Evaluation of an integrated yoga intervention in modulating psychological stress and radiation induced genotoxic stress in breast cancer patients undergoing radiotherapy

By

Birendra Nath Bannerjee

Abstract

Background

Stress in any form both psychological and physiological has been known to be associated with cancer from the time of diagnosis. Yoga is an ancient eastern system which is being extensively used in research worldwide to study the effects in reducing the stress at the psychological and physiological levels.

Aim

In the current study an attempt has been made to find the effects of an integrated yoga program in modulating the perceived stress levels, the anxiety as well as depression levels and radiation induced DNA damage in breast cancer patients undergoing radiotherapy

Method

A total of 78 patients were recruited and randomized in the study cohort. The Yoga group had n=35 patients and control group had n=23 patients with 20drop outs Two Psychological questionnaires (HADS and PSS) were taken pre and post radiotherapy. DNA damage was measured to find the effect of radiation on the peripheral blood lymphocytes (PBLs) of the patients both pre and post radiotherapy. The intervention group was given an integrated Yoga program which included guided relaxation, meditation, breathing practices, asanas and imageries with positive group practices for six weeks. The control group was given supportive counseling and waitlisted to be given yoga after the trial period.

Results

There was a significant decrease in the anxiety levels in yoga intervention group from Mean=8.5(baseline) to Mean=4.1 (48.2%) after the 6 weeks of yoga program, where as in the control group the Mean anxiety score increased to 10.5 (28%) p<0.001 for ANCOVA. The post depression score for the intervention group decreased from Mean= 8.0(baseline) to Mean= 3.4(57.5%) where as the in the control group the score increased from 7.8(baseline) to 9.7(24%). In the yoga group the mean perceived stress score (PSS) decreased from 20.4(baseline) to14.9 (26.9%) where as the control group showed no change in pre and post radiotherapy Mean= 19.0 and Mean=20.4. The DNA damage due to radiation was significantly high in both the yoga and control group after radiotherapy But the post radiotherapy DNA damage was slightly lesser Mean=24.3 when compared to the control group Mean=28.8.p<0.001.The baseline DNA damage being 2.6 and 2.8 respectively.

Conclusion

In our study we found that the integrated Yoga program was effective in reducing the stress significantly at both physiological and psychological levels when compared to the supportive counseling.

Key words: Yoga, meditation, radiotherapy, stress, DNA damage.

REFERENCES:

- Tapasyananda, Swami. (2003). Srimad Bhagavad Gita. Madras: Sri Ramakrishna Math.
- Prabhavananda , Swami. (2003). Patanjali Yoga Sutras. Madras: Sri Ramakrishna Math.
- Sarvananda, Swami. (1965). Taittiriyopanisad. . Madras: Sri Ramakrishna Math.
- Madhavanäëòä,Swami.(May2004).TheBrahadäraëyaka.Upaniñada Kolkata: Advaita Ashrama,
- Gambhiränandä, Swami (April 2003). Chändogya Upanisad. Kolkata Advaita Ashrama.

- Geden, Rev A.S. (1999). The Philosophy of the Upanishads. Delhi: Motilal Banarsidas.
- Nagendra, H. R. (August 2004). Yoga Its Basis and Applications. Bangalore, Swami Vivekananda Yoga Prakashana.
- Vidyaranya, Swami. (1995). Païcadaçé . Madras: Sri Ramakrishna Math.
- Anubhavananda, Swami. (1995). Païcadaçé Vol.V. Bombay: Central Chinmaya Mission Trust.
- Paramahamsa Niranjanānanda.(1994). Nirvanoniñad (Yoga Siddanta Bhashya). Bihar: Sri Panch Dashnam Paramhamsa Alakh Bara.
- Muktibodhananda, Swami. (2004). Hatha Yoga Pradipika. Bihar: Yoga Publications Trust. Suddhabodhananda, Swami. (1994). Vedānta Païcadaçé .Bombay: Sri Visweswar Trust.
- Ghosh, Manmohan. (2002). NATYASASTRA. Varanasi: Chowkhambasanskrit series Office.
- Bhuteshananda, Swami. (April 2003). Narada Bhakti Sutras. Calcutta.: Advaita Ashram.
- Gruol, D.L., Netzeband, J.G., Schneeloch, J., Gullette, C.E. L-type Ca2+ channels contribute to current-evoked spike firing and associated Ca2+ signals in cerebellar Purkinje neurons. Developmental Regulation of Ion Channel Function in CNS Neurons Cerebellum 5:146, 2006
- Lijun Zhou Johns Hopkins study: Scientific American Mind- April 14, 2006
- David R Hamilton <u>It's the Thought That Counts</u>, 2005, published by Hay House.
- Carey, Benedict. (28 September 2007). Genes Tied to Bad Reactions to Antidepressant Drug. New York Times.
- Courtney M, Cognition, August 1999 Am J Psychiatry 156:1137,
- Bruce McNaughton 'Speed Of Thought' Guides Brain's Memory Consolidation Science Daily (Nov. 16, 2007)
- Damodaran A, Malathi A, Patil N, Shah N, Suryavansihi, Marathe S.Therapeutic potential of yoga practices in modifying cardiovascular risk profile in middle aged men and women. J Assoc Physicians India. 2002 May;50(5):631-2
- Bijlani RL, Vempati RP, Yadav RK, Ray RB, Gupta V, Sharma R, Mehta N,
 Mahapatra SC.A brief but comprehensive lifestyle education program based on yoga reduces risk factors for cardiovascular disease and diabetes

- mellitus. J Altern Complement Med. 2005 Apr;11(2):267-74
- Qigong practitioners: a pilot study in gene regulation by mind-body interaction. Microarray Core, Center for Immunology, University of Texas Southwestern Medical Center, Dallas, TX, Li QZ, Li P, Garcia GE, Johnson RJ, Feng L Genomic profiling of neutrophil transcripts in Asian USA.
- Touitou Y, Bogdan A, Levi F, Benavides M, Auzeby A. Disruption of the circadian patterns of serum cortisol in breast and ovarian **cancer** patients: relationships with tumour marker antigens. Br JCancer 1996; 74: 1248–52.
- Van der Pompe G, Antoni MH, Heijnen CJ. Elevated basal cortisol levels and attenuated ACTH and cortisol responses to a behavioral challenge in women with metastatic breast cancer. Psychoneuroendocrinology 1996; 21: 361–74.
- Stanton AL, Snider PR. Coping with a breast **cancer** diagnosis: a prospective study. Health Psychol 1993; 12: 16–23.
- Carver CS, Pozo-Kaderman C, Price AA, Noriega V, Harris SD, Derhagopian RP, Robinson DS, Moffatt FL. Concern about aspects of body image and adjustment to early stage breast **cancer**. Psychosom Med 1998; 60: 168–74
- Andersen BL. Psychological interventions for cancer patients to enhance the quality of life. J Consult Clin Psychol 1992; 60: 552–68.
- Trijsburg RW, van Knippenberg FCE, Rijpma SE. Effects of psychological treatment on **cancer**patients: a critical review. Psychosom Med 1992; 54: 489–517.
- van der Pompe G, Duivenvoorden HJ, Antoni MH, Visser A, Heijnen CJ. Effectiveness of a short-term group psychotherapy program on endocrine and immune function in breast **cancer** patients: an exploratory study. J Psychosom Res 1997; 42: 453–66.
- Schedlowski M, Jung C, Schimanski G, Tewes U, Schmoll HJ. Effects of behavioral intervention on plasma cortisol and lymphocytes in breast cancer patients: an exploratory study. Psychooncology 1994; 3: 181–7.
- Fromm K, Andrykowski MA, Hunt J. Positive and negative psychosocial sequelae of bone marrow transplantation: implications for quality of life assessment. J Behav Med 1996; 19: 221–40.
- Tedeschi RG, Calhoun LG. Trauma and transformation: growing in the aftermath of suffering. Thousand Oaks (CA): Sage; 1995.
- Antoni MH, Carver C, Boyers A, McGregor B, Arena P, Lehman J, Harris S,

Price A, Alferi S, Culver J, Cruess DG. Cognitivebehavioral **stress** management intervention enhances optimism and the sense of positive contribution among women under treatment for early stage breast **cancer**[abstract]. Psychosom Med 1999; 61: 94.

- McNair D, Lorr M, Droppleman L. Manual for the Profile of Mood States.
 San Diego (CA): Educational and Industrial Testing Service; 1981.
- Carver CS, Pozo C, Harris SD, Noriega V, Scheier MF, Robinson DS, Ketcham AS, Moffat FL, Clark KC. How coping mediates the effect of optimism on distress: a study of women with early stage breast cancer. J Pers Social Psychol 1993; 65: 375–90.
- Baron RM, Kenny DA. The moderator-mediator variable distinction in social psychological research: conceptual, strategic, and statistical considerations.
 J Pers Soc Psychol 1986; 51: 1173–82.
- Berk L, Bittmen B, Covington T, Bickford K, Tom S, Westengard J. A video presentation of music, nature's imagery, and positive affirmations as a combined eustress paradigm modulates neuroendocrine hormones [abstract]. Ann Behav Med 1997; 19: 201.
- Buchanan T al'Absi M Lovallo W Cortisol fluctuates with increases and decreases in negative affect. Psychoneuroendocrinology 1999; 24: 227–41.
- Smyth J, Ockenfels M, Porter L, Kirschbaum C, Hellhammer D, Stone AA. Stressors and mood measured on a momentary basis are associated with salivary cortisol secretion. Psychoneuroendocrinology 1998; 23: 353– 70
- McEwen B. Protective and damaging effects of stress mediators. N Engl J Med 1998; 338: 171–9.
- Antoni MH, Wagner S, Cruess DG, Kumar M, Ironson G, Schneiderman N. Cognitive behavioralstress management effects on distress and adrenal hormones in HIV-infected gay men [abstract]. Psychosom Med 1999; 61: 127.
- Antoni MH, Baggett L, Ironson G, LaPerriere A, August S, Klimas N, Schneiderman N, Fletcher MA. Cognitive-behavioral stress management intervention buffers distress responses and immunologic changes following notification of HIV-1 seropositivity. J Consult Clin Psychol 1991; 59: 906–15.
- Lutgendorf SK, Antoni MH, Ironson G, Klimas N, Kumar M, Starr K, McCabe
 P, Cleven K, Fletcher MA, Schneiderman N. Cognitive-

- behavioral **stress** management decreases dysphoric mood and herpes simplex virus-type 2 antibody titers in symptomatic HIV-seronegative gay men. J Consult Clin Psychol 1997; 65: 31–43.
- Stone AA, Neale JM, Cox DS, Napoli A, Valdimarsdottir H, Kennedy-Moore E. Daily events are associated with a secretory immune response to an oral antigen in men. Health Psychol 1994; 13: 440–6.
- Bower J, Kemeny M, Taylor S, Fahey J. Cognitive processing, discovery of meaning, CD4 decline, and AIDS-related mortality among bereaved HIVseropositive men. J Consult Clin Psychol 1998; 66: 979–86.
- Hanahan D, Weinberg RA. Cell. 2000 The hallmarks of cancer. Jan 7;100(1):57-70.
- Parshad R, Price FM, Bohr VA, Cowans KH, Zujewski JA, Sanford KK Deficient DNA repair capacity, a predisposing factor in breast cancer. Br J Cancer. 1996 Jul;74(1):1-5.
- <u>Pinar B</u>, <u>Lara PC</u>, <u>Lloret M</u>, <u>Bordón E</u>, <u>Núñez MI</u>, <u>Villalobos M</u>, <u>Guerrero R</u>, <u>Luna JD</u>, <u>Ruiz de Almodóvar JM</u> Radiation-induced DNA damage as a predictor of long-term toxicity in locally advanced breast cancer patients treated with high-dose hyper fractionated radical radiotherapy. <u>Radiat Res. 2007 Oct; 68(4):415-22</u>
- Ashikaga T, Bosompra K, O'Brien P et al (2002) Use of complimentary and alternative medicine by breast cancer patients: prevalence, patterns and communication with physicians. Support Care Cancer 10(7):542–548
- Burstein HJ, Gelber S, Guadagnoli E et al (1999) Use of alternative medicine by women with early-stage breast cancer. N Engl J Med 340(22):1733–1739
- Edgar L, Remmer J, Rosberger Z et al (2000) Resource use in women completing treatment for breast cancer. Psychooncology 9(5):428–438
- Henderson JW, Donatelle RJ (2004) Complementary and alternative medicine use by women after completion of allopathic treatment for breast cancer. Altern Ther HealthMed 10(1):52–57
- Lee MM, Lin SS, Wrensch MR et al (2000) Alternative therapies used by women with breast cancer in four ethnic populations. J Natl Cancer Inst 92(1):42–47
- Lengacher CA, Bennett MP, Kipp KE et al (2003) Design and testing of the use of a complementary and alternative therapies survey in women with breast cancer. Oncol Nurs Forum 30(5):811–821
- Nagel G, Hoyer H, Katenkamp D (2004) Use of complementary and alternative medicine by patients with breast cancer: observations from a

- health-care survey. Support Care Cancer 12(11):789–796
- Navo MA, Phan J, Vaughan C et al (2004) An assessment of the utilization of complementary and alternative medication in women with gynecologic or breast malignancies. J Clin Oncol 22(4):671–677
- Richardson MA, Sanders T, Palmer JL et al (2000) Complementary/ alternative medicine use in a comprehensive cancer center and the implications for oncology. J Clin Oncol 18(13):2505–2514
- Moschen R, Kemmler G, Schweigkofler H et al (2001) Use of alternative/complementary therapy in breast cancer patients—a psychological perspective. Support Care Cancer 9(4):267–274.
- Shen J, Andersen R, Albert PS et al (2002) Use of complementary/ alternative therapies by women with advanced stage breast cancer. BMC Complement Altern Med 2:8
- Rakovitch E, Pignol JP, Chartier C et al (2005) Complementary and alternative medicine use is associated with an increased perception of breast cancer risk and death. Breast Cancer Res Treat 90(2):139–148
- Gray RE, Fitch M, Goel V et al (2003) Utilization of omplementary alternative services by women with breast cancer. J Health Soc Policy 16(4):75–84
- Boon H, Stewart M, Kennard MA et al (2000) Use of complementary/alternative medicine by breast cancer survivors in Ontario: prevalence and perceptions. J Clin Oncol 8(13):2515–2521
- Casso D, Buist DS, Taplin S (2004) Quality of life of 5–10 year breast cancer survivors diagnosed between age 40 and 49. Health Qual Life Outcomes 2:25.
- Paltiel O, Avitzour M, Peretz T et al (2001) Determinants of the use of complementary therapies by patients with cancer. J Clin Oncol 19(9):2439– 2448
- Barnes PM, Powell-Griner E, McFann K et al (2004) Complementary and alternative medicine use among adults: United States, 2002. Adv Data 2004(343):1–19
- Alferi SM, Antoni MH, Ironson G et al (2001) Factors predicting the use of complementary therapies in a multiethnic sample of early-stage breast cancer patients. J Am Med Womens Assoc 56(3):120–123, 126
- Swisher EM, Cohn DE, Goff BA et al (2002) Use of complementary and alternative medicine among women with gynecologic cancers. Gynecol

- Oncol 84(3):363-367
- Wyatt GK, Friedman LL, Given CW et al (1999) Complementary therapy use among older cancer patients. Cancer Pract 7(3):136–144
- Davidson R, Geoghegan L, McLaughlin L et al (2005) Psychological characteristics of cancer patients who use complementary therapies. Psychooncology 14(3):187–195.
- Eisenberg DM, Davis RB, Ettner SL et al (1998) Trends in alternative medicine use in the United States, 1990–1997: results of a follow-up national survey. Jama 280(18):1569–1575
- Brook RH, Ware JE Jr, Davies-Avery A et al (1979) Overview of adult health measures fielded in Rand's health insurance study. Med Care 17(7 Suppl):iii–x, 1–131
- Ware JE Jr, Sherbourne CD (1992) The MOS 36-item short form health survey (SF-36). I. Conceptual framework and item selection. Med Care 30(6):473–483
- McHorney CA, Ware JE Jr, Lu JF et al (1994) The MOS 36-item Short-Form Health Survey (SF-36): III. Tests of data quality, scaling assumptions, and reliability across diverse patient groups. Med Care 32(1):40–66
- McHorney CA, Ware JE Jr, Raczek AE (1993) The MOS 36-Item Short-Form Health Survey (SF-36): II. Psychometric and clinical tests of validity in measuring physical and mental health constructs. Med Care 31(3):247–263
- Scheier MF, Carver CS, Bridges MW (1994) Distinguishing optimism from neuroticism (and trait anxiety, self-mastery, and self-esteem): a reevaluation of the Life Orientation Test. J Pers Soc Psychol 67(6):1063– 1078
- Carver CS, Pozo C, Harris SD et al (1993) How coping mediates the effect of optimism on distress: a study of women with early stage breast cancer. J Pers Soc Psychol 65(2):375–390
- Osoba D, Bezjak A, Brundage M et al (2005) Analysis and interpretation of health-related quality-of-life data from clinical trials: basic approach of The National Cancer Institute of Canada Clinical Trials Group. Eur J Cancer 41(2):280–287
- Bairati I, Meyer F, Gelinas M et al (2005) Randomized trial of antioxidant vitamins to prevent acute adverse effects of radiation therapy in head and neck cancer patients. J ClinOncol 23(24):5805–5813 Breast Cancer Res Treat (2006) 100:219–227 227

- Cohen, S., & Williamson, G. (1988). Perceived stress in a probability sample of the United States. In S. Spacapam & S. Oskamp (Eds.), The social psychology of health: Claremont Symposium on applied social psychology. Newbury Park, CA: Sage.
- Johansen, J., Bentzen, S.M., Overgaard, J., and Overgaard, M. (1996) Relationship between the in vitro radiosensitivity of skin fibroblasts and the expression of subcutaneous fibrosis, telangiectasia, and skin erythema after radiotherapy. *Radiother.Oncol.*, **40**, 101-109.
- Johansen, J., Streffer, C., Fuhrmann, C., Bentzen, S.M., Stausbol-Gron, B., Overgaard, M., and Overgaard, J. (1998) Radiosensitivity of normal fibroblasts from breast cancer patients assessed by the micronucleus and colony assays. *Int. J. Radiat. Biol.*, **73**, 671-678.
- Bonassi,S., Fenech,M., Lando,C., Lin,Y.P., Ceppi,M., Chang,W.P., Holland,N., Kirsch-Volders,M., Zeiger,E., Ban,S., Barale,R., Bigatti,M.P., Bolognesi,C., Jia,C., Di,G.M., Ferguson,L.R., Fucic,A., Lima,O.G., Hrelia,P., Krishnaja,A.P., Lee,T.K., Migliore,L., Mikhalevich,L., Mirkova,E., Mosesso,P., Muller,W.U., Odagiri,Y., Scarffi,M.R., Szabova,E., Vorobtsova,I., Vral,A., and Zijno,A. (2001) HUman MicroNucleus project: international database comparison for results with the cytokinesis-block micronucleus assay in human lymphocytes: I. Effect of laboratory protocol, scoring criteria, and host factors on the frequency of micronuclei. *Environ.Mol.Mutagen.*, 37, 31-45.
- Fenech, M. (2002) Chromosomal biomarkers of genomic instability relevant to cancer. *Drug Discov.Today*, **7**, 1128-1137.
- Fenech, M. (2002) Biomarkers of genetic damage for cancer epidemiology. *Toxicology*, **181-182:411-6.**, 411-416.
- Fenech, M. (2006) Cytokinesis-block micronucleus assay evolves into a "cytome" assay of chromosomal instability, mitotic dysfunction and cell death. *Mutat.Res.*, **600**, 58-66.
- Oppitz, U., Baier, K., Wulf, J., Schakowski, R., and Flentje, M. (2001) The in vitro colony assay: a predictor of clinical outcome. *Int. J. Radiat. Biol.*, 77, 105-110.
- Lee,T.K., Allison,R.R., O'Brien,K.F., Naves,J.L., Karlsson,U.L., and Wiley,A.L. (2002) Persistence of micronuclei in lymphocytes of cancer patients after radiotherapy. *Radiat.Res.*, **157**, 678-684.
- Blackburn, E.H. (1991) Telomeres. *Trends Biochem. Sci.*, **16**, 378-381.
- Greider, C.W. (1996) Telomere length regulation. *Annu.Rev.Biochem.*, **65:337-65.**, 337-365.

- Greider, C.W. (1998) Telomerase activity, cell proliferation, and cancer. *Proc.Natl.Acad.Sci.U.S.A.*, **95**, 90-92.
- Moyzis,R.K., Buckingham,J.M., Cram,L.S., Dani,M., Deaven,L.L., Jones,M.D., Meyne,J., Ratliff,R.L., and Wu,J.R. (1988) A highly conserved repetitive DNA sequence, (TTAGGG)n, present at the telomeres of human chromosomes. *Proc.Natl.Acad.Sci.U.S.A.*, 85, 6622-6626.
- Blasco, M.A. (2003) Telomeres and cancer: a tale with many endings. *Curr.Opin.Genet.Dev.*, **13**, 70-76.
- Slijepcevic, P. (2004) Is there a link between telomere maintenance and radiosensitivity? *Radiat.Res.*, **161**, 82-86.
- Hande, M.P., Balajee, A.S., Tchirkov, A., Wynshaw-Boris, A., and Lansdorp, P.M. (2001) Extra-chromosomal telomeric DNA in cells from Atm(-/-) mice and patients with ataxia-telangiectasia. *Hum. Mol. Genet.*, 10, 519-528.
- d'Adda di Fagagna, F., Hande, M.P., Tong, W.M., Lansdorp, P.M., Wang, Z.Q., and Jackson, S.P. (1999) Functions of poly (ADP-ribose) polymerase in controlling telomere length and chromosomal stability. *Nat Genet*, **23**, 76-80.
- d'Adda di Fagagna, F., Hande, M.P., Tong, W.M., Roth, D., Lansdorp, P.M., Wang, Z.Q., and Jackson, S.P. (2001) Effects of DNA nonhomologous endjoining factors on telomere length and chromosomal stability in mammalian cells. *Curr Biol*, **11**, 1192-6.
- Gilley, D., Tanaka, H., Hande, M.P., Kurimasa, A., Li, G.C., Oshimura, M., and Chen, D.J. (2001) DNA-PKcs is critical for telomere capping. *Proc.Natl.Acad.Sci.U.S.A.*, **98**, 15084-15088.
- Hande, M.P. (2004) DNA repair factors and telomere-chromosome integrity in mammalian cells. *Cytogenet. Genome Res.*, **104**, 116-122.
- Hande, P., Slijepcevic, P., Silver, A., Bouffler, S., van, B.P., Bryant, P., and Lansdorp, P. (1999) Elongated telomeres in scid mice. *Genomics*, 56, 221-223.
- Hsu,H.L., Gilley,D., Galande,S.A., Hande,M.P., Allen,B., Kim,S.H., Li,G.C., Campisi,J., Kohwi-Shigematsu,T., and Chen,D.J. (2000) Ku acts in a unique way at the mammalian telomere to prevent end joining. *Genes Dev.*, 14, 2807-2812.
- McPherson, J.P., Hande, M.P., Poonepalli, A., Lemmers, B., Zablocki, E., Migon, E., Shehabeldin, A., Porras, A., Karaskova, J., Vukovic, B., Squire, J., and Hakem, R. (2006) A role for Brca1 in chromosome end

- maintenance. Hum. Mol. Genet., 15, 831-838.
- Fenech, M. and Morley, A.A. (1985) Measurement of micronuclei in lymphocytes. *Mutat.Res.*, **147**, 29-36.
- Bender, M.A., Awa, A.A., Brooks, A.L., Evans, H.J., Groer, P.G., Littlefield, L.G., Pereira, C., Preston, R.J., and Wachholz, B.W. (1988) Current status of cytogenetic procedures to detect and quantify previous exposures to radiation. *Mutat. Res.*, 196, 103-159.
- Venkatachalam,P., Solomon,F.D., Prabhu,B.K., Mohankumar,M.N., Gajendiran,N., and Jeevanram,R.K. (1999) Estimation of dose in cancer patients treated with fractionated radiotherapy using translocation, dicentrics and micronuclei frequency in peripheral blood lymphocytes. *Mutat.Res.*, 429, 1-12.
- Venkatachalam, P., Paul, S.F., Mohankumar, M.N., Prabhu, B.K.,
 Gajendiran, N., Kathiresan, A., and Jeevanram, R.K. (1999) Higher frequency of dicentrics and micronuclei in peripheral blood lymphocytes of cancer patients. *Mutat. Res.*, 425, 1-8.
- Mozdarani, H., Mansouri, Z., and Haeri, S.A. (2005) Cytogenetic radiosensitivity of g0-lymphocytes of breast and esophageal cancer patients as determined by micronucleus assay. *J. Radiat. Res. (Tokyo).*, **46**, 111-116.
- Scott,D., Barber,J.B., Levine,E.L., Burrill,W., and Roberts,S.A. (1998) Radiation-induced micronucleus induction in lymphocytes identifies a high frequency of radiosensitive cases among breast cancer patients: a test for predisposition? *Br.J.Cancer.*, **77**, 614-620.
- Scott,D., Barber,J.B., Spreadborough,A.R., Burrill,W., and Roberts,S.A. (1999) Increased chromosomal radiosensitivity in breast cancer patients: a comparison of two assays.*Int.J.Radiat.Biol.*, **75**, 1-10.
- Scott, D. (2004) Chromosomal radiosensitivity and low penetrance predisposition to cancer. *Cytogenet. Genome Res.*, **104**, 365-370.
- Goytisolo, F.A., Samper, E., Martin-Caballero, J., Finnon, P., Herrera, E., Flores, J.M., Bouffler, S.D., and Blasco, M.A. (2000) Short telomeres result in organismal hypersensitivity to ionizing radiation in mammals. *J. Exp. Med.*, **192**, 1625-1636.
- Poonepalli, A., Balakrishnan, L., Khaw, A.K., Low, G.K., Jayapal, M., Bhattacharjee, R.N., Akira, S., Balajee, A.S., and Hande, M.P. (2005) Lack of poly(ADP-ribose) polymerase-1 gene product enhances cellular sensitivity to arsenite. *Cancer Res.*, **65**, 10977-10983.
- Acar, H., Caliskan, U., Demirel, S., and Largaespada, D.A. (2001) Micronucleus

- incidence and their chromosomal origin related to therapy in acute lymphoblastic leukemia (ALL) patients: detection by micronucleus and FISH techniques. *Teratog.Carcinog.Mutagen.*, **21**, 341-347.
- Norppa,H. and Falck,G.C. (2003) What do human micronuclei contain? *Mutagenesis.*, **18**, 221-233.
- Desmaze, C., Soria, J.C., Freulet-Marriere, M.A., Mathieu, N., and Sabatier, L. (2003) Telomere-driven genomic instability in cancer cells. *Cancer Lett.*, 194, 173-182.
- Desmaze, C., Pirzio, L.M., Blaise, R., Mondello, C., Giulotto, E., Murnane, J.P., and Sabatier, L. (2004) Interstitial telomeric repeats are not preferentially involved in radiation-induced chromosome aberrations in human cells. Cytogenet. Genome Res., 104, 123-130.
- Slijepcevic, P., Natarajan, A.T., and Bryant, P.E. (1998) Telomeres and radiation-induced chromosome breakage. *Mutagenesis.*, **13**, 45-49.
- Gimbel MA (1998). Yoga, meditation, and imagery: clinical applications. NursePract Forum Dec; **9**(4): 243-55.
- Telles S, Nagarathna R, Nagendra HR. (1998) Autonomic changes while mentally repeating two syllables—one meaningful and the other neutral. Indian J Physiol Pharmacol; 42: 57–63.
- Nagarathna R, Nagendra HR (1985). Yoga for bronchial asthma: a controlled study. *Br Med J (Clin Res Ed)*; **291**:1077–1079.
- Henderson LE. (1989). Characteristics and immune function of meditators and nonmeditators: an exploratory study. Calgary, Canada: University of Calgary;
- Derogatis LR, Morrow GR, Fetting J. The prevalence of psychiatric disorders among cancer patients. JAMA 1983; 249: 751–7
- Stefanek M, Derogatis L, Shaw (1987) A. Psychological distress among oncology patients. Psychosomatics; 28: 530–8
- Farber JM, Weinerman BH, Kuypers JA. (1983). Psychosocial distress in oncology outpatients. J Psychosoc Oncol; 2: 109–18
- Spiegel D. The Fox article reviewed. Oncology (1995); 9: 253–5.
- Fox BH. (1995). The role of psychological factors in cancer incidence and prognosis. Oncology; **9**: 245–53.
- Holmes MD (2006) Correlates of use of different types of complementary and alternative medicine by breast cancer survivors in the nurses' health study.
 - Breast Cancer Res Treat. Nov; 100(2):219-27.

- Cassileth BR, Chapman CC. (1998) Alternative and complementary cancer therapies. Cancer; **77**: 1026–34.
- Benson H. The relaxation response. New York: Morrow; 1975.
- Johnson NA, Heller RF. (1998). Prediction of patient non-adherence with home-based exercise for cardiac rehabilitation: The role of perceived barriers and perceived benefits. Prev Med **27**: 56–64.
- Brawley LR, Culos-Reed SN, Angove J, Hoffman- Goetz L. (2002).
 Understanding the barriers to physical activity for cancer patients: Review and recommendations. JPsychosoc Oncol 20: 1–21.
- Solberg EE, Halvorsen R, Sundgot-Borgen J, Ingjer F, Holen A. (1995) Meditation: a modulator of the immune response to physical stress? A brief report. Br J Sports Med; **29**: 255–7.
- Walton KG, Pugh ND, Gelderloos P, Macrae P. (1995). Stress reduction and preventing hypertension: preliminary support for a psychoneuroendocrine mechanism. J Altern Complement Med; 1: 263–83.
- Schneider RH, Staggers F, Alexander CN, Sheppard W, Rainforth M, Kondwani B, Smith S, King CG. (1995). A randomized controlled trial of stress reduction for hypertension in older African Americans.
 Hypertension; 26: 820–
- Raub JA (2002). Psychophysiologic effects of Hatha Yoga on musculoskeletal and cardiopulmonary function: a literature review. J Altern Complement Med.Dec; **8**(6): 797-812.
- Jayasinghe SR (2004). Yoga in cardiac health (a review). Eur J Cardiovasc PrevRehabil Oct; **11**(5): 369-75.
- Wenneberg SR, Schneider RH, Walton KG, MacLean CR, Levitsky DK, Salerno JW, Wallace RK, Mandarino JV, Rainforth MV, Waziri R. A controlled study of the effects of the transcendental meditation program on cardiovascular reactivity and ambulatory blood pressure. Int J Neurosci 1997; **89**: 15–28.
- Sahay BK, Sahay RK (2002). Lifestyle modification in management of diabetes mellitus. J Indian Med Assoc Mar; **100**(3): 178-80.
- Sainani GS (2003). Non-drug therapy in prevention and control of hypertension. J Assoc Physicians India Oct; **51**: 1001-6.
- Sudsuang R, Chentanez V, Veluvan K. Effect of Buddhist meditation on serum cortisol and total protein levels, blood pressure, pulse rate, lung volume and reaction time. Physiol Behav 1991; **50**: 543–8.
- Burstein HJ, Gelber S, Guadagnoli E et al (1999) Use of alternative medicine by women with early-stage breast cancer. N Engl J Med 340(22):1733–1739

- Carlson LE, Ursuliak Z, Goodey E, Angen M, Speca M. (2001). The effects of a mindfulness meditation based stress reduction program on mood and symptoms of stress in cancer outpatients: 6-month follow-up. Support Care Cancer; 9:112–23.
- <u>Carlson LE, Speca M, Patel KD, Goodey E, (2004)</u>. Mindfulness-based stress reduction in relation to quality of life, mood, symptoms of stress and levels of cortisol, dehydroepiandrosterone sulfate (DHEAS) and melatonin in breast and prostate cancer outpatients. Psychoneuroendocrinology. May; 29(4):448 -74.
- <u>Carlson LE, Speca M, Patel KD, Goodey E, (2003)</u>. Mindfulness-based stress reduction in relation to quality of life, mood, symptoms of stress, and immune parameters in breast and prostate cancer outpatients. Psychosom Med. Jul-Aug; 65(4):571-8
- Rosenbaum E, Gautier H, Fobair P, Neri E, Festa B, Hawn M, Andrews A,
 Hirshberger N, Selim S, Spiegel D.2004 Cancer supportive care, improving
 the quality of life for cancer patients. A program evaluation report. Support
 Care Cancer. May; 12(5):293-301.
- Felten DL, Cohen N, Ader R, et al. Central neural circuits involved in
- neural-immune interactions. In: Ader R, Cohen N, Felten DL, eds.
- Psychoneuroimmunology, second ed. New York: Academic, 1991: 3-25.
- Renoux G, Bizierre K. Neocortex lateralization of immune function and
- of the activities of imuthiol, a T-cell specific immunopotentiator. In:
- Ader R, Cohen N, Felten DL, eds. Psychoneuroimmunology, second ed. New
- York: Academic, 1991: 127-48.
- Besedovsky HO, delRey A. Physiologic implications of the
- immunoneuro-endocrine network. In: Ader R, Cohen N, Felten DL, eds.
- Psychoneuroimmunology, second ed. New York: Academic, 1991: 589-608.
- Berkenbosch J, Van Oers J, del Rey A, et al. Corticotropin-releasing
- factor-producing neurons in the rat activated by inteleukin-1. Science
- 1987; 238: 524-26.
- Dantzer R, Kelley KW. Stress and immunity: an integrated view of
- relationships between the brain and the immune system. Life Sci 1989;
- 44: 1995-2008.
- Kelley KW. Growth hormone in immunobiology. In: Ader R, Cohen N,
- Felten DL, eds. Psychoneuroimmunology, second ed. New York: Academic,
- 1991: 377-402.
- Bernton EW, Bryant HU, Holaday JW. Prolactin and immune function. In:

- Ader R, Cohen N, Felten DL, eds. Psychoneuroimmunology, second ed. NewYork: Academic, 1991: 403-28.
- Casso D, Buist DS, Taplin S (2004) Quality of life of 5–10 year breast cancer survivors diagnosed between age 40 and 49. Health Qual Life Outcomes 2:25
- <u>Carson JW, Carson KM, Porter LS, Keefe FJ, Shaw H, Miller JM.</u> (2007) Yoga for women with metastatic breast cancer: results from a pilot study. J Pain Symptom Manage Mar; 33(3):331-4
- <u>Culos-Reed SN, Carlson LE, Daroux LM, Hately-Aldous S (2006)</u>. A pilot study of yoga for breast cancer survivors: physical and psychological benefits.
 Psychooncology. Oct; **15**(10):891-7.
- Raghavendra Rao M, Nagarathna Raghuram, Nagendra HR, Gopinath KS, Ravi B Diwakar, Shekar Patil, Ramesh Bilimagga S, Nalini Rao. (2007). Effects of an integrated yoga program on chemotherapy induced nausea and emesis in breast cancer patients. Eur J Cancer care (In press)
- Scott D barber JBP Levine El BursiloW Robert SA, (1998) Radiation induced micronucleus induction in lymphocytes identifies a high frequency of radiosensitive cases among breast cancer patients: a test for predisposition. British .J.Cancer 7:614-620.
- Hossein Mozdarani, ZahraMansouri and Abolghasem HAER (2005)
 Cytogenetic Radiosensitivity of G-lymphocyte of Breast and Esophageal
 Carrier Patients as Determined by Micronuleus Assay. J. Radiat. Res 46, 111
 116
- <u>Banerjee B</u>, <u>Sharma S</u>, <u>Hegde S</u>, <u>Hande MP</u> (2007) Analysis of telomere damage by fluorescence in situ hybridisation on micronuclei in lymphocytes of breast carcinoma patients after radiotherapy<u>Breast Cancer Res Treat.</u> Feb 28; (in print)
- Kiecolt-Glaser, J.K., Stephens R.E, Lipetz P.D, Speicher C.E and Glaser R (1985). Distress and DNA repair in human Lymphocytes. J Behav Med.8 (4):311-320
- Cohen L, Marshall G.D,Jr., Cheng L, Agarwal SK, Wei Q. (2000) DNA Repair capacity in healthy medical students during and after exam stress. J Behav Med. Dec; **23**(6):531-44.
- Glaser R, Thorn B.E, Tarr K.L, Kiecolt-Glaser, J.K., D'Ambrosio, S.M. (1985).
 Effects of stress on methyltransferase synthesis. An important DNA repair enzyme. Health Psychol. 4(5): 403-412.
- E. Epel and E. Blackburn et al., (2004) Accelerated telomere shortening in

response to exposure to life stress, PNAS 101, pp. 17312–17315.

- Dr R.Nagaratna and Dr H.R Nagendra; Integrated approach of yoga therapy for positive health Swami Vivekananda yoga prakashana India
- Elliott, G.R., Eisdorfer, C. 1982. Stress and Human Health. New York: Springer Publishing Company.
- Manuck, S.B., Cohen, S., Rabin, B.S., Muldoon, M.F., Bachen "Individual _differences in cellular immune response to stress. Psychological Science. 2: 111- 114.
- Cancer Research UK What's New Clinical Trials Donate About Access Keys NHS Information Partners
- McEwen BS. Stressed or stressed out: what is the difference? J Psychiatry Neurosci. 2005 Sep; 30(5):315-8.
- Ballieux, R.E. Breakdown in human adaptation to 'stress'. 1984. Boston: Martinus Nijhoff Publishers.
- Brosschot, J.F., Benschop, R.J., Godaert, G.L.R., Olff, M., De Smet, M., Heijnen, C.J., Ballieux, R.E. 1994. Influence of life stress on immunological reactivity to mild psychological stress. Psychosomatic Medicine. 56: 216-224.
- Glaser, W.D., Anderson, K.N, Anderson, L.E. 1992. The Mosby Medical Encyclopedia. New York: Plume.
- Kiecolt-Glaser, J.K., Cacioppo, J.T., Malarkey, W.B., Glaser, R. 1992. Acute psychological stressors and short-term immune changes: What, why, for whom, and to what extent? *Psychosomatic Medicine*. 54: 680-685.
- Ader R, Cohen N. Psychoneuroimmunology: conditioning and stress.
 AnnRev Psychol 1993; 44: 53-85.
- Ader R, Cohen N. The influence of conditioning on immune responses.
- In: Ader R, Cohen N, Felten DL, eds. Psychoneuroimmunology, second ed.New York: Academic, 1991: 611-46.
- Bruce D. Naliboff, , Minou Mayer, , Ronnie Fass, , Leah Z. Fitzgerald, , Lin Chang, , Roger Bolus, and Emeran A. Mayer "The Effect of Life Stress on Symptoms of Heartburn Psychosomatic Medicine 66:426-434 (2004)
- Naliboff BD, Benton D, Solomon GF, Morley JE, Fahey JL, Bloom
 ET, Makinodan T, Gilmore SL. Immunological changes in young and old adults during brief laboratory stress. Psychosom Med. 1991 Mar-Apr; 53(2):121-32
- Naliboff BD, Mayer M, Fass R, Fitzgerald LZ, Chang L, Bolus R, Mayer EA, The

- effect of life stress on symptoms of heartburn. <u>Psychosom Med.</u> 2004 May-Jun; 66(3):426-34.
- Walton KG, Pugh ND, Gelderloos P, Macrae P. Stress reduction and preventing hypertension: preliminary support for a psychoneuroendocrine mechanism. *J Altern Complement Med.* 1995; 1:263-283.
- <u>McEwen BS</u>. From molecules to mind. Stress, individual differences, and the social environment. *Ann N Y Acad Sci.* 2001 May; 935:42-9.
- Debouck C, Goodfellow PN. DNA microarrays in drug discovery and development. *Nat Genet*. 1999; **21**: 48–50.
- Chuaqui RF, Bonner RF, Best CJ *et al*. The genetics of cancer-a 3D model. *Nat Genet*. 1999; **21**: 38–41.
- Gerhold DL, Jensen RV, Gullans SR: Better therapeutics through microarrays. *Nat Genet*. 2002;**32**: 547–51.
- Jayapal M, Melendez AJ. DNA microarray technology for target identification and validation. *Clinical and Experimental Pharmacology & Physiology*. 2006; 33:496-503.
- Julian White. The Future of Microarray Readers. Pharmaceutical Discovery. 2004: 30-34