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Preservatives and the Link with Adverse Health Outcomes

Due to their ability to extend shelf life and prevent spoilage, synthetic and industrialized

preservatives are common in U.S. processed foods and drinks—both on the grocers' shelves and at

conventional and fast food restaurants. Unfortunately, despite their ability to retard spoilage a number

of the most commonly used preservatives have been linked in clinical trials and empirical studies to

potential adverse health outcomes for some consumers and several have been shown in scientific

studies to be a potential health threat for the general population.

Sodium Benzoate

Empirical studies and clinical trials indicate sodium benzoate is linked in sensitive individuals to

skin reactions including urticaria, pruritus and atopic dermatitis, and intestinal disturbances such as

gastritis, nasal polyps, rhinitis, migraine headaches and arthralgia, shortness of breath,

broncoconstriction, asthma, and with behavioral, mood and psychiatric disorders. Clinical and

consumer reports indicate this additive may cause adverse reactions for people with chemical

sensitivities (including people with asthma and other respiratory conditions) and allergies (including to

polyethylene glycol); symptoms may include skin reactions (including rash/urticaria), headaches, breathing problems, swelling of face, throat, tongue, hands, and feet, swollen/painful lymph nodes, earaches, sinus problems, edema and subsequent weight gain, digestive problems, rhinitis, difficulty concentrating, insomnia, anxiousness, hyperactivity, and depression, to name a few. This additive is commonly used as an antibacterial and antifungal preservative (as well as flavor enhancer) in a wide variety of processed foods including soft drinks, fruit juices, milk and milk products, cheeses, margarine, meat products, maple syrup, relishes, pickles, jarred peppers, bottled lemon juice, salad dressings, sauces, mixes, gravies, soups, condiments, flour, baked goods, confectionery and snacks. Sodium benzoate may also be present in conventional and fast food restaurants and can also be found in toothpastes, mouth washes, and other personal care products, as well as prescription and over the counter drugs. While food items containing sodium benzoate must list its presence, there are no required warnings for consumers.

Gallates (Propyl Gallate, Dodecyl Gallate, Octyl Gallate, and EGCG)

Gallates, like industrialized phosphates, have the distinction of not only being linked to adverse symptoms for sensitive groups, but also having the potential to harm well-being for the larger consumer population base as well. Propyl gallate is commonly used as a preservative in a variety of processed foods, and may be present with other preservatives such as BHT/BHA (two other food chemicals of concern) and has been linked with adverse reactions such as respiratory problems (especially for people with asthma), stomach irritation, swelling, weight gain, adverse skin reactions, insomnia/sleep disturbances, behavioral/mood disturbances, and more. Those people with chemical sensitivities (including to aspirin), people with asthma and other respiratory ailments, as well as those with liver and

kidney problems may be advised by health care professionals to avoid this additive. Additionally, chellitis (dry, scaly, cracked lips) and stomatis (oral inflammation/inflammation of mucous lining of the mouth including cheeks, gums, tongue, lips, throat, etc.) have been reported by some consumers following exposure to propyl gallate. Diagnosis is frequently complicated by the fact that adverse reactions to gallates may be delayed for as long as 5 days. On an even more serious note, gallates have been linked in animal studies to anemia, as well as changes in the kidney and liver. Additionally, recent study findings have linked the food additive propyl gallate with hormonal disruptions in women. Propyl gallate is also a possible carcinogen. This additive can be found in a wide range of processed, packaged, commercially prepared meals, dried milk, dried meats, sauces, snacks (especially potatobased), sports drinks, vegetable oils and dressings, soups (especially chicken), condiments and chewing gum.

Dodecyl gallate is a synthetic antioxidant used as a preservative in processed foods and commonly found in salad dressings, margarine, oils, fats/lard, rice, bakery goods, cake mixes, dehydrated meats, soups, potatoes and broths, candy, cereals, processed nuts, spices, condiments, and a variety of snack foods, etc. Dodecyl gallate may cause allergic reactions in some people, especially swelling of the lips, dermatitis, chronic eczema or ulcerations. Chellitis (dry, scaly, cracked lips) and stomatis (oral inflammation/inflammation of mucous lining of the mouth including cheeks, gums, tongue, lips, throat, etc.) have been reported with exposure to dodecyl gallate.

Octyl gallate is commonly used as a preservative (and synergistic antioxidant with other preservatives, see: BHA/BHT) and can also be used as a bulking agent, thickener or emulsifier in processed foods such as peanut butter, margarine, bakery products, cake mixes, candy, cereals, processed nuts, processed and dehydrated meats, soups and broths, enriched rice, sauces, dips, seasonings, condiments, and snack foods. Octyl gallate may also be present in food packaging materials. This additive can cause allergic reactions in some people, sometimes severe. People with

asthma, or sensitivity to aspirin are frequently warned by health care professionals to avoid foods containing this additive. Octyl gallate may cause digestive/gastric problems in some people and can cause dermatitis in sensitive individuals. Reported adverse skin reactions to octyl gallate include redness, swelling, itching and fluid-filled blisters. Chellitis (dry, scaly, cracked lips) and stomatis (oral inflammation/inflammation of mucous lining of the mouth including cheeks, gums, tongue, lips, throat, etc.) have also been reported with exposure to octyl gallate. This additive is not recommended for infants or children.

# Parabens

Parabens are highly controversial due to scientific study findings suggesting a link with possible carcinogenicity and estrogenic and reproductive effects. Because the majority of news reports on parabens have focused on their presence in personal care products many people are unaware that they are also used as food preservatives. The industrialized version of this naturally occurring chemical is widely used as a preservative in processed foods including soft drinks, juices, baked goods, jellies and jams, candies, salad dressings, mayonnaise, mustard, sauces, marinated fish products, foods containing processed vegetables, frozen dairy products, and snack food, to name a few. Though rare, some people may develop hypersensitivity or allergic reactions to this chemical when ingested as a food preservative. Symptoms may include swelling of lips, face, tongue and throat, difficulty swallowing, breathing difficulties, dermatitis, skin sensitivity, rash, itching, edema.

Other common food additives frequently used as preservatives such as EDTA (Ethylenediamine Tetraacetic Acid/Tetra-acetate) have been linked with allergic reactions in some people and TBHQ (Tert-Butylhydroquinone) has been linked with allergic reactions and possible toxicity. EDTA in particular may worsen conditions for people with asthma, heart rhythm problems, kidney disease, epilepsy, liver disease, hepatitis, tuberculosis, allergies, diabetes, and electrolyte imbalance, including hypomagnesemia (low magnesium levels), hypocalcemia (low calcium levels), hypokalemia (low potassium levels), and people with these health conditions or taking certain medications may be advised by health care professionals to avoid processed foods containing this additive. Reports of adverse reactions to this additive include allergic reactions, respiratory problems, skin reactions (including rash, redness, urticaria), weight gain, edema, swelling of face, hands, feet, tongue, excessive thirst, insomnia, and headaches, among others.

EDTA is a salt produced from synthesizing ethylenediamine, formaldehyde and sodium cyanide; it is used as a fortifier, texture enhancing agent, and food preservative and is commonly found in a wide variety of commercially-processed foods and restaurant foods. It can be found in breakfast cereals and breakfast bars, salad dressing, sauces, dips, toppings, canned fish and seafood (including shrimp, crab meat, etc.), canned beans, processed vegetables, including pickled cabbage and pickled cucumbers, canned potatoes (and products such as soups containing white potatoes), sandwich spreads, canned gravy, canned mushrooms, potato salad, processed fruit, bakery goods, margarine, some alcoholic/malt beverages, and soft drinks, among many other items.

Reported adverse effects associated with the additive TBHQ include headaches, stomach/digestive problems, allergic reactions, ringing in ears, edema/swelling of face, feet, hands, ADHD, skin reactions/dermatitis, urticaria, angioedema, sleepiness, rhinitis, sinus problems, breathing difficulties,

asthma, allergies, vasculitis, flushing, excessive sweating, vision/eye problems, and joint pain. Some researchers have argued that chronic exposure to this additive may induce carcinogenicity. TBHQ is a petroleum-based chemical used as a preservative for vegetable oils and animal fats. It is commonly found in packaged breads and baked goods products, breakfast cereals, frozen waffles, toaster pastries, snack crackers, conventional and fast food restaurant items, and baked goods in grocery store bakery sections, among others.

### Nitrates/Nitrites

Nitrates/Nitrites is a possible carcinogen and has been linked with the formation of carcinogenic nitrosamines, bladder cancer, chronic liver disease, as well as respiratory, skin and intestinal disturbances, and Alzheimer's disease, diabetes mellitus and Parkinson's disease. Adverse reactions may include headaches, migraines, swelling, weight gain, and respiratory problems (including asthma problems). Research has linked nitrates/nitrites to bladder, colon and pancreatic cancers. It has also been linked to leukemia. Sensitive individuals (especially those with Alzheimer's disease, Diabetes, Blood Disorders, Parkinson's disease, and Thyroid Disease) and pregnant women may be advised by health care professionals to avoid or restrict consumption of processed foods containing nitrates/nitrites. Not to be confused with nitrates/nitrites which occur naturally in vegetables and fruits, industrialized versions of these preservatives are commonly added to processed lunch meats, hot dogs, sausages, bacon, ham, pork, jerky, deli meats, and other food items containing processed meat items (frequently listed on ingredients labels as "sodium nitrite", "potassium nitrite", etc.). As with many of

the food additives found in scientific studies and clinical trials to be potentially problematic for consumers, there are no required warnings on food ingredients labels.

# Sulfites

Sulfites have been linked in scientific studies and clinical trials with several idiosyncratic and allergic reactions including respiratory tract irritation, bronchospasm, oculonasal symptoms, skin reactions including urticaria and angioedema, flushing, hypotension and intestinal disturbances, especially in sulfite-sensitive individuals and those with asthma. Reported reactions to this additive range from mild to severe and even life threatening, especially for those people with sulfite allergies or asthma (where it may provoke life-threatening asthmatic complications), bronchitis, or other respiratory illnesses, as well as for those individuals with certain medical conditions (such as liver enzyme deficiencies) or sensitivities to food chemicals. Reported symptoms include asthmatic reactions/lung irritation/difficulty breathing, swelling of face, throat, tongue, difficulty swallowing, excessive thirst, edema (and subsequent weight gain), swelling of hands, feet, legs, vision problems, headaches, earaches, elevated blood pressure, skin reactions such as urticaria/rash/hives, insomnia, nervousness, anxiousness, difficulty concentrating, painful, swollen lymph nodes, and anaphylaxis.

Sulfites are commonly used as a preservative against spoilage and color changes, and as a bleaching agent and dough conditioner, and may be present at varying levels in a variety of processed foods including dried/dehydrated fruits (such as raisins and apricots), dehydrated/dried and pickled vegetables, prepared toppings, dips (including processed avocado/guacamole), gravies and sauces, condiments, olives, pepperoncini, pickles, (including relish/pickled peppers/horseradish, etc.), salad

dressings and mixes, processed cheese mixtures and pastes, soy sauce, tomato paste and purée, molasses, maple syrup, brown sugar, tea, wine, beer, sparkling grape juice, wine vinegar, malt vinegar, fruit juices, soft drinks, cocktail mixes, cider/cider vinegar, canned vegetables and fruit (including potatoes, mushrooms, tomatoes, hominy grits), fruit pie filling, canned sauerkraut, corn starch (and processed foods containing corn starch, cornmeal, corn syrup (and processed foods containing corn syrup), baked goods (including pastries, cookies and crackers), frosting, dessert toppings, breading mixes, dried noodle and rice mixes, cereals, frozen pizza, frozen pizza dough/pie dough/cookie dough, high fructose corn syrup (and processed foods containing HFCS), trail mix, jams/jellies, bottled lemon juice/lime juice, vinegars and vinegar-based sauces and salad dressings, maraschino cherries, fresh and frozen shrimp, canned tuna, canned and dried seafood, canned clams/clam chowder, sausages/processed meats, foods containing pectin, beet sugars, or textured vegetable protein (TVP), gelatin, canned soups and dried soup mixes, coconut toppings/flavorings/dried coconut, instant drink mixes, cookie mixes, non-organic loose bulk foods, diet foods, baby formula, packaged salads (including fruit salads, lettuce-based salads and coleslaw), and conventional, "health food", and fast food restaurant foods (including dehydrated or precut potatoes such as those used for French fries, hash browns, and other dishes containing potatoes), as well as various salad bar items.

While prohibited on fresh, raw produce sold in U.S. grocery stores, restaurants are another matter. Sulfite agents—frequently sulfur dioxide—can be present on restaurant and catered salad bar offerings, sandwich sauces and toppings, as well as any number of food ingredients that arrive to the restaurant kitchen either packaged and/or partially or fully prepared. This additive may also be present in potato chips/snack food, hard candy/caramel candy, foods and drinks with caramel coloring, pet food, and some medications.

It is important to note that several forms of sulfites exist and are allowed for use in foods: sodium sulfite, sodium bisulfite, sodium metabisulfite, potassium sulfite, potassium bisulfite, potassium metabisulfite, and sulfur dioxide.

In 1986 the FDA banned the use of sulfites on fresh produce and began requiring that foods containing more than 10 parts per million (ppm) concentration of sulfites be declared on U.S. food labels. However, there remains no requirement for explicit warnings on processed or packaged food labels of the potential dangers from sulfites. Individuals who are highly sensitive to sulfites may experience adverse reactions to foods containing even low levels of sulfites. As lower levels of sulfites are not required to appear on U.S. food labels, individuals with the potential for severe reactions to sulfites may be advised by health care professionals to avoid such processed foods altogether.

# Formaldehyde

While U.S. consumers may be hard pressed to find anything obvious about it on the label, the food additive formaldehyde is used in preserved foods, medicines and vitamins, sugar production, as a preservative for grain and seed dressings and as a disinfectant for seeds, as well as being present in the artificial sweetener Aspartame, and has been detected in sources that can leach or migrate into food such as food packaging, tableware and cooking utensils. The presence of formaldehyde in food is usually unlabeled. Studies have uncovered its presence in some sodas, packaged food, and beer. Formaldehyde often presents either as a byproduct of other synthetic food additives (such as EDTA/Disodium EDTA, etc.) or during digestive/metabolic processes as a byproduct of certain synthetic food additives such as the artificial sweetener Aspartame. Formaldehyde can occur naturally

and can be produced via metabolism of some amino acids and a variety of xenobiotics. Formaldehyde can also result from smoking/cooking foods (especially fish, seafood, and smoked ham), and it is also used/added as a preservative, biocide, disinfectant, or bacteriostatic agent in some processed food items.

Classified as a known human carcinogen (cancer-causing substance) by the International Agency for Research on Cancer (IARC) and the National Toxicology Program Report, U.S. Department of Health and Human Services, this chemical is long known to have mutagenic properties. According to a recent report from the National Toxicology Program, ingestion of food can be a significant source of exposure to formaldehyde—in addition to low levels of formaldehyde occurring naturally in a variety of foods such as fruit, food may contain small amounts of formaldehyde from its use as a fumigant, fertilizer and preservative.

It is important to note that much of the research on formaldehyde has centered on inhalation (scientific findings are based primarily on inhalation and oral exposure in experimental animal studies); epidemiological findings (from both cohort & nested case control studies) are based primarily on occupational exposure (humans working directly and indirectly with this chemical) and under these conditions it has been linked to a variety of adverse reactions including respiratory problems for people with asthma, migraine headaches, insomnia, memory loss, mood alterations, nausea, fatigue and leukemia/cancer. The FDA position on formaldehyde is that "The Food and Drug Administration (FDA) do not believe that the very low levels that are used in food and cosmetics present a significant safety concern," (Scheuplein 1985, 245).

"Although the FDA seeks to reassure us they are keeping a close watch over our food, the job of rigorously regulating thousands of food additives is simply too big for an underfunded agency. Brominated vegetable oil (BVO), for instance, the subject of a well-circulated petition by a 15-year-old in Alabama, was flagged for further study in the 1970s—testing that was never done. And BHA, a "probable carcinogen" according to the Department of Health and Human Services, is still allowed in food."

~Melanie Warner

#### BHT/BHA

The synthetic antioxidants and preservatives BHT (Butylated hydroxytoluene) and BHA (Butylated hydroxyanisole) are commonly found in breakfast cereals (where it is commonly used in packaging and leaches into the food—though two of the largest breakfast cereal producers have been working on alternatives for the past few years in an effort to phase out BHT), butter, margarine, vegetable oil, shortening, meats, grains, cake mixes, baked goods, beer, and numerous other processed foods and snacks (including potato chips and nuts), spices, and candy, as well as chewing gum, and various types of food packaging. BHT and BHA may also be present in pet food. The oxidative characteristics and/or metabolites of the commonly used preservatives BHA and BHT have been found in animal studies to contribute to carcinogenicity and tumorigenicity, as tumor promoters. It should be noted that there exist some evidence suggesting that under certain conditions these additives may also have the opposite effect in that they may be anti-carcinogenic.

Clinical reports suggest that BHT may complicate health conditions for those individuals with bladder problems such as interstitial cystitis ("painful bladder syndrome"), sensitive bladders, or who suffer from frequent bladder infections or irregularities. Other adverse effects associated with this

preservative include intestinal and digestive problems, headaches, allergic reactions, swelling of face, feet, hands, weight gain, skin reactions/dermatitis, urticaria, angioedema, sleep abnormalities, rhinitis, respiratory disturbances, asthma, allergies, vasculitis, flushing, diaphoresis (excessive sweating), vision/eye problems, and joint pain. Long-term exposure may create changes in lungs, liver, kidneys, and thyroid. Additionally, there is evidence that certain individuals may have difficulty metabolizing BHT, resulting in health and behavioral changes, respiratory, skin and intestinal disturbances. Also of note, in animal studies BHT has been linked to having a toxic effect on the lungs, having a significant adverse effect on body weight to developing fetuses and later during the lactation period, and adverse effects to adipose cells, as well as acting as a developmental neurobehavioral toxin, including triggering behavioral abnormalities in women during pregnancy, as well as to offspring (which can lead to severe deficits in learning, decreases in sleeping, as well as increases in aggression and social isolation), promoting liver abnormalities and toxicity, and promoting cancerous tumors.

BHA is now considered as "reasonably anticipated to be a human carcinogen" by the National Institutes of Health Report on Carcinogens, Public Health Service National Toxicology Program Report on Carcinogens and the U.S. Department of Health and Human Services' Report on Carcinogens (BHA), Eleventh Report on Carcinogens. BHA has been linked in animal studies as a developmental toxin, and as promoting decreases in sleeping, orientation reflex and learning. BHA has also be shown to potentially adversely affect liver and kidney functions. Additionally, like BHT, there is evidence that certain people may have difficulty metabolizing BHA, resulting in health and behavioral changes, respiratory, skin and intestinal disturbances. Clinical and consumer reports indicate other adverse effects associated with this preservative include intestinal and digestive problems, headaches, allergic reactions, swelling of face, feet, hands, skin reactions/dermatitis, urticaria, edema/angioedema, weight gain, sleep abnormalities, rhinitis, respiratory disturbances, asthma, allergies, sinus problems, vasculitis, flushing, excessive sweating, vision/eye problems, and joint pain. Citing numerous animal

studies linking BHA to serious adverse health consequences, Glenn Scott, M.D. filed a Citizens Regulatory Petition with the FDA back in 1990, asking the agency to prohibit the use of BHA in food. To date BHA continues to be used as a food additive/preservative in the U.S. While both BHA and BHT appear on the ingredients listings, there are no explicit warnings required on food products that contain these additives to alert consumers.

# BVO

Brominated Vegetable Oil (BVO) contains bromine and is used in certain processed food items to maintain consistency and prevent separation. It is commonly found in processed citrus- and fruit-flavored drinks including juices, health drinks, sports drinks and soda. Bromine is a preservative used in certain processed foods to prevent them from going stale and to extend their shelf life. Bromine is a base element from which bromide and bromate are derived. Bromine and its derivatives Bromine, Bromate, Bromide: Potassium Bromate and Brominated Vegetable Oil (BVO) may cause nausea or diarrhea in some people. Other adverse reactions may include headaches, confusion, swelling, skin problems, fatigue, memory loss, ulcers, and loss of muscle coordination. In some cases excessive intake may lead to neurological or reproductive problems. Bromates have been linked in animal studies with the development of cancerous tumors. Potassium Bromate, for example, (most commonly present in commercial breads, bread products, including fast food breads and buns, and processed baked goods) is a known toxin and carcinogen and has been linked with thyroid and kidney cancers in animal studies. This additive competes with iodine in the body and may be a concern for people with iodine deficiencies and hypothyroidism. Brominated Vegetable Oil (BVO) contains bromine (a substance also

used as a flame retardant) which can build up in fatty tissues. Animal studies have linked bromine to the development of heart lesions. Bromine has also been linked with neurological impairment, reduced fertility, changes in thyroid hormones, and early onset puberty. Recent studies have linked excessive use of BVO to skin lesions, nerve disorders, depression, irritability, confusion, cognitive problems, slurred speech, and memory loss. This additive and its derivatives have been banned in some countries. In the U.S. the FDA maintains that BVO and other bromine derivatives are safe for consumers and it is legal for food manufacturers to use these additives, though pressure from recent consumer and activism efforts, including one particularly newsworthy campaign and petition spearheaded by a teenage girl, has led the two leading soda manufacturers to voluntarily remove BVO from their products.

Growing consumer skepticism as the ultimate agent of change

These commonly used preservatives can be found in hundreds and hundreds of packaged food and drink items in local grocery markets as well as in the food served at both conventional and fast food restaurants, catered food, hotel and hospital foods, school lunchrooms, and the list goes on. But food items are not the only place some of these preservatives make an appearance. They can also be found in many personal care products such as mouthwash and over the counter medications. With all the ways that consumers can be exposed to these preservatives they hold the potential for some real problems, especially in those groups of people who are the most vulnerable. Fortunately, there is finally a growing awareness among consumers about the potential adverse health effects of some of the

most commonly used food additives, preservatives among them. According to a recent study, the number of people in all age groups who are reading ingredients labels before choosing to purchase is at an all-time high.

"In a 2014 online survey about food safety...
88 percent of respondents were concerned about preservatives."
~University of Florida's Public Issues in Education Center (PIE) Survey

As a result of the growing consumer skepticism about synthetic and industrialized food additives such as preservatives, in order to remain competitive food industry corporations have had to shift their current models. Packaged food producers and fast food restaurants have begun the process of removing and replacing questionable food preservatives in some food items and are making obvious attempts through labeling and packaging, advertising and submitting press releases to the media to alert consumers of those changes. Likewise, the sale of organic, preservative-free foods and products are at an all-time high, with conventional Big Food corporations—the ones who have been putting all the preservatives in the processed foods for decades—owning substantial shares of the organic food market.

But are all these shifts enough to protect consumers from preservatives of concern in their food and personal care products? Not really. To begin with, there are large numbers of U.S. consumers who simply cannot afford to purchase organic or preservative-free foods each week. Additionally, there are over 10,000 additives in U.S. foods with more than a thousand new chemicals entering the food supply every month—some, and perhaps even many lacking proper safety testing to assure they will not pose problems to those who consume them. With all of these additives it is impossible to expect consumers

to become knowledgeable about all of the potentially problematic preservatives in the food they put into their carts and ultimately, their bodies. The only real power U.S. consumers currently have is to communicate with Big Food producers, to prepare as many meals as possible from scratch using fresh, whole ingredients, and to vote with their dollars by refusing to purchase brands that contain unwanted preservatives and choosing preservative-free options for themselves and their family. Fortunately, given enough time, that power will likely be enough to push the changes needed forward.

#### References:

Aalto-Korte, K. 2000. Allergic contact dermatitis from Tertiary-Butylhydroquinone (TBHQ) in a vegetable hydraulic oil. *Contact Dermatitis*, 43 (5): 303–315.

Abdo, K., et al. 1983. Carcinogenesis bioassay of Propyl Gallate in F-344 rats and B6C3F1 mice. *Journal of American College of Toxicology*, 3(6): 425-433.

Abegaz, E. & Bursey, R. 2009. Formaldehyde, Aspartame, migraines: A possible connection. Dermatitis, 20(3):176-179.

Agency for Toxic Substances and Disease Registry (ATSDR). 2008. Toxicological profile for Formaldehyde. U.S. Department of Health and Human Services, Public Health Service. Public Health Statement, Atlanta, GA.

Anon., Two preservatives to avoid? BHT and BHA. Berkeley Wellness Center, February 1, 2011.

Anon., Online Survey: Floridians Concerned About Food Safety, GMO Foods, Food Safety News, December 12, 2014.

Arai Y., et al. 1998. Food and food additives hypersensitivity in adult asthmatics. III. Adverse reaction to Sulfites in adult asthmatics, *Arerugi*, 47(11):1163-1167.

Asero, R. 2006. Sodium Benzoate-induced pruritus. Allergy, 61(10):1240-1241.

Atkinson, D., Sim, T., & Grant, J. 1993. Sodium Metabisulfite and SO2 release: An under-recognized hazard among shrimp fishermen. Department of Internal Medicine, University of Texas Medical Branch, Galveston, National Institute of Health. *Annals of Allergy*, 71(6):563-566.

Buist, B., Ph.D., Food chemical sensitivity, New York: Prism Press, 1987.

Carlisle, P. The Food Hacker's Handbook: A Guide to Breaking the Processed Foods and Additives Addiction, New York, REP Books, Publication expected: 2015.

Center for Food Safety and Applied Nutrition.. 2009. Formaldehyde in food. Risk in brief. The Center for Food Safety, Hong Kong. Corder, E. & Buckley, C. 1995. Aspirin, Salicylate, Sulfite and Tartrazine induced bronchoconstriction: Safe doses and case definition in epidemiological studies. *Journal of Clinical Epidemiology*, 48(10):1269-1275.

Dacre, J. 1974. Long-term toxicity study of n-propyl gallate in mice. Food and Cosmetics Toxicology, 12, 125-129.

de Bie, A., & Ommen, B. 1992. Study on the *in vitro* biotransformation of Propyl, Octyl, and Dodecyl Gallate. TNO-report. V92.101, TNO Toxicology and Nutrition Institute, Zeist, The Netherlands.

Dubik, M. 2004. Food colorings, preservatives, and hyperactivity. *American Academy of Pediatrics, Grand Rounds*, 1295): 54-55. European Food Safety Authority (EFSA). 2004. Opinion of the scientific panel on food additives, flavourings, processing aids and materials in contact with food on a request from the Commission related to Tertiary-Butylhydroquinone (TBHQ). *The EFSA Journal*,84: 1-50

Environmental Health & Safety. 2008. Sodium Metabisulfite Material Safety Data Sheet, MSDS Number: S4378.

European Medicines Agency (EMA), Reflection paper on the use of Methyl and Propylparaben as excipients in human medicinal products for oral use. April 25, 2013.

Fisherman, F. & Cohen, G. 1973. Chemical intolerance to Butylated Hydroxyanisole (BHA) and Butylated Hydroxytoluene and (BHT) and vascular response as an indicator and monitor of drug intolerance. *Annals of Allergy*, 31:126-133.

Formaldehyde Council Fact Sheets http://formaldehyde.org/

Freedman, N., et al. 2010. Association of meat [Nitrates-Nitrites] and fat intake with liver disease and hepatocellular carcinoma in the NIH-AARP cohort. *Journal of the National Cancer Institute*, 102(17): 1354-1365.

Garcia-Melganes, M., et al. 2007. Sensitization to Gallates: Review of 46 cases. Actas Dermosifiligor, 98(10): 688-693.

Goodman D., et al. 1990. Chronic urticaria exacerbated by the antioxidant food additives, Butylated Hydroxyanisole (BHA) and Butylated Hydroxytoluene (BHT). *Journal of Allergy and Clinical Immunology*, 86(4) (Part 1): 570-575.

IBM. 2009. Less than 20 percent of consumers trust food they buy is safe and healthy, IBM Survey Reveals, Survey Sampling International (SSI).

International Agency for Research on Cancer (IARC). IARC Monographs on the evaluation of carcinogenic risks to humans: Formaldehyde, 2-Butoxyethanol and 1-tert-Butoxypropan-2-ol. Volume 88, June, 2004.

International Food Information Council (IFIC). Food and health survey: Consumer attitudes toward food safety. Cogent Research Institute, Cambridge, MA. 2010.

International Journal of Toxicology. 2007. Final Report on the amended safety assessment of Propyl Gallate. 26 Suppl 3:89-118. Johnson, A. & Hewgill, F. 1961. The effect of the antioxidants, Butylated Hydroxy Anisole (BHA), Butylated Hydroxytoluene (BHT) and Propyl Gallate on growth, liver and serum lipids and serum sodium levels of the rat. *Australian Journal of Experimental Biology and Medical Science*, 39: 353-360.

Kahl, R. & Kappus, H. 1993. Toxicology of the synthetic antioxidants BHA and BHT in comparison with the natural antioxidant vitamin E. Zeitschrift für Lebensmittel-Untersuchung und-Forschung, 196(4):329-338.

King, M. & McCay, P. 1980. DMBA-induced mammary tumour incidence: Effect of Propyl Gallate supplementation in purified diets containing different types and amounts of fat. *Proceedings of the American Association for Cancer Research*, (21):113.

Lawrence J. & Iyengar, J. 1983. The determination of Formaldehyde in beer and soft drinks by HPLC of the 2,4-Dinitrophenylhydrazone derivative. *International Journal of Environmental and Analytical Chemistry*, 15(1):47-52.

Lester, M. 1995. Sulfite sensitivity: Significance in human health. Journal of the American College of Nutrition, 14(3):229-232.

Malkinson, A. 1999. Lung tumor promotion by BHT. *National Library of Medicine, Crisp Data Base, National Institutes* of *Health*, CRISP/99/CA33497-14.

Material Safety Data Sheet (MSDS). Sodium Metabisulfite Material Safety Data Sheet, MSDS Number: S4378.

McCann *et al.* (2007) on the effect of some colours and Sodium Benzoate on children's behaviour: Scientific opinion of the panel on food additives, flavourings, processing aids and food contact materials (AFC). *The EFSA Journal*, 660:5-5.

McGwin, G, Lienert, J. & Kennedy, J. 2010. Formaldehyde exposure and asthma in children: A systematic review. *Environmental Health Perspectives*, 118: 313-317.

Meyer, O. & Hansen, E. 1980. Behavioural and developmental effects of Butylated Hydroxytoluene (BHT) dosed to rats *in utero* and in the lactation period. *Toxicology*, 16(3):247-258.

Michaelsson, G. & Juhlin, L. 1973. Urticaria induced by preservatives and dye additives in food and drugs. *British Journal of Dermatology*, 88(6): 525-532.

Mikhailova, Z., et al. 1985. Toxicological studies of the long term effects of the antioxidant Dodecyl Gallate on albino rats. *Voprosy Pitaniia*, (2):49-52.

Moneret-Vautrin D., Einhorn, C., & Tisserand, J. 1980. The role of Sodium Nitrite in allergic urticaria. (Le role du nitrite de sodium dans les urticaires histaminiques d'origine alimentaire.) *Annals of Nutritional Aliment*, 34(5-6):1125–1132.

Mpountoukas, P., et al. 2008. Cytogenetic study in cultured human lymphocytes treated with three commonly used preservatives. *Food and Chemical Toxicology*, 46:2390–2393.

Msagati, T. Chemistry of food additives and preservatives. Oxford, UK: Blackwell Publishing Ltd., 2012.

National Institutes of Health Report on Carcinogens (ROC). 2002. Report on Butylated Hydroxyanisole (BHA) in the Tenth Annual Report on Carcinogens (2002) of the Public Health Service National Toxicology Program. [BHA is "reasonably anticipated to be a human carcinogen..."]

National Organic Standards Board (NOSB). Organic food labels survey, 2014. Consumer Reports National Research Center Survey Research Report.

National Toxicology Program (NTP). 2010. Final report on carcinogens background document for Formaldehyde, (10-5981i): 512.

National Toxicology Program (NTP). Propyl-4-Hydroxybenzoate, Propylparaben, and n-Propyl Paraben: Human toxicity excerpts.

Selected toxicity information from HSDB, National Library of Medicine's database. CAS Registry Number: 94-13-3 Toxicity Effects, ChemIDPlus/HSDB 94-13-3 Toxicity.

National Toxicology Program, (NTP). 2010. Report on Carcinogens Background Document. Final Report on Carcinogens Background Document for Formaldehyde, January (10-5981i): 512.

Neltner, T. & Maffini, M. 2014. Generally recognized as secret: Chemicals added to food in the United States. *National Resource Defense Council (NRDC)Report.* 

Nera, E., et al. 1984. Short-term pathological and proliferative effects of BHA and other phenolic antioxidants in the forestomach of Fischer 344 rats. *Toxicology*, 32:197-205.

OECD SIDS Report on 2, 6-di-tert-butyl-p-cresol (BHT). Cas No: 128-37-0. UNEP Publications 2002.

Orten, J., Kuyper, A. & Smith, A. 1948. Studies on the toxicity of Propyl Gallate. Food Technology, 2:308-316.

Pacor, M., et al. 2004. Monosodium Benzoate hypersensitivity in subjects with persistent rhinitis. Allergy, 59(2):192-197.

Parke, D. & Lewis, D. 1992. Safety aspects of food preservatives. Food Additives and Contaminants, 9(5):561-577.

Perez, A., et al. 2008. Positive rates to Propyl Gallate on patch testing: a change in trend. Contact Dermatitis, 58(1): 47-48.

Product Stewardship Summary, 2010. Formaldehyde: Product Stewardship Summary, Ashland Inc.

Race, S. Antioxidants: The truth about BHA, BHT, TBHQ and other antioxidants used as food additives. Tigmor Books, 2009.

Rangan, C. & Barceloux, D. 2009. Food additives and sensitivities. Disease-a-Month, 55(5): 292-311.

Rangan C., et al. Food additives and sensitivities. In: D. Barceloux, (*Ed.*), Medical Toxicology of Natural Substances. Hoboken, N.J.: Wiley & Sons, 2008.

Ribeiro, F., et al. 2010. Sulfite hypersensitivity in an atopic patient. [Hipersensibilidade a sulfitos em doente atopica]. *Revista Portuguesa de Imunoalergologia*, July-August issue, 373-381.

Rosin, M. & Stich, H. 1980. Enhancing and inhibiting effects of Propyl Gallate on carcinogen-induced mutagenesis. *Journal of Environmental Pathology and Toxicology*, 4: 159-167.

Safer, A. & Al-Nughamish. A. 1999. Hepatotoxicity induced by the anti-oxidant food additive, Butylated Hydroxytoluene (BHT), in rats: An electron microscopical study. *Histology and Histopathology*, 14(2):391-406.

Scheuplein, R. Formaldehyde: The Food and Drug Administration's perspective. In Formaldehyde: Analytical chemistry and toxicology, V. Turoski (*ed.*) Office of Toxicological Sciences, Bureau of Foods, Food and Drug Administration, Washington, DC 20204, 1985. Vol 210, Chpt. 16, pp. 237–245.

Schecter, A., et al. 2011. Contamination of U.S. butter with Polybrominated Diphenyl Ethers (PBDEs) from wrapping paper. *Environmental Health Perspectives*, 119(2): 151–154.

ScienceLab.com. Material Safety Data Sheet on BHA. 2008.

Scientific Committee on Consumer Products (SCCP). Opinion on Benzoic Acid and Sodium Benzoate. SCCP/0891/05, European Commission for Health and Consumer Protection General Directorate C - Public Health and Risk Assessment C7.

Scott, MD, Glenn. 2011. Citizens Regulatory Petition to U.S. Food and Drug Administration (FDA) requesting Food and Color Additive changes and FAP 0A4216 Petition to Prohibit the Use of BHA in Food. Petitions Under Review as of March, 2011.

Scott, G. 1990. Citizens Regulatory Petition FDA Requesting the U.S. Food and Drug Administration (FDA) to prohibit the use of BHA in food

Shim, S., et al. 2011. Consumers' knowledge and safety perceptions of food additives: Evaluation on the effectiveness of transmitting information on preservatives. *Food Control*, 22(7): 1054-1060.

Siman, C. & Eriksson, U. 1996. Effect of Butylated Hydroxytoluene (BHT) on alpha-tocopherol content in liver and adipose tissue of rats. *Toxicology Letters*, 87(2-3):103-108.

Speijers, G. & van Apeldoorn, M. Gallates (Propyl, Octyl & Dodecyl). Toxicological evaluation of certain food additives and contaminants. National Institute of Public Health and Environmental Protection Laboratory for Toxicology, Bilthoven, The Netherlands. World Health Organization Series 32, 1993.

Statista, The Statistics Portal, Organic Food Sales in the U.S., 2000-2014.

Stokes, J. & Scudder, C. 1974. The effect of Butylated Hydroxyanisole (BHA) and Butylated Hydroxytoluene (BHT) on behavioral development of mice. *Developmental Psychobiology*, 7(4):343-350.

Tamano, S., et al. 1987. Modification by O-Tocopherol, Propyl Gallate and Tertiary Butylhydroquinone (TBHQ) of urinary bladder carcinogenesis in Fischer 344 rats pretreated with N-Butyl-N-(4-Hydroxybutyl) nitrosamine. *Cancer Letters*, 35:39-46.

Tang, X., et al. 2009. Formaldehyde in China: Production, consumption, exposure levels, and health effects. *Environment International*, 35 (8):1210-1224.

Technical University of Denmark, "Less nitrite in meat products reduces levels of nitrosamines." *ScienceDaily*,6 November 2014. Thomson Reuter poll on consumer preferences for organic food, 2011.

Towns, S. & Mellis, C. 1984. Role of Acetyl Salicylic Acid and Sodium Metabisulfite in chronic childhood asthma. *Pediatrics*, 73(5):631-637.

Tosti, A., et al. 1987. Contact dermatitis from Butylated Hydroxyanisole (BHA). Contact Dermatitis, 17:257-258.

Trocho, C., et al. 1998. Formaldehyde derived from dietary Aspartame binds to tissue components *in vivo*. *Life Sciences*, 63(5): 337-349. U.S. Code of Federal Regulations (CFR).. 2010. BHA. Code of Federal Regulations, Title 21, Volume 3. CITE: 21CFR173.340. Revised April 1.

U.S. Code of Federal Regulations (CFR). 2010. BHT (Butylated Hydroxytoluene). Code of Federal Regulations, Title 21, Volume 6. CITE: 21CFR573.940.

U.S. Environmental Protection Agency (EPA). Integrated Research Information Center (IRIS) Summary Document for Formaldehyde, 1991.

U.S. Food and Drug Administration (FDA). 2011. Food and Color Additive Petitions Under Review as of March 2011, Glenn Scott, FAP 0A4216 Petition to Prohibit the Use of BHA in Food.

U.S. Environmental Protection Agency (EPA). Integrated Research Information Center (IRIS) Summary Document for Formaldehyde, 1991.

van der Heijden, C., Janssen, P. & Strik, J. 1986. Toxicology of Gallates: A review and evaluation. *Food Chemical Toxicology*, 24(10-11): 1067-1070.

van Esch, G. 1986. Toxicology of Tert-Butylhydroquinone (TBHQ). Food and Chemical Toxicology, 24(10-11): 1063-1065.

van Esch, G. 1955. The toxicity of the antioxidants Propyl-, Octyl- and Dodecyl Gallate, Voeding, 16: 683-686.

van Laar, T., et al. 1998. The role of EDTA in provoking allergic reactions to subcutaneous infusion of apomorphine in patients with Parkinson's Disease: A histologic study. *Movement Disorders*, 13(1):52-55.

Vorhees, C., et al. 1981. Developmental neurobehavioral toxicity of Butylated Hydroxyanisole (BHA) in rats. *Neurobehavioral Toxicology and Teratology*, 3(3):321-329.

Warner, Melanie. Pandora's Lunchbox: How Processed Food Took Over the American Meal, Scribner, 2013. Grist.org interview, March, 2013.

Williams, G., Iatropoulos, M. & Whysner, J. 1999. Safety assessment of Butylated Hydroxyanisole (BHA) and Butylated Hydroxytoluene (BHT) as antioxidant food additives. *Food and Chemical Toxicology*, 37(9-10): 1027-1038.

Wong, G. & Shear, N. 2003. Melkersson-Rosenthal syndrome associated with allergic contact dermatitis from Octyl and Dodecyl Gallates. *Contact Dermatitis*, 49(5): 266-267.

World Health Organization (WHO). Gallates: Dodecyl, Octyl, and Propyl. Food Additives Series No. 5. Seventeenth Report of the Joint FAO/WHO Expert Committee on Food Additives, World Health Organization Technical Report Series No. 539 and FAO Nutrition Meetings Report Series No. 53, 1974.

World Health Organization (WHO). n-Octyl Gallate. Sixteenth Report of the Joint FAO/WHO Expert Committee on Food Additives, World Health Organization Technical Report Series, No. 505 and FAO Nutrition Meetings Report Series, No. 51, 1972.

World Health Organization (WHO). Evaluation of Mercury, Lead, Cadmium and the Food Additives Amaranth, Diethylpyrocarbonate and Octyl Gallate. Food Additives Series, No. 4. The evaluations contained in this publication were prepared by the Joint FAO/WHO Expert Committee on Food Additives which met in Geneva, 4-12 April 1972.

Zengin, N., et al. 2011. The evaluation of the genotoxicity of two food preservatives: Sodium Benzoate and Potassium Bezoate. *Food and Chemical Toxicology*, 49(4):763–769.

Zhang, L., et al. 2010. Formaldehyde and leukemia: Epidemiology, potential mechanisms, and implications for risk assessment. *Environmental and Molecular Mutagenesis*, 51 (3):181–191.

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