	91146	500 mL	Chemically-defined, animal component-free formula. Does not contain antibiotics or phenol red.

## Related Products

Item	Catalog #	Size*	Additional Information
PRIME-XV T Cell CDM	91154	1 L	Chemically-defined, animal component-free formula. Does not contain antibiotics or phenol red.
Recombinant Human GM-CSF ACF	95112	20 µg	Animal component-free. Accession Number: P04141
Recombinant Human TNF-α ACF	95117	10 µg	Animal component-free. Accession Number: P01375
Recombinant Human IL-6 ACF	95121	20 µg	Animal component-free. Accession Number: P05231
Recombinant Human IL-4 ACF	95114	20 µg	Animal component-free. Accession Number: P05112
Recombinant Human IL-2 ACF	95118	10 µg	Animal component-free. Accession Number: P60568
PRIME-XV FreezIS	91139	100 mL 10 mL	Chemically-defined, free from animal components and proteins. Contains 10% DMSO.

\*Custom sizes and packaging available on request.



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