



## FOOD SCIENCES

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NEW YORK STATE AGRICULTURAL EXPERIMENT STATION, GENEVA, A DIVISION OF THE NEW YORK STATE COLLEGE OF AGRICULTURE AND LIFE SCIENCES, A STATUTORY COLLEGE OF THE STATE UNIVERSITY, CORNELL UNIVERSITY, ITHACA

# Dietary vegetables and environmental health

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## INTRODUCTION

In 1941 Kensler, et al observed that the B-vitamin riboflavin protected rats against the potent hepatocarcinogen aminoazo dyes (1). Ten years later Mueller and Miller showed that this vitamin enzymatically cleaved these dyes to noncarcinogenic metabolites in the liver (2). Further studies in the laboratory of Miller showed enhanced hepatic metabolism of these dyes when rats were fed a practical ingredient type of diet as compared to a more purified diet composed of casein, sugar, corn oil, cellulose, vitamins, and minerals (3).

These pioneering studies on the influence of nutritional and dietary factors and the metabolism of dyes revealed that changes in diets can have dramatic effects on the metabolism and action of foreign chemicals. Extension of these studies by many investigators have shown that changes in specific foods, as well as specific nutrients, can have an effect on multifunctional tissue enzymes collectively known as microsomal mono-oxygenases, or mixed-function oxidases (MFO) (4,5,6). These enzymes are induced by many foreign chemicals including drugs, environmental carcinogens etc. Induction of these enzymes indicate that increased metabolism of exogenous chemicals are occurring. However, this increased metabolism could mean either the chemicals are being detoxified for rapid excretion, or being metabolized to more potent metabolites known as the "ultimate carcinogen" (7).

A number of common vegetables in the botanical family *Cruciferae*, e.g. cabbage, brussels sprouts, cauliflower, kale, etc. have been observed to stimulate increased MFO activity in the intestines of experimental animals (8,9). Work in our laboratory (10) showed that this increased activity also occurred in the liver of rodents. This report is a brief

overview of some recent investigation of ours (11) showing that experimental diets containing cauliflower reduces the hepatic toxicities of two chemically dissimilar food contaminants, i.e. the flame retardant, polybrominated biphenyls (PBB's) and aflatoxin B1 (AFB1). PBB's have contaminated meat and milk, especially in Michigan, and AFB1 is a potent carcinogenic metabolite of the mold *Aspergillus flavus* observed growing in peanuts, grains, and other foods stored under relatively high moisture conditions.

## RESULTS

PBB is not an acutely toxic chemical. However, feeding rats diets containing 50 ppm of PBB's for 3 weeks produced fatty livers, as well as increased residues of the compound in this organ. Male rats fed diets containing 25 per cent freeze-dried cauliflower exhibited markedly reduced levels of liver fat, as well as residual liver PBB depression of approximately 50 per cent.

Fischer rats fed 2 ppm AFB1 for 26 weeks exhibited 100 per cent mortality with large hepatocarcinomas by the end of 41 weeks when consuming a purified diet. Animals fed this level of AFB1, combined with a 25 per cent cauliflower diet all survived. However, these animals did show lowered body weights and small grade I and grade II adenomas.

*In-vitro* experiments on the tissues of cauliflower fed rats indicated that MFO activity was moderately activated with increased enzyme-substrate reaction rates in the liver as well as in the intestine and kidney.

## CONCLUSIONS

These data indicate that dietary cauliflower inhibited the

toxic responses, including carcinogenicity, of two dissimilar chemical contaminants. Various sulfur containing compounds, known as glucosinolates, are naturally present in relatively high quantities in *Cruciferae* vegetables. Metabolite products of glucosinolates, indoles, have been attributed to inducing MFO activity in animals (12). Investigations in our laboratory have shown that indoles can stimulate liver MFO activity, but apparently only at exceedingly high dietary levels (13). Recent 10-year epidemiological studies with over 120,000 Japanese men showed a significantly lowered death rate due to prostate cancer in men consuming "green-yellow vegetables" as compared with men rarely consuming these vegetables (14).

The mechanisms for this cancer inhibitory effect of dietary vegetables appears to be involved with tissue MFO induction. However, naturally-occurring vitamins, fiber, minerals, or non-nutrients may be directly, or indirectly responsible for this effect. It appears crucial to know if other common dietary vegetables besides *Cruciferae* exert similar effects.

#### REFERENCES

1. Kensler, C. J., K. Sugiura, N. F. Young, C. R. Halter and C. P. Rhoads. Partial protection of rats by riboflavin with casein against liver cancer by dimethylaminoazobenzene. *Science* 93, 308-310. 1941.
2. Mueller, G. C. and J. A. Miller. The reductive cleavage of 4-dimethyl-aminoazobenzene by rat liver. Reactivation of carbon dioxide-treated homogenates by riboflavin-adenine dinucleotide. *J. Biol. Chem.* 185, 145-154. 1950.
3. Brown, R. R., J. A. Miller, and E. C. Miller. The metabolism of methylated aminoazo dyes. IV. Dietary factors enhancing demethylation in vitro. *J. Biol. Chem.* 209, 211-222. 1954.
4. Campbell, T. C. Nutrition and drug metabolizing enzymes. *Clin. Pharmacol. and Ther.* 22, 699-706. 1977.
5. Becking, G. C. Hepatic drug metabolism in iron-, magnesium- and potassium-deficient rats. *Fed. Proc.* 35, 2480-2485. 1976.
6. Zannoni, V. G. and P. H. Sato. The effect of certain vitamin deficiencies on hepatic drug metabolism. *Fed. Proc.* 35, 2464-2469. 1976.
7. Parke, D. V. Induction of the Drug-Metabolizing Enzymes. In: *Enzyme Induction* (ed. D. V. Parke), pp. 207-271. Plenum Press, London. 1975.
8. Wattenberg, L. W. Studies of polycyclic hydrocarbon hydroxylases of the intestine possibly related to cancer. Effect of diet on benzpyrene hydroxylase activity. *Cancer* 28, 99-102. 1971.
9. Wattenberg, L. W. Enzymatic reactions and carcinogenesis. In: *Environment and Cancer* (ed. R. D. Cumley) pp. 241-255. Williams and Wilkins Co., Baltimore. 1972.
10. Babish, J. G. and G. S. Stoewsand. Hepatic microsomal enzyme induction in rats fed varietal cauliflower leaves. *J. Nur.* 105, 1592-1599. 1972.
11. Stoewsand, G. S., J. B. Babish, and H. C. Wimberly. Inhibition of hepatic toxicities from polybrominated biphenyls and aflatoxin B<sub>1</sub> in rats fed cauliflower. *J. Environ. Path. and Toxicol.* 2, 399-406. 1978.
12. Loub, W. B., L. W. Wattenberg, and D. W. Davis. Aryl hydrocarbon hydroxylase induction of rat tissues by naturally occurring indoles of cruciferous plants. *J. Natn. Cancer Inst.* 54, 985-989. 1975.
13. Babish, J. G. and G. S. Stoewsand. Effect of dietary indole-3-carbinol on the induction of the mixed-function oxidases of rat tissue. *Fd. and Cosmet. Toxicol.* 16, 151-155. 1978.
14. Hirayama, T. Epidemiology of prostatic cancer with special reference to the risk lowering effect of green-yellow vegetable intake. *Proc. 37th Ann. Meeting of the Japanese Cancer Assn., Tokyo, Japan. August, 1978*



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