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IMMEDIATELY DANGEROUS TO LIFE OR HEALTH (IDLH) VALUE PROFILE

FOR

NITROGEN DIOXIDE

[CAS[®] No. 10102-44-0]

Department of Health and Human Services
Centers for Disease Control and Prevention
National Institute for Occupational Safety and Health

**External Review Draft
April 2017**

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1 **Foreword**

2 Chemicals are a ubiquitous component of the modern workplace. Occupational exposures to chemicals have the
3 potential to adversely affect the health and lives of workers. Acute or short-term exposures to high concentrations
4 of some airborne chemicals have the ability to quickly overwhelm workers, resulting in a spectrum of undesirable
5 health outcomes that may inhibit the ability to escape from the exposure environment (e.g., irritation of the eyes
6 and respiratory tract or cognitive impairment), cause severe irreversible effects (e.g., damage to the respiratory
7 tract or reproductive toxicity), and in extreme cases, cause death. Airborne concentrations of chemicals capable of
8 causing such adverse health effects or of impeding escape from high-risk conditions may arise from a variety of
9 nonroutine workplace situations, including special work procedures (e.g., in confined spaces), industrial
10 accidents (e.g., chemical spills or explosions), and chemical releases into the community (e.g., during
11 transportation incidents or other uncontrolled-release scenarios).

12
13 The immediately dangerous to life or health (IDLH) air concentration values developed by the National Institute
14 for Occupational Safety and Health (NIOSH) characterize these high-risk exposure concentrations and conditions
15 [NIOSH 2013]. IDLH values are based on a 30-minute exposure duration and have traditionally served as a key
16 component of the decision logic for the selection of respiratory protection devices [NIOSH 2004].

17
18 Occupational health professionals have employed these values beyond their initial purpose as a component of the
19 *NIOSH Respirator Selection Logic* to assist in developing risk management plans for nonroutine work practices
20 governing operations in high-risk environments (e.g., confined spaces) and the development of emergency
21 preparedness plans.

22
23 The approach used to derive IDLH values for high priority chemicals is outlined in the *NIOSH Current*
24 *Intelligence Bulletin (CIB) 66: Derivation of Immediately Dangerous to Life or Health Values* [NIOSH 2013].
25 CIB 66 provides (1) an update on the scientific basis and risk assessment methodology used to derive IDLH
26 values, (2) the rationale and derivation process for IDLH values, and (3) a demonstration of the derivation of
27 scientifically credible IDLH values using available data resources.

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1 The purpose of this technical report is to present the IDLH value for Nitrogen Dioxide (CAS[®] No.
2 10102-44-0). The scientific basis, toxicologic data, and risk assessment approach used to derive the IDLH value
3 are summarized to ensure transparency and scientific credibility.

4
5

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1 Abbreviations

2		
3	ACGIH®	American Conference of Governmental Industrial Hygienists
4	AEGLs	Acute Exposure Guideline Levels
5	AIHA®	American Industrial Hygiene Association
6	BMC	benchmark concentration
7	BMD	benchmark dose
8	BMCL	benchmark concentration lower confidence limit
9	C	ceiling value
10	°C	degrees Celsius
11	CAS®	Chemical Abstracts Service, a division of the American Chemical Society
12	ERPGs™	Emergency Response Planning Guidelines
13	°F	degrees Fahrenheit
14	IDLH	immediately dangerous to life or health
15	IFA	Institut für Arbeitsschutz der Deutschen Gesetzlichen Unfallversicherung (Institute for Occupational Safety and Health of the German Social Accident Insurance)
16		
17	LC	lethal concentration
18	LC ₅₀	median lethal concentration
19	LC _{LO}	lowest concentration that caused death in humans or animals
20	LEL	lower explosive limit
21	LOAEL	lowest observed adverse effect level
22	mg/m ³	milligram(s) per cubic meter
23	min	minutes
24	mmHg	millimeter(s) of mercury
25	NAC	National Advisory Committee
26	NAS	National Academy of Sciences
27	NIOSH	National Institute for Occupational Safety and Health
28	NLM	National Library of Medicine
29	NOAEL	no observed adverse effect level
30	NOEL	no observed effect level
31	NR	not recommended
32	OSHA	Occupational Safety and Health Administration
33	PEL	permissible exposure limit
34	ppm	parts per million
35	RD ₅₀	concentration of a chemical in the air that is estimated to cause a 50% decrease in the respiratory rate
36		
37	REL	recommended exposure limit
38	SCP	Standards Completion Program (joint effort of NIOSH and OSHA)
39	STEL	short-term exposure limit
40	TLV®	Threshold Limit Value
41	TWA	time-weighted average
42	UEL	upper explosive limit
43	WEELs®	Workplace Environmental Exposure Levels
44	µg/kg	microgram(s) per kilogram of body weight
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1 **Glossary**

- 2
- 3 **Acute exposure:** Exposure by the oral, dermal, or inhalation route for 24 hours or less.
- 4 **Acute Exposure Guideline Levels (AEGLs):** Threshold exposure limits for the general public, applicable to
5 emergency exposure periods ranging from 10 minutes to 8 hours. AEGL-1, AEGL 2, and AEGL-3 are
6 developed for five exposure periods (10 and 30 minutes, 1 hour, 4 hours, and 8 hours) and are distinguished
7 by varying degrees of severity of toxic effects, ranging from transient, reversible effects to life-threatening
8 effects [NAS 2001]. AEGLs are intended to be guideline levels used during rare events or single once-in-a-
9 lifetime exposures to airborne concentrations of acutely toxic, high-priority chemicals [NAS 2001]. The
10 threshold exposure limits are designed to protect the general population, including the elderly, children, and
11 other potentially sensitive groups that are generally not considered in the development of workplace exposure
12 recommendations (additional information available at <http://www.epa.gov/oppt/aegl/>).
- 13 **Acute reference concentration (Acute RfC):** An estimate (with uncertainty spanning perhaps an order of
14 magnitude) of a continuous inhalation exposure for an acute duration (24 hours or less) of the human
15 population (including sensitive subgroups) that is likely to be without an appreciable risk of deleterious
16 effects during a lifetime. It can be derived from a NOAEL, LOAEL, or benchmark concentration, with
17 uncertainty factors (UFs) generally applied to reflect limitations of the data used. Generally used in U.S. EPA
18 noncancer health assessments [U.S. EPA 2016].
- 19 **Acute toxicity:** Any poisonous effect produced within a short period of time following an exposure, usually 24 to
20 96 hours [U.S. EPA 2016].
- 21 **Adverse effect:** A substance-related biochemical change, functional impairment, or pathologic lesion that affects
22 the performance of an organ or system or alters the ability to respond to additional environmental challenges.
- 23 **Benchmark dose/concentration (BMD/BMC):** A dose or concentration that produces a predetermined change in
24 response rate of an effect (called the benchmark response, or BMR) compared to background [U.S. EPA
25 2016] (additional information available at <http://www.epa.gov/ncea/bmds/>).
- 26 **Benchmark response (BMR):** A predetermined change in response rate of an effect. Common defaults for the
27 BMR are 10% or 5%, reflecting study design, data variability, and sensitivity limits used.
- 28 **BMCL:** A statistical lower confidence limit on the concentration at the BMC [U.S. EPA 2016].
- 29 **Bolus exposure:** A single, relatively large dose.
- 30 **Ceiling value (“C”):** U.S. term in occupational exposure indicating the airborne concentration of a potentially
31 toxic substance that should never be exceeded in a worker’s breathing zone.
- 32 **Chronic exposure:** Repeated exposure for an extended period of time. Typically exposures are more than
33 approximately 10% of life span for humans and >90 days to 2 years for laboratory species.
- 34 **Critical study:** The study that contributes most significantly to the qualitative and quantitative assessment of risk
35 [U.S. EPA 2016].
- 36
- 37 **Dose:** The amount of a substance available for interactions with metabolic processes or biologically significant
38 receptors after crossing the outer boundary of an organism [U.S. EPA 2016].
- 39 **EC₅₀:** A combination of the effective concentration of a substance in the air and the exposure duration that is
40 predicted to cause an effect in 50% (one half) of the experimental test subjects.

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- 1 **Emergency Response Planning Guidelines (ERPGsTM):** Maximum airborne concentrations below which nearly
2 all individuals can be exposed without experiencing health effects for 1-hour exposure. ERPGs are presented
3 in a tiered fashion, with health effects ranging from mild or transient to serious, irreversible, or life
4 threatening (depending on the tier). ERPGs are developed by the American Industrial Hygiene Association
5 [AIHA 2006].
- 6 **Endpoint:** An observable or measurable biological event or sign of toxicity, ranging from biomarkers of initial
7 response to gross manifestations of clinical toxicity.
- 8 **Exposure:** Contact made between a chemical, physical, or biological agent and the outer boundary of an
9 organism. Exposure is quantified as the amount of an agent available at the exchange boundaries of the
10 organism (e.g., skin, lungs, gut).
- 11 **Extrapolation:** An estimate of the response at a point outside the range of the experimental data, generally
12 through the use of a mathematical model, although qualitative extrapolation may also be conducted. The
13 model may then be used to extrapolate to response levels that cannot be directly observed.
- 14 **Hazard:** A potential source of harm. Hazard is distinguished from risk, which is the probability of harm under
15 specific exposure conditions.
- 16 **Immediately dangerous to life or health (IDLH) condition:** A condition that poses a threat of exposure to
17 airborne contaminants when that exposure is likely to cause death or immediate or delayed permanent adverse
18 health effects or prevent escape from such an environment [NIOSH 2004, 2013].
- 19 **IDLH value:** A maximum (airborne concentration) level above which only a highly reliable breathing apparatus
20 providing maximum worker protection is permitted [NIOSH 2004, 2013]. IDLH values are based on a 30-
21 minute exposure duration.
- 22 **LC₀₁:** The statistically determined concentration of a substance in the air that is estimated to cause death in 1% of
23 the test animals.
- 24 **LC₅₀:** The statistically determined concentration of a substance in the air that is estimated to cause death in 50%
25 (one half) of the test animals; median lethal concentration.
- 26 **LC_{LO}:** The lowest lethal concentration of a substance in the air reported to cause death, usually for a small
27 percentage of the test animals.
- 28
- 29 **LD₅₀:** The statistically determined lethal dose of a substance that is estimated to cause death in 50% (one half) of
30 the test animals; median lethal concentration.
- 31 **LD_{LO}:** The lowest dose of a substance that causes death, usually for a small percentage of the test animals.
- 32 **LEL:** The minimum concentration of a gas or vapor in air, below which propagation of a flame does not occur in
33 the presence of an ignition source.
- 34 **Lethality:** Pertaining to or causing death; fatal; referring to the deaths resulting from acute toxicity studies. May
35 also be used in lethality threshold to describe the point of sufficient substance concentration to begin to cause
36 death.
- 37 **Lowest observed adverse effect level (LOAEL):** The lowest tested dose or concentration of a substance that has
38 been reported to cause harmful (adverse) health effects in people or animals.

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- 1 **Mode of action:** The sequence of significant events and processes that describes how a substance causes a toxic
2 outcome. By contrast, the term *mechanism of action* implies a more detailed understanding on a molecular
3 level.
- 4 **No observed adverse effect level (NOAEL):** The highest tested dose or concentration of a substance that has
5 been reported to cause no harmful (adverse) health effects in people or animals.
- 6 **Occupational exposure limit (OEL):** Workplace exposure recommendations developed by governmental
7 agencies and nongovernmental organizations. OELs are intended to represent the maximum airborne
8 concentrations of a chemical substance below which workplace exposures should not cause adverse health
9 effects. OELs may apply to ceiling limits, STELs, or TWA limits.
- 10 **Peak concentration:** Highest concentration of a substance recorded during a certain period of observation.
- 11 **Permissible exposure limits (PELs):** Occupational exposure limits developed by OSHA (29 CFR 1910.1000) or
12 MSHA (30 CFR 57.5001) for allowable occupational airborne exposure concentrations. PELs are legally
13 enforceable and may be designated as ceiling limits, STELs, or TWA limits.
- 14
- 15 **Point of departure (POD):** The point on the dose–response curve from which dose extrapolation is initiated. This
16 point can be the lower bound on dose for an estimated incidence or a change in response level from a
17 concentration-response model (BMC), or it can be a NOAEL or LOAEL for an observed effect selected from
18 a dose evaluated in a health effects or toxicology study.
- 19 **RD₅₀:** The statistically determined concentration of a substance in the air that is estimated to cause a 50% (one
20 half) decrease in the respiratory rate.
- 21 **Recommended exposure limit (REL):** Recommended maximum exposure limit to prevent adverse health
22 effects, based on human and animal studies and established for occupational (up to 10-hour shift, 40-hour
23 week) inhalation exposure by NIOSH. RELs may be designated as ceiling limits, STELs, or TWA limits.
- 24 **Short-term exposure limit (STEL):** A worker’s 15-minute time-weighted average exposure concentration that
25 shall not be exceeded at any time during a work day.
- 26 **Target organ:** Organ in which the toxic injury manifests in terms of dysfunction or overt disease.
- 27 **Threshold Limit Values (TLVs[®]):** Recommended guidelines for occupational exposure to airborne
28 contaminants, published by the American Conference of Governmental Industrial Hygienists (ACGIH[®]).
29 TLVs refer to airborne concentrations of chemical substances and represent conditions under which it is
30 believed that nearly all workers may be repeatedly exposed, day after day, over a working lifetime, without
31 adverse effects. TLVs may be designated as ceiling limits, STELs, or 8-hr TWA limits.
- 32 **Time-weighted average (TWA):** A worker’s 8-hour (or up to 10-hour) time-weighted average exposure
33 concentration that shall not be exceeded during an 8-hour (or up to 10-hour) work shift of a 40-hour week.
34 The average concentration is weighted to take into account the duration of different exposure concentrations.
- 35 **Toxicity:** The degree to which a substance is able to cause an adverse effect on an exposed organism.
- 36
- 37 **Uncertainty factors (UFs):** Mathematical adjustments applied to the POD when developing IDLH values. The
38 UFs for IDLH value derivation are determined by considering the study and effect used for the POD, with
39 further modification based on the overall database.

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- 1 **Workplace Environmental Exposure Levels (WEELs[®]):** Exposure levels developed by the American Industrial
2 Hygiene Association (AIHA[®]) that provide guidance for protecting most workers from adverse health
3 effects related to occupational chemical exposures, expressed as TWA or ceiling limits.
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2
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1.0 Introduction

1.1 Overview of the IDLH Value for Nitrogen Dioxide

IDLH Value: 11 ppm (21 mg/m³)

Basis for IDLH Value: The IDLH value for nitrogen dioxide is based on a LOAEL of 30 ppm for development of latent severe respiratory effects observed in humans following a 40 minute exposure; the duration adjusted LOAEL for a 30-minute exposure is 33 ppm [Henschler et al. 1960; Morley and Silk 1970]. An uncertainty factor of 3 was applied to account for extrapolation from a threshold for the onset of severe effects and human variability, resulting in an IDLH value of 11 ppm.

1.2 Purpose

This *IDLH Value Profile* presents (1) a brief summary of technical data associated with acute inhalation exposures to nitrogen dioxide and (2) the rationale behind the immediately dangerous to life or health (IDLH) value for nitrogen dioxide. IDLH values are developed on the basis of scientific rationale and logic outlined in the *NIOSH Current Intelligence Bulletin (CIB) 66: Derivation of Immediately Dangerous to Life or Health (IDLH) Values* [NIOSH 2013]. As described in CIB 66, NIOSH performs in-depth literature searches to ensure that all relevant data from human and animal studies with acute exposures to the substance are identified. Information included in CIB 66 on the literature search includes pertinent databases, key terms, and guides for evaluating data quality and relevance for the establishment of an IDLH value. The information that is identified in the in-depth literature search is evaluated with general considerations that include description of studies (i.e., species, study protocol, exposure concentration and duration), health endpoint evaluated, and critical effect levels (e.g., NOAELs, LOAELs, and LC₅₀ values). For nitrogen dioxide, the in-depth literature search was conducted through September 2016.

1.3 General Substance Information

Chemical: Nitrogen dioxide

CAS No: 10102-44-0

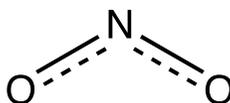
Synonyms: Nitrogen oxide (NO₂); Nitrogen peroxide*

Chemical category: Nitrogen oxides; Inorganic gases[†]

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1 **References:** * NLM [2017], † IFA [2017]

2
3 **Structural formula:**



7
8 Table 1 highlights selected physiochemical properties of nitrogen dioxide relevant to IDLH conditions. Table 2
9 provides alternative exposure guidelines for nitrogen dioxide. Table 3 summarizes the Acute Exposure Guidelines
10 Level (AEGL) values for nitrogen dioxide.

11
12 **Table 1: Physiochemical Properties of Nitrogen Dioxide**

13

Property	Value
Molecular weight	46.006*
Chemical formula	NO ₂ †
Description	Reddish-brown gas above 21.15 °C, pale yellow/yellowish-brown liquid above 21.15 °C
Odor	Pungent, acrid
Odor Threshold	0.058 ppm‡
UEL	Not applicable
LEL	Not applicable
Vapor pressure	900 mmHg at 25°C (77°F)§
Flash point	Not flammable
Ignition temperature	Not flammable
Solubility	Rapid hydrolysis§
Reactivity	Decomposes in water forming nitric oxide and nitric acid¶

14 *HSDB [2017]; †NLM [2017]; ‡Murnane et al. [2013]; §IFA [2017]; ¶NAS [2012]

15
16
17 **Table 2: Alternative Exposure Values for Nitrogen Dioxide**

18

Organization	Value
NIOSH (1994) IDLH value*	20 ppm
NIOSH REL†	1 ppm - STEL
OSHA PEL‡	5 ppm - Ceiling
CA OSHA PEL§	1 ppm - STEL
ACGIH TLV¶	0.2 ppm TWA
AIHA ERPGs TM **	ERPG-1: 1 ppm; ERPG-2: 15 ppm; ERPG-3: 30 ppm
AIHA WEELs [@] ***	Not available

19 **References:** *NIOSH [1994]; †NIOSH [2017]; ‡OSHA [2017]; §ACGIH [2016]; ¶CA OSHA [2017]; **AIHA [2014]

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Table 3: AEGL Values for Nitrogen Dioxide

Classification	10-min	30-min	1-hour	4-hour	8-hour	Endpoint [reference]
AEGL-1	0.50 ppm 0.94 mg/m ³	Slight burning of the eyes, slight headache, chest tightness or labored breathing with exercise in 7/13 asthmatics [Kerr et al. 1978, 1979]				
AEGL-2	20 ppm 38 mg/m ³	15 ppm 28 mg/m ³	12 ppm 23 mg/m ³	8.2 ppm 15 mg/m ³	6.7 ppm 13 mg/m ³	Burning sensation in nose and chest, cough, dyspnea, sputum production in normal volunteers [Henschler et al. 1960]
AEGL-3	34 ppm 64 mg/m ³	25 ppm 47 mg/m ³	20 ppm 38 mg/m ³	14 ppm 26 mg/m ³	11 ppm 21 mg/m ³	Marked irritation, histopathologic changes in lungs, fibrosis and edema of cardiac tissue, necrosis in liver, no deaths in monkeys [Henry et al. 1969]

*Reference: NAS [2012].

2.0 Animal Toxicity Data

Nitrogen dioxide is an irritant to the mucous membranes and has been shown to cause coughing and dyspnea during exposure. Severe exposure pulmonary edema with symptoms of chest pain, cough, dyspnea, and cyanosis have been reported [NIOSH 1976; Douglas et al. 1989]. Lethality from nitrogen dioxide exposure is reported to be due to bronchospasm and pulmonary edema occurring with hypoxemia and respiratory acidosis, metabolic acidosis, decreased oxygenation of hemoglobin and low arterial blood pressure [Douglas et al. 1989].

Furthermore, after acute nitrogen dioxide intoxication and an apparent recovery, a late-onset bronchiolar injury in the form of bronchiolitis fibrosa obliterans is observed [NIOSH 1976; Hamilton 1983; Douglas et al. 1989].

Five- to 60-minute LC₅₀ values for nitrogen dioxide in the rat ranged from 416 to 115 ppm, respectively in one study [Carson et al. 1962] and from 833 to 168 ppm in another study [Gray et al. 1954]. Hine et al. [1970] studied the effects of varying concentrations (50-200 ppm) and durations (5 min to 24 hours) of nitrogen dioxide exposure in several species (rat, mouse, guinea pig, rabbit and dog). During this study lethality in one guinea pig (n = 6) was reported following the lowest exposure of 50 ppm for one hour (which was the shortest duration for this concentration). The highest exposure of 200 ppm for 5 minutes caused lethality in 6/12 rats, 4/6 mice and 2/2 guinea pigs. In squirrel monkeys exposed to 10-50 ppm nitrogen dioxide for 2 hours [Henry et al. 1969], exposure to 35 or 50 ppm resulted in a markedly increased respiratory rate and decreased tidal volume.

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1 Table 4 summarizes the lethal concentration (LC) data identified in animal studies and provides 30-minute
2 equivalent derived values for nitrogen dioxide. Table 5 provides non-lethal data reported in animal studies with
3 30-minute equivalent derived values. Information in these tables includes species of test animals, toxicological
4 metrics (i.e., LC, BMCL, NOAEL, LOAEL), adjusted 30-minute concentration, and the justification for the
5 composite uncertainty factors applied to calculate the derived values.

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Table 5: Non-lethal Concentration Data for Nitrogen Dioxide

Reference	Species	Critical adverse health effects	LOAEL (ppm)	Time (min)	Adjusted 30-min Concentration* (ppm)	Composite Uncertainty Factor	Derived Value (ppm) [†]	Final Value (ppm) [‡]
Morley and Silk [1970]	Human	Cyanosis, dyspnea and pulmonary edema	30	40	33	3 [§]	10.9	11
Henschler et al. [1960]	Human	Burning sensation in upper respiratory tract and severe cough followed by dyspnea	30	70	38	3 [§]	12.7	13
Norwood et al. [1966]	Human	Shortness of breath, chest discomfort, and pulmonary edema	90	40	98	3 [§]	32.6	33
Henry et al. [1969]	Monkey	Increased respiratory rate and decreased tidal volume	35	120	52	10 [¶]	5.2	5.0

*ten Berge et al. [1986] empirically estimated n = 3.5. The n = 3.5 was used during all duration adjustments for nitrogen dioxide. Additional information on the calculation of duration-adjusted concentrations can be found in NIOSH [2013].

[†]The derived value is the result of the adjusted 30-min value divided by the composite uncertainty factor. The composite uncertainty factor used varies for each study on the basis of the nature and severity of the endpoint observed.

[‡]Values rounded to the appropriate significant figure.

[§]Composite uncertainty factor assigned to account for human variability.

[¶]Composite uncertainty factor assigned to account for interspecies differences and human variability.

3.0 Human Data

Numerous human studies are available that describe effects of nitrogen dioxide in humans. Overall, the pattern of respiratory tract effects includes initial signs of respiratory tract irritation, with latent tracheobronchial and pulmonary involvement. Douglas et al. [1989] reported that death from the inhalation of nitrogen dioxide is caused by bronchospasm and pulmonary edema. NAS [2012] stated that a characteristic of nitrogen dioxide intoxication after the acute phase is a period of apparent recovery. A second phase of symptoms may occur after several hours or several days with the development of fever with progressively more severe dyspnea, cyanosis, and cough. An estimation of the concentration causing death in humans is approximately ≥ 150 ppm, but no duration of exposure was given.

In one controlled exposure study [Henschler et al. 1960], three healthy male volunteers were exposed to 30 ppm for 2 hours. Effects observed after 30 to 40 minutes of exposure were limited to minimal signs of irritation in the nose and throat. Effects following exposure for 70 minutes were reported as a burning sensation with increasingly severe cough. With continued exposure, coughing decreased, but the burning sensation moved to deeper portions of the lung, accompanied by sputum secretion and dyspnea. The effects of this exposure were reversible. This study design cannot provide information on the degree to which latent respiratory tract effects would have occurred following exposure for only 30 minutes in the subjects. Welders exposed to fumes containing nitrogen dioxide developed a variety of symptoms including cough, headache, tightness/pain in chest, and nausea. Two of seven workers were hospitalized with cyanosis, dyspnea and pulmonary edema following exposure to 30 ppm of nitrogen dioxide for 40 minutes [Morley and Silk 1970]. Another welder developed shortness of breath, chest discomfort, and pulmonary edema while using an acetylene torch for metal-cutting in a poorly ventilated enclosed space for 30 minutes. The authors noted that seven other workers were exposed during this incident, but were unaffected. This indicates the potential for human variability in response to nitrogen dioxide. Simulation of the accident produced a nitrogen dioxide concentration of 90 ppm [Norwood et al. 1966]. The reports of effects in welders as the basis for IDLH value derivation are limited since the effects of the complex mix of materials released during welding need to be considered.

4.0 Summary

Nitrogen dioxide is an irritant to the mucous membranes and has been shown to cause coughing and dyspnea during exposure. Severe exposure pulmonary edema with symptoms of chest pain, cough, dyspnea, and cyanosis

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1 have been reported [NIOSH 1976; Douglas et al. 1989]. Human data indicate that acute inhalation exposures may
2 induce potential escape-impairing effects of dyspnea, cyanosis, and pulmonary edema [Henschler et al. 1960;
3 Norwood et al. 1969; Morley and Silk 1970]. Morley and Silk [1970] identified a LOAEL for severe respiratory
4 effects following exposure to 30 ppm for a 40 minute exposure. This result is supported by the observations of
5 Henschler et al. [1960], where 70 minutes of exposure at 40 ppm nitrogen dioxide caused severe cough and an
6 increased burning sensation of the upper respiratory tract. Duration adjustment of the LOAEL reported by Morley
7 and Silk [1970] resulted in a 30-minute equivalent concentration of 33 ppm. An uncertainty factor of 3 was
8 applied to account for the potential for human variability in response to nitrogen dioxide documented in Morley
9 and Silk [1970]. This resulted in an IDLH value of 11 ppm.

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