



THE FLAVOR AND EXTRACT
MANUFACTURERS ASSOCIATION
OF THE UNITED STATES

APRIL 2012

Respiratory Health and Safety in the Flavor Manufacturing Workplace

2012 Update

Page

2	Background
2	Workplace Safety Regulatory Requirements and Advice
2	Federal
3	California
3	Washington
4	Occupational Exposure Guidelines
4	Management and Employee Awareness Through Education, Training, and Hazard Communication
5	Hazard identification
5	Hazard communication - MSDSs
6	Hazard communication - labeling
7	Exposure Assessment and Control
7	Assessment of potential exposures
7	Exposure Control
8	Product substitution
8	Heating of Flavors
8	Facility structure and organization
8	Ventilation
9	Material handling - flavor compounding and packing activities
9	Material storage
9	Cleaning of vessels and work areas
10	Personal respiratory protection
10	Medical surveillance
12	Developing a Respiratory Safety Program for Flavor Manufacturing Facilities
12	References and Other Relevant Publications
15	Table 1
29	List of Acronyms

Background

In August, 2004, FEMA published the report [*Respiratory Health and Safety in the Flavor Manufacturing Workplace*](#) containing information to help flavor manufacturers maintain safe workplaces. This web-based report updates and supplements the August, 2004, report. Many documents referred to in this report may be accessed through the links provided, while other copyrighted reports may be retrieved through the usual appropriate means.

Maintaining safe and healthy workplaces is a matter of utmost importance to FEMA and its members. FEMA consulted with experts on the development of this report and made extensive use of a wide variety of information resources. The application of the information in this report to flavor manufacturing workplaces is a function of the specific aspects of workplaces and the products handled and manufactured therein. Because of the unique nature of individual flavor manufacturing workplaces, the information in this report should be used only as a general guide. FEMA strongly urges users of this report to consult with appropriate experts regarding the circumstances relevant to respiratory health and safety in flavor manufacturing facilities.

Like the August, 2004, report, this report is not a standard but is intended to share information that flavor manufacturers may use as they choose. FEMA is not responsible for the use or non-use of the information, or any action or failure to act, in any specific workplace based on reliance on this report. It is the responsibility of users of this report to verify information as it applies to specific workplaces before acting and to comply with all relevant local, state, and

federal laws and regulations.

The original 2004 report described three areas of emphasis for flavor manufacturers in maintaining the safest workplaces possible.

- Management and employee awareness through education and hazard communication
- Exposure assessment and control
- Medical surveillance

Since the 2004 report was published, important new information has become available on each of these areas of emphasis. This update provides a summary of important, new information but is not comprehensive in including every report or publication on any of the three subjects. Other information may be available, and FEMA urges flavor manufacturers to consult with appropriate experts in developing and implementing a respiratory health and safety program for their facilities.

Workplace Safety Regulatory Requirements and Advice

Federal

Although there is no federal standard specifically for the flavor manufacturing industry, the industry is subject to a number of OSHA regulatory requirements. As OSHA has made clear in its [National Emphasis Program for flavor manufacturing](#), the agency will rely heavily on the General Duty Clause of the Occupational

Safety and Health Act (Section 5(a)(1) of the Act) for broad authority over flavor manufacturing in general, and specifically for possible exposure to diacetyl. OSHA also implemented a [National Emphasis Program for microwave popcorn manufacturing plants](#).

Flavor manufacturers should be familiar with, and comply with, the [OSHA respirator standard](#). This standard remains the most common source of OSHA citations among flavor manufacturers.

The federal [Hazard Communication Standard](#) applies to many areas of flavor manufacturing. [OSHA issued specific guidance in 2007](#) on hazard communication as it relates to diacetyl. OSHA has informally stated that this guidance also applies to 2,3-pentanedione, although OSHA has not yet amended the 2007 guidance to include this flavoring substance.

[OSHA's confined space regulations](#) may apply to some flavor manufacturers employing large vessels or tanks in their operations.

OSHA and NIOSH maintain web pages on flavorings-related lung disease. The OSHA page can be found at www.osha.gov/SLTC/flavoringlung/index.html. The NIOSH page can be found at www.cdc.gov/niosh/topics/flavorings/current-niosh.html. NIOSH also maintains a science blog on diacetyl and food flavorings at www.cdc.gov/niosh/blog/nsb111008_diacetyl.html.

Important information is available through the NIOSH program of Health Hazard Evaluations (HHEs). These are evaluations conducted by NIOSH staff on a wide variety of workplaces. Several types of facilities relevant to the flavor industry have been evaluated by NIOSH, including microwave popcorn manufacturing plants and flavor manufacturing fa-

cilities. Reports of NIOSH HHEs are available on the NIOSH website at www.cdc.gov/niosh/hhe/.

California

In 2006, the California Division of Occupational Safety and Health (Cal/OSHA) implemented the innovative program the Flavor Industry Safety and Health Evaluation Program (FISHEP) with the assistance of FEMA. This program resulted in workplace safety consultations by Cal/OSHA with flavor manufacturers operating in California. FISHEP also resulted in the completion of serial spirometry testing of several hundred California flavor manufacturing workers. FISHEP assisted California flavor manufacturers in maintaining safe workplaces and also provided important information used in the development and adoption of the California flavor manufacturing workplace regulations.

In 2010, California implemented a [workplace safety regulation](#) specifically for flavor manufacturing in California. The regulation contains a “1% cut-off” for diacetyl in compounded flavors and contains provisions related to flavoring substances considered as “alternatives” to diacetyl – acetoin, 2,3-pentanedione, diacetyl trimer, 2,3-hexanedione, and 2,3-heptanedione. The state’s [final statement of reasons](#) explains the rationale for the regulations.

Washington

In 2008, the Washington State Department of Labor and Industries published an [information sheet for restaurant owners and workers about diacetyl](#). The sheet states that it “provides information to cooks and restaurant workers about possibly harmful exposures to diacetyl during cooking.”

Occupational Exposure Guidelines

Permissible Exposure Limits and Occupational Exposure Guidelines

Few flavoring substances have permissible exposure limits (PELs) promulgated by OSHA. PELs have the force of regulation and are an important way that OSHA regulates workplace safety. Table 1 lists priority flavoring substances with OSHA PELs. Diacetyl and related substances such as 2,3-pentanedione and acetoin have not been assigned PELs by OSHA.

Three groups have published voluntary occupational exposure guidelines for diacetyl, and in one case for 2,3-pentanedione. While they don't have the regulatory authority of an OSHA PEL, these guidelines can be helpful to flavor manufacturers.

The American Conference of Government Industrial Hygienists (ACGIH) publishes threshold limit values (TLVs) and short-term exposure limits (STELs) for a wide variety of commercial substances including a small number of flavoring substances. [In 2010, ACGIH proposed a TLV and STEL for diacetyl that were adopted in January, 2012.](#) The ACGIH TLV for diacetyl is 0.01 ppm/8 hr. time-weighted average (TWA) and the STEL is 0.02 ppm/15 minutes.

In 2010, the non-governmental non-profit organization Toxicology Excellence for Risk Assessment (TERA) proposed an occupational exposure limit for diacetyl of 0.2 ppm/8 hr. TWA (Maier et al., 2010).

[In its draft criteria document issued in 2011,](#) NIOSH recommended exposure limits for diacetyl and 2,3-pentanedione of 5 ppb/8

hr. TWA and 9.3 ppb/8 hr. TWA, respectively. NIOSH also recommended short-term exposure limits for each substance in the draft criteria document – 25 ppb/15 minutes for diacetyl and 31 ppb/15 minutes for 2,3-pentanedione.

The three groups publishing occupational exposure guidelines (TERA, ACGIH, and NIOSH) all employed different methods and data in arriving at their conclusions. Brosseau (2012) commented on the need for the application of consistent best practices in the development of occupational exposure guidelines.

Management and Employee Awareness Through Education, Training and Hazard Communication

A variety of information resources are available for management from both OSHA and NIOSH on their websites on flavorings-related lung disease. The OSHA page can be found at www.osha.gov/SLTC/flavoringlung/index.html. The NIOSH page can be found at www.cdc.gov/niosh/topics/flavoring. Among the available resources are an OSHA [worker alert](#) published in 2010 and a NIOSH [brochure for healthcare providers](#) published in 2011.

Thorough education and communication among both management and employees of flavor manufacturers are critical to the success of any workplace health and safety program. Formal, mandatory hazard communication and

training sessions assure that employees have the appropriate awareness of respiratory safety issues. Personnel in task areas that merit focus include:

- Personnel who blend and mix flavors in bulk quantities, especially those exposed to heated flavors, and the powder flavor and spray-dry manufacturing processes.
- Personnel who pack flavors (liquid or dry).
- Quality assurance personnel and flavorists who may have repeated exposure to flavors, even though exposure may be in smaller amounts.

Hazard identification

Over 2,700 chemically-defined flavoring substances and natural flavoring complexes are commonly used to formulate flavors. The U.S. Food and Drug Administration (FDA) is primarily responsible for the regulation of the addition of flavoring substances to food through several regulatory pathways (Hallagan and Hall, 2009). The vast majority of these materials have chemical and physical characteristics that would make it highly unlikely that they would pose a risk of respiratory injury in the workplace. Most of the materials are not very volatile and do not have a significant degree of reactivity. However, some low molecular weight chemically-defined flavoring substances may have sufficient volatility, and possibly reactivity, to pose a risk of respiratory injury when improperly handled. Table 1 contains a list of chemically-defined flavoring substances that may pose potential respiratory hazards when improperly handled.

The priority levels in Table 1 were assigned based on available inhalation exposure data in animals and humans, chemical structure,

volatility, and volume of use. In many instances, data on one flavoring substance was used to evaluate the priority level for other structurally-related substances. In many cases, relevant inhalation exposure information is limited and, on occasion, is available for flavoring substances that have other, larger-scale industrial uses. The assignment of priority levels required the application of significant judgment and also took into account anecdotal information provided by workers in the flavor industry who shared their valuable experience related to the handling of flavoring substances.

Since the original edition of this report was published in 2004, new information has become available on substances structurally-related to diacetyl resulting in their addition to the table as high priority substances. These substances include 2,3-pentanedione, 2,3-hexandione, 3,4-hexanedione, 2,3-heptandione, and diacetyl trimer. A few substances were reassigned from high priority to low priority based on new information. No substances were removed from Table 1.

Table 1 contains updated information on each flavoring substance from the FEMA Poundage Survey. Data from the 1995, 2005, and 2010 surveys are included and represent the amount of each flavoring substance estimated to “disappear” into the U.S. food supply in a given year. This information allows a rough estimate on the trends of use for the listed substances.

Hazard communication - MSDSs

Clear communication of potential hazards is of critical importance. [The OSHA Hazard Communications Standard](#) establishes a minimum for hazard communication through its material safety data sheet (MSDS) requirements.

The development and provision of MSDSs is an individual company obligation. FEMA does not, and has never, compiled or published MSDSs, or reviewed or approved company MSDSs. FEMA has assisted members in meeting their hazard communications obligations only through serving as a “library” for information.

FEMA members have access to the Flavor and Fragrance Ingredient Data Sheet (FFIDS) program that provides information on workplace hazards and that can be used as one source of information to formulate MSDSs on flavoring substances.

Another resource available to FEMA members is the IFRA/IOFI GHS Labeling Manual issued jointly by the International Fragrance Association (IFRA) and the International Organization of the Flavor Industry (IOFI). FEMA is a member of IOFI. The Labeling Manual provides information on the harmonization of hazard statements for hazard communications purposes according to the Globally Harmonized System of Classification and Labeling of Chemicals (the “GHS”). [OSHA published its final rule adopting the GHS in March 2012.](#)

Hazard communication - labeling

In addition to the various types of labeling requirements under relevant regulations, the labeling of bulk flavors may also be appropriate in certain circumstances to alert workers to potential respiratory hazards. It is recommended that the following bulk flavors bear a label using the language described below, or language that conveys a similar warning.

- Containers of “high priority” neat substances listed in Table 1 such as diacetyl, 2,3-pentanedione and acetaldehyde.

- Containers of compounded flavors (liquid and dry or powdered) or natural flavoring complexes that contain “high priority” flavoring substances in concentrations >1.0%.

- Any compounded flavors (liquid and dry or powdered) containing any flavoring substances listed in Table 1 in any concentrations if the compounded flavor or any of its individual flavoring substances will be heated during processing.

WARNING – This flavor may pose an inhalation hazard if improperly handled. Please contact your workplace safety officer before opening and handling, and read the MSDS. Handling of this flavor that results in inhalation of fumes, especially if the flavor is heated, may cause severe adverse health effects.

It is recommended that the following bulk flavors bear a label using the language described below, or language that conveys a similar warning.

- Containers of neat “low priority” chemically-defined substances as listed in Table 1.

- Containers of natural flavoring complexes known to contain chemically-defined flavoring substances listed in Table 1.

- Containers of compounded flavors (liquid and dry or powdered) containing “high priority” chemically-defined substances at concentrations <1.0%, or “low priority” chemically-defined substances at any level.

ATTENTION - Safe flavors can be used in an unsafe manner. Please contact your workplace safety officer before opening and handling this flavor, and read the MSDS.

Flavor manufacturers and suppliers cannot in all instances know how their customer will use a flavoring material. In many instances, the customer chooses to keep information related to how they will use a flavoring material confidential to protect valuable trade secret information related to their products. In other instances, customers may communicate to a supplier how they plan to use a flavor but then modify their plans. The warning statements suggested in this section provide a means for flavor manufacturers to assure their customers receive helpful information related to the safe handling and use of flavors.

Important resources on the subject of management and employee awareness include:

- [The OSHA Hazard Communications Standard.](#)
- [OSHA GHS Implementation Program.](#)
- [Hazard Communication Guidance for Diacetyl and Food Flavorings Containing Diacetyl \(OSHA, 2007\).](#)
- [Occupational Exposure to Flavoring Substances: Health Effects and Hazard Control \(OSHA, 2010\).](#)
- [OSHA National Emphasis Program for Facilities that Manufacture Food Flavorings Containing Diacetyl \(OSHA, 2011\).](#)

Exposure Assessment and Control

Assessment of potential exposures

Environmental monitoring is a necessary component of a sound respiratory health and safety program. Monitoring at regular intervals can provide valuable information on potential exposures. In addition to monitoring for the presence of certain vapors in air, monitoring of particulates, when appropriate, should also be conducted if reliable methods are available. A critical issue in any monitoring program is identifying the appropriate substances to monitor. NIOSH has focused on diacetyl and 2,3-pentanedione and provides a thorough discussion of analytical methods and sampling in the draft report [Occupational Exposure to Diacetyl and 2,3-Pentanedione \(NIOSH, 2011\)](#). Also relevant are reports by Cox-Gansler et al. (2011) and Day et al. (2011).

Exposure control

Flavor manufacturing facilities vary greatly in size, structure, age, manufacturing technologies employed, flavoring substances stocked and handled, types of flavors manufactured, and many other characteristics. Manufacturing processes may range from simple blending and packaging to more complicated processes that include repeated heating of flavoring substances that are mixed and heated again, extraction at room temperature and with heat, and other processes that may result in significant opportunities for exposure.

Some facilities have extensive automated processes that minimize opportunities for exposure while others have little automation and

rely on workers manipulating large quantities of flavors and other materials by hand during the formulation and packing processes. Because of this great variability, “one size fits all” solutions to material handling strategies and engineering controls is inappropriate. However, a number of basic principles can greatly reduce opportunities for hazardous exposures.

Product substitution

Awareness that specific substances are hazardous often results in the application of a simple industrial hygiene strategy – product substitution. Product substitution may be employed when it is known that a substance is hazardous and can be replaced with one that isn’t. The unique nature of the flavor imparted by certain substances, many of which are naturally occurring constituents of food, makes it difficult to identify substitutes that are effective.

In 2010 OSHA published a [“Worker Alert” on diacetyl and substitutes](#) noting that “some diacetyl substitutes may also cause harm.” The Alert stated, “Diacetyl substitutes that have not been proven to be safe include diacetyl trimer, 2,3-hexanedione, 2,3-heptanedione, and 2,3-pentanedione.” In January, 2011, NIOSH requested information on flavoring substances that may be used as substitutes for diacetyl ([76 Fed. Reg. 1434, 10 January 2011](#)). [FEMA provided information to NIOSH](#) on six flavoring substances that it was aware of that may serve as substitutes for diacetyl:

- Four alpha-diketone substances: 2,3-pentanedione, 2,3-hexanedione, 3,4-hexanedione, and 2,3-heptanedione.
- Two other related substances: acetoin and diacetyl trimer.

NIOSH published results from environmental monitoring at a microwave popcorn production plant focusing on the possible presence of “diacetyl substitutes” (Boylstein, 2012).

Heating of flavors

Heating of flavors is of particular concern with regard to potential hazardous exposures. Heating will increase volatility and greatly increase air concentrations of flavoring substances. Mixing of heated flavors should be conducted in closed vessels with local ventilation. Workers should not open heated vessels to conduct visual inspections in such a way as to create an opportunity for exposure. In instances when workers must work near open vessels that are heated and cannot be closed or do not have local ventilation, their exposures should be promptly evaluated by environmental sampling. If exposures are elevated then the proper personal protective equipment should be employed.

Facility structure and organization

Opportunities for exposure can be greatly decreased by segregating functions that involve the handling of flavors from functions that do not. For example, a flavor compounding, packing, or shipping area should not share space with a sales office. Flavor production areas should be separate from non-production areas and they should not share the same air handler.

Ventilation

Flavoring substances and mixtures, whether liquid or dry, must be handled in such a way as to minimize the creation of airborne aerosols or particulate matter. This means that mixing, blending, and other physical manipulation activities should be performed in closed

systems when possible. When systems must remain open, local (“spot”) ventilation (e.g. “elephant trunks”) should be used. Fume hoods are commonly used in research and development laboratories. Dilution through general room ventilation seldom results in exposure reduction unless extremely high volumes of air are circulated.

Material handling – flavor compounding and packing activities

Simple flavor compounding activities such as mixing or pouring can result in significant exposures. In most instances, mixing of liquid and dry flavors should be conducted in fully or partially closed vessels with local ventilation. Opportunities for the generation of airborne particles and aerosols should be minimized.

Proper pouring techniques for liquid and dry flavors can greatly reduce opportunities for exposure. For liquid flavoring substances, techniques can be adopted that pipe material into mixing vessels so that workers do not have to pour. In some instances it is appropriate to pipe in liquids below the surface of solutions in vessels to minimize splashing. This is particularly important for volatile substances.

For dry and powdered flavors, pouring should be conducted in such a way that the generation of airborne particulates is minimized. Simple, proper pouring techniques such as pouring slowly close to the mixing vessel can greatly minimize airborne particulates. Mixing ingredients in an order in which dry ingredients are added last to liquid mixtures also can minimize particulate generation. Local exhaust is the most effective control for these operations. Systems can be designed that will allow easy pouring and at the same time control exposures.

Packaging activities can result in significant opportunities for exposure, especially when dry flavors are filled into bags, boxes, or drums under pressure. Closed systems should be used when possible. But unless there is an unusually high degree of automation, workers will have opportunities for exposure as filled containers must be replaced with empty ones. The use of personal protective equipment may then need to be considered to minimize exposure.

Material storage

Flavoring substances that are volatile should be stored in cooled storage areas. Substances such as acetaldehyde are often stored in cooled rooms, and are often also used in flavor manufacture in a cooled state. Liquid and dry and powdered flavors should ideally be stored in store-rooms with their own air handler that has minimum recirculation. In some instances, flavor facilities have negative air flow in storage areas to reduce opportunities for exposure.

Cleaning of vessels and work areas

Cleaning of process vessels that contained liquid flavors or viscous mixtures, or work areas with spilled material, especially with steam or heated water, may create opportunities for exposure to flavoring substances. Similarly, cleaning vessels or areas used to manufacture or mix powdered flavors with compressed air may also result in airborne particulates.

It is important that cleaning activities be conducted in a manner that does not result in significant air concentrations of flavors and other materials present in the vessel. Cleaning areas should be isolated and contained to prevent the dissemination of airborne flavors. Automated cleaning processes will greatly reduce opportu-

nities for exposure. In some instances, the most effective way to protect workers responsible for cleaning activities will be to use respirators.

It is also important that adequate care be exercised if workers are to enter or partially enter equipment in order to clean it. In addition to concerns about possible respiratory exposures, in some instances, cleaning activities involving vessel entry may be subject to the requirements of OSHA's confined space regulations (29 CFR 1910.146).

Personal respiratory protection

The implementation of appropriate process and engineering controls is preferable to simply providing employees with personal respiratory protection. However, respirators do have a role in many respiratory health and safety programs. Critical to their success is the selection of the proper respirator for the conditions present in a given facility, the proper fit of that respirator to the person using it, and the training in its use, maintenance, and storage. OSHA also requires that employees wearing most types of respirators undergo medical clearance prior to their use.

In terms of specific duty requirements relevant to protection against respiratory hazards, OSHA regulations require that personal protective equipment must be provided to employees whenever necessary to address chemical or other hazards which are "capable of causing injury or impairment in the function of any part of the body through absorption, inhalation or physical contact." Under OSHA's [Respiratory Protection Standard](#), the "primary objective shall be to prevent atmospheric contamination." Where, however, that is not feasible through engineering and process controls, respirators shall

be used. The regulations contain a range of requirements, including the proper selection of respirators, standard procedures for use, training of employees, respirator maintenance, and other safety measures. The standard and relevant background information were published in the *Federal Register* notice announcing the standard ([63 Fed. Reg. 1152, 8 January 1998](#)).

Important resources on the subjects of exposure assessment and control include:

- Diacetyl exposures in the flavor manufacturing industry (Martyny et al., 2008).
- Evaluation of a local exhaust ventilation system for controlling exposures during liquid flavor production (Dunn et al., 2008).
- [Occupational Exposure to Flavoring Substances: Health Effects and Hazard Control \(OSHA, 2010\)](#).
- [OSHA National Emphasis Program for Facilities that Manufacture Food Flavorings Containing Diacetyl \(OSHA, 2011\)](#).
- [OSHA National Emphasis Program for Microwave Popcorn Processing Plants \(OSHA, 2011\)](#).
- [Draft Criteria Document - Occupational Exposure to Diacetyl and 2,3-Pentanedione \(NIOSH, 2011\)](#).
- Occupational lung disease risk and exposure to butter-flavoring chemicals after implementation of controls at a microwave popcorn plant (Kanwal et al., 2011).

Medical surveillance

Medical surveillance is a key component of an effective respiratory health and safety program in the flavor industry. This is especially the case when it is difficult to identify a specific causative agent for an observed effect and when

symptoms and/or lung function abnormalities may be the first clue to an exposure-related problem. Appropriate medical surveillance can identify health issues before progression to severe illness occurs and when opportunities for reducing or eliminating exposure exist.

Medical surveillance should include an evaluation at the time of hire and at least annually thereafter. The exam should include both a medical and occupational history and a pulmonary function component. Spirometry is a simple and inexpensive way to monitor pulmonary function status and should be included in the exam at hire and in follow-up exams thereafter. It is important that spirometry testing follow the most recent American Thoracic Society guidelines for accurate testing.

A sound medical surveillance program will facilitate the identification of respiratory symptoms and lung function abnormalities. As reported by NIOSH, some workers in microwave popcorn manufacturing facilities, and in a few flavor manufacturing facilities, exhibited findings of fixed airway obstruction manifested by symptoms of cough (often without the production of phlegm) and shortness of breath after exertion as well as spirometric abnormalities (e.g. decreased FEV-1, a parameter of airflow). Frequent or persistent symptoms of eye, nose, throat or skin irritation have also been reported by NIOSH in some affected workers. A plan should be in place to refer employees for further medical follow-up and evaluation if such symptoms and lung function abnormalities are identified in the surveillance examinations or if there are significant unexplained declines in employee lung function as measured by periodic spirometry.

It is particularly important to note that

bronchiolitis obliterans, a very serious lung disease, has been implicated in cases of respiratory illness seen in microwave popcorn manufacturing plants and in flavor manufacturing facilities. Early detection of symptoms and spirometric abnormalities through a medical surveillance program will allow workers to seek timely follow-up and may prevent progression of disease. Early detection is especially important with bronchiolitis obliterans because the disease is difficult to treat.

Important information resources on the subject of medical surveillance include:

[Establishing a quality medical surveillance program in your flavor company.](#) Rose C. and Sells T. National Jewish Health. Presentation at the conference “Respiratory Health and Safety in the Flavor Manufacturing Workplace – A Training Session for Flavor Plant Staff.” FEMA. 2005.

[Occupational Exposure to Flavoring Substances: Health Effects and Hazard Control](#) (OSHA, 2010).

[Occupational Exposure to Diacetyl and 2,3-Pentanedione](#) (NIOSH, 2011).

Developing a Respiratory Safety Program for Flavor Manufacturing Facilities

FEMA and a number of its members have worked with National Jewish Health in Denver, Colorado (NJH) through Cecile Rose, M.D., M.P.H., and Mike VanDyke, Ph.D., C.I.H. Dr. Rose is an expert in pulmonary diseases, and Dr. VanDyke is a certified industrial hygienist. Both have worked extensively with the flavor industry and have significant experience with flavors and flavor manufacturing. Other sources of expert assistance include occupational medicine physicians and industrial hygiene consultants located in the communities where flavor manufacturing facilities are located. Among other sources of information, consultants can be identified through the consultant directory maintained by the American Industrial Hygiene Association (www.aiha.org).

National Jewish Health was awarded a grant by OSHA in late 2011 to conduct training on respiratory health and safety programs for flavor manufacturers. NJH has been provided funding by OSHA to train 500 people working in flavor manufacturing. The focus is expected to be on workers employed in direct flavor manufacturing, i.e. workers responsible for handling and formulating flavorings. Please contact John Hallagan of FEMA for information (202.331.2333; Hondobear@aol.com).

References and Other Relevant Publications

Since 2004, a number of reports have been published relevant to the safety of possible exposure to flavoring substances in the workplace and other aspects of respiratory health and safety in flavor manufacturing. References for publications cited in the body of this report may be found in this list in addition to publications not cited but of interest.

Auttachoat W., White K.L. and Guo T.L. Diacetyl, an artificial flavoring compound in popcorn causes contact sensitization in female BALB/c mice. *The Toxicologist*, 96, 239. 2007.

Boylstein R. Identification of diacetyl substitutes at a microwave popcorn production plant. *Journal of Occupational and Environmental Hygiene*. 9, D33. 2012.

Brosseau L.M. Occupational exposure guidelines - why process is as important as science. *Journal of Occupational and Environmental Hygiene*. 8, D109. 2012.

Cox-Ganser J., Ganser G., Saito R., Hobbs G., Boylstein R., Hendricks W., Simmons M., Eide M., Kullman G. and Piacitelli C. Correcting diacetyl concentrations from air samples collected with NIOSH method 2557. *Journal of Occupational and Environmental Hygiene*. 8, 59. 2011.

Day G., LeBouf R., Grote A., Pendergrass S., Cummings K., Kreiss K. and Kullman G. Identification and measurement of diacetyl substitutes

in dry bakery mix production. *Journal of Occupational and Environmental Hygiene*. 8, 93. 2011.

Dunn K.H., Garcia A. and Echt A. Evaluation of a local exhaust ventilation system for controlling exposures during liquid flavoring production. *Journal of Occupational and Environmental Hygiene*. 5, D103. 2008.

Fedan J.S., Dowdy J.A., Fedan K.B. and Hubbs, A.F. Popcorn worker's lung: In vitro exposure to diacetyl, an ingredient in microwave popcorn butter flavoring, increases reactivity to methacholine. *Toxicol. Appl. Pharmacol.*, 215, 17. 2006

Flake G.P., Kirby P.J., Price H.C., Gage P.S., Kelly F.L., Palmer S.M., Foley J.F. and Morgan D.L. Bronchiolitis obliterans-like lesions in rats treated with diacetyl, acetoin, or acetyl propionyl by intratracheal instillation. *The Toxicologist*, 114, 316. 2010.

Hallagan J.B. and Hall R.L. Under the conditions of intended use – new developments in the FEMA GRAS program and the safety assessment of flavor ingredients. *Food & Chemical Toxicology*. 47, 267. 2009.

Harber P., Saechao K. and Boomus C. Diacetyl-induced lung disease. *Toxicol. Rev.* 25, 261. 2006.

Hubbs A.F., Battelli L.A., Goldsmith W.T., Porter D.W., Frazer D., Friend S., Schwegler-Berry D., Mercer R.R., Reynolds J.S., Grote A., Castranova V., Kullman G., Fedan J.S., Dowdy J., and Jones, W.G. Necrosis of nasal and airway

epithelium in rats inhaling vapors of artificial butter flavoring. *Toxicol. Appl. Pharmacol.* 185, 128. 2002.

Hubbs A.F., Battelli L.A., Mercer R.R., Kashon M., Friend S., Schwegler-Berry D., and Goldsmith W.T. Inhalation toxicity of the flavoring agent, diacetyl (2,3-butanedione) in the upper respiratory tract of rats. *The Toxicologist* 78(S-1), 438. 2004.

Hubbs AF, Goldsmith W.T., Kashon M.L., Frazer D., Mercer R.R., Battelli L.A., Kullman G.J., Schwegler-Berry D., Friend, S. and Castranova V. Respiratory toxicologic pathology of inhaled diacetyl in Sprague-Dawley rats. *Toxicol. Pathol.* 36, 330. 2008.

Hubbs A.F., Moseley A.E., Goldsmith W.T., Jackson M.C., & Kashon M.L. Airway epithelial toxicity of the flavoring agent, 2,3-pentanedione. *The Toxicologist* 114(1): 319. 2010.

Kanwal R., Kullman G., Piacitelli C., Boylstein R., Sahakian N., Martin S., Fedan K. and Kreiss K. Evaluation of flavorings-related lung disease risk at six microwave popcorn plants. *J. Occup. Environ. Med.*, 48, 149. 2006.

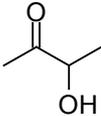
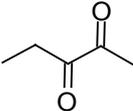
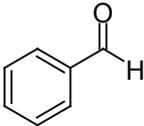
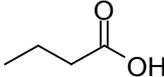
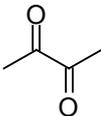
King T.E. and Kinder B.W. Bronchiolitis. In *Fishman's Pulmonary Diseases and Disorders*, Vol. 1, Chapter 52. Fishman et al. Eds. 4th Ed. 2008.

Kreiss K., Gomaa A., Kullman G., Fedan K., Simoes E.J. and Enright P.L. (2002) Clinical bronchiolitis obliterans in workers at a microwave-popcorn plant. *N. Engl. J. Med.* 347, 330. 2002.

- Kovacic P. and Cooksy A.L. Electron transfer as a potential cause of diacetyl toxicity in popcorn lung disease. *Rev. Environ. Contam. Toxicol.*, 204,133. 2010.
- Maier A., Kohrman-Vincent M., Parker A., and Haber L. Evaluation of concentration-response options for diacetyl in support of occupational risk assessment. *Regulatory Toxicology & Pharmacology*. 58, 285. 2010.
- Mathews J. M., Burgess J. P., Snyder R. W., Etheridge A. S., Burka L. T. Reaction of the butter flavorant butanedione with N-R-acetylglycine. *Drug Metab. Rev.*, 37, 273. 2005.
- Mathews J.M., Watson S.C., Snyder R.W., Burgess J.P. and Morgan D.L. Reaction of the butter flavorant diacetyl (2,3-butanedione) with N-r-acetylglycine: A model for epitope formation with pulmonary proteins in the etiology of obliterative bronchiolitis. *J. Agric. Food Chem.*, 58,12761. 2010.
- Morgan D.L., Flake G.P., Kirby P.J. and Palmer S.M. Respiratory toxicity of diacetyl in C57B1/6 mice. *Toxicol. Sci.*, 103(1): 169-180. 2008.
- Morgan D.L., Kirby P.J., Price H.C., Bousquet R.W., Taylor G.J., Gage N. and Flake G.P. Inhalation toxicity of acetyl propionyl in rats and mice. *The Toxicologist*, 114, 316. 2010.
- Morris J.B. and Hubbs A.F. Inhalation dosimetry of diacetyl and butyric acid, two components of butter flavoring vapors. *Toxicol. Sci.* 108(1): 173. 2009.
- Gloede E., Cichoki J.A., Baldino J.B. and Morris J.B. A validated hybrid computational fluid dynamics-physiologically based pharmacokinetic model for respiratory tract vapor absorption in the human and rat and its application to inhalation dosimetry of diacetyl. *Toxicological Sciences*. Advanced access. 24 June 2011.
- VanRooy F.G.B.G.J., Rooyackers J.M., Prokop M., Houba R., Smit A.M. and Heederik D.J.J. Bronchiolitis obliterans syndrome in chemical workers producing diacetyl for food flavorings. *Am. J. Respir. Crit. Care Med.*, 176(5): 498-504. 2007.
- VanRooy F.G.B.G.J., Smit L.A.M., Houba R., Zaat V.A.C., Rooyackers J.M. and Heederik D.J.J. A cross sectional study on lung function and respiratory symptoms among chemical workers producing diacetyl for food flavourings. *Occup Environ Med*, 66,105. 2009.
- Zviely M. Molecule of the month: 2,3-pentanedione. *Perfumer & Flavorist*. 34(7), 20. 2009.

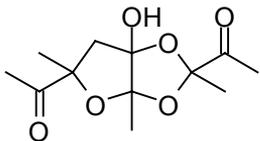
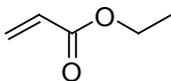
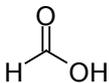
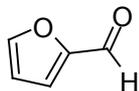
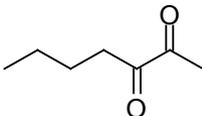
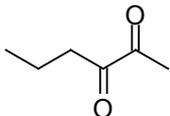
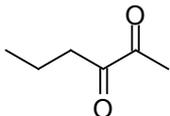
THE FLAVOR AND EXTRACT MANUFACTURERS ASSOCIATION

TABLE 1

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
High Priority						
2003	75-07-0	Acetaldehyde 	44.05	759 mm Hg 20 °C 910 mm Hg 25 °C	OSHA PEL - TWA 200 ppm, 360 mg/m3	1995: 321,000 2005: 70,200 2010: 59,000
2006	64-19-7	Acetic acid 	60.05	12.9 mm Hg 20 °C 17.2 mm Hg 25 °C	OSHA PEL - TWA 10 ppm, 25 mg/m3	1995: 310,000 2005: 271,000 2010: 373,000
2008	513-86-0	Acetoin 	88.11	1.36 mm Hg 20 °C 2 mm Hg 25 °C	NA	1995: 116,000 2005: 133,000 2010: 150,000
2841	600-14-6	2,3-Pentanedione 	100.12	23.8 mm Hg 20 °C 31.1 mm Hg 25 °C	NIOSH REL- TWA 9.3 ppb NIOSH STEL 31 ppb	1995: 2,600 2005: 4,590 2010: 38,000
2127	100-52-7	Benzaldehyde 	106.13	0.705 mm Hg 20 °C 1.01 mm Hg 25 °C	NA	1995: 603,000 2005: 1,410,000 2010: 578,000
2221	107-92-6	Butyric Acid 	88.11	1.5 mm Hg 20 °C 2.11 mm Hg 25 °C	NA	1995: 180,000 2005: 130,000 2010: 229,000
2370	431-03-8	Diacetyl 	86.09	55 mm Hg 20 °C 70.2 mm Hg 25 °C	NIOSH REL- TWA 5 ppb NIOSH STEL 25 ppb TERA OEL- TWA 0.2 ppm	1995: 211,000 2005: 228,000 2010: 85,000

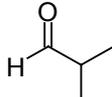
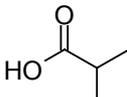
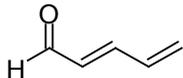
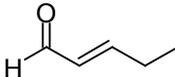
RESPIRATORY HEALTH AND SAFETY IN THE FLAVOR MANUFACTURING WORKPLACE

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
4303	18114-49-3	Diacetyl trimer	258.27	NA	See Diacetyl	1995: NA 2005: NA 2010: 0
						
2418	140-88-5	Ethyl acrylate	100.12	29.4 mm Hg 20 °C 38.4 mm Hg 25 °C	PEL - Skin TWA 25 ppm, 100 mg/m ³	1995: 11 2005: 3 2010: 59
						
2487	64-18-6	Formic Acid	46.03	27.4 mm Hg 20° C 35.9 mm Hg 25 °C	OSHA PEL - TWA 5 ppm, 9 mg/m ³	1995: 20,600 2005: 7,150 2010: 9,500
						
2489	98-01-1	Furfural	96.09	1.65 mm Hg 20 °C 2.32 mm Hg 25 °C	OSHA PEL - Skin TWA 5 ppm, 20 mg/m ³	1995: 7,710 2005: 8,240 2010: 9,300
						
2543	96-04-8	2,3-Heptanedione	128.17	0.785 mm Hg 20 °C 1.11 mm Hg 25 °C	NA	1995: 100 2005: 78 2010: 120
						
2558	3848-24-6	2,3-Hexanedione	114.14	2.11 mm Hg 20 °C 2.93 mm Hg 25 °C	NA	1995: 620 2005: 400 2010: 2,500
						
3168	4437-51-8	3,4-Hexanedione	114.14	9.2 mm Hg 20 °C 12.3 mm Hg 25 °C	NA	1995: 39 2005: 13 2010: 200
						

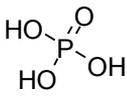
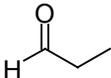
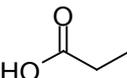
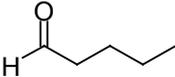
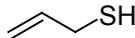
THE FLAVOR AND EXTRACT MANUFACTURERS ASSOCIATION

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
3779	7783 - 06 4	Hydrogen sulfide 	34.08	3.81 x 10 ⁻⁹ mm Hg 25 °C	OSHA PEL - Exposures shall not exceed 20 ppm (ceiling) with the following exception: if no other measurable exposure occurs during the 8-hour work shift, exposures may exceed 20 ppm, but not more than 50 ppm (peak), for a single time period up to 10 minutes NIOSH REL - 10 ppm, ceiling limit 15 mg/m ³ (10 minutes)	1995: 3 2005: 210 2010: 22
2220	78-84-2	Isobutyraldehyde 	72.11	131 mm Hg 20 °C 164 mm Hg 25 °C	NA	1995: 1,100 2005: 1,010 2010: 1,400
2222	79-31-2	Isobutyric acid 	88.11	2.35 mm Hg 20 °C 3.27 mm Hg 25 °C	NA	1995: 8,750 2005: 6,840 2010: 8,700
2716	74-93-1	Methyl mercaptan 	48.11	1.28 X 10 ³ mm Hg 20 °C 1510 mm Hg 25 °C	OSHA PEL - ceiling limit TWA 10 ppm, 20 mg/m ³	1995: 52 2005: 240 2010: 400
2746	75-18-3	Methyl sulfide 	62.13	391 mm Hg 20 °C 479 mm Hg 25 °C	NA	1995: 8,200 2005: 22,900 2010: 17,000
3217	764-40-9	2,4-Pentadienal 	82.1	20 mm Hg 25 °C	NA	1995: 0 2005: 0 2010: 0
3218	764-39-6	2-Pentenal 	84.12	13.9 mm Hg 20 °C 18.5 mm Hg 25 °C	NA	1995: 3 2005: 24 2010: 54

RESPIRATORY HEALTH AND SAFETY IN THE FLAVOR MANUFACTURING WORKPLACE

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
2900	7664-38-2	Phosphoric acid 	67.02	6.09 x 10 ⁻¹¹ mm Hg 25 °C	OSHA PEL - TWA 1 mg/ m ³	1995: 4,840,000 2005: 5,780,000 2010: 423,000
2923	123-38-6	Propionaldehyde 	58.08	256 mm Hg 20 °C 317 mm Hg 25 °C	NA	1995: 3,870 2005: 590 2010: 700
2924	79-09-4	Propionic acid 	74.08	4.41 mm Hg 20 °C 6.04 mm Hg 25 °C	NIOSH REL- TWA 10 ppm, 30 mg/m ³	1995: 44,400 2005: 24,500 2010: 33,000
3039	9/5/7446	Sulfur dioxide 	64.06	2600 mm Hg 25 °C	OSHA PEL - TWA 5 ppm, 13 mg/m ³	1995: 4,100 2005: 470 2010: 330
3241	75-50-3	Trimethylamine 	59.11	1680 mm Hg 25 °C	NIOSH REL - TWA 10 ppm; STEL 15 ppm	1995: 870 2005: 640 2010: 3200
3098	110-62-3	Valeraldehyde 	86.13	25.1 mm Hg 20 °C 32.9 mm Hg 25 °C	NOSH REL -TWA 50 ppm, 176 mg/m ³	1995: 140 2005: 150 2010: 1600
Low Priority						
3326	67-64-1	Acetone 	58.08	249 mm Hg 25 °C	OSHA PEL - TWA 1000 ppm, 2400 mg/m ³	1995: 400 2005: 360 2010: 370
2035	870-23-5	Allyl mercaptan 	74.14	161 mm Hg 25 °C	NA	1995: 1 2005: 1 2010: 1

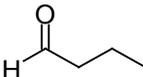
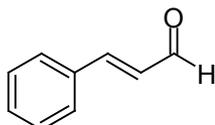
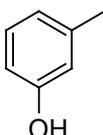
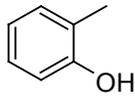
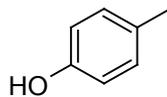
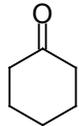
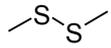
THE FLAVOR AND EXTRACT MANUFACTURERS ASSOCIATION

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
2053	12124-99-1	Ammonium sulfide <chem>(NH4)2S</chem>	68.15	9.2 x 10-15 mm Hg 25 °C	NA	1995: 660 2005: 600 2010: 1300
3965	78-96-6	1-Amino-2-propanol <chem>CC(N)CO</chem>	75.13	0.768 mm Hg 25 °C	NA	1995: NA 2005: 0 2010: 0
3616	108-98-5	Benzenethiol <chem>c1ccccc1S</chem>	110.18	1.63 mm Hg 25 °C	"NIOSH REL-ceiling limit TWA 0.1 ppm, 0.5 mg/m3 (15 minutes)"	1995: 4 2005: 1 2010: 3800
2147	100-53-8	Benzyl mercaptan <chem>c1ccccc1CS</chem>	124.21	0.3 mm Hg 20 °C 0.474 mm Hg 25 °C	NA	1995: 0 2005: 0.1 2010: 0.1
3129	92-52-4	Biphenyl <chem>c1ccc(cc1)-c2ccccc2</chem>	154.21	0.00419 mm Hg 20 °C 0.00749 mm Hg 25 °C	OSHA PEL - TWA 0.2 ppm, 1mg/m3	1995: 0 2005: 0 2010: 0
2170	78-93-3	2-Butanone <chem>CCC(=O)C</chem>	72.11	77.9 mm Hg 20 °C 98.5 mm Hg 25 °C	OSHA PEL - TWA 200 ppm, 590 mg/m3	1995: 530 2005: 960 2010: 1200
2174	123-86-4	Butyl acetate <chem>CCCC(=O)OC</chem>	116.16	8.85 mm Hg 20 °C 11.9 mm Hg 25 °C	OSHA PEL- TWA 150 ppm, 710 mg/m3	1995: 26,300 2005: 36,700 2010: 37,000
2178	71-36-3	Butyl Alcohol <chem>CCCCO</chem>	74.12	5.49 mm Hg 20 °C 7.78 mm Hg 25 °C	OSHA PEL - TWA 100 ppm, 300 mg/m3	1995: 13,300 2005: 6,580 2010: 7,200
3130	109-73-9	Butylamine <chem>CCCCN</chem>	73.14	71 mm Hg 20 °C 94.7 mm Hg 25 °C	OSHA PEL - TWA 5 ppm, 15 mg/m3 ceiling; skin	1995: 0 2005: 0 2010: 0

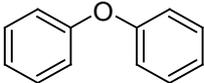
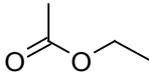
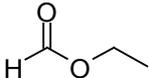
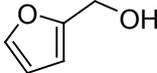
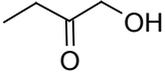
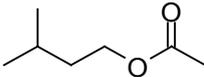
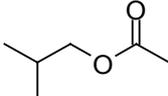
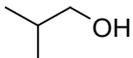
RESPIRATORY HEALTH AND SAFETY IN THE FLAVOR MANUFACTURING WORKPLACE

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
2219	123-72-8	Butyraldehyde 	72.11	84.9 mm Hg 20 °C 108 mm Hg 25 °C	NA	1995: 340 2005: 240 2010: 100
2286	104-55-2	Cinnamaldehyde 	132.16	0.0214 mm Hg 20 °C 0.0337 mm Hg 25 °C	NA	1995: 993,000 2005: 683,000 2010: 421,000
3530	108-39-4	m-Cresol 	108.14	0.109 mm Hg 20 °C 0.167 mm Hg 25 °C	OSHA PEL - Skin TWA 5 ppm, 22 mg/m ³ . OSHA standard is for all cresols combined under CAS Number 1319-77-3 (Federal Register 7/5/89).	1995: 0 2005: 0.1 2010: 0
3480	95-48-7	o-Cresol 	108.14	0.15 mm Hg 20 °C 0.25 mm Hg 25 °C	OSHA PEL - Skin TWA 5 ppm, 22 mg/m ³ . OSHA standard is for all cresols combined under CAS Number 1319-77-3 (Federal Register 7/5/89).	1995: 1 2005: 0.1 2010: 2
2337	106-44-5	p-Cresol 	108.14	0.073 mm Hg 20 °C 0.124 mm Hg 25 °C	OSHA PEL - Skin TWA 5 ppm, 22 mg/m ³ . OSHA standard is for all cresols combined under CAS Number 1319-77-3 (Federal Register 7/5/89).	1995: 17 2005: 170 2010: 810
3909	108-94-1	Cyclohexanone 	98.15	2.94 mm Hg 20 °C 4.04 mm Hg 25 °C	OSHA PEL - TWA 50 ppm, 200 mg/m ³	1995: NA 2005: 0 2010: 0
3536	624-92-0	Dimethyl disulfide 	94.2	18.5 mm Hg 20 °C 24.5 mm Hg 25 °C	NA	1995: 4,170 2005: 32 2010: 3,600

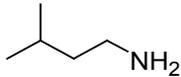
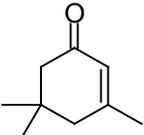
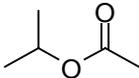
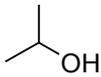
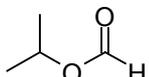
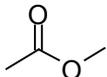
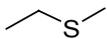
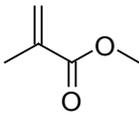
THE FLAVOR AND EXTRACT MANUFACTURERS ASSOCIATION

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
3667	101-84-8	Diphenyl ether 	170.21	0.00973 mm Hg 20 °C 0.017 mm Hg 25 °C	OSHA PEL - Vapor TWA 1ppm, 7 mg/m ³	1995: 86 2005: 280 2010: 4,000
2414	141-78-6	Ethyl acetate 	88.11	77 mm Hg 20 °C 98.3 mm Hg 25 °C	OSHA PEL - TWA 400 ppm, 1400 mg/m ³	1995: 462,000 2005: 514,000 2010: 460,000
2419	64-17-5	Ethyl alcohol 	46.07	45.3 mm Hg 20 °C 60.9 mm Hg 25 °C	OSHA PEL - TWA 1000 ppm, 1900 mg/m ³	1995: 23,500,000 2005: 30,600,000 2010: 20,200,000
2434	109-94-4	Ethyl formate 	74.08	198 mm Hg 20 °C 246 mm Hg 25 °C	OSHA PEL - TWA 100 ppm, 300 mg/m ³	1995: 18,600 2005: 21,400 2010: 19,000
2491	98-00-0	Furfuryl alcohol 	98.1	0.267 mm Hg 20 °C 0.409 mm Hg 25 °C	OSHA PEL - TWA 50 ppm, 200 mg/m ³	1995: 410 2005: 560 2010: 1,000
3173	5077-67-8	1-Hydroxy-2-butanone 	88.11	0.768 mm Hg 25 °C	NA	1995: 0 2005: 0 2010: 0
2055	123-92-2	Isoamyl acetate 	130.19	4.13 mm Hg 20 °C 5.67 mm Hg 25 °C	OSHA PEL - TWA 100 ppm, 525 mg/m ³	1995: 441,000 2005: 372,000 2010: 270,000
2175	110-19-0	Isobutyl acetate 	116.16	13.7 mm Hg 20 °C 18.3 mm Hg 25 °C	OSHA PEL - TWA 150 ppm, 700 mg/m ³	1995: 35,600 2005: 39,300 2010: 60,000
2179	78-83-1	Isobutyl alcohol 	74.12	9.55 mm Hg 20 °C 13.4 mm Hg 25 °C	OSHA PEL - TWA 100 ppm, 300 mg/m ³	1995: 27,700 2005: 20,000 2010: 8,800

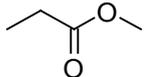
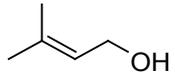
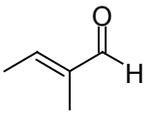
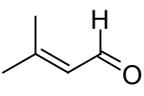
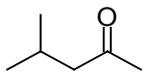
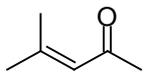
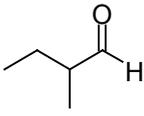
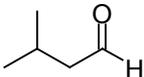
RESPIRATORY HEALTH AND SAFETY IN THE FLAVOR MANUFACTURING WORKPLACE

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
3219	107-85-7	Isopentylamine 	87.17	40 mm Hg 20 °C 44.5 mm Hg 25 °C	NA	1995: 1 2005: 0 2010: 0.4
3553	78-59-1	Isophorone 	138.2	0.169 mm Hg 20 °C 0.249 mm Hg 25 °C	OSHA PEL - TWA 25 ppm, 140 mg/m3	1995: 2 2005: 220 2010: 3
2926	108-21-4	Isopropyl acetate 	102.13	47.1 mm Hg 20 °C 60.8 mm Hg 25 °C	OSHA PEL - TWA 250 ppm, 950 mg/m3	1995: 1,180 2005: 70 2010: 190
2929	67-63-0	Isopropyl alcohol 	60.1	36.7 mm Hg 20 °C 49.6 mm Hg 25 °C	OSHA PEL - TWA 400 ppm, 980 mg/m3	1995: 176,000 2005: 14,300 2010: 96,000
2944	625-55-8	Isopropyl formate 	88.11	110 mm Hg 20 °C 142 mm Hg 25 °C	NA	1995: 0 2005: 0 2010: 0
2676	79-20-9	Methyl acetate 	74.08	40.6 mm Hg 20 °C 52.7 mm Hg 25 °C	OSHA PEL - TWA 200 ppm, 610 mg/m3	1995: 1,060 2005: 1,370 2010: 1,500
3860	624-89-5	Methyl ethyl sulfide 	76.16	150 mm Hg 25 °C	NA	1995: NA 2005: 0 2010: 0
4002	80-62-6	Methyl methacrylate 	100.12	36.7 mm Hg 25 °C	OSHA PEL - TWA 100 ppm, 410 mg/m3	1995: NA 2005: 0 2010: 0

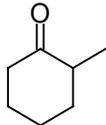
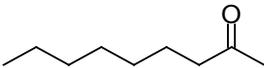
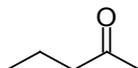
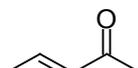
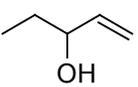
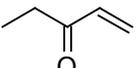
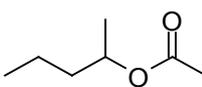
THE FLAVOR AND EXTRACT MANUFACTURERS ASSOCIATION

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
2742	554-12-1	Methyl propionate 	88.11	68.8 mm Hg 20 °C 87.9 mm Hg 25 °C	NA	1995: 270 2005: 400 2010: 1,000
3647	556-82-1	3-Methyl-2-buten-1-ol 	86.13	1.61 mm Hg 20 °C 2.36 mm Hg 25 °C	NA	1995: 65 2005: 1 2010: 6
3407	497-03-0	2-Methyl-2-butenal 	84.12	13.4 mm Hg 20 °C 17.9 mm Hg 25 °C	NA	1995: 3 2005: 0.7 2010: 0.1
3646	107-86-8	3-Methyl-2-butenal 	84.12	6.14 mm Hg 20 °C 8.35 mm Hg 25 °C	NA	1995: 0 2005: 4 2010: 3
2731	108-10-1	4-Methyl-2-pentanone 	100.16	16.6 mm Hg 20 °C 21.8 mm Hg 25 °C	OSHA PEL -TWA 100 ppm, 410 mg/m3	1995: 19 2005: 5 2010: 1
3368	141-79-7	4-Methyl-3-penten-2-one 	98.15	7.9 mm Hg 20 °C 12.3 mm Hg 25 °C	OSHA PEL - TWA 25 ppm, 100 mg/m3	1995: 0 2005: 0 2010: 15
2691	96-17-3	2-Methylbutyraldehyde 	86.13	6.9 mm Hg 20 °C 10.4 mm Hg 25 °C	NA	1995: 33 2005: 87 2010: 38
2692	590-86-3	3-Methylbutyraldehyde 	86.13	39.8 mm Hg 20 °C 51.6 mm Hg 25 °C	NA	1995: 3,290 2005: 6,480 2010: 6,700

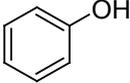
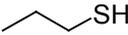
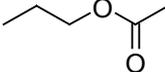
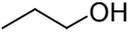
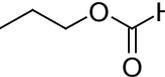
RESPIRATORY HEALTH AND SAFETY IN THE FLAVOR MANUFACTURING WORKPLACE

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
3946	583-60-8	2-Methylcyclohexanone 	112.17	2.17 mm Hg 20 °C 3.01 mm Hg 25 °C	OSHA PEL - Skin TWA 100 ppm, 460 mg/m ³	1995: NA 2005: 0 2010: 0
3875	67-68-5	Methylsulfinylmethane 	78.14	0.427 mm Hg 20 °C 0.622 mm Hg 25 °C	NA	1995: NA 2005: 0.9 2010: 0.8
2785	821-55-6	2-Nonanone 	142.24	0.449 mm Hg 20 °C 0.647 mm Hg 25 °C	NA	1995: 3,520 2005: 1,000 2010: 3,500
2842	107-87-9	2-Pentanone 	86.13	30.5 mm Hg 20 °C 39.4 mm Hg 25 °C	OSHA PEL - TWA 200 ppm, 700 mg/m ³	1995: 640 2005: 1,140 2010: 1,900
3417	625-33-2	3-Penten-2-one 	84.12	17.3 mm Hg 25 °C	NA	1995: 0 2005: 0 2010: 0.1
3584	616-25-1	1-Penten-3-ol 	86.13	6.47 mm Hg 20 °C 9.13 mm Hg 25 °C	NA	1995: 20 2005: 160 2010: 150
3382	1629-58-9	1-Penten-3-one 	84.12	29.5 mm Hg 20 °C 38.2 mm Hg 25 °C	NA	1995: 2 2005: 340 2010: 600
4012	626-38-0	2-Pentyl acetate 	130.19	7.22 mm Hg 20 °C 9.78 mm Hg 25 °C	OSHA PEL - TWA 125 ppm, 650 mg/m ³	1995: NA 2005: 0 2010: 0

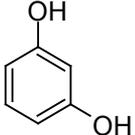
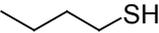
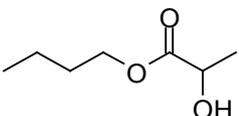
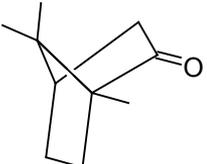
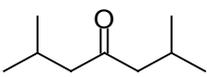
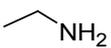
THE FLAVOR AND EXTRACT MANUFACTURERS ASSOCIATION

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
3223	108-95-2	Phenol 	94.11	0.193 mm Hg 20 °C 0.323 mm Hg 25 °C	OSHA PEL - Skin TWA 5 ppm, 19 mg/m ³	1995: 9 2005: 12 2010: 2400
2908	110-89-4	Piperidine 	85.15	22 mm Hg 20° C 28.9 mm Hg 25 °C	NA	1995: 1,610 2005: 200 2010: 52
3521	107-03-9	Propanethiol 	76.16	114 mm Hg 20 °C 144 mm Hg 25 °C	NA	1995: 120 2005: 12 2010: 5
3897	75-33-2	2-Propanethiol 	76.16	265 mm Hg 25 °C	NA	1995: NA 2005: 0 2010: 0.1
2925	109-60-4	Propyl acetate 	102.13	26.8 mm Hg 20 °C 35.1 mm Hg 25 °C	OSHA PEL - TWA 200 ppm, 840 mg/m ³	1995: 3,110 2005: 2,610 2010: 7,200
2928	71-23-8	Propyl alcohol 	60.1	16.9 mm Hg 20 °C 23.2 mm Hg 25 °C	OSHA PEL - TWA 200 ppm, 500 mg/m ³	1995: 8,470 2005: 9,010 2010: 8,000
2943	110-74-7	Propyl formate 	88.11	64 mm Hg 20 °C 84 mm Hg 25 °C	NA	1995: 110 2005: 890 2010: 1,400
2966	110-86-1	Pyridine 	79.10	14.5 mm Hg 20 °C 19.3 mm Hg 25 °C	OSHA PEL - TWA 5 ppm, 15 mg/m ³	1995: 71 2005: 120 2010: 1,100
3523	123-75-1	Pyrrolidine 	71.12	49 mm Hg 20 °C 66.5 mm Hg 25 °C	NA	1995: 28 2005: 12 2010: 9

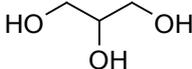
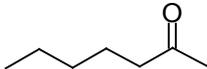
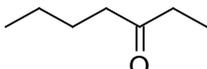
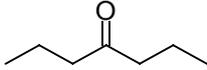
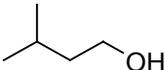
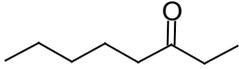
RESPIRATORY HEALTH AND SAFETY IN THE FLAVOR MANUFACTURING WORKPLACE

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
3898	5724-81-2	1-Pyrroline 	69.1	80.7 mm Hg 25 °C	NA	1995: NA 2005: 0 2010: 0
3589	108-46-3	Resorcinol 	110.1	5.90 x 10 ⁻⁵ mm Hg 20 °C 1.2 x 10 ⁻⁴ mm Hg 25 °C	PEL-TWA 10 ppm; STEL 20 ppm (see Footnote d)	1995: 5 2005: 0 2010: 0
Other Flavoring Substances with OSHA PEL's						
3478	109-79-5	1-Butanethiol 	90.19	39.9 mm Hg 25 °C	OSHA PEL - TWA 10 ppm, 35 mg/m ³	1995: 0.1 2005: 1 2010: 2
2205	138-22-7	Butyl lactate 	146.19	0.109 mm Hg 20 °C 0.17 mm Hg 25 °C	NIOSH REL-TWA 5 ppm	1995: 410 2005: 1,060 2010: 2,000
2230	464-49-3	d-Camphor 	152.24	0.00604 mm Hg 20 °C 0.0107 mm Hg 25 °C	OSHA PEL - TWA 2 mg/m ³	1995: 6,630 2005: 140 2010: 160
3537	108-83-8	2,6-Dimethyl-4-heptanone 	142.24	1.54 mm Hg 20 °C 2.15 mm Hg 25 °C	OSHA PEL - TWA 50 ppm, 290 mg/m ³	1995: 0 2005: 0 2010: 0
4236	75-04-7	Ethylamine 	45.09	153 mm Hg 25 °C	OSHA PEL - TWA 10 ppm, 18 mg/m ³	1995: NA 2005: 0 2010: 0

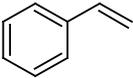
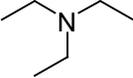
THE FLAVOR AND EXTRACT MANUFACTURERS ASSOCIATION

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
2433	75-21-8	Ethylene oxide 	44.05	1250 mm Hg 25 °C	OSHA PEL- TWA 1ppm, OSHA STEL 5 ppm/15 min	1995: 0 2005: 0 2010: 0
2525	56-81-5	Glycerol 	92.10	7.98 x 10 ⁻⁵ mm Hg 25 °C	OSHA PEL - Glycerol mist - TWA 15 mg/m ³ (total dust); 5 mg/m ³ (respi- rable fraction)	1995: 3,620,000 2005: 2,030,000 2010: 1,640,000
2544	110-43-0	2-Heptanone 	114.19	3.59 mm Hg 20 °C 4.91 mm Hg 25 °C	OSHA PEL - TWA 100ppm, 465 mg/m ³	1995: 2,280 2005: 3,100 2010: 3,600
2545	106-35-4	3-Heptanone 	114.19	4.31 mm Hg 20 °C 5.86 mm Hg 25 °C	OSHA PEL - TWA 50ppm, 230 mg/m ³	1995: 120 2005: 100 2010: 170
2546	123-19-3	4-Heptanone 	114.19	4.93 mm Hg 20 °C 6.69 mm Hg 25 °C	REL-TWA 50 ppm	1995: 130 2005: 57 2010: 37
2057	123-51-3	Isoamyl alcohol 	88.15	2.66 mm Hg 20 °C 3.84 mm Hg 25 °C	OSHA PEL - TWA 100 ppm, 360 mg/m ³ (for primary and secondary)	1995: 36,600 2005: 29,900 2010: 42,000
2803	106-68-3	3-Octanone 	128.22	1.68 mm Hg 20 °C 2.34 mm Hg 25 °C	OSHA PEL - TWA 25 ppm, 130 mg/m ³ . OSHA lists standard for this ma- terial under CAS Number 541-85-5 (Federal Regis- ter 7/5/89).	1995: 12 2005: 75 2010: 11

RESPIRATORY HEALTH AND SAFETY IN THE FLAVOR MANUFACTURING WORKPLACE

CONTINUED

FEMA	CAS	Principal Name	Molecular Weight	Calculated Vapor Pressure Value	PEL Data ^{a,b}	Reported Poundage ^c (lbs)
3233	100-42-5	Styrene 	104.15	3.67 mm Hg 20 °C 5.05 mm Hg 25 °C	OSHA PEL - TWA 100 ppm; Acceptable ceiling concentration - 200 ppm; Acceptable maximum peak above the acceptable ceiling concentration for an 8-hr shift is 600 ppm for 5 minutes in any 2 hours	1995: 9 2005: 2 2010: 0.6
4246	121-44-8	Triethylamine 	101.19	59.1 mm Hg 25 °C	OSHA PEL - TWA 25ppm, 100mg/m3	1995: NA 2005: 9 2010: 0

^a Additional occupational exposure guidance may be available from the American Conference of Governmental Industrial Hygienists (ACGIH®).

^b PEL=OSHA Permissible Exposure Limit; REL=NIOSH Recommended Exposure Limit; STEL=Short Term Exposure Limit;TWA=8 hour Time Weighted Average Exposure

^c NA=Not available

^d PEL for resorcinol subsequently rescinded and is currently not in force.

List of Acronyms

ACGIH	American Conference of Government Industrial Hygienists
AIHA	American Industrial Hygiene Association
Cal/OSHA	California Division of Occupational Safety and Health
CAS	Chemical Abstract Service
FDA	Food and Drug Administration
FEMA	Flavor and Extract Manufacturers Association
FFIDS	Flavor and Fragrance Ingredient Data Sheet
FISHEP	Flavor Industry Safety and Health Evaluation Program
GHS	Globally Harmonized System of Classification and Labeling of Chemicals
HCS	OSHA Hazard Communication Standard
HHE	NIOSH Health Hazard Evaluation
IFRA	International Fragrance Association
IOFI	International Organization of the Flavor Industry
MSDS	Material Safety Data Sheet
NEP	OSHA National Emphasis Program
NJH	National Jewish Health
NIOSH	National Institute of Occupational Safety and Health
OEL	Occupational Exposure Limit
OSHA	Occupational Safety and Health Administration
PEL	Permissible Exposure Limit
REL	Recommended Exposure Limit
STEL	Short-Term Exposure Limit
TERA	Toxicology Excellence for Risk Assessment
TLV	Threshold Limit Value
TWA	Time-Weighted Average

Maintaining safe and healthy workplaces is a matter of utmost importance to FEMA and its members. FEMA consulted with experts on the development of this document and made extensive use of a wide variety of information resources. As described in detail in their report, the application of this information to your workplace is a function of specific aspects of your workplace and the products manufactured and handled. Because of the unique nature of each workplace, the information in this report should be considered only a general guide. FEMA is not responsible for either the use or nonuse of the information, or any actions, or failure to act, in any specific workplace based on reliance on the report. It is your individual responsibility to verify this information as it applies to your workplace before acting, and to comply with all relevant federal, state, and local laws and ordinances. We strongly urge you to consult with appropriate experts regarding the circumstances relevant to respiratory health and safety in your facilities.

© Copyright 2012. The Flavor and Extract Manufacturers Association of the United States. 1620 I Street, N.W., Suite 925, Washington, D.C. 20006



**The Flavor and Extract
Manufacturers Association of
the United States**

1620 I Street, N.W., Suite 925
Washington, D.C., 20006
www.femaflavor.org