

U.S. DEPARTMENT OF HEALTH, EDUCATION, AND WELFARE  
CENTER FOR DISEASE CONTROL  
NATIONAL INSTITUTE FOR OCCUPATIONAL SAFETY AND HEALTH  
CINCINNATI, OHIO 45226

HEALTH HAZARD EVALUATION DETERMINATION  
REPORT NO. 78-103-561

WELD COUNTY HOSPITAL  
GREELEY, COLORADO

FEBRUARY 1979

I. TOXICITY DETERMINATION

A health hazard evaluation was conducted by the National Institute for Occupational Safety and Health (NIOSH) at Weld County Hospital, Greeley, Colorado, on June 29, 1978. Environmental breathing zone samples were collected to determine concentrations of halothane, ethrane, and nitrous oxide in the operating rooms.

Atmospheric concentrations of halothane, ethrane, and nitrous oxide indicate that a potential health hazard existed during this evaluation when compared to NIOSH recommended levels. Since information on adverse health effects due to exposure to halothane, ethrane, and nitrous oxide are not completely defined, and many unknown factors still exist, recommended permissible levels of exposure are not defined as safe levels but rather as levels which are attainable under current technology. NIOSH recommends an 8-hour exposure limit of 25 parts per million (ppm) for nitrous oxide and 2 ppm for the halogenated anesthetics (halothane and ethrane). Throughout this evaluation, nitrous oxide was used in conjunction with either ethrane or halothane. When used in this manner, the recommended standard of 0.5 ppm for halogenated anesthetic agents should be followed. These levels should prevent both chronic and acute effects. Two out of 28 halothane breathing zone samples exceeded the NIOSH recommended level of 0.5 ppm; 3 out of 28 ethrane breathing zone samples exceeded the NIOSH recommended level of 0.5 ppm. The nitrous oxide NIOSH recommended level of 25 ppm was equal to or exceeded in approximately 20% of the breathing zone samples. The slightly excessive levels of waste anesthetic gases found during this survey were presumably caused by an inadequate venting system coming from the scavenging system, difficulty in administering the gas to small patients, improperly fitted face masks, and the technique of administration by the anesthesiologist.

II. DISTRIBUTION AND AVAILABILITY

Copies of this determination report are currently available upon request from NIOSH, Division of Technical Services, Information Resources and Dissemination Section, 4676 Columbia Parkway,

Cincinnati, Ohio 45226. After 90 days the report will be available through the National Technical Information Service (NTIS), Springfield, Virginia. Information regarding its availability through NTIS can be obtained from NIOSH, Publications Office, at the Cincinnati address.

Copies of this report have been sent to:

1. Weld County Hospital
2. U.S. Department of Labor/OSHA - Region VIII
3. NIOSH - Region VIII

For the purpose of informing the approximately 30 affected employees, a copy of this report shall be posted in a prominent place accessible to the employees for a period of 30 calendar days.

### III. INTRODUCTION

Section 20(a)(6) of the Occupational Safety and Health Act of 1970, 29 U.S.C. 669(a)(6), authorizes the Secretary of Health, Education, and Welfare, following a written request by any employer or authorized representative of employees, to determine whether any substance normally found in the place of employment has potentially toxic effects in such concentrations as used or found.

NIOSH received such a request from hospital management at Weld County Hospital. There were no specific health problems at the time of this request. The recognition by operating room personnel of the potential health hazards associated with chronic exposures to anesthetic gases was responsible for the health hazard evaluation request.

### IV. HEALTH HAZARD EVALUATION

#### A. Process Description

At the time of this evaluation, Weld County Hospital had 5 operating rooms. These operating rooms operate basically from 7:00 a.m. until 5:00 or 6:00 p.m. However, they are open 24 hours a day, and emergency surgery is performed in these rooms. During this evaluation all operating rooms were monitored for waste anesthetic gas exposures. There are 30 workers in the operating rooms. Surgery performed in these operating rooms includes orthopedic surgery, hysterectomies, colostomies, eye surgery, and tonsillectomies. The workload during the two days of this evaluation was typical. The scavenging system on each anesthetic cart in the operating room is vented to the outside

by gravity flow, but since there is no vacuum pump to assist in evacuation of waste gases, this system may be inadequate.

B. Evaluation Design

Nitrous oxide samples were collected in 20 liter mylar bags using a vacuum pump operated at 300 cubic centimeters (cc) per minute. These samples were analyzed immediately on the surgical floor by infrared spectrometry using a Wilks Miran 1A with a sensitivity of 5 ppm. Instrument settings were wave length 4.47 microns, path length 5.25 meters, and slit width 0.5 millimeters (mm). Halothane and ethrane samples were collected on charcoal tubes using vacuum pumps operated at 200 cc per minute. Analysis of these samples was performed using gas chromatography and P&CAM Method No. 127. Breathing zone air samples of operating room personnel were collected during each surgical procedure for the above anesthetic gases.

C. Evaluation Criteria

In the NIOSH criteria document for a recommended standard for occupational exposure to anesthetic gases, NIOSH states: (Reference 1) "Current scientific evidence obtained from human and animal studies suggests that chronic exposure to anesthetic gases increases the risk of both spontaneous abortion among female workers and congenital abnormalities in the offspring of female workers and the wives of male workers. Risks of hepatic and renal diseases are also increased among exposed personnel. In addition, physiological function may be impaired. A few studies have suggested increased risk of cancer. Effects on the central nervous system due to acute exposures of anesthetic gases have been associated with headaches, nausea, fatigue, irritability, etc." Control procedures and work practices presented in that document, however, should prevent the effects caused by acute exposure and significantly reduce the risk associated with long-term, low level exposure. A dose response relationship for halogenated anesthetic toxicity has not been defined.

That same NIOSH publication recommends maximum exposures of 25 ppm nitrous oxide (8-hour time weighted average) and 2 ppm halogenated anesthetic when used alone, or 0.5 ppm when used with nitrous oxide. These recommendations are based upon available technology in reducing waste anesthetic gas levels.

Reports by Vaisman (Reference 2) and Askrog and Harvald (Reference 3) were among the first to identify increased incidence of spontaneous abortion in women exposed to anesthetic

gases and in wives of men exposed to anesthetic gases. Results of a more recent and comprehensive nationwide survey of occupational disease among operating personnel were published in 1974 by the American Society of Anesthesiologists (ASA). (Reference 4) The results of this study indicate "that female members of the operating room-exposed group were subject to increased risks of spontaneous abortion, congenital abnormalities in their children, cancer and hepatic and renal disease. This increased risk of congenital abnormalities was also present among the unexposed wives of male operating room personnel. No increase in cancer was found among the exposed males, but an increased incidence of hepatic disease similar to that in the female was found."

While several investigators have reported increased rates of resorption in animals, particularly rats, most of these studies involved concentrations of anesthetic gases well above the levels found in occupational exposure. One investigator (Reference 5) showed increased fetal death rates in two groups of rats following exposure of 1,000 and 100 ppm of nitrous oxide. Doenicke, et al., (Reference 6) concluded from their study of anesthetized pregnant rats that halothane demonstrates an abortive effect directly proportional to the concentration inhaled, again referring to anesthetic concentrations; but nitrous oxide does not produce an abortive effect. Bruce (Reference 7) reports no significant difference, including implantations and resorptions per pregnancy, in his exposure of rats to 16 ppm halothane.

Several epidemiological studies that indicate increased spontaneous abortions also indicate an increased rate of congenital abnormalities. The ASA study (Reference 4) (as well as surveys by Knill-Jones, et al., (Reference 8) and Corbett, et al. (Reference 9) indicated an increased rate of congenital abnormalities in children of women with occupational exposures to anesthetic gases and to wives of men with similar exposures. While most animal exposure studies have been conducted at anesthetic levels, one study (References 10, 11, 12) indicated liver, kidney, and brain tissue changes in pups born to rats exposed to sub-anesthetic concentrations of halothane during pregnancy.

The same epidemiological and toxicological studies that indicated an increase in spontaneous abortion and congenital abnormalities also indicated an increase in liver and kidney abnormalities. This increase, however, was less pronounced in both rate and severity.

In a study published by NIOSH, (Reference 13) "nitrous oxide and halothane in respective concentrations as low as 50 ppm and 1.0 ppm caused measurable decrements in performance on some psychological tests taken by healthy male graduate students. Nitrous oxide alone caused similar effects. The functions apparently most sensitive to these low concentrations on anesthetics were visual perception, immediate memory, and a combination of perception, cognition, and motor responses required in a task of divided attention to simultaneous visual and auditory stimuli." Headache, fatigue, irritability, and disturbance of sleep have also been reported; (References 2, 14) and damage to cerebral cortical neurons has been seen in rats after sub-anesthetic exposure to halothane. (Reference 15) Quimby, et. al., (Reference 16) reported permanent learning deficits in rats exposed to anesthetic concentrations of halothane during early development (from conception).

Mortality and epidemiological studies have raised the questions of possible carcinogenicity of anesthetic gases, but sufficient data are lacking to list nitrous oxide, halothane, or ethrane as suspected carcinogens.

Literature reviews regarding halothane (References 17, 18, 19, 20) indicate the most widely accepted mechanism of bio-transformation is the production of trifluoroacetic acid with resulting urinary excretion of trifluoroacetic acid and bromide. The literature regarding ethrane (References 21, 22) does not indicate any one accepted mechanism, but increased serum and urinary fluoride levels were found in patients receiving ethrane anesthesia. While epidemiological and toxicological studies have indicated several symptoms apparently related to sub-anesthetic exposure to anesthetic gases, no cause and effect relationship has yet been shown.

#### D. Evaluation Results

Table I shows the results of 28 samples taken for halothane and ethrane. Only 3 of the ethrane samples exceeded the recommended level of 0.5 ppm. Two out of 28 samples exceeded the evaluation criteria for halothane. Nitrous oxide concentrations exceeded the NIOSH recommended level of 25 ppm in 8 out of 34 samples. Nitrous oxide levels found during this evaluation may be reviewed in Table II. Table III is an attempt to evaluate the time weighted average (TWA) exposures for the various operating rooms. It should be noted that any time the evaluation criteria is exceeded, a hazard exists since these evaluations are based on ceiling values.



#### E. Results and Discussion

Findings during this evaluation indicate that this hospital has done a lot of work in eliminating exposures to employees from waste anesthetic gases, even though the NIOSH recommended levels for nitrous oxide and the halogenated anesthetics were exceeded. The recommended standards were never grossly exceeded. Reasons for the elevated levels of these gases were perhaps due to an inadequate exhaust system coming from the anesthetic carts' scavenging system. It is doubtful that there was a leak in any of the anesthetic gas-administering machinery, since levels during this survey were relatively low when compared with other NIOSH studies. (Reference 23) If there had been a leak in any of the anesthetic gas-administering machinery, concentrations would have been much higher than those found during this evaluation. Most of the employees, including surgeons, anesthesiologists, nurses, and operating room technicians, seemed to think that they were working in a safe place and that their work had no ill effects on their health.

#### F. Conclusions

There were scavenging systems on all anesthetic carts. All operating room personnel were aware of the hazards associated with chronic exposures to anesthetic gases. Concentrations found during this survey do not pose an immediate danger to the health of the operating room personnel. Every effort should be made to lower these concentrations to levels that are less than those recommended by NIOSH.

#### V. RECOMMENDATIONS

1. Anesthetic equipment should be checked and maintained on a regular basis.
2. The ventilation exhaust coming from the scavenging system on the anesthetic cart should be plugged into a vacuum system which would actually draw the waste gas out of the operating room.
3. Face masks, tubing, and breathing bags should all be checked for cracks and other leaks.
4. All high pressure connections and valves should be checked periodically. Care should be taken when pouring the halogenated agents into the anesthetic gas-administering machinery so that they are not spilled on the floor.

5. Operating rooms should be monitored at least once a year to make sure that workers are not being overexposed to waste anesthetic gases.

#### VI. REFERENCES

1. Criteria for a Recommended Standard...Occupational Exposure to Waste Anesthetic Gases and Vapors. NIOSH, 1977.
2. Vaisman, A. I. (Working conditions in surgery and their effect on the health of anesthesiologists). Fksp Khir Anest 3:44-49, 1967 (Rus).
3. Adkrog, V., and Harvald, B. (Teratogenic effect of inhalation anesthetics). Nord Med 83:498-504, 1970.
4. Cohen, E.N., Brown, B.W., Bruce, D.K., Cascorbi, H.F., Corbett, T.H., Jones, T.H., and Whitcher, C.E. Occupational Disease Among Operating Room Personnel--A National Study. Anesthesiology 41:321-40, 1974.
5. Corbett, T.H., Cornell, R.G., Endres, J.L., and Millard, R.I. Effects of Low Concentrations of Nitrous Oxide on Rat Pregnancy. Anesthesiology 39:299-301, 1973.
6. Doenicke, A., Wittmann, R., Heinrich, H., and Pausch, H. (Abortive effect of halothane). Anesth Analg (Paris) 32:47-51, 1975 (Fre).
7. Bruce, D.L. Murine Fertility Unaffected by Traces of Halothane. Anesthesiology 38:473-77, 1973.
8. Knill-Jones, R.P., Moir, D.D., Rodrigues, L.V., and Spence, A.A. Anesthetic Practice and Pregnancy--Controlled Survey.
9. Corbett, T.H., Cornell, R.G., Lieding, K., and Endres, J.L. Incidence of Cancer Among Michigan Nurse-Anesthetists. Anesthesiology 41:34-44, 1974.
10. Chang, L.W., Lee, Y.K., Dudley, A.W., Jr., and Katz, J. Ultrastructural Evidence of the Hepatotoxic Effect of Halothane in Rats Following In-Utero Exposure. Can Anaesth Soc J. 22:330-37, 1975.
11. Chang, L.W., Dudley, A.W., Jr., Lee, Y.K., and Katz, J. Ultrastructural studies on the Pathological Changes in the Neonatal Kidney Following In-Utero Exposure to Halothane. Environ. Res 10:174-89, 1975.

12. Chang, L.W., Dudley, A.W., Jr., Katz, J., and Martin, A.H. Nervous System Development Following In-Utero Exposure to Trace Amounts of Halothane. *Teratology* 9:A-15, 1974.
13. Bruce, D.L., and Bach, M.J. Trace Effects of Anesthetic Gases on Behavioral Performance of Operating Room Personnel. HFW Publication No. NIOSH 76-169, 1976, 33 pp.
14. Uhlirova, A., and Polorny, J. (Results of questionnaire survey of health damage to anesthesiologists). *Rozhl Chir* 53:761-70, 1976 (Cze).
15. Chang, L.W., Dudley, A.W., Jr., Lee, Y.K., and Katz, J. Ultrastructural Changes in the Nervous System After Chronic Exposure to Halothane. *Exp Neurol* 45:209-19, 1974.
16. Ouimby, K.L., Aschkenase, L.J., Bowman, R.E., Katz, J., and Chang, L.W. Enduring Learning Deficits and Cerebral Synaptic Malformation From Exposure to Ten Parts of Halothane Per Million. *Science* 185:615-27, 1974.
17. Rehder, K., and Sessler, A.D. Biotransformation of Halothane. *Int Anesthesiol Clin* 12:41-53, 1974.
18. Sawyer, D., and Eger, E., II. Hepatic Metabolism of Halothane. *Int Anesthesiol Clin* 12:55-62, 1974.
19. Cascorbi, H.F. Factors Causing Differences in Halothane Biotransformation. *Int Anesthesiol Clin* 12:63-71, 1974.
20. Van Dyke, R.A. Biotransformation of Volatile Anesthetics With Special Emphasis on the Role of Metabolism in the Toxicity of Anesthetics. *Can Anesth Soc J* 20:21-33, 1973.
21. Mazze, R.I., and Cousins, M.J. Biotransformation of Methoxyflurane. *Int Anesthesiol Clin* 12:93-105, 1974.
22. Cousins, M.D., and Mazze, R.I. Biotransformation of Enflurane (Ethrane) and Isoflurane (Forane). *Int Anesthesiol Clin* 12:111-119, 1974.
23. Hazard Evaluation Report 77-85, Mesa Veterinary Hospital, Golden, Colorado, 1977 (NIOSH). Technical Assistance Report 77-31, Boulder Memorial Hospital, Boulder, Colorado, 1977 (NIOSH). Hazard Evaluation Report 77-100, Porter Memorial Hospital, Denver, Colorado, 1977, (NIOSH). Hazard Evaluation Report 78-80, Denver General Hospital, Denver, Colorado, 1977 (NIOSH).



VII. AUTHORSHIP AND ACKNOWLEDGMENTS

Report Prepared By:

Bobby J. Gunter, Ph.D.  
Regional Industrial Hygienist  
NIOSH - Region VIII  
Denver, Colorado

Originating Office and  
Evaluation Assistance:

Jerome P. Flesch, Acting Chief  
Hazard Evaluation and Technical  
Assistance Branch (HETAB)  
NIOSH - Cincinnati, Ohio

Charles L. Geraci, Ph.D.  
Management Support Branch  
NIOSH - Cincinnati, Ohio

Report Typed By:

Marilyn K. Schulenberg  
NIOSH - Region VIII  
Denver, Colorado

TABLE I

Breathing Zone and General Room Air Concentrations of  
Halothane and Ethrane

Weld County Hospital  
Greeley, Colorado  
June 29, 1978

Sample Number	Job Classification	Operating Room No.	Time of Sample	Halothane PPM	Ethrane
1	Anesthetist	3	7:30 AM - 12:00 PM	*	*
2	Scrub Nurse	1	7:30 AM - 11:50 AM	*	*
3	Scrub Nurse	3	7:30 AM - 11:30 AM	0.53	*
4	Scrub Nurse	4	7:36 AM - 11:50 AM	0.46	*
5	Scrub Nurse	5	7:40 AM - 11:50 AM	0.41	*
6	Anesthetist	5	7:50 AM - 11:30 AM	*	*
7	Anesthesiologist	1	7:40 AM - 11:56 AM	1.73	*
8	Anesthesiologist	4	7:40 AM - 11:30 AM	0.30	*
9	Anesthetist	3	12:30 PM - 1:30 PM	*	*
10	Scrub Nurse	3	8:30 AM - 12:00 PM	*	16
11	-----	Recovery	8:00 AM - 12:00 PM	*	0.37
12	Scrub Nurse	5	11:50 AM - 2:00 PM	*	*
13	Scrub Nurse	4	11:50 AM - 1:00 PM	*	0.12
14	Scrub Nurse	1	11:50 AM - 1:30 PM	*	*
15	Scrub Nurse	3	12:30 PM - 1:35 PM	*	*
16	Anesthetist	5	11:40 AM - 1:00 PM	*	*
17	Anesthetist	3	12:10 PM - 1:40 PM	*	*
22	-----	Recovery	1:00 PM - 2:00 PM	*	0.20
25	Scrub Nurse	3	7:36 AM - 12:00 PM	*	0.28
26	Scrub Nurse	1	7:35 AM - 12:04 PM	*	*
27	Anesthesiologist	4	7:35 AM - 12:10 PM	0.13	*
28	Anesthetist	3	7:36 AM - 12:16 PM	*	0.33
29	Anesthetist	Recovery	11:25 AM - 1:20 PM	*	0.74
30	Anesthetist	5	7:36 AM - 12:06 PM	*	0.12
31	Anesthetist	2	9:20 AM - 12:30 PM	*	1.5
32	Anesthesiologist	1	7:35 AM - 12:45 PM	*	0.37
33	Scrub Nurse	5	7:35 AM - 12:05 PM	*	0.33
34	Scrub Nurse	4	7:35 AM - 12:01 PM	*	*
EVALUATION CRITERIA				0.5	0.5
LABORATORY LIMIT OF DETECTION IN mg/sample				0.01	0.01

\* = below laboratory limit of detection

TABLE II

Breathing Zone Air Concentrations of  
Nitrous Oxide (N<sub>2</sub>O)

Weld County Hospital  
Greeley, Colorado

June 29, 1978

Sample Number	Job Classification	Operating Room	N <sub>2</sub> O ppm
B	Anesthetist	3	65
R-1	Scrub Nurse	1	5
A-3	Scrub Nurse	3	30
F-3	Scrub Nurse	4	5
C-2	Scrub Nurse	5	25
4	Anesthetist	5	65
W	Anesthesiologist-1	1	20
E1	Anesthesiologist-2	4	10
U	Anesthesiologist-1	1	10
W	Anesthetist	3	60
D2	Scrub Nurse	3	30
G	Anesthesiologist-2	4	5
5	General Room	Recovery	25
Y	Scrub Nurse	5	15
B	Scrub Nurse	1	5
D-3	Scrub Nurse	4	5
A-3	Anesthetist	5	25
E-1	Anesthesiologist-1	1	25
B-1	Anesthetist	2	15
4	Anesthesiologist-2	4	5
C-2	Scrub Nurse	1	5
U	Anesthetist	2	10
5	Anesthetist	5	25
D-2	Anesthesiologist-1	1	5
W	Scrub Nurse	5	10
G	Anesthesiologist-1	1	10
Y	Anesthetist	2	50
D3	Scrub Nurse	3	15
B	Anesthetist	3	40
E-1	Scrub Nurse	4	5
A3	Anesthesiologist-2	4	20
B1	Anesthetist	5	25
D2	Anesthesiologist-1	1	5
U	Anesthetist	2	55

TABLE III

Breathing Zone Air Concentrations of Nitrous Oxide ( $N_2O$ )  
in Five Operating Rooms

Weld County Hospital  
Greeley, Colorado

June 29, 1978

OPERATING ROOM 1

Job Classification	$N_2O$ ppm
Circulating Nurse	20
Anesthesiologist-1	20
Anesthesiologist-1	10
Scrub Nurse	5
Anesthesiologist-1	25
Scrub Nurse	5
Anesthesiologist-1	5
Anesthesiologist-1	10
Anesthesiologist-1	5
<hr/>	
EVALUATION CRITERIA	25
LIMIT OF DETECTION IN MICROGRAMS/SAMPLE	0.8

Approximate 6 hour average exposure for anesthesiologist = 12.5 ppm

Approximate 6 hour average exposure for scrub nurse = 5.0 ppm

OPERATING ROOM 2

Job Classification	$N_2O$ ppm
Anesthetist	15
Anesthetist	10
Anesthetist	50
Anesthetist	55

EVALUATION CRITERIA 25

Approximate 6 hour average exposure = 32.50 ppm

OPERATING ROOM 3

Job Classification	N <sub>2</sub> O ppm
Anesthetist	65
Scrub Nurse	30
Anesthetist	60
Scrub Nurse	30
Scrub Nurse	15
Anesthetist	40

EVALUATION CRITERIA 25

Approximate 6 hour average exposure for anesthetist = 55 ppm  
Approximate 6 hour average exposure for scrub nurse = 25 ppm

OPERATING ROOM 4

Job Classification	N <sub>2</sub> O ppm
Scrub Nurse	5
Anesthesiologist-2	10
Anesthesiologist-2	5
Scrub Nurse	5
Anesthesiologist-2	5
Scrub Nurse	5
Anesthesiologist-2	20

EVALUATION CRITERIA 25

Approximate 6 hour average exposure for anesthesiologist = 10 ppm  
Approximate 6 hour average exposure for scrub nurse = 5 ppm



OPERATING ROOM 5

Job Classification	N2O ppm
Scrub Nurse	25
Anesthetist	65
Scrub Nurse	15
Anesthetist	25
Anesthetist	25
Scrub Nurse	10
Anesthetist	25

EVALUATION CRITERIA 25

Approximate 6 hour average exposure for anesthetist = 35 ppm  
Approximate 6 hour average exposure for scrub nurse = 16.7 ppm