Fix Your Body Glucosamine; Friend or Foe By Bryon Verhaeghe March, November 2004

-Without prejudice-

Glucosamine is an amino acid derivative of glucose. It is a constituent of polysaccharide (large sugar) in animal supporting structures and the cell walls of bacteria and fungus (yeast). In the late 1980's many articles appeared regarding glucosamine inhibitors as having anti-fungal, anti-Candida, anti-leukemia, anti-salmonella, anti-malarial and anti-bacterial properties.

In the early 1990's a company found that glucosamine sulfate relieved joint pain. A double blind study where the study group was divided into three groups were given either a sugar pill, glucosamine HCL, or glucosamine sulfate without anyone knowing which group got which "medication". The study group filled out extensive questionnaires about their symptoms and pain levels. The improvement in symptoms between the sugar and glucosamine HCL groups was identical. The improvement in the glucosamine sulfate group was distinctly better. The report concluded that only the sulfate was responsible for the benefit and not the glucosamine.

In another study it was found that glucosamine increased juvenile arthritis by 18%. Many of the studies state that the most common reason that people dropped out of a study was because of increased bloating and gas. Some reported an increase in headache and migraine. Many people that I have talked to tell me that their bloating and gas subsides when they stop taking glucosamine.

Another reported side effect of glucosamine is weight gain. Some find that they are 15-20 pounds heavier after about 2 years of glucosamine without any other dietary of life style changes. Weight gain with glucosamine makes perfect sense to me because a healthy person carries about 4.5 pounds of bacteria and if we supplement something that enhances their growth we would get heavier. Greater numbers of bacteria in our body would also consume more of our energy. This weight gain also causes an increase in fatigue and tiredness. An increase in depression, anxiety and panic would result as the fungal levels went up.

On the recent medical side of glucosamine causing fatigue there are studies showing that elevated glucosamine levels are associated with mitochondrial dysfunction. The mitochondrial are responsible for producing the energy for life (ATP). It is also strongly noted that mitochondrial dysfunction is directly related to rapid ageing.

The research seems to be coming back to life since the middle of 2003 in regard to glucosamine. These more recent studies reiterate the diabetic risk but now also include cancer and neuro-degeneration. A molecule closely related to glucosamine is the standard agent used to induce diabetes in lab animals. The internet is loaded with slick sites to lure people into glucosamine. Who would be so interested in you taking glucosamine?

United States Senate Special Committee on Aging

Hearing on Swindlers, Hucksters and Snake Oil Salesmen: The Hype and Hope of Marketing Anti-Aging Products to Seniors September 10, 2001

Testimony of Timothy N. Gorski, M.D., F.A.C.O.G. Assistant Clinical Professor, University of North Texas Health Science Center President, Dallas/Fort Worth Council Against Health Fraud Board Member, National Council Against Health Fraud Associate Editor, *Scientific Review of Alternative Medicine*

> Current Issues in Protecting the Public from Health Fraud: "Dietary Supplements" as a Public Health Problem

• Another potential public health threat is that of glucosamine, which is widely promoted for the treatment of arthritis on the basis of very scanty evidence. It is probably among the top ten best-selling "dietary supplements." Yet glucosamine is known to increase resistance to insulin at doses comparable to those recommended for these products. In layman's terms, glucosamine tends to cause diabetes, a disorder that many older Americans have or are susceptible to. Diabetes, in turn, is a risk factor for heart disease.

References:

Horal M, Zhang Z, Stanton R, Virkamaki A, Loeken MR.
Activation of the hexosamine pathway causes oxidative stress and abnormal embryo gene expression: involvement in diabetic teratogenesis.
Birth Defects Res Part
A Clin Mol Teratol. 2004 Aug;70(8):519-27.
PMID: 15329829 [PubMed - in process]

Zachara NE, Hart GW. O-GlcNAc a sensor of cellular state: the role of nucleocytoplasmic glycosylation in modulating cellular function in response to nutrition and stress. Biochim Biophys Acta. 2004 Jul 6;1673(1-2):13-28. PMID: 15238246 [PubMed - indexed for MEDLINE]

Akimoto Y, Kawakami H, Yamamoto K, Munetomo E, Hida T, Hirano H. Elevated expression of O-GlcNAc-modified proteins and O-GlcNAc transferase in corneas of diabetic Goto-Kakizaki rats. Invest Ophthalmol Vis Sci. 2003 Sep;44(9):3802-9. PMID: 12939295 [PubMed - indexed for MEDLINE]

Slawson C, Hart GW. Dynamic interplay between O-GlcNAc and O-phosphate: the sweet side of protein regulation. Curr Opin Struct Biol. 2003 Oct;13(5):631-6. Review. PMID: 14568619 [PubMed - indexed for MEDLINE]

Janiak A, Cybulska B, Szlinder-Richert J, Borowski E, Milewski S. Facilitated diffusion of glucosamine-6-phosphate synthase inhibitors enhances their antifungal activity. Acta Biochim Pol. 2002;49(1):77-86. PMID: 12136959 [PubMed - indexed for MEDLINE]

Zgodka D, Milewski S, Borowski E. A diffusible analogue of N(3)-(4-methoxyfumaroyl)-L-2,3-diaminopropanoic acid with antifungal activity. Microbiology. 2001 Jul;147(Pt 7):1955-9. PMID: 11429472 [PubMed - indexed for MEDLINE]

Milewski S, Mignini F, Prasad R, Borowski E. Unusual susceptibility of a multidrug-resistant yeast strain to peptidic antifungals. Antimicrob Agents Chemother. 2001 Jan;45(1):223-8. PMID: 11120970 [PubMed - indexed for MEDLINE]

Zgodka D, Jedrzejczak R, Milewski S, Borowski E. Amide and ester derivatives of N3-(4-methoxyfumaroyl)-(S)-2,3-diaminopropanoic acid: the selective inhibitor of glucosamine-6-phosphate synthase. Bioorg Med Chem. 2001 Apr;9(4):931-8. PMID: 11354676 [PubMed - indexed for MEDLINE] Andruszkiewicz R, Jedrzejczak R, Zieniawa T, Wojciechowski M, Borowski E. N3-oxoacyl derivatives of L-2,3-diaminopropanoic acid and their peptides; novel inhibitors of glucosamine-6-phosphate synthase. J Enzyme Inhib. 2000;15(5):429-41. PMID: 11030083 [PubMed - indexed for MEDLINE]

Milewski S, Mignini F, Micossi L, Borowski E. Antihistoplasmal in vitro and in vivo effect of Lys-Nva-FMDP. Med Mycol. 1998 Jun;36(3):177-80. PMID: 9776831 [PubMed - indexed for MEDLINE]

Shankar RR, Zhu JS, Baron AD.
Glucosamine infusion in rats mimics the beta-cell dysfunction of non-insulin-dependent diabetes mellitus.
Metabolism. 1998 May;47(5):573-7.
PMID: 9591749 [PubMed - indexed for MEDLINE]

Chmara H, Milewski S, Andruszkiewicz R, Mignini F, Borowski E. Antibacterial action of dipeptides containing an inhibitor of glucosamine-6-phosphate isomerase. Microbiology. 1998 May;144 (Pt 5):1349-58. PMID: 9660640 [PubMed - indexed for MEDLINE]

Wojciechowski M, Mazerski J, Borowski E. Constrained search of conformational hyperspace of inactivators of glucosamine-6phosphate synthase. J Enzyme Inhib. 1996;10(1):17-26. PMID: 8835927 [PubMed - indexed for MEDLINE]

Milewski S, Mignini F, Covelli I, Borowski E. Specific inhibition of acid proteinase secretion in Candida albicans by Lys-Nva-FMDP. J Med Vet Mycol. 1994;32(1):1-11. PMID: 8207618 [PubMed - indexed for MEDLINE]

Kasprzak L, Milewski S, Gumieniak J, Borowski E.
The influence of serum proteins on biological activity of anticandidal peptides containing N3-(4-methoxyfumaroyl)-L-2,3-diaminopropanoic acid.
J Chemother. 1992 Apr;4(2):88-94.
PMID: 1629751 [PubMed - indexed for MEDLINE]

Tarnowska M, Oldziej S, Liwo A, Grzonka Z, Borowski E. Investigation of the inhibition pathway of glucosamine synthase by N3-(4methoxyfumaroyl)-L-2,3-diaminopropanoic acid by semiempirical quantum mechanical and molecular mechanics methods. Eur Biophys J. 1992;21(4):273-80. PMID: 1425480 [PubMed - indexed for MEDLINE]

Milewski S, Chmara H, Andruszkiewicz R, Borowski E. N3-haloacetyl derivatives of L-2,3-diaminopropanoic acid: novel inactivators of glucosamine-6-phosphate synthase. Biochim Biophys Acta. 1992 Jan 23;1115(3):225-9. PMID: 1739736 [PubMed - indexed for MEDLINE]

Milewski S, Andruszkiewicz R, Kasprzak L, Mazerski J, Mignini F, Borowski E. Mechanism of action of anticandidal dipeptides containing inhibitors of glucosamine-6-phosphate synthase.

Antimicrob Agents Chemother. 1991 Jan;35(1):36-43. PMID: 1901701 [PubMed - indexed for MEDLINE]

Andruszkiewicz R, Chmara H, Milewski S, Zieniawa T, Borowski E. Antimicrobial properties of N3-(iodoacetyl)-L-2,3-diaminopropanoic acid-peptide conjugates. J Med Chem. 1990 Oct;33(10):2755-9. PMID: 2120441 [PubMed - indexed for MEDLINE]

Andruszkiewicz R, Milewski S, Zieniawa T, Borowski E.
Anticandidal properties of N3-(4-methoxyfumaroyl)-L-2,3-diaminopropanoic acid oligopeptides.
J Med Chem. 1990 Jan;33(1):132-5.
PMID: 2104933 [PubMed - indexed for MEDLINE]

Kucharczyk N, Denisot MA, Le Goffic F, Badet B. Glucosamine-6-phosphate synthase from Escherichia coli: determination of the mechanism of inactivation by N3-fumaroyl-L-2,3-diaminopropionic derivatives. Biochemistry. 1990 Apr 17;29(15):3668-76. PMID: 2111163 [PubMed - indexed for MEDLINE]

Milewski S, Chmara H, Andruszkiewicz R, Borowski E, Zaremba M, Borowski J. Antifungal peptides with novel specific inhibitors of glucosamine 6-phosphate synthase. Drugs Exp Clin Res. 1988;14(7):461-5. PMID: 3149235 [PubMed - indexed for MEDLINE]

Andruszkiewicz R, Chmara H, Milewski S, Borowski E. Synthesis and biological properties of N3-(4-methoxyfumaroyl)-L-2,3-diaminopropanoic acid dipeptides, a novel group of antimicrobial agents. J Med Chem. 1987 Oct;30(10):1715-9. PMID: 3309312 [PubMed - indexed for MEDLINE]

Milewski S, Chmara H, Andruszkiewicz R, Borowski E. Synthetic derivatives of N3-fumaroyl-L-2,3-diaminopropanoic acid inactivate glucosamine synthetase from Candida albicans. Biochim Biophys Acta. 1985 Apr 29;828(3):247-54. PMID: 3921053 [PubMed - indexed for MEDLINE]

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