

12 May 2009

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President Barack Obama
The White House
1600 Pennsylvania Avenue, N.W.
Washington, D.C. 20500

Dear Mr. President and Staff,

We present to you a modest proposal for your consideration. If adopted, we believe it will have a number of positive effects on our country and the world:

- improving the physical and psychological health and well being of the next generations of our children, and
- reducing the terrible costs of cancer in our society and health care system.

It has been well established that breastfeeding of infants is the natural and superior approach to creating healthy individuals. Breastfed babies have stronger immune systems, fewer allergies, less probability of developing childhood (and likely adult) cancers, half the chance of dying of SIDS [1], lower probabilities of developing diabetes and other chronic diseases, and, in breastfeeding, have the opportunity to bond with their mothers. All medical groups advocate breastfeeding above any other approach to infant nutrition. [2] Recent studies [3, 4] have also shown that women who have breast-fed are at lower risk than mothers who have not for developing high blood pressure, diabetes and cardiovascular disease decades later, when they are in menopause. Any program that could increase the number of infants breastfed and the length of time they are breastfed would be advantageous to the individuals involved as well as to society at large because of the downstream consequences.

Likewise, it has been known since 1995 that human mothers' milk (unlike that of other mammals) contains substances that can kill cancer cells. This has been demonstrated in vitro[5] and in pilot animal [6] and human [7] studies. Ongoing research has mostly been pursued in Sweden to date. [8-29] The main substance, a form of the protein alpha-lactalbumin, is modified in the gut to a new molecular shape, called HAMLET, which is selectively taken up by cancer cells and causes them to undergo programmed cell death (*apoptosis*), while leaving normal cells unaffected. In fact, ingestion of human mothers' milk has led to the prostate cancer of one of us becoming undetectable and remaining so for almost ten years since he was biopsy diagnosed. [30]

Our proposal has several aspects:

1. A program should be established to encourage women to breastfeed their infants for at least one year. In order to enable this, a federal subsidy should be put in place for these women, much as is done in Europe. As a *quid pro quo*, they would need to pump their extra milk and donate it to a set of milk banks. (These may be an expansion of the extant non-profit and under-funded

Human Milk Banking Association of North America facilities. [31]) Here, the milk would be screened and then distributed to those who need it, other infants, cancer patients, and others needing immune system boosting, as determined by their doctors. The milk banks within this system already collect milk from screened volunteer donors, screen the milk, and distribute it to infants in need of human milk (and some cancer patients). This life saving activity and organization would need to be taken much more seriously from the health care establishment for the success of this program.

2. Cancer patients, in consultation with their doctors, would use mothers' milk as a monotherapy or an adjuvant to their other therapies.
3. Concurrently, and as a preliminary to setting up this system, NIH would fund academic and intramural research programs to further elucidate the nature of the cancer fighting capabilities of human mothers' milk, its pharmacokinetics, and other aspects of the best practices in its medical use. Since mothers' milk is a natural substance, abundantly available, it is unpatentable and therefore of negative interest to the profit oriented pharmaceutical medical industrial complex, and so requires public funding and advocacy for the vast potential public benefit to be realized.

Another positive aspect of this program would be the bonding experienced by mothers and their children, which could increase family and social cohesiveness, thereby decreasing the amount of social disconnection, mental problems and crime later in the infants' lives. Women who today are forced to return to work mere weeks after giving birth cannot fully emotionally bond with their children, and *vice versa*.

We encourage the creation of a multi-disciplinary task force to further explore and flesh out this program and its social and medical underpinnings. We see it as one small but significant aspect of improving the social and physical health of America and reining in the escalating health care costs, direct and indirect, that are draining our nation and its people.

Thank you for your consideration.

Sincerely,

Howard J. Cohen, Ph.D.
Barbara L. Cohen, M.A., LMFT

cc:

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Notes

[1] *Does Breastfeeding Reduce the Risk of Sudden Infant Death Syndrome?*, M.M. Vennemann, MD, MPH, PD, T. Bajanowski, MD, PD, B. Brinkmann, MD, PD, G. Jorch, MD, PD, K. Yucesan, MD, C. Sauerland, MScd, E.A. Mitchell, FRACP, DSc and the GeSID Study Group, *Pediatrics*, Vol. 123 No. 3 March 2009, pp. e406-e410

<http://pediatrics.aappublications.org/cgi/content/full/123/3/e406>

[2] The American Academy of Pediatrics Policy Statement on Breastfeeding and the Use of Human Milk (from *Pediatrics*, Vol. 115, No. 2, 2 Feb 2005, pp 496-406).

<http://aappolicy.aappublications.org/cgi/content/full/pediatrics;115/2/496>

See also the references therein and many of the articles that cite this publication (list and links available on the web site).

[3] *Breast-Feeding Benefits Mothers, Study Finds*, By Roni Caryn Rabin, New York Times, April 21, 2009

<http://www.nytimes.com/2009/04/22/health/research/22breast.html>

[4] *Duration of Lactation and Risk Factors for Maternal Cardiovascular Disease*, Schwarz, Eleanor Bimla MD, MS; Ray, Roberta M. MS; Stuebe, Alison M. MD, MSc; Allison, Matthew A. MD, MPH; Ness, Roberta B. MD, MPH; Freiberg, Matthew S. MD, MSc; Cauley, Jane A. DrPH, *Obstetrics & Gynecology*: May 2009 - Volume 113 - Issue 5 - pp 974-982

http://journals.lww.com/greenjournal/Fulltext/2009/05000/Duration_of_Lactation_and_Risk_Factors_for.5.aspx

[5] *Apoptosis Induced by a Human Milk Protein*, Anders Hakansson, Boris Zhivotovsky, Sten Orrenius, Hemant Sabharwal, and Catharina Svanborg, *Proceedings of the National Academy of Sciences, USA*, 92: 8064-8068 (1995)

[6] *Human Alpha-lactalbumin Made Lethal to Tumor Cells (HAMLET) Kills Human Glioblastoma Cells in Brain Xenografts by an Apoptosis-Like Mechanism and Prolongs Survival*, Walter Fischer, Lotta Gustafsson, Ann-Kristin Mossberg, Janne Gronli, Sverre Mork, Rolf Bjerkvig, and Catharina Svanborg, *Cancer Research* 64, 2105-2112, March 15, 2004. This research was partially funded by the American Cancer Society.

<ftp://www.cohensw.com/pub/pca/Fischer-braintumor-HAMLET.pdf>

[7] *Bladder Cancers Respond to Intravesical instillation of HAMLET (Human α-Lactalbumin Made Lethal to Tumor Cells)*, Ann-Kristin Mossberg, Bjorn Wullt, Lotta Gustafsson, Wiking Mansson, Eva Ljunggren, and Catharina Svanborg, *Int. J. Cancer*: Vol. 121, pp 1352-1359 (2007) [15 September 2007]

<ftp://www.cohensw.com/pub/pca/Mossberg-bladder-cancer-HAMLET.pdf>

Nine bladder cancer patients received 5 daily instillations of HAMLET during the week before surgery. HAMLET stimulated a rapid increase in the shedding of tumor cells into the urine, daily. Most of the shed cells were dead and 6 of 9 patients showed an apoptotic response. At surgery 8 of 9 patients showed a reduction in tumor size or change of tumor character. Adjacent healthy tissue showed no negative changes.

- [8] Peter Radetsky, *Got Cancer Killers?*, Discover Magazine, June 1999, pp 68-75
http://www.findarticles.com/cf_0/m1511/6_20/55926784/p1/article.jhtml?term=22Catharina+Svanborg%22
- [9] *Milk Therapy: Breast-milk compounds could be a tonic for adult ills*, Julie J. Rehmeyer, Science News Online, Week of Dec. 9, 2006; Vol. 170, No. 24 , p. 376
<http://www.sciencenews.org/articles/20061209/bob8.asp>
- [10] *Multimeric Alpha-Lactalbumin from Human Milk Induces Apoptosis through a Direct Effect on Cell Nuclei*, Anders Hakansson, Jesper Andreasson, Boris Zhivotovsky, Diana Karpman, Sten Orrenius, and Catharina Svanborg, Experimental Cell Research, 246: 451-460 (1999)
ftp://www.cohensw.com/pub/pca/AH_ECR.pdf
- [11] *Molecular Characterization of Alpha-Lactalbumin Folding Variants That Induce Apoptosis in Tumor Cells*, Malin Svensson, Hemant Sabharwal, Anders Hakansson, Ann-Kristin Mossberg, Peter Lipniunas, Hakon Leffler, Catharina Svanborg, and Sara Linse, Journal of Biological Chemistry, vol 275, no 10, pp 6388-6396 (5 March 1999)
ftp://www.cohensw.com/pub/pca/Malin_JBC.pdf
- [12] *Protease Activation in Apoptosis Induced by MAL*, Camilla Kohler, Anders Hakansson, Catharina Svanborg, Sten Orrenius, and Boris Zhivotovsky, Experimental Cell Research, 249: 260-268 (1999)
<ftp://www.cohensw.com/pub/pca/MAL-caspases.pdf>
- [13] *Conversion of Alpha-Lactalbumin to a Protein Inducing Apoptosis*, M. Svensson, A. Hakansson, A.-K. Mossberg, S. Linse, C. Svanborg, Proceedings of the National Academy of Sciences, USA, 97: 4221-4226 (2000)
ftp://www.cohensw.com/pub/pca/PNAS_2000-97-4221.pdf
- [14] *A Folding Variant of alpha-lactalbumin With Bactericidal Activity Against **Streptococcus pneumoniae***, Anders Hakansson, Malin Svensson, Ann-Kristin Mossberg, Hemant Sabharwal, Sara Linse, Irene Lazou, Bo Lonnerdal, and Catharina Svanborg, Molecular Microbiology (2000), 35 (3), 589-600
<ftp://www.cohensw.com/pub/pca/MolMicro2000-35-589.pdf>
- [15] *A Folding Variant of Human Alpha-Lactalbumin Induces Mitochondrial Permeability Transition in Isolated Mitochondria*, Camilla Kohler, Vladimir Gogvadze, Anders Hakansson, Catharina Svanborg, Sten Orrenius, Boris Zhivotovsky, European Journal of Biochemistry, 268, 186-191 (Feb 2001)
<ftp://www.cohensw.com/pub/pca/EurJBiochem2001-268-186.pdf>
- [16] *Hamlet -- A Complex From Human Milk That Induces Apoptosis in Tumor Cells But Sparing Healthy Cells*, Malin Svensson, Caroline Durringer, Oskar Hallgren, Ann-Kristin Mossberg, Anders Hakansson, Sara Linse, Catharina Svanborg, Advances in Experimental and Medical Biology (US), 503, 125-132 (2002)
ftp://www.cohensw.com/pub/pca/ISRHML_review.doc
- [17] *Alpha-lactalbumin Unfolding is not Sufficient to Cause Apoptosis, but is Required for the*

Conversion to HAMLET (Human Alpha-lactalbumin Made Lethal to Tumor cells), Malin Svensson, Jonas Fast, Ann-Kristin Mossberg, Caroline Durringer, Lotta Gustafsson, Oskar Hallgren, Charles L. Brooks, Lawrence Berliner, Sara Linse, and Catharina Svanborg, *Protein Science* (2003), 12:2794-2804.

<http://www.proteinscience.org/cgi/doi/10.1110/ps.0231003>

[18] *Lipids as Cofactors in Protein Folding: Stereo-Specific Lipid-Protein Interactions are Required to Form HAMLET (Human Alpha-lactalbumin Made Lethal to Tumor cells)*, Malin Svensson, Ann-Kristin Mossberg, Jenny Pettersson, Sara Linse, and Catharina Svanborg, *Protein Science* (2003), 12:2805-2814.

<http://www.proteinscience.org/cgi/doi/10.1110/ps.0231103>

[19] *HAMLET Kills Tumor Cells by Apoptosis - Cellular, Molecular and Therapeutic Aspects*, Catharina Svanborg, Helena Agerstam, Caroline Durringer, Walter Fischer, Lotta Gustafsson, Oskar Hallgren, Irene Leijonhuvud, Sara Linse, Ann-Kristin Mossberg, Hanna Nilsson, Jenny Peterson, Malin Svensson, Annika Aronson, Rolf Bjerkvig, *Advances in Cancer Research* 88:1-29 (2003)

ftp://www.cohensw.com/pub/pca/Svanborg_apoptosis_AdvCancRes_2003.pdf

[20] *Treatment of Skin Papillomas with Topical Alpha-Lactalbumin-Oleic Acid*, Lotta Gustafsson, Irene Leijonhufvud, Annika Aronsson, Ann-Kristin Mossberg, and Catharina Svanborg, *New England Journal of Medicine*, 2004; 350:2663-72 (June 24, 2004)

<ftp://www.cohensw.com/pub/pca/Gustafsson-papilloma-HAMLET.pdf>

[21] *Apoptotic Cell Death in the Lactating Mammary Gland is Enhanced by a Folding Variant of Alpha-Lactalbumin*, A. Baltzer, C. Svanborg and R. Jaggi, *Cellular and Molecular Life Sciences*, 61 (2004) 1221-1228

<ftp://www.cohensw.com/pub/pca/Baltzer-lactation-HAMLET.pdf>

[22] *HAMLET triggers apoptosis but tumor cell death is independent of caspases, Bcl-2 and p53*, O. Hallgren, L. Gustafsson, H. Irjala, G. Selivanova, S. Orrenius and C. Svanborg, *Apoptosis*, vol 11, No 2, 2006. pp 221-233.

<ftp://www.cohensw.com/pub/pca/Hallgren-bcl2-p53-HAMLET.pdf>

Indicates that HAMLET may have several pathways to induce tumor cell death (apoptosis), including avoiding pathways that tumor cells often block.

[23] *Structure-function analysis of HAMLET*, Jenny Pettersson's Ph.D. thesis at Lund University, Institute of Laboratory Medicine, Department of Microbiology, Immunology, and Glycobiology (MIG), 3 December 2007

<http://luur.lub.lu.se/luur?func=downloadFile&fileOID=605650>

[24] *Mini review: HAMLET, protein folding, and tumor cell death*, K. Hun Mok, Jenny Pettersson, Sten Orrenius and Catharina Svanborg, *Biochemical and Biophysical Research Communications*, Volume 354, Issue 1, 2 March 2007, Pages 1-7.

ftp://www.cohensw.com/pub/pca/Mok_BiochemBiophysResComm_2007.pdf

[25] *Heat-treatment method for producing fatty acid-bound alpha-lactalbumin that induces tumor cell death*, Kamijima T, Ohmura A, Sato T, Akimoto K, Itabashi M, Mizuguchi M, Kamiya M, Kikukawa T,

Aizawa T, Takahashi M, Kawano K, and Demura M, Biochemical and Biophysical Research Communications, Volume 376, 5 September 2008 (online), Pages 211-214.
ftp://www.cohensw.com/pub/pca/Kamijima_BiochemBiophysResComm_2008_18774773.pdf

[26] *Can misfolded proteins be beneficial? The HAMLET case*, Pettersson-Kastberg J, Aits S, Gustafsson L, Mossberg A, Storm P, Trulsson M, Persson F, Hun Mok K, and Svanborg C, Annals of Medicine, (2008), Pages 1-15.
ftp://www.cohensw.com/pub/pca/Pettersson-Kastberg_AnnMed_2008.pdf

[27] *HAMLET (human alpha-lactalbumin made lethal to tumor cells) triggers autophagic tumor cell death*, Aits S, Gustafsson L, Hallgren O, Brest P, Gustafsson M, Trulsson M, Mossberg AK, Simon HU, Mograbi B, and Svanborg C, Int. J. Cancer, 124, Pages 1008-1019 (2009).
ftp://www.cohensw.com/pub/pca/Aits_IntJCancer_2009_19048621.pdf

[28] *Who Is Mr. HAMLET? Interaction of Human alpha-Lactalbumin with Monomeric Oleic Acid*, Ekaterina L. Knyazeva, Valery M. Grishchenko, Roman S. Fadeev, Vladimir S. Akatov, Sergei E. Permyakov and Eugene A. Permyako, Biochemistry, 2008, 47 (49), pp 13127-13137 (online publication 12 Nov 2008).
ftp://www.cohensw.com/pub/pca/Knyazeva_Biochem_20081112.pdf

[29] <http://www.hakanssonlab.com>
Professor Anders Hakansson's research web site, which includes a discussion of the HAMLET research he initiated in 1995 as a graduate student.

[30] http://www.cohensw.com/mvpcsg_nov99_text.html

[31] <http://www.hmbana.com/>
Human Milk Banking Association of North America