What's in Our Food?

The Science and Safety of Food Additives

right. Academy of Nutrition and Dietetics
Foundation

Future of Food Initiative

Amy Myrdal Miller, MS, RDN, FAND

Founder and President
Farmer's Daughter Consulting, Inc.

Roger Clemens, DrPH, CFS, CNS, FACN, FIFT, FIAFST

Adjunct Professor
USC School of Pharmacy, International
Center for Regulatory Science





Session Objectives

- 1. Describe the history of food additives and their specific function in food systems.
- 2. Describe major contributions of food additives as well as their risks and benefits.
- 3. Identify sources of credible, science-based information for the public about food additives.

Speakers

Future of Food Initiative



Roger Clemens, DrPH, CFS, CNS, FACN, FIFT, FIAFST

Adjunct Professor

USC School of Pharmacy, International Center for Regulatory Science



Amy Myrdal Miller, MS, RDN, FAND Founder and President Farmer's Daughter Consulting, LLC



Lisa Medrow, RDN, LD Foundation Project Specialist Academy of Nutrition and Dietetics Foundation

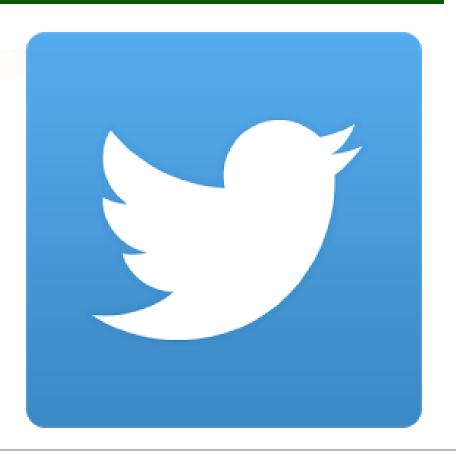


Tweeting Today?

@eatrightPRO

@AmyMyrdalMiller

@FoodGeekRoger





Future of Food Initiative

Academy of Nutrition and Dietetics Foundation





Future of Food Resources for Members

- Toolkits <u>www.eatrightfoundation.org/toolkits-webinars</u>
 - Hunger in Our Community. What We Can Do.
 - Smart Choices. For a Healthy Planet. (English/Spanish!)
 - Tossed Treasures. How We All Can Waste Less Food. (English/Spanish!)
- Supervised Practice Concentrations:
 - Food Insecurity and Food Banking—available now! www.healthyfoodbankhub.org
 - Food Systems—under development!
- Webinars and Infographics <u>www.eatrightfoundation.org</u>
- Affiliate Presentations:
 - "Changing the Way We Look at Agriculture" 32 affiliates/DPGs (2015)
 - Food waste, food additives, and GMO presentations 10 affiliates (2016)
 - Foods of future, farming tools, and food preservation presentations
 10 affiliates (2017)

Last year our donors' generosity helped us award:

\$446,900 in student scholarships to 194 students

\$14,000 in student stipends to help 140 students attend FNCE.

\$40,000 through Home Food Safety Challenge grants to dietetics students.

www.eatrightfoundation.org



Roger Clemens, DrPH, CFS, CNS, FACN, FIFT, FIAFST

Adjunct Professor, Pharmacology & Pharmaceutical Sciences USC School of Pharmacy, International Center for Regulatory Science, Los Angeles, CA clemens@usc.edu

@FoodGeekRoger

Disclosures (2010-2016)

right. Academy of Nutrition and Dietetics
Foundation

- Abbott Nutrition
- Almond Board of California
- American Society for Quality
- Aspire Food Group
- Assure Water
- Authen Technologies
- Barilla
- Bayer
- Biothera
- California Walnut Commission
- Coca-Cola (honorarium directly given to charity)
- Corn Refiners Association
- Danish Agriculture and Food Council
- Dairy Council of California
- Dentons LLP; E.T. Horn
- FMC (honorarium directly given to charity)
- Food Minds
- HyCite
- Jenner & Block LLP

- Karma Agency (honorarium directly given to charity)
- Kellogg
- Kerry
- Malaysian Palm Oil Council
- McDonalds
- Mead Johnson
- Mushroom Council
- National Aeronautical Space Agency
- National Fisheries Institute
- National Restaurant Association
- Nestlé SA
- Petcurean
- Pfizer
- Quaker Oats
- Schwann Foods
- Senomyx (honorarium directly given to charity)
- Spherix Consulting
- U.S. Dept. of Agriculture
- Whitewave
- Yakult

What is a Food?¹

Future of Food Initiative

Food

- Defined as "articles or components of articles used for food or drink for humans or animals"
- Safety standard:
 - Presumption of safety (unless it contains a poisonous or deleterious substance in an amount which is shown to make it *ordinarily injurious* to health)

What is a Food Additive?²



Food Additive (See Food Additive Amendment, 1958) Any substance, the intended use of which may reasonably be expected to, directly or indirectly, becoming a component or otherwise affecting the characteristics of any food

Exclusions:

- Pesticides in or on raw agricultural commodities
- Color additives
- Prior sanctioned substances
- Substances that are generally recognized as safe (GRAS)

Safety burden on the manufacturer

Added Substances to Foods³



Future of Food Initiative

In cases where the substance is not naturally present in food but is a contaminant or added ingredient, the safety standard is quite different

A food is adulterated if it contains any poisonous or deleterious substance that *may render it injurious*



http://www.ars.usda.gov/is/graphics/photos/k7191-4.htm



www.freeimages.com/helenb jork



www.freeimages.co m/williamstadler

- Part of the 1958 Food Additives Amendment; invoked in 1959
- Governs regulation of pesticide residues in processed foods (zero-tolerance)
- Pesticide Residue Amendment (focus on raw commodities) of 1954
 - EPA must conduct risk/benefit analysis
- Many subsequent conflicting laws
 - Regulation of new / old pesticides in raw and processed foods
 - "Negligible risk" (1/1,000,000)
 - Pesticide registration
- "De minimis" risk policy in 1988; over turned 1992

- To preserve the nutritional quality of the food
- To provide necessary ingredients or constituents for foods manufactured for groups of consumers having special dietary needs
- To enhance the keeping quality or stability of a food or to improve its organoleptic properties
- To provide aids in the manufacture, processing, preparation, treatment, packing, transport or storage of food





Food without Food Additives

- Consider bread, e.g., baking soda, folic acid, B vitamins
- Consider ice cream, e.g., colors, flavors and consistency
- Consider medical foods, e.g., vitamins and mineral
- Consider foods with standards of identity, e.g., vitamin A and vitamin D in milk and other dairy products
- Consider product stability, e.g., preservatives
- Consider food sustainability, e.g., re-worked foods and ingredients → reduce waste

Why Food Preservatives?8



Fundamental functions

- Promote food safety
- Provide greater food choices
- Promote food conveniences

Example:

- Sulfites reduce lipid oxidation (rancidity; <10 ppm-1000 ppm)
- Nitrate/nitrite to inhibit C. botulinum (natural in some vegetables, such as beets)
- Benzoic acid (salts) inhibits bacteria and molds (natural in some vegetables, such as strawberries and tomatoes)
- Sorbic acid (K, Na, Ca) antimicrobial originally isolated from berries (often used in wine < 300 ppm)



Future of Food Initiative

Direct

- Added during processing
- Functions
 - Provide nutrients
 - Help process or prepare the food
 - Keep the product fresh
 - Make the food more appealing

Indirect

- May be found in food during or after it is processed
- Substances found in trace amounts due to
 - Packaging migration
 - Storage environment
 - Handling procedures

- Food Additives Amendment of 1958
- Substances subject to prior sanction
- GRAS substances are exempt from food additive requirements
 - If a substance did not have a common use prior to 1958, the substance is subject to scientific review procedures (same criteria for food additive)
 - Exemptions to GRAS → commodities; separate regulations primarily under FDA and USDA jurisdiction

Hazard

- Types of toxic effects caused by the chemical
- Manifestation depends on route, amount, duration and frequency of exposure

Risk

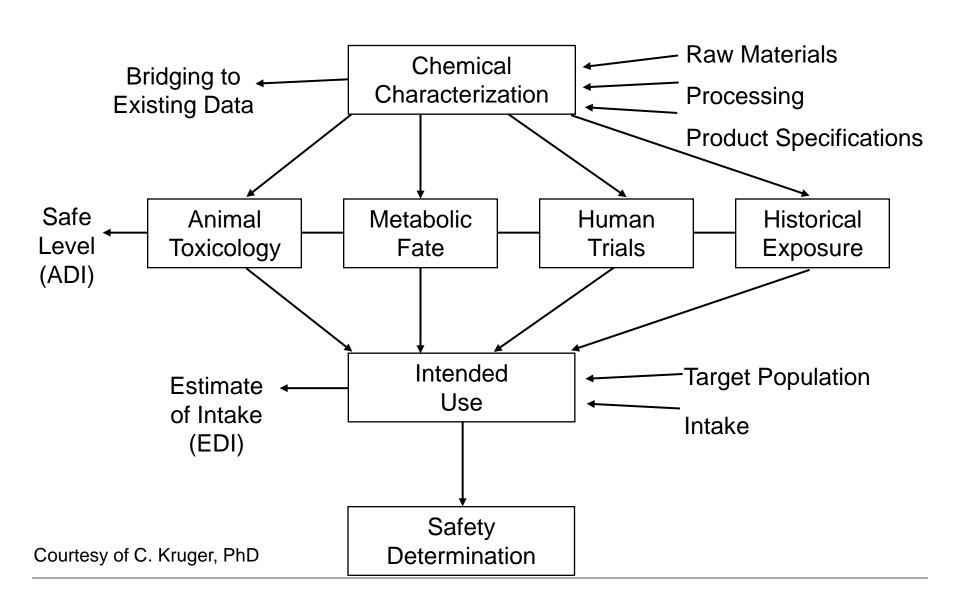
• Likelihood that the toxic properties of a chemical will be produced in populations of individuals under their actual conditions of exposure; exposure must precede adverse event

Safety

- Little or no harm will result from chemical under given set of exposure circumstances
- It is not the absolute absence of risk; it is the inverse of risk

Safety Analysis¹⁵





- Human trumps animal data
- In the absence of human data, animal data are used
- In the absence of information to demonstrate that the selection is incorrect, data from the animal species showing the greatest sensitivity are selected for the risk assessment
- Animal data using the same route of exposure as relevant to human exposure are preferred
- For all toxic effects other than carcinogenicity, a threshold is assumed
- Thresholds in humans are established by application of safety factors to data
- For carcinogens, a linear, no threshold dose-response model is assumed to apply

Direct Food Additives¹⁸



1.	Anticaking agents and free-flow agents	18.	Leavening agents
2.	Antimicrobial agents	19.	Lubricants and release agents
3.	Antioxidants	20.	Nonnutritive sweeteners
4.	Colors and color adjuncts	21.	Nutrient supplements
5.	Curing and pickling agents	22.	Nutritive sweeteners
6.	Dough strengtheners	23.	Oxidizing and reducing agents
7.	Drying agents	24.	pH control agents
8.	Emulsifiers and emulsifier salts	25.	Processing aids
9.	Enzymes	26.	Propellants, aerating agents, and gases
10.	Firming agents	27.	Sequestrants
11.	Flavor enhancers	28.	Solvents and vehicles
12.	Flavor agents and adjuvants	29.	Stabilizers and thickeners
13.	Flour-treating agents	30.	Surface-active agents
14.	Food preservatives	31.	Surface-finishing agents
15.	Formulation aids	32.	Synergists
16.	Fumigants	33.	Texturizers
17.	Humectants		

Food Preservatives^{8,19,20}



- Substances added to foods to inhibit microbial growth or retard product deterioration
- Prior sanctioned GRAS
 - Ascorbic acid
 - Erythorbic acid
 - Sorbic acid
 - Thiodipropionic acid
 - Ascorbyl palmitate
 - Butylated hydroxyanisole (BHA)
 - Butylated hydroxytoluene (BHT)
 - Calcium ascorbate
 - Calcium sorbate
 - Dilauryl thiodipropionate

- Potassium bisulfite
- Potassium metabisulfite
- Potassium sorbate
- Sodium ascrobate
- Sodium bisulfite
- Sodium metabisulfite
- Sodium sorbate
- Sodium sulfite
- Sulfur dioxide
- Tocopherols

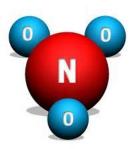
Applications

- Depend on regulatory statutes and ingredient (often establish upper limits, e.g., < 0.1% [1000 ppm])
 - Most regulations only stipulate usage levels within Good Manufacturing Practice
- Depend on food matrix (e.g., liquid, dry, pH)
- Depend on processing conditions (e.g., temperature), storage environment, transportation demands

- JECFA (Joint FAO/WHO Expert Committee on Food Additives) has evaluated more than 2,500 food additives
- Risk assessment/safety evaluation of:
 - Food additives (intentionally added)
 - Processing aids (considered as food additives)
 - Flavoring agents (by functional groups)
 - Contaminants
 - Natural toxins
 - Exposure assessment

Future of Food Initiative

Nitrate



Nitrite=NO₂

- Naturally generated by plants during nitrogen fixation process, and produced in humans
- Naturally occur in soils (part of organic matter)
- Naturally found in vegetables (e.g., celery, lettuce, spinach [500-1900 ppm]), fruits, meats, fish, dairy products, fermented beverages (e.g., beers), and cereals
- Typical usage level < 500 ppm (food matrix dependent)
- IARC (2010) classified these substances as probable carcinogens (Group 2A) to humans → ingested nitrate or nitrite under conditions that result in endogenous nitrosation → N-nitroso compounds
- Reported toxicity by association; limited direct evidence except through nitrosation

Potential adverse effects → methemoglobinemia Image Credit: uswatersystems.com

Precursor to nitric oxide → limits gastric pathogens (antimicrobial effects, reduces pulmonary hypertension, promotes tissue perfusion

Processing Aids²³



Definition:

- Codex: Substances that fulfil a certain technological purpose during treatment or processing and which may result in the non-intentional but unavoidable presence of residues or derivatives in the final product
- FDA: Substances that are added to a food during the processing of such food but are removed in some manner from the food before it is packaged in its finished form
 - Exempt from food labeling requirements in USA
 - Under CVM, required labeling for pet food/livestock feed in USA

Food (inorganic) Phosphates²⁴

- Classification: monovalent salts, divalent salts, ammonium salts and aluminum salts
- Regulatory: GRAS
- Examples (usage levels)
 - Mono, di and trisodium phosphate (< 0.5%)
 - Mono, di and tripotassium phosphate (< 0.5%)
 - Sodium hexametaphosphate (GMP)
 - Ammonium polyphosphate (GMP)
 - Sodium aluminum phosphate (GMP)

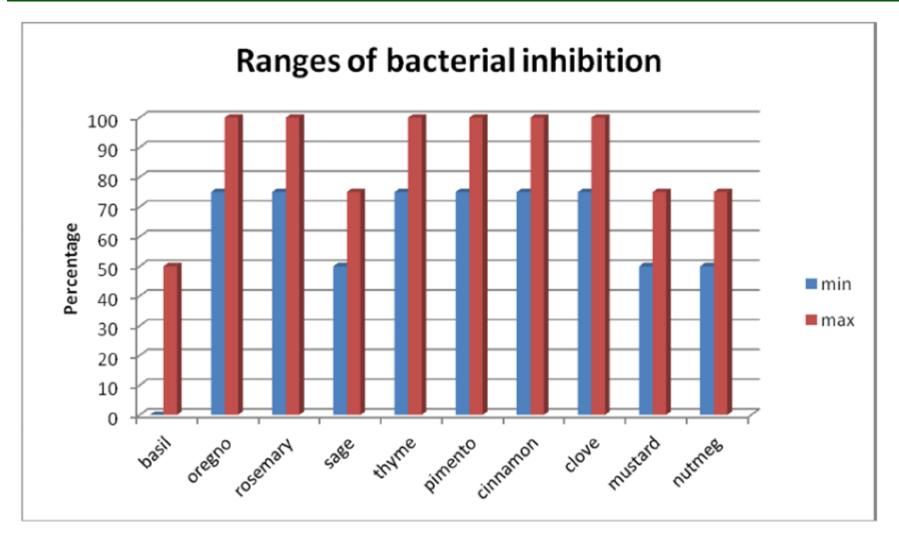
Food (inorganic) Phosphates²¹

- Functional Properties (examples)
 - TSP, 19.5% phosphorus,— sanitizer (cleaning reagents); emulsifier (pasteurized cheese)
 - Sodium aluminum phosphate, 31% phosphorus— chemical leavening (baking powder)
- Labeling
 - Ingredient declared in descending order of prevalence
 - Specific phosphorus contributions in Nutrition Facts Panel not quantified or required

Future Role of Spices²⁵



Future of Food Initiative



Typical tested extract doses: 500-1000 ppm (0.05-0.1%); effectiveness depends on cultivar, extract, organism and food matrix.

Conclusion



Photo credit: www.freeimages.com/zernliew

- Public Health and Safety is the over-arching priority
- Vigilance and good data on safety trump every other factor
- Regulatory future

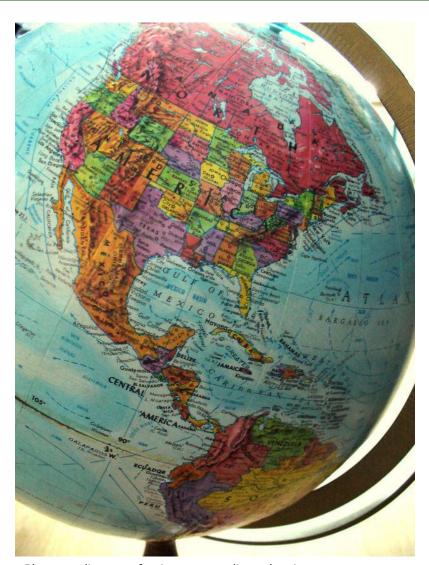


Photo credit: www.freeimages.com/jessetherrien

- US has one of the safest and most abundant food supplies in the world.
- Food safety is everyone's responsibility.
- Challenge to assure safe global food supply.
- Challenge to harmonize safety assessment of food additives.



Thank you!

Roger Clemens, DrPH, CFS, CNS, FACN, FIFT, FIAFST

Adjunct Professor, Pharmacology & Pharmaceutical Sciences
USC School of Pharmacy, International Center for Regulatory Science, Los Angeles, CA clemens@usc.edu



Roger Clemens, DrPH Quito, Ecuador



Amy Myrdal Miller, MS, RDN, FAND

Founder and President
Farmer's Daughter Consulting, Inc.
Carmichael, CA

@AmyMyrdalMiller

Disclosures



- Academy of Nutrition and Dietetics Foundation honoraria for speaking
- American Beverage Association *consultant*
- California Leafy Greens Marketing Agreement *consultant*
- The Culinary Institute of America *consultant*
- Dairy Management, Inc. *honoraria for speaking*
- Dairy MAX *consultant*
- Florida Department of Citrus consultant
- Great Valley Publishing Company *blogger, honoraria for speaking*
- Monsanto Company *consultant*
- Monsanto Company *L.E.A.D. Network Member*
- Mushroom Council consultant/research funding
- National Dairy Council Ambassador
- Naturipe® Farms *consultant*
- Northarvest Bean Growers Association consultant
- Northern California Golden Arches Association consultant
- *Produce Business* / Phoenix Media Network *consultant/columnist*
- Produce Marketing Association *honoraria for speaking*
- University of California, Davis Honey and Pollination Center honoraria for speaking

Where are Consumers?^{26,27,28}



Future of Food Initiative

- 51% report that additives have a strong influence on purchase
- Over 30% "cautious about serving foods with preservatives" compared to 24% 10 years ago; trend for additives follows same progression
- Shoppers sought products made without
 - Preservatives 33%
 - Chemical additives 32%
 - High fructose corn syrup 28%
- 25% sought products only with recognizable ingredients or shortest ingredient list

Nutrition	Amount/Serving	%DV*	Amount/Serving	%DV*	Amount/Serving %DV*
Facts Serving Size 1 Container (28g) Calories 100 Calories from Fat 0 * Percent Daily Values (DV) are based on a 2,000 calorie diet.		• Vit		g 8 % Calcium Riboflav	0% • Iron 20% rin 20% • Niacin 20%
INGREDIENTS: MILLED CORN, SUG. VITAMINS AND MINERALS: IRON, V (PYRIDOXINE HYDROCHLORIDE), V PALMITATE, FOLIC ACID, VITAMIN D	ITAMIN C (ASCORBI	CACID	AND SODIUM ASCO	RBATE),	NIACINAMIDE, VITAMIN B6

Image source: Kellogg's

Where are Consumers?²⁹



Future of Food Initiative

59% believe fewer ingredients means a healthier product

<u>Top claims</u> among "free-from" claim product seekers:

- Trans-fat-free 78%
- Preservative-free 71%



Photo Credit: USDA ARS Image Number D1830-30. http://www.ars.usda.gov/is/graphics/photos/mar12/d1830-30.htm?pf=1

Where are Consumers?30,31



Future of Food Initiative

"Very important" - absence of:

- High fructose corn syrup 32% of respondents
- No artificial colors 29%
- No artificial flavors 30%

More than 60% of Americans say that the absence of artificial colors or flavors is important in their food-buying decisions.

Artificially sweetened "diet/light" products
 12%

Where are Consumers?³²

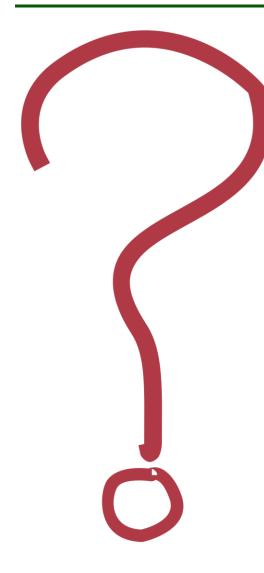


Future of Food Initiative

According to research from The Hartman Group, consumers deliberately avoid the following ingredients:

- High fructose corn syrup 56% Artificial flavors 49%
- Saccharin 52%
- Growth hormones 52%
- MSG 51%
- Aspartame 49%

- Artificial colors or dyes 49%
- Artificial preservatives 45%
- Sucralose 42%
- Partially hydrogenated vegetable oils 37%



- Confusion and fear of the unknown continue to drive interest in simplified labels and more natural ingredients
- "Clean" "Simple" "Real" used on labels to introduce natural, no additives or preservatives
- Looking for removal of 'artificial' ingredients
- Looking to avoid ingredients they know are additives, preservatives, or words they don't know

Additives and Labeling^{37,38}



Current Food Labeling Ingredient List Regulation Requirements:

Flavors – have definitions for compositional criteria and must be labeled "Artificial" unless Natural

Chemical Preservatives – identity must include function

Colors – identity must include function, even if of natural origin, when added solely to color

<u>Functional labeling</u> (ingredient purpose) - is permitted but not required for other additives

<u>Incidental additives</u> - insignificant levels, no technical/functional effect, introduced by another ingredient processing aids, equipment or packaging substances are not required to be labeled

Future of Food Initiative

- Consumers are willing to pay a premium when a product label says "free of" something
 - ✓ But only if the package includes "negative" information on whatever the product is "free of"

When provided more information about ingredients, consumers are more confident about their decisions and value the product more.



Photo credit: ww.freeimages.com/billyalexander

Commonly listed additives at top ranked consumer-advocacy styled sites:

MSG BHA/BHT

Nitrates Trans Fat

Potassium Bromate Color Additives

Sulfites Artificial Sweeteners

Facts About Function



Future of Food Initiative

To improve **safety**, **freshness** and keep **quality** (preservatives)

- Nitrates/Nitrites
- Sulfites
- BHA/BHT



Photo credit: www.freeimages.com/tes

To enhance consumer acceptability including flavor and color

MSG (flavor enhancer)

To improve texture and flavor stability

Trans fat from partially hydrogenated oils

To maintain product **consistency**:

Potassium bromate (dough conditioner)

Additives in Perspective



For Food Preservation

- Prevented spoilage foodborne illness was once the #1 cause of death
- Enabled geographic distribution greater variety of food for more people in areas with less diverse agriculture

For Sensory Characteristics

Consumer **appeal** and **acceptability** to increase sales – some "conditioning" has resulted in establishing consumer expectations – e.g. "red" beverages like "fruit punch"; pink hot-dogs (might naturally be gray)

For Industry (internal) Cost-Savings

Some additives **reduce labor costs**, replace or **minimize more expensive ingredients**



- Consumers tend to:
 - ✓ amplify the risk when a food or a technology is unknown

OR

- ✓ minimize the risk in familiar foods or home preparation
- Difficult-to-pronounce additive names lead to the impression of unfamiliarity, perceptions of higher health risk
- Consumer perceptions and resulting actions determine the commercial future of any food ingredient

The goal of risk communication should be to consolidate these views and enable consumers to make informed decisions about food additives.



Future of Food Initiative



How Industry is Responding



Future of Food Initiative

- Several restaurant chains are eliminating additives such as artificial ingredients
- Food manufacturers and grocers are replacing artificial colors and flavors with natural ingredients; working with suppliers and producers that avoid antibiotics



Photo credit: www.freeimages.com/simonstratford

na '

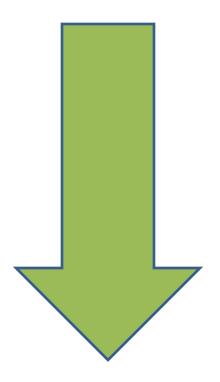
* Academy of Nutrition right and Dietetics Foundation

Future of Food Initiative

How Industry is Responding

- ↓ artificial colors & flavors
- ↓ preservatives/artificial preservatives
- ↓ artificial sweeteners
- ↓ corn syrup/high fructose corn syrup
- **↓ MSG**





Consumer's "Right to Know"26



Future of Food Initiative

What can Academy members do?

- Provide more information on the benefits (functions) of certain additives
- Refute misinformation
- Understand consumer's perception of risks

Consumers should be able to <u>share concerns</u> and Academy members should be able to provide <u>accessible</u> and <u>accurate</u> information.



Consumer's Right to Choose

 Role of RD/RDN = sciencebased facts to enable informed decisions

 Each consumer has the individual right to decide what they put in their body and choose foods based on their preferences.



right. Academy of Nutrition and Dietetics Foundation

Consumer-Friendly Resources

Future of Food Initiative

- FDA Food Additives & Ingredients
 www.fda.gov/Food/IngredientsPackagingLabeling/FoodAdditivesIngredients/
 default.htm
- FDA Consumer Info About Additives & Ingredients
 www.fda.gov/Food/IngredientsPackagingLabeling/FoodAdditivesIngredients/ucm094210.htm
- U.S. Dept. of Agriculture (USDA) Food Safety & Inspection Service (USDA-FSIS regulates meat and poultry products) - Additives in Meat and Poultry Products
 - <u>www.fsis.usda.gov/wps/portal/fsis/topics/food-safety-education/get-answers/food-safety-fact-sheets/food-labeling/additives-in-meat-and-poultry-products/additives-in-meat-and-poultry-products</u>
- Medline Plus Food Additives
 www.nlm.nih.gov/medlineplus/ency/article/002435.htm
- International Food Information Council www.foodinsight.org/Food Ingredients Colors



Thank you!

QUESTIONS?

Amy Myrdal Miller, MS, RDN, FAND

Roger Clemens, DrPH, CFS, CNS, FACN, FIFT, FIAFST

Please complete this short online feedback survey:

www.surveymonkey.com/r/fdaddweb





- Food and Drug Administration Web site. <u>http://www.fda.gov/regulatoryinformation/legislation/federalfooddrugandcosmeticactfdcact/</u>. Accessed February 3, 2016.
- 2. U.S. Public Law 85-929. September 6, 1958. http://uscode.house.gov/statutes/pl/85/929.pdf. Accessed February 3, 2016.
- 3. U.S. Food and Drug Administration Web site. FD&C Act, Section 402. http://www.fda.gov/RegulatoryInformation/Legislation/FederalFoodDrugandCosmeticActFDCAct/FDCActChapterIVFood/. Accessed February 4, 2016.
- 4. The Free Dictionary Web site. http://legal-dictionary.thefreedictionary.com/De+Minimis. Accessed February 4, 2016.
- 5. Merrill, RA. Food Safety Regulation: Reforming the Delaney Clause. *Annual Review of Public Health.* 1997;18:313-40.
- 6. Fortin ND. Food Regulation Law, Science, Policy and Practice. John Wiley & Sons, Inc.; 2009.
- 7. Codex Alimentarius Web site. Codex General Standard for Food Additives (GSFA) Online Database. http://www.fao.org/fao-who-codexalimentarius/standards/gsfa/en/. Accessed February 4, 2016.
- 8. Food and Drug Administration Web site. 21CFR101.22 http://www.fda.gov/Food/IngredientsPackagingLabeling/FoodAdditivesIngredients/ucm094211.htm. Accessed February 3, 2016.



- U.S. Food and Drug Administration Web site. Loss of Indirect Additives Used in Food Contact Substances. http://www.fda.gov/Food/IngredientsPackagingLabeling/PackagingFCS/Indirect-Additives/default.htm. Accessed February 4, 2016.
- 10. U.S. Food and Drug Administration Web site. Food Additive Status List. http://www.fda.gov/Food/IngredientsPackagingLabeling/FoodAdditivesIngredients/ucm091048.htm. Accessed February 4, 2016.
- 11.U.S. Government Publishing Office Web site. Electronic Code of Federal Regulations. (21CFR181) http://www.ecfr.gov/cgi-bin/text-idx?rgn=div5&node=21:3.0.1.1.12. Accessed February 4, 2016.
- 12.U.S. Food and Drug Administration Web site. 21CFR170.30(b). http://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfcfr/CFRSearch.cfm?fr=17 0.30. Accessed February 4, 2016.
- 13. U.S. Food and Drug Administration Web site. 21CFR170.35(c)(1). http://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm?fr=170.3 5. Accessed February 4, 2016.
- 14. U.S. Food and Drug Administration. FDA Food Code 2009. http://www.fda.gov/Food/GuidanceRegulation/RetailFoodProtection/FoodCode/ucm186464.htm. Accessed February 4, 2016.



- 15. U.S. Food and Drug Administration Web site. Generally Recognized as Safe (GRAS). http://www.fda.gov/Food/IngredientsPackagingLabeling/GRAS/. Accessed February 4, 2016.
- 16. Williams, latropoulos, Enzmann, et al. *Hayes' Principles and Methods of Toxicology* . 6th ed. CRC Press; 2014:25.
- 17. Klaassen C, Watkins JB. Casarett & Doull's Essentials of Toxicology. 7th ed. McGraw-Hill; 2010:8.
- 18. U.S. Food and Drug Administration Web site. 21CFR184.

 https://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfcfr/CFRSearch.cfm?CFR

 Part=184. Accessed February 5, 2016.
- 19. U.S. Food and Drug Administration Web site. 21CFR182. Accessed February 5, 2016.
- 20. U.S. Food and Drug Administration Web site. 21CFR172. Accessed February 5, 2016.
- 21. U.S. Food and Drug Administration Web site. Code of Federal Regulations Title 21. http://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm. Accessed February 5, 2016.
- 22. Food and Agriculture Organization of the United Nations Web site. Chemical risks and JECFA. http://www.fao.org/food/food-safety-quality/scientific-advice/jecfa/en/. Accessed January 12, 2016.



- 23. U.S. Food and Drug Administration Web site. 21 CFR 101.100 (a) (3) (ii) http://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm?fr=101.1 http://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm?fr=101.1 http://occessed.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm?fr=101.1 https://occessed.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm?fr=101.1 https://occessed.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm?fr=101.1 <a href="https://occessed.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.cfm?fr=101.1 <a href="https://occessed.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cdrh/cfdocs/cfcfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scripts/cfr/cfrsearch.gov/scrip
- 24. Weiner ML, Salminen WF, Larson PR, et al. Toxicological Review of Inorganic Phosphates. *Food and Chemical Toxicology.* 2001;39(8):759-86.
- 25. De La Torre Torres, JE, Gassara F, Kouassi AP, et al. Critical Reviews in Food Science and Nutrition. 2013; DOI:10.1080/10408398.2013.858235.
- 26. Petrun L, Flood A, Sellnow T, . et al. Shaping health perceptions: Communicating effectively about chemicals in food. *Food Protection Trends*. 2014:35(1):24–35
- 27. The NPD Group Web site. www.npd.com/wps/portal/npd/us/news/press-releases/2015/consumer-concern-about-food-ingredients-changes-their-view-on-how-natural-should-be-defined-on-food-labels-reports-npd-group/.

 Accessed January 22, 2016.
- 28. Sloan AE. The Top Ten Food Trends. Food Technology . 2015:69(4).
- 29. Mintel Web site. www.mintel.com/press-centre/food-and-drink/84-of-americans-buy-free-from-foods-because-they-believe-them-to-be-more-natural-or-less-processed. Accessed January 22, 2016.
- 30. Nielsen Web site. www.nielsen.com/us/en/insights/reports/2015/we-are-what-we-eat.html. Accessed January 22, 2016.



- 31. Gasparro A. Nestlé USA to Remove Artificial Ingredients From Candy. *Wall Street Journal*. Feb. 17, 2015.
- 32. The Hartman Group Web site. www.hartman-group.com/acumenPdfs/ingredients-avoid-v-seek-infographic-2015-05-14.pdf. Accessed January 22, 2016.
- 33. Mintel Web site. www.mintel.com/blog/food-market-news/like-oil-and-water-brominated-vegetable-oil-being-phased-out-in-the-us. Accessed January 22, 2016.
- 34. Mintel Web site. www.mintel.com/press-centre/food-and-drink/mintel-serves-up-5-key-food-and-drink-trends-for-2015. Accessed January 22, 2016.
- 35. Food Business News Web site.

 <u>www.foodbusinessnews.net/articles/news_home/Consumer_Trends/2015/09/How_consumers_define_clean_eat.aspx?ID=%7BA2B69947-B22D-48E1-AC70-C32011064786%7D</u>. Accessed January 22, 2016.
- 36. NPD Web site. www.npd.com/wps/portal/npd/us/news/press-releases/2015/real-clean-and-fresh-are-the-attributes-consumers-will-look-for-in-the-foods-they-eat-in-2016/. Accessed January 22, 2016.
- 37.U.S. Food and Drug Administration Web site. 21CFR101.22. www.gpo.gov/fdsys/pkg/CFR-2015-title21-vol2/xml/CFR-2015-title21-vol2-sec101-22.xml. Accessed January 22, 2016.



38. U.S. Food and Drug Administration Web site. 21CFR101.22. www.gpo.gov/fdsys/pkg/CFR-2015-title21-vol2/xml/CFR-2015-title21-vol2-sec101-100.xml. Accessed January 22, 2016.

- 39. Liaukonyte J, Streletskaya NA, Kaiser HM, et al. Consumer Response to "Contains" and "Free of" Labeling: Evidence from Lab Experiments. *Applied Economic Perspectives and Policy.* 2013 doi: 10.1093/aepp/ppt015.
- 40. Council For Agricultural Science And Technology. Process Labeling of Food: Consumer Behavior, the Agricultural Sector, and Policy Recommendations. October 2015. www.cast-science.org/download.cfm?PublicationID=283819&File=1030ac46417e576660 c87b6b2553352b6624TR. Accessed January 22, 2016.
- 41. Delish Web site. www.delish.com/cooking/g1586/food-additives/. Accessed January 22, 2016.
- 42. The Richest Web site. www.therichest.com/rich-list/the-biggest/unnatural-10-most-common-artificial-additives-to-avoid. Accessed January 22, 2016.
- 43. Kaptan B, Kayisoglu S. Consumers' Attitude towards Food Additives. *American Journal of Food Science and Nutrition Research*. 2015;2(2):21-25.