

Cytokine	Cell source	Target(s)	What It does at the Target
IL-1	Macrophages	Macrophages, Neutrophils, Endothelial Cells.	<p>**Know that along with TNF, it has a huge role in inflammation.</p> <p><b>Phagocytes:</b> Activates 'em, or enhances their activity.</p> <p><b>Endothelial cells:</b> they get damaged when macrophages come "eat." IL-1 causes expression of ICAM-1, VCAM-1, E-Selectin. In addition, it causes them to secrete IL-8, and MCP-1.</p> <p><b>Physiologically,</b> it induces low BP (possible shock), and on the hypothalamus to cause fevers (higher set-point)</p>
IL-2	CD4+ T-cells, specifically Thp, Th0, and Th1 (notice no Th2)	CD8+ T-cells, NK Cells	<p><b>Naïve CD8+ T-cells (pCTLs):</b> along with IFN<math>\gamma</math>, IL-2 allows them to mature into Cytotoxic T-cells (CTLs).</p> <p><b>Natural Killer cells:</b> enhances their activity in killing virally infected cells (their KIR recognizes Class I MHCs of infected cells, while their Fc<math>\gamma</math>R recognizes viral antigens bound to IgG)</p>
IL-3	T-cells (and mast cells)	Hematopoietic cells, and Myeloid Progenitors	<p>Thus... <b>Hematopoiesis.</b> It helps when the Pluripotent Stem Cells (the cells you get during a bone marrow transplant) need to split into the 2 lineages of Myeloid progenitor cells, and Lymphoid Progenitor Cells</p>
IL-4	Mast cells, Th0 cells, and Th2 cells (CD4+)	B-cells, Th0 CD4+ T-cells, Th1 CD4+ T-cells, Phagocytes	<p><b>B-cells:</b> causes them to undergo isotype switching to IgE.</p> <p><b>Th0 cells:</b> lotsa IL-4 prefers the Th2 CD4+ T-cell population.</p> <p><b>Th1 cells:</b> inhibits/downregulates them (while increasing Th2 type).</p> <p><b>Phagocytes:</b> downregulates the production of iNOS.</p>
IL-5	Th2 cells, and Mast Cells	automatically think <b>Eosinophils!</b> and Bone Marrow too...	<p><b>Eosinophils:</b> Activates them for growth and differentiation, causing them to express the adhesion molecule VLA-4 (which is needed to bind to VCAM-1 of the endothelium so it can enter the endothelial cells), also to express the Chemokine Receptor CCR3</p> <p><b>Bone Marrow:</b> takes that 1st lineage of the Pluripotent Stem Cells, the Myeloid Progenitor and allows it to mature into either Eosinophils, Basophils, or GM Progenitor.</p>
IL-6	Th2 cells, and Macrophages	Hepatocytes and Bone marrow	<p><b>Hepatocytes:</b> causes them to secrete C-Reactive Protein (CRP), which is an Opsonin for phagocytes.</p> <p><b>Bone Marrow:</b> promotes Hematopoiesis b/c it stimulates the Pluripotent Stem Cells to be more responsive to IL-3, IL-5, and GM-CSF which are all cytokines for Hematopoiesis as well</p>

IL-7	Bone Marrow Stromal Cells	Bone Marrow and Thymus	<p><b>Bone Marrow:</b> takes the 2nd lineage of the Pluripotent Stem Cells, the Lymphoid Progenitor and allows it to mature to either B-cell Progenitor, T-cell Progenitor, or NK Cells. Also, within the Bone Marrow, it promotes B-cell Maturation.</p> <p><b>Thymus:</b> In addition, it allows the T-cell Progenitors to mature into their respective T-cells (that is, either they gonna be CD4+ or CD8+ after the maturation process).</p>
IL-8	Endothelial Cells, activated macrophages	causes inflammation of tissues	It is a chemokine, and <b>NOT a cytokine</b> . Chemokines are chemoattractants, and thus IL-8 is <b>chemotactic</b> for Neutrophils.
IL-9	Th2 cells	Activated T-cells and Activated B-cells	????
IL-10	Th2 cells	Dendritic Cells and Macrophages	<p><b>Dendritic Cells:</b> causes them to secrete LESS IL-12. And thus less IL-12 leads to less Th1 proliferation (since both IFN<math>\gamma</math> and IL-12 are both needed for Th1 proliferation)</p> <p><b>Macrophages:</b> along with IL-4 and TNF<math>\beta</math>, downregulate iNOS</p>
IL-11	Bone Marrow again	Hepatocytes and Bone marrow	<b>same as IL-6</b>
IL-12	Dendritic Cells and Macrophages	NK Cells and Th0 cells	<p><b>NK Cells:</b> It activates NK cells so that they can secrete IFN<math>\gamma</math>.</p> <p><b>Th0 cells:</b> along with IFN<math>\gamma</math> allows Th1 cells to proliferate</p>
IL-15	Epithelial Cells, Fibroblasts	CD8+ T-cells, NK Cells	????
GM-CSF	T-cells and Macrophages	Bone Marrow	<p><b>Bone Marrow-</b> allows the 1st lineage of Pluripotent Stem Cells, the Myeloid Progenitor to become the GM Progenitor. Easy enough... GM-CSF = GM Progenitor</p>
all other CSF	GM-CSF is the only one secreted by both Macrophages AND T-cells. Otherwise, all other CSF's (including GM-CSF) is secreted by Macrophages	Bone Marrow	<p><b>Bone Marrow:</b> all the CSF's are involved in Hematopoiesis:</p> <p><b>GM-CSF</b> --&gt; said earlier it promotes Myeloid Progenitor (1st lineage) to mature into the GM Progenitor. Again think the letter "GM" promotes production of the GM Progenitor.</p> <p><b>G-CSF</b> --&gt; Think the letter "G" thus it promotes the GM Progenitor to mature into Granulocytes (neutrophils).</p> <p><b>M-CSF</b> --&gt; Think the letter "M" thus it promotes the GM Progenitor to mature into Monocytes.</p>

IFN $\gamma$	NK Cells	Phagocytes and T-cells
TNF	Th1 Cells and Macrophages	Endothelial cells
TGF $\beta$ MCP-1	Th2 Cells Endothelial Cells, activated macrophages	Macrophages causes inflammation of tissues
IFN $\alpha/\beta$	all cells infected by a virus	all cells

**Phagocytes:** Activates iNOS (along with TNF. Both TNF and IFN $\gamma$  are needed in order for iNOS to be activated), and also enhances activation of NADPH Oxidase

**T-cells:** Along with IL-12, allows Th0 cells to become Th1, thus it also downregulates Th2 cells and Type 2 cytokines. AND when it is combined with IL-2, it promotes pCTLs to mature into CTLs

Know that it has the **EXACT SAME effects as IL-1** (they both work together anyway)

Thus, Along with IL-1, it tells the endothelial cells to express VCAM-1 and ICAM-1. And again, the VCAM-1 is what the VLA-4 expressed by the Eosinophils will bind to in order to enter the endothelium.

**\*\*Main difference between TNF and IL-1** to know is that TNF is an activator of iNOS (along with IFN $\gamma$ . In fact you need both TNF and IFN $\gamma$  in order to iNOS to be activated)

**Macrophages:** just like IL-4 and IL-10. Job is to downregulate iNOS.

**same as IL-8**

**\*aka Type 1 Interferons (but this is NOT a Type 1 Cytokine as the name would imply).** A shitload of activity:

- 1) Inhibits T-cell proliferation (whereas IFN $\gamma$  promoted T-cell proliferation and maturation [like with IL-2, it promoted pCTL--> CTL, and like with IL-12 it promoted Th0 --> Th1])
- 2) Increase in number of soluble adhesion molecules
- 3) Triggers enzymes such as **2'-5' Oligoadenylate Synthetase**, which interferes with Viral reproduction
- 4) Decreases B-cells in one population
- 5) is therapeutic for Hairy Cell Leukemia, Genital Warts, Kaposi's Sarcoma, Multiple Sclerosis, and Hepatitis B and Hepatitis C

Pyrogenic Cytokines: IL-1, 6, TNF

IL – 13 – inhibits IL- 1, 6, 8, 10, MCP

Macrophages release : IL-1, 6, 12, TNF

Th1 – IL-2, TNF, IFN

Th2 – IL-4, 5, 6, 10, 13, TGF

Tho – IL-2, 4, IFN