

Cardiovascular Disease and the Traditional Japanese Diet

Are some Japanese foods effective for the treatment of Cardiovascular Disease?

Cardiovascular disease (CVD) is the second largest “disease burden” to Australia, with the latest statistics showing over 469,000 individuals were hospitalised in 2006 for CVD conditions.¹ Mortality caused from CVD was recorded at over 45,600 individuals.¹ The chief preventable risk factors have been identified as cigarette smoking, obesity, high blood pressure, high cholesterol, insufficient exercise, poor nutrition and diabetes.²

Studies spanning decades investigate several Japanese staple foods as being potentially responsible for lower cases of CVD in Japan when compared with CVD in other countries such as Australia.³ Evidence suggests that diets predominantly high in soybean products, fish, seaweeds, vegetables, fruits and green tea, in addition to reducing chief risk factors, may contribute to a decreased risk of CVD.³ Furthermore, data suggests that lowered risk of CVD still exists with Japan’s high sodium intake and significant hypertensive population.³

Green Tea

One study of 1371 Japanese men aged over 40 years old showed increased green tea consumption decreased serum

concentrations of total cholesterol and triglyceride in blood samples taken from participants. Furthermore the results indicated an increased proportion of high density lipoprotein (HDL) cholesterol together with a decreased proportion of low and very low lipoprotein (LDL) cholesterols.⁴ One eleven year study of over 40,500 Japanese adults concluded that although the exact effects were still unclear, green tea consumption contributed to a reduction in CVD mortality. Moreover the strongest association expressed reduced stroke mortality in tea drinkers.⁵

More recent studies have identified and studied catechin and epigallocatechin gallate (EGCG) as two chief components of green tea which contribute to not only cardiovascular health but also anti-cancer effects and general metabolic health.⁶

One 12-week double-blind parallel multicenter trial of 240 individuals concluded that a green tea extract which contained 583 mg of catechin decreased body fat, systolic blood pressure (SBP) and low density lipoprotein (LDL) cholesterol when taken everyday.⁷ A study this year investigated the effects of catechin on 40



obese or overweight children through measurements of waist circumference, SBP, and LDL cholesterol. The data presented a definite observed benefit to health associated with the consumption of the catechin-rich green tea.⁸

The latest research on EGCG indicates that the mechanisms involve reducing vascular endothelial Monocyte chemotactic protein-1 (MCP-1) which is a fundamental element in the development of CVD.⁹

Natto

Fermented soya beans, or more commonly known by Japanese as *natto*, contains a unique bacteria strain identified as *Bacillus natto*.¹⁰ Nattokinase, which is an enzyme found in natto, has been investigated for its health effects and studies conclude a significant health benefit of the enzyme.¹⁰⁻¹⁴ Studies indicate positive in vitro hemorheological effects of nattokinase¹¹, particularly reducing thrombotic activity.¹²⁻¹⁴ Anti-thrombotic effects of nattokinase supplements are so effective that it has been suggested haematologists should be aware of interaction with anticoagulant treatment.¹⁵

In vitro studies of natto have shown it to have useful antioxidant properties, principally inhibiting effects on LDL oxidation in rats.¹⁶

Natto can be purchased, usually in 55 gram packs, within most large asian supermarkets who stock Japanese foods



Shiitake Mushrooms

Shiitake mushrooms (*Lentinus edodes*), as a wholefood, have numerous positive health effects within the body. Specifically in the treatment of CVD, shiitake are hypocholesterolemic.¹⁷⁻¹⁸ In one in vivo study, rats were fed diets containing different levels of corn fat and 2% shiitake mushrooms. The results indicated a definite positive effect on fat metabolism in the rats which were fed the mushrooms.¹⁷ In a different in vivo study, the cholesterol lowering effects of three different mushrooms were compared with cellulose powder. The fecal cholesterol excretion in the shiitake was significantly higher than the cellulose powder.¹⁸



The anti-atherogenic properties of *Lentinus edodes* mycelia were established in a studying which involved feeding cholesterol to 32 japanese rabbits for a period of 8 weeks. The rabbits were split into four groups and it was demonstrated that atherosclerotic development was significantly inhibited in the shiitake consuming rabbits.¹⁹

Research has identified eritadenine as most likely the active component of shiitake mushrooms which facilitates a hypocholesterolemic effect in the body.²⁰⁻²² The exact mechanisms of action are still not clearly understood and more research must be conducted.

Nattokinase is an extremely effective anti-thrombotic and antioxidant.

Fish

The Japanese diet consists of a large component of fish. *Dashi*, or stock used to make soups, is often made with small sardines or bonito fish flakes. *Sashimi* sushi, or raw fish sushi, is enjoyed over the world.

There has been an abundant amount of research and investigations into the relationship of fish consumption with lowered incidences of CVD. Evidence continues to suggest that regular consumption of fish, high in omega-3 oil, reduces the risk of CVD and can even be effective as a therapeutic treatment protocol.²⁴

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One important study surveyed 672 men and 676 women in Japan and Hawaii and was conducted over three years. The *INTERLIPID* study concluded regular fish consumption decreased CVD and mortality in the participants.³¹

Leptin, a hormone involved in the suppression of appetite²⁵, is considered to be involved in the development of CVD.²⁶⁻²⁸ A population eating a diet rich in marine fish was shown to have a significant low level of plasma leptin.²⁹ A further clinical trial of 69 overweight, hypertensive men and women were treated with a daily component of fish and exercise. The results again indicated a significant decrease in serum leptin in addition to weight loss.³⁰

The Australian Heart foundation recommends individuals should eat a 150g serve of fish two to three times a week in order to achieve an average recommended consumption of 500mg of combined docosahexaenoic acid (DHA) and eicosapentaenoic acid (EPA) a day.²³ For individuals with documented CVD it is recommended that the consumption of EPA and DHA is doubled.²³



Considerations

Salt

Hypertension is considered one of the most common causes of CVD and in a large number of cases it can be minimised through a reduced intake of salt.³³⁻³⁵ The relationship of salt intake and hypertension has long been understood and a reason for the development of the Dietary Approaches to Stop Hypertension (DASH) diet.³⁶

Some Japanese food can contain a high content of salt and should therefore eaten in moderation. Miso, Kombu seaweed, salted seaweed sushi sheets and salted fish can include significantly high levels of sodium and should therefore be eaten in moderation.

The National Heart Foundation of Australia gives recommendations for salt intake at 2300 mg a day for the general population and 1550 mg a day for individuals at risk of CVD.³⁷



Fish contaminants

Fish can contain varying levels of contaminants such as mercury, dioxins or Polychlorinated biphenyls (PCBs).^{38, 42-43} The most significant contaminant in fish is methylmercury (MeHg) which is absorbed from water and bound to proteins within the fish.³⁸ Large fish such as swordfish or sharks predominately contain higher contaminants although exact levels vary between species and environment.³⁹

One study in Japan examined mercury levels in individuals to compare male and females consuming moderate amounts of fish. Hair samples were taken from over 3600 individuals, spanning five prefectures through Japan and tested specifically for mercury levels. The data indicated a direct correlation to the amount of fish consumption and mercury contamination.⁴⁰

PCBs and dioxin are toxic organic chemicals often found in fish⁴⁶, accumulated through the food chain and agricultural contamination.⁴²⁻⁴³ Similar to MeHg, organic chemicals found in fish are toxic to individuals eating them, especially pregnant women or children.⁴¹ PCB content was recorded higher in farmed Atlantic salmon compared with wild Atlantic salmon in one study and was contributed to feed contamination.⁴⁴ A later study of 58 individuals examined PCBs and other dioxins in a range of farmed Atlantic salmon which had been given different fish feeds. Plasma samples were examined from the participants and results indicated higher levels of PCBs and other dioxins in those that ate fish fed on marine-based oil foods.⁴⁵

Besides the potential for contaminants, fish is still considered essential for cardiovascular health.⁴⁷ FSANZ suggests that to avoid excessive exposure to contaminants, high risk fish should only be consumed occasionally. It is recommended that swordfish, shark, marlin and broadbill should be consumed no more than one serve per fortnight.⁴⁸ Orange roughy and catfish (imported basa) should be consumed no more than once a week.⁴⁸

Because these varieties of fish contain such high levels of contaminants, FSANZ recommends that no other fish should be consumed in the same period.⁴⁸



References

¹Australian Institute of Health and Welfare. (2008) *Cardiovascular Disease in Australia* (online) available <<http://www.aihw.gov.au/cvd/index.cfm>> accessed on September 3, 2009

²Australian Institute of Health and Welfare. (2008) *Australia's Health 2009* (online), pg 182, available <<http://www.aihw.gov.au/cvd/index.cfm>> accessed on September 3, 2009

³Shimazu, T., Kuriyama, S., Hozawa, A., Ohmori, K., Sato, Y., Nakaya, N., et al. (2007). *Dietary patterns and cardiovascular disease mortality in Japan: a prospective cohort study*. *International Journal Of Epidemiology*, 36(3), 600-609. Retrieved September 18, 2009, from MEDLINE database.

⁴Imai, K., & Nakachi, K. (1995). *Cross sectional study of effects of drinking green tea on cardiovascular and liver diseases*. *BMJ (Clinical Research Ed.)*, 310(6981), 693-696. Retrieved September 18, 2009, from MEDLINE database

⁵Kuriyama, S., Shimazu, T., Ohmori, K., Kikuchi, N., Nakaya, N., Nishino, Y., et al. (2006). *Green tea consumption and mortality due to cardiovascular disease, cancer, and all causes in Japan: the Ohsaki study*. *JAMA: The Journal Of The American Medical Association*, 296(10), 1255-1265. Retrieved September 18, 2009, from MEDLINE database

If you require the other references, please ask Shannon Burford.

- ⁶Wolfram, S. (2007). *Effects of green tea and EGCG on cardiovascular and metabolic health*. Journal Of The American College Of Nutrition, 26(4), 373S-388s. Retrieved September 18, 2008, from MEDLINE database(abstract)
- ⁷Nagao, T., Hase, T., & Tokimitsu, I. (2007). *A green tea extract high in catechins reduces body fat and cardiovascular risks in humans*. Obesity (Silver Spring, Md.), 15(6), 1473-1483. Retrieved October 18, 2008, from MEDLINE database. (abstract)
- ⁸Matsuyama, T., Tanaka, Y., Kamimaki, I., Nagao, T., & Tokimitsu, I. (2008). *Catechin safely improved higher levels of fatness, blood pressure, and cholesterol in children*. Obesity (Silver Spring, Md.), 16(6), 1338-1348. Retrieved September 18, 2008, from MEDLINE database. (abstract)
- ⁹Ahn, H., Xu, Y., & Davidge, S. (2008). *Epigallocatechin-3-O-gallate inhibits TNFalpha-induced monocyte chemotactic protein-1 production from vascular endothelial cells*. Life Sciences, 82(17-18), 964-968. Retrieved October 18, 2008, from MEDLINE database. (abstract)
- ¹⁰Fujita, M., Nomura, K., Hong, K., Ito, Y., Asada, A., & Nishimuro, S. (1993). *Purification and characterization of a strong fibrinolytic enzyme (nattokinase) in the vegetable cheese natto, a popular soybean fermented food in Japan*. Biochemical And Biophysical Research Communications, 197(3), 1340-1347. Retrieved October 19, 2008, from MEDLINE database(abstract)
- ¹¹Pais, E., Alexy, T., Holsworth, R., & Meiselman, H. (2006). *Effects of nattokinase, a pro-fibrinolytic enzyme, on red blood cell aggregation and whole blood viscosity*. Clinical Hemorheology And Microcirculation, 35(1-2), 139-142. Retrieved October 17, 2008, from MEDLINE database.
- ¹²Yamashita, T., Oda, E., Giddings, J., & Yamamoto, J. (2003). *The effect of dietary bacillus natto productive protein on in vivo endogenous thrombolysis* Pathophysiology Of Haemostasis And Thrombosis, 33. (3), 138-143. Retrieved August 9, 2008, from MEDLINE database (abstract)
- ¹³Suzuki, Y., Kondo, K., Ichise, H., Tsukamoto, Y., Urano, T., & Umemura, K. (2003). *Dietary supplementation with fermented soybeans suppresses intimal thickening*. Nutrition (Burbank, Los Angeles County, Calif.), 19(3), 261-264. Retrieved September 17, 2008, from MEDLINE database.
- ¹⁴Peng Y, Yang X, Zhang Y (2005) *Microbial fibrinolytic enzymes: an overview of source, production, properties, and thrombolytic activity in vivo*, Applied Microbiology And Biotechnology [Appl Microbiol Biotechnol] Nov; Vol. 69 (2), pp. 126-32.Nov 12. Retrieved September 17, 2008, from MEDLINE database
- ¹⁵Schurgers, L., Teunissen, K., Hamulyák, K., Knapen, M., Vik, H., & Vermeer, C. (2007). *Vitamin K-containing dietary supplements: comparison of synthetic vitamin K1 and natto-derived menaquinone-7*. Blood, 109(8), 3279-3283. Retrieved August 19, 2008, from MEDLINE database (abstract)
- ¹⁶Iwai, K., Nakaya, N., Kawasaki, Y., & Matsue, H. (2002). *Inhibitory effect of natto, a kind of fermented soybeans, on LDL oxidation in vitro*. Journal Of Agricultural And Food Chemistry, 50(12), 3592-3596. Retrieved September 19, 2008, from MEDLINE database. (abstract)
- ¹⁷Shimada Y; Morita T; Sugiyama K (2002) *Effects of Lentinus edodes on fatty acid and molecular species profiles of phosphatidylcholine in rats fed different levels of corn oil*. Bioscience, Biotechnology, And Biochemistry Aug; Vol. 66 (8), pp. 1759-63. Retrieved October 17, 2008, from MEDLINE database (abstract)
- ¹⁸Fukushima, M., Ohashi, T., Fujiwara, Y., Sonoyama, K., & Nakano, M. (2001). *Cholesterol-lowering effects of maitake (Grifola frondosa) fiber, shiitake (Lentinus edodes) fiber, and enokitake (Flammulina velutipes) fiber in rats*. Experimental Biology And Medicine (Maywood, N.J.), 226(8), 758-765. Retrieved August 17, 2008, from MEDLINE database(abstract)

- ¹⁹Yamada T; Oinuma T; Niihashi M; Mitsumata M; Fujioka T; Hasegawa K; Nagaoka H; Itakura H (2002) *Effects of Lentinus edodes mycelia on dietary-induced atherosclerotic involvement in rabbit aorta*. Journal Of Atherosclerosis And Thrombosis [J Atheroscler Thromb] 2002; Vol. 9 (3), pp. 149-56. Retrieved August 7, 2008, from MEDLINE database
- ²⁰Enman, J., Rova, U., & Berglund, K. (2007). Quantification of the bioactive compound eritadenine in selected strains of shiitake mushroom (*Lentinus edodes*). *Journal Of Agricultural And Food Chemistry*, 55(4), 1177-1180. Retrieved October 17, 2008, from MEDLINE database
- ²¹Sugiyama, K., Akachi, T., & Yamakawa, A. (1995). *Hypocholesterolemic action of eritadenine is mediated by a modification of hepatic phospholipid metabolism in rats*. The Journal Of Nutrition, 125(8), 2134-2144. Retrieved September 19, 2008, from MEDLINE database.
- ²²Shimada, Y., Morita, T., & Sugiyama, K. (2002). *Effects of dietary eritadenine on delta6-desaturase activity and fatty acid profiles of several lipids in rats fed different fats*. Bioscience, Biotechnology, And Biochemistry, 66(7), 1605-1609. Retrieved August 22, 2008, from MEDLINE database (abstract)
- ²³Colquhoun, D., Ferreira-Jardim, A., Udell, T., Eden, B. (2008) *Fish, fish oils, n-3 polyunsaturated fatty acids and cardiovascular health*, Heart Foundation, Nutrition and Metabolism Committee of the Heart Foundation, pg 4, available <http://www.heartfoundation.org.au/document/NHF/HW_FS_FishOil_RevEv_FINAL.pdf> accessed September 24, 2008
- ²⁴Mozaffarian D and Rimm EB (2006) *Fish intake, contaminants, and human health: evaluating the risks and the benefits*. JAMA. 296: 1885-1899. Retrieved August 20, 2008, from MEDLINE database (abstract)
- ²⁵Tortora, G., Grabowski, S. (2003) Principles of Anatomy & Physiology, Wiley International Edition, USA
- ²⁶Söderberg, S., Ahrén, B., Jansson, J., Johnson, O., Hallmans, G., Asplund, K., et al. (1999). *Leptin is associated with increased risk of myocardial infarction*. Journal Of Internal Medicine, 246(4), 409-418. Retrieved October 20, 2008, from MEDLINE database.
- ²⁷Wallace, A., McMahon, A., Packard, C., Kelly, A., Shepherd, J., Gaw, A., Sattar, N. (2001) *Plasma leptin and the risk of cardiovascular disease in the west of Scotland coronary prevention study*, Circulation, 104: 3052-6 Available <<http://circ.ahajournals.org/>> accessed September 12, 2008
- ²⁸Ren, J. (2004). *Leptin and hyperleptinemia - from friend to foe for cardiovascular function*. The Journal Of Endocrinology, 181(1), 1-10. Retrieved October 2, 2008, from MEDLINE database (abstract)
- ²⁹Winnicki, M., Somers, V., Accurso, V., Phillips, B., Puato, M., Palatini, P., Pauletto, P. (2002), *Fish-rich diet, leptin, and body mass*. Circulation. 106: 289-91, 2002. Available <<http://circ.ahajournals.org/>> accessed September 12, 2008
- ³⁰Mori, T., Burke, V., Puddey, I., Shaw, J., & Beilin, L. (2004). *Effect of fish diets and weight loss on serum leptin concentration in overweight, treated-hypertensive subjects*. Journal Of Hypertension, 22(10), 1983-1990. Retrieved September 12, 2008, from MEDLINE database (abstract)
- ³¹Okuda, N., Ueshima, H., Okayama, A., Saitoh, S., Nakagawa, H., Rodriguez, B., et al. (2005). *Relation of long chain n-3 polyunsaturated fatty acid intake to serum high density lipoprotein cholesterol among Japanese men in Japan and Japanese-American men in Hawaii: the INTERLIPID study*. Atherosclerosis, 178(2), 371-379. Retrieved September 20, 2008, from MEDLINE database (abstract)
- ³²He, K., Liu, K., Daviglius, M., Mayer-Davis, E., Jenny, N., Jiang, R., et al. (2008). *Intakes of long-chain n-3 polyunsaturated fatty acids and fish in relation to measurements of subclinical atherosclerosis*. The American Journal Of Clinical Nutrition, 88(4), 1111-1118. Retrieved September 20, 2008, from MEDLINE database (abstract)

- ³³He, F., & MacGregor, G. (2002). *Effect of modest salt reduction on blood pressure: a meta-analysis of randomized trials. Implications for public health.* *Journal Of Human Hypertension*, 16(11), 761-770. Retrieved September 11, 2008, from MEDLINE database (abstract)
- ³⁴Midgley, J., Matthew, A., Greenwood, C., & Logan, A. (1996). *Effect of reduced dietary sodium on blood pressure: a meta-analysis of randomized controlled trials.* *JAMA: The Journal Of The American Medical Association*, 275(20), 1590-1597. Retrieved September 11, 2008, from MEDLINE database(abstract)
- ³⁵Hooper, L., Bartlett, C., Davey Smith, G., & Ebrahim, S. (2002). *Systematic review of long term effects of advice to reduce dietary salt in adults.* *BMJ (Clinical Research Ed.)*, 325(7365), 628-628. Retrieved September 11, 2008, from MEDLINE database(abstract)
- ³⁶Sacks, F., Svetkey, L., Vollmer, W., Appel, L., Bray, G., Harsha, D., et al. (2001). *Effects on blood pressure of reduced dietary sodium and the Dietary Approaches to Stop Hypertension (DASH) diet.* *DASH-Sodium Collaborative Research Group.* *The New England Journal Of Medicine*, 344(1), 3-10. Retrieved September 11, 2008, from MEDLINE database.(abstract)
- ³⁷National Heart Foundation of Australia (2008) Position statement on the relationships between dietary electrolytes and cardiovascular disease (online), NHFA, available < http://www.heartfoundation.org.au/document/NHF/NHFA_DietaryElectrolytes_CVD_PositionStatement_2006-07_FINAL.pdf> accessed September 11, 2008
- ³⁸National Heart Foundation of Australia (2008)*Fish, fish oils, n-3 polyunsaturated fatty acids and cardiovascular health* (online), NHFA, available < http://www.heartfoundation.org.au/document/NHF/HW_FS_FishOil_RevEv_FINAL.pdf> accessed September 11, 2008
- ³⁹Kales, S., & Goldman, R. (2002). *Mercury exposure: current concepts, controversies, and a clinic's experience.* *Journal Of Occupational And Environmental Medicine / American College Of Occupational And Environmental Medicine*, 44(2), 143-154. Retrieved September 21, 2008, from MEDLINE database.(abstract)
- ⁴⁰Yasutake, A., Matsumoto, M., Yamaguchi, M., & Hachiya, N. (2003). *Current hair mercury levels in Japanese: survey in five districts.* *The Tohoku Journal Of Experimental Medicine*, 199(3), 161-169. Retrieved September 21, 2008, from MEDLINE database.(abstract)
- ⁴¹Schantz, S., Widholm, J., & Rice, D. (2003). *Effects of PCB exposure on neuropsychological function in children.* *Environmental Health Perspectives*, 111(3), 357-576. Retrieved October 21, 2008, from MEDLINE database(abstract)
- ⁴²Bhavsar, S., Fletcher, R., Hayton, A., Reiner, E., & Jackson, D. (2007). *Composition of dioxin-like PCBs in fish: an application for risk assessment.* *Environmental Science & Technology*, 41(9), 3096-3102. Retrieved September 19, 2008, from MEDLINE database.(abstract)
- ⁴³Bhavsar, S., Hayton, A., & Jackson, D. (2008). *Uncertainty analysis of dioxin-like polychlorinated biphenyls-related toxic equivalents in fish.* *Environmental Toxicology And Chemistry / SETAC*, 27(4), 997-1005. Retrieved September 19, 2008, from MEDLINE database.(abstract)
- ⁴⁴Easton, M., Luszniak, D., & Von der, G. (2002). *Preliminary examination of contaminant loadings in farmed salmon, wild salmon and commercial salmon feed.* *Chemosphere*, 46(7), 1053-1074. Retrieved September 19, 2008, from MEDLINE database.(abstract)
- ⁴⁵Bethune, C., Seierstad, S., Seljeflot, I., Johansen, O., Arnesen, H., Meltzer, H., et al. (2006). *Dietary intake of differently fed salmon: a preliminary study on contaminants.* *European Journal Of Clinical Investigation*, 36(3), 193-201. Retrieved September 19, 2008, from MEDLINE database.(abstract)

⁴⁶Hites, R., Foran, J., Carpenter, D., Hamilton, M., Knuth, B., & Schwager, S. (2004). *Global assessment of organic contaminants in farmed salmon*. Science (New York, N.Y.), 303(5655), 226-229. Retrieved September 11, 2008, from MEDLINE database.(abstract)

⁴⁷Mozaffarian, D., & Rimm, E. (2006). *Fish intake, contaminants, and human health: evaluating the risks and the benefits*. JAMA: The Journal Of The American Medical Association, 296(15), 1885-1899.

⁴⁸Food Standards Australia New Zealand (2004) *Mercury in fish: fact sheet*. Available <www.foodstandards.gov.au/_srcfiles/FS_Mercury_in_fish_final.pdf> accessed September 1, 2008