

Immudyne Nutritional NQ<sup>®</sup>

ImmuDyne Nutritional

\*NEW DATA\* The UNIQUE immune ingredient

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#### Immudyne Nutritional NQ The UNIQUE immune support ingredient

#### **Produces UNIQUE results**

#### STRUCTURE OF IMMUDYNE NQ YEAST B-1,3 1,6-GLUCAN

DEGREE OF BRANCHING shown with m = 15 as average number of ß-1,3 1,6-glucan units separating each ß-1,6-glucose unit



#### Immudyne Nutritional NQ The UNIQUE immune support ingredient

**Connects with innate immunity multiple ways: 1) macrophage cells** 



The UNIQUE immune support ingredient: increases phagocytosis



#### Immudyne Nutritional NQ The UNIQUE immune support ingredient

Connects with innate immunity multiple ways: 2) NK cells



The UNIQUE immune support ingredient: NK cell activity



#### The UNIQUE immune support ingredient: NK Cell importance



NK cells generally require signals from accessory cells in order to respond to pathogens. Accessory cells become activated following ligation of pathogen-recognition receptors (PRRs) by pathogen-encoded ligands.



Nature Reviews | Immunology

Immudyne Nutritional NQ

Nature Reviews Immunology 7, 279-291 (April 2007)

#### The UNIQUE immune support ingredient: More NK Cell importance



The natural killer (NK)-cell response to tumour cells

Expert Reviews in Molecular Medicine © 2003 Cambridge University Press

The natural killer (NK)-cell response to tumor cells. NK cells kill tumor cells through mechanisms that involve: antibody (Ab)-dependent cellular cytotoxicity (ADCC), in which the Fc portion of an Ab bound to antigen (Ag) on the tumor cell surface binds to Fc receptor (FcR) on the NK cell; Fas (CD95)–Fas ligand (CD95L) interaction; and release of perforin and granzyme B molecules, which cause apoptosis/necrosis of the tumor cell.

#### Immudyne Nutritional NQ The UNIQUE immune support ingredient

**Connects with innate immunity multiple ways: 3) Complement** 



#### The UNIQUE immune support ingredient: activates Complement



#### \*NEW DATA\* Immudyne Nutritional NQ

Whole Blood Assay: TNF alpha levels at 5 hrs in the presence or abssence of Beta Glucan Preparations



### \*NEW DATA\* Immudyne Nutritional NQ

#### The UNIQUE immune support ingredient

Whole Blood Assay: IL-10 levels at 5 hrs in the presence or abssence of Beta Glucan preparations



#### \*NEW DATA\*

#### **Immudyne Nutritional NQ**

#### The UNIQUE immune support ingredient

Whole Blood Assay: IL-6 levels at 3 hrs in the presence or abssence of Beta Glucan preparations



#### \*NEW DATA\* Immudyne Nutritional NQ

#### The UNIQUE immune support ingredient

Whole Blood Assay: IL-6 levels at 5 hrs in the presence or abssence of Beta Glucan preparations



#### \*NEW DATA\* Immudyne Nutritional NQ

#### The UNIQUE immune support ingredient

Whole Blood Assay: IFN gamma levels at 5 hrs in the presence or abssence of Beta Glucan preparations



#### Immudyne Nutritional NQ The UNIQUE immune ingredient

#### Importance of the Complement activation and immune health

These tests were conducted at National Jewish Health in Denver, Colorado and led by Dr. Manuel Galvan, Dr. Jaclyn Dennis and their team, considered to be the leading Complement Laboratory in the world. Their Advanced Diagnostic Laboratories (ADx) are fully accredited by CAP, CLIA, ISO15189 and compliant with U.S. Food and Drug Administration Good Manufacturing Laboratory Practices (GLP), 21 CFR 58. National Jewish conducted these test to measure Complement Activation. According to National Jewish, "Complement is the immune system's first line of defense against infection. In addition to killing invading microbes, Complement also is a strong modulator of inflammation." The Activation of the Complement system is critically important for overall immune health. It is a benchmark for how well a particular compound demonstrates its capacity to modulate the immune system.

This very recent (June, 2019) data from National Jewish above confirms that Immudyne Nutritional NQ is best in class.

#### Immudyne Nutritional NQ The UNIQUE immune support ingredient

What the Doctors Say: Joseph DiTrolio M.D., Senior Professor of Surgery and prominent Urologist at The Stone Center of New Jersey, has been using Immudyne Nutritional for 7 years adds: "I take Immudyne Nutritional every day and recommend it my patients. Immudyne helps modulate the immune system to prioritize what's important in life. It brings the mindset to the immune system to normalize immune response for overall immune health."

Alan B. Weitberg, M.D., and Professor of Medicine at Boston University, Boston, MA., who conducted an Immudyne Nutritional independent Clinical Study:

"My patients and I have been taking Immudyne Nuritional for over 10 years. It is a safe, well tolerated daily ingredient."

Sven Rohmann, M.D. PhD, who has served as Global Head Oncology at Merck Serono and is a world-wide expert on the immune system:

"I have personally observed all Immudyne NQ batches for over 10 years, and theirs is a track record that always, consistently produces the best. Only Immudyne has the pure, reliable beta glucan, with the proper clean molecular structure necessary to achieve the desired reliable result." Dr. Rohmann is now the CMO of Immudyne Nutritional.

#### The UNIQUE immune support ingredient

#### **Clinical evidence**

Treatment	Cancer Type	# Patients	# Relapse
None	Breast Stage II	9	2 (8.7%)
	Ovarian Stage II	4	0
	Cervical Stage II	10	3 (13%)
Beta Glucan	Breast Stage II	10	0
	Ovarian Stage II	4	0
	Cervical Stage II	12	0

"We have treated patients with breast, ovarian and cervical with 7.5 mg of Immudyne Nutritional/day and found that in treated patients there was no recurrence of cancer, whereas in the non-treated group there was a recurrence rate of 22%. We also treated terminal cancer patients with 15 mg of Immudyne Nutritional/day and demonstrated a decrease in mortality with time. " **Hiroiku Ueno, MD President of Asahi Alternative Clinic, Japan** 

The UNIQUE immune support ingredient



#### Immudyne Nutritional NQ The UNIQUE immune ingredient

#### **Clinical Phase I Study at Boston University**

This trial (n=20) demonstrates that  $\beta$ -(1,3)/(1,6) D-glucan is extremely well-tolerated in patients with advanced malignancies receiving chemotherapy. No adverse effects or toxicities were reported by any of the patients.

A significant number of patients reported a sense of well-being while taking the glucan.

There clearly was some amelioration of the blood counts in patients taking the glucan as compared to the pre-treatment mean counts indicating an improvement in haematopoiesis.

A phase I/II trial of beta-(1,3)/(1,6) D-glucan in the treatment of patients with advanced malignancies receiving chemotherapy . J Exp Clin Cancer Res. 2008; 27(1): 40.

Alan B Weitberg Address: Department of Medicine, Roger Williams Medical Center, Providence, Rhode Island and Boston University School of Medicine, Boston, Massachusetts, USA

#### The UNIQUE immune support ingredient

15 mg/d applied to two

terminally-ill cancer patients



#### The UNIQUE immune support ingredient

15 mg/d applied to 54 terminally-ill cancer patients



#### The UNIQUE immune support ingredient

**Effective Dosing** 



## The UNIQUE immune support ingredient: in sum

- Immudyne Nutritional NQ: UNIQUE, scientifically proven ingredient that potentiates, modulates, and helps to normalize the immune response.
- Enabling the immune system to fully support overall immune health.

Additional Immudyne references and studies are available upon request.

Immudyne Nutritional owns its manufacturing facility in the USA. Tasteless and odorless.

Kosher, Halal, organic compliant, contains NO yeast by-products or stimulants. GRAS status in the USA, and Food Approval in Europe.

## Proof of concept – IBD

YBG in the treatment of ulcerative colitis



YBG protects against DSS-induced colitis, and has effect on epithelial proliferation and intestinal restitution

Oral YBG treatment stimulates expansion of Peyer's 2 patches and mesenteric lymph nodes (not shown)



Ctr

SBG

DSS



### **Immudyne Nutritional NQ: Selected References**

- 1. Kadish, J. L., C. C. Choi, and J. K. Czop. 1986. Phagocytosis of unopsonized zymosan particles by trypsin-sensitive and -glucan-inhibitable receptors on bone marrow-derived murine macrophages. *Immunol. Res. 5:129.*
- 2. Czop, J. K., D. T. Fearon, and K. F. Austen. 1978. Opsonin-independent phagocytosis of activators of the alternative complement pathway by human monocytes. *J. Immunol.* 120:1132.
- 3. Williams, D. L., A. Mueller, and W. Browder. 1996. Glucan-based macrophage stimulators. *Clin. Immunother. 5:392.*
- 4. Ross, G. D., V. Vetvicka, J. Yan, Y. Xia, and J. Vetvickova. 1999. Therapeutic intervention with complement and -glucan in cancer. *Immunopharmacology* 42:61.
- 5. Williams, D. L. 1997. Overview of (1,3)--D-glucan immunobiology. *Mediat. Inflamm. 6:247.*
- 6. Ross, G. D., J. A. Cain, and P. J. Lachmann. 1985. Membrane complement receptor type three (CR3) has lectin-like properties analogous to bovine conglutinin as functions as a receptor for zymosan and rabbit erythrocytes as well as a receptor for iC3b. *J. Immunol. 134:3307.*
- 7. Thornton, B. P., V. Vetvicka, M. Pitman, R. C. Goldman, and G. D. Ross. 1996. Analysis of the sugar specificity and molecular location of the glucan-binding lectin site of complement receptor type 3 (CD11b/CD18). *J. Immunol. 156:1235.*
- 8. Brown, G. D., and S. Gordon. 2001. Immune recognition: a new receptor for glucans. Nature 413:36.
- 9. Ariizumi, K., G. L. Shen, S. Shikano, S. Xu, R. Ritter III, T. Kumamoto, D. Edelbaum, A. Morita, P. R. Bergstresser, and A. Takashima. 2000. Identification of a novel, dendritic cell-associated molecule, dectin-1, by subtractive cDNA cloning. *J. Biol. Chem.* 275; 20157.
- 10. Hermanz-Falcon, P., I. Arce, P. Roda-Navarro, and E. Fernandez-Ruiz. 2001. Cloning of human DECTIN-1, a novel C-type lectin-like receptor gene expressed on dendritic cells. *Immunogenetics 53:288.*
- 11. Yokota, K., A. Takashima, P. R. Bergstresser, and K. Ariizumi. 2001. Identification of a human homologue of the dendritic cell-associated C-type lectin-1, dectin-1. *Gene 272:51.*
- 12. Willment, J. A., S. Gordon, and G. D. Brown. 2001. Characterisation of the human -glucan receptor and its alternatively spliced isoforms. *J. Biol. Chem.* 20:20.
- Sobanov, Y., A. Bernreiter, S. Derdak, D. Mechtcheriakova, B. Schweighofer, M. Duchler, F. Kalthoff, and E. Hofer. 2001. A novel cluster of lectin-like receptor genes expressed in monocytic, dendritic and endothelial cells maps close to the NK receptor genes in the human NK gene complex. *Eur. J. Immunol.* 31:3493.
- 14. Brown, G. D., P. R. Taylor, D. M. Reid, J. A. Willment, D. L. Williams, L. Martinez-Pomares, S. Y. C. Wong, and S. Gordon. 2002. Dectin-1 is a major glucan receptor on macrophages. J. Exp. Med. 196:407.

### Immudyne Nutritional NQ: Selected References

- 1. Ezekowitz, R. A., R. B. Sim, M. Hill, and S. Gordon. 1984. Local opsonization by secreted macrophage complement components: role of receptors for complement in uptake of zymosan. *J. Exp. Med.* 159:244.
- 2. Ezekowitz, R. A., R. B. Sim, G. G. MacPherson, and S. Gordon. 1985. Interaction of human monocytes, macrophages, and polymorphonuclear leukocytes with zymosan in vitro: role of type 3 complement receptors and macrophage-derived complement. *J. Clin. Invest.* 76:2368.
- 3. Arase, H., T. Saito, J. H. Phillips, and L. L. Lanier. 2001. Cutting edge: the mouse NK cell-associated antigen recognized by DX5 monoclonal antibody is CD49b (2 integrin, very late antigen-2). *J. Immunol.* 167:1141.
- 4. Ross, G. D., and V. Vetvicka. 1993. CR3 (CD11b, CD18): a phagocyte and NK cell membrane receptor with multiple ligand specificities and functions. *Clin.Exp. Immunol.* 92:181.
- 5. McMahon, C. W., and D. H. Raulet. 2001. Expression and function of NK cell receptors in CD8 T cells. *Curr. Opin. Immunol.* 13:465.
- 6. Fernandes, M. J., A. A. Finnegan, L. D. Siracusa, C. Brenner, N. N. Iscove, and B. Calabretta. 1999. Characterization of a novel receptor that maps near the natural killer gene complex: demonstration of carbohydrate binding and expression in hematopoietic cells. *Cancer Res.* 59:2709.

#### Reviews:

- Lanier, LL. 1998. NK cell receptors. Ann. Rev. Immunol. 16:359.
- II. Lanier, LL. 2005. NK cell recognition. Ann. Rev. Immunol. 23:225.
- III. Gordon, GD. 2006. Dectin-1: a signalling non-TLR pattern recognition receptor. *Nature Review Immunol. 6:33.*