

## Technology Offer

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TO 02-0089

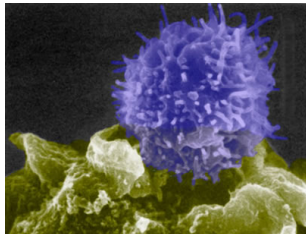
## MALP-2-based Modulation of Dendritic Cells for Stimulation of Th1 Immune Response

### The Challenge

Dendritic cells are the most important antigen-presenting cells of the immune system. In principle, two types of immune response can be distinguished; a) the cytotoxic Th1-mediated reaction and b) the Th2-immune response induced by antibodies, whereas the later one rather tends to provoke allergic reactions. For therapeutic purposes, the controlled generation of a Th1-type immune response is desired, e.g. for the defence of tumors and viruses or for the treatment of severe allergic reactions. However, a central problem is to influence dendritic cells to selectively induce a Th1-type immune response.

### The Technology

The MALP-2 lipopeptide (macrophage activating lipopeptide-2) isolated from *Mycoplasma fermentans* modulates the activity of antigen processing cells.



T Cell (blue) binding a Dendritic Cell (green).  
Source: Lawrence Berkeley National Laboratory

Among other effects, it stimulates murine as well as human macrophages and monocytes to liberate cytokines and prostaglandines and to induce a high titer of chemokines in vivo. MALP-2 is a ligand for the 'toll-like receptor 2'.

In combination with interferon  $\gamma$ , MALP-2 biases dendritic cells to preferentially induce a Th1-type immune response.

### Commercial Opportunity

The generation of a Th1-type immune reaction is of prime importance in a variety of therapeutic fields. Immunotherapy via pre-treated dendritic cells is more and more considered to be a promising strategy especially for the treatment of various types of cancer, infections, allergies and autoimmune diseases. Already to date, clinical trials (Phase I/II) for the treatment of cancer are on-going with pre-treated dendritic cells (loaded with either antigen or nucleic acids) being used to stimulate the immune system.

The advantages are:

- Handy and easy possibility to influence dendritic cells in a way that they induce a Th1-type immune response
- Availability of a high potential substance, the macrophage activating lipopeptide MALP-2, for development of new innovative therapeutic products in the field of immunotherapy
- Easy and reproducible isolation of natural macrophage activating lipopeptide; synthetic macrophage activating lipopeptide available by simple chemical synthesis

### Patent situation

Pending European patent application filed at EPO 2003, international application (PCT) will follow.

### Further Reading

Weigt H. et al. Immunobiology. 2003;207(3):223-33.

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