

Vinyl Chloride Cytogenetics

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This report presents cytogenetic findings from a group of 209 workers employed for up to 28 years in the manufacture of vinyl chloride monomer at the Texas Division of Dow Chemical U.S.A. Cytogenetic evaluation results from this group were compared to results found in examination of individuals being considered for employment. Statistical analyses were performed on a group basis for chromatid aberrations, chromosome aberrations and proportion of abnormal cells; no statistical difference of significance was found between the two groups. Comparison of these results with reported studies suggests that the level of cytogenetic aberrations in vinyl chloride workers is probably related to the length and level of exposure, and that risk of adverse genetic effect can be avoided in controlled, minimal-exposure environments.

Vinyl chloride (VC) has been reported to be carcinogenic in animals^{1,2} and human beings³ and mutagenic in bacterial test systems^{4,5}. There have been reports of increased frequencies of chromosomal aberrations in workers exposed to vinyl chloride at polyvinyl chloride (PVC) facilities^{7,8,9}. Similar increases were not found upon cytogenetic evaluation of German PVC workers¹⁰ nor in our own preliminary investigation of a group of American VC workers¹¹. One study based on interviews with male employees has indicated an increased rate of fetal wastage among wives of VC-exposed workmen involved in polymerization operations¹².

We have recently completed cytogenetic evaluation of all currently-employed workers exposed to vinyl chloride at the Dow (Freeport) Texas Division. Both vinyl chloride and vinylidene chloride are produced in this plant. Vinylidene chloride has also been reported as mutagenic in bacterial test systems¹³. However, while these Dow workers are involved with the production of both compounds, the degree of contact with vinylidene chloride is far less than the exposure to vinyl chloride.

Vinyl chloride monomer has been produced at the Texas Division of Dow Chemical U.S.A. since 1948. There are no vinyl chloride polymerization operations. Average exposure levels, as is the case for almost all vinyl chloride monomer plants, have been generally lower than those reported for facilities involved in polymerization operations¹⁴. Although the threshold limit values for vinyl chloride had been 500 parts per million (ppm) until 1974, Dow had established a goal of 50 ppm or below in 1959.¹⁵ The current Occupational Health and Safety Administration (OSHA) standard for vinyl chloride is one ppm as a time-weighted-average over an eight-hour workday. We have found that the TWA concentration of vinyl chloride for all VC-related job classifications was approximately 5 ppm from 1968 to 1973; in 1974, TWA concentrations were 1 to 2 ppm. Since 1975, average levels of exposure have been lowered to less than 1 ppm. Estimates of average exposure levels for the years prior to 1968 are less reliable. For the purposes of this study, estimates of exposure were calculated for specific job classifications, based on records of both personnel and area monitoring (Table 1). However, actual exposure may vary from individual to individual within the same job classification and accidental, short-term exposures of some workers to concentrations in excess of the standard probably occur from time to time. Documentation of such incidents is difficult.

Methods

Our study group was composed of 209 workers who had worked in the vinyl chloride plant for periods ranging between one and 332 months (average, 48.3 months) at the time of this investigation. As part of the medical surveillance program for this group, peripheral blood samples were obtained, and evaluation of lymphocyte chromosomes was performed. Lymphocytes were cultured using a modification of the Moorhead technique.¹⁶ Standard procedures, as previously described,¹⁷ were used for incubation, processing, and analysis.

Findings were compared to cytogenetic data from a group of 295 "preemployment examinees" who had chromosome evaluation done as part of routine preemployment examination and who, at that time, had no known exposure to chromosome-break-

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- DA. Chromosome preparations of leukocytes cultured from human peripheral blood. *Exp. Cell Res.* **20**: 613-616, 1960.
17. Kilian DJ and Picciano D. Cytogenetic surveillance of industrial populations. In *Chemical Mutagens: Principles and Methods for their Detection*, Vol. 4. New York: A. Hillaender, ed. Plenum Press, 1976, pp. 321-339.
18. Funes-Cravioto et al. Reference 7.
19. Ducatman et al. Reference 8.
20. Heig & Thiess. Reference 10.
21. Purchase et al. Reference 9.
22. Xian et al. Reference 11.
23. Court Brown WM. Human population cytogenetics. In *Frontiers of Biology*, Vol. V. A. Neuberger and EL Tatum, eds. North Holland Publishing, 1967, pp. 1-31.
24. Buckton KE, Jacobs PA, Court Brown WM, and Doll R. A study of chromosome damage persisting after x-ray therapy for ankylosing spondylitis. *Lancet* **2**: 676-682, 1962.
25. Bloom AD, Nakagome Y, Awa AA, and Nerishi S. Chromosome aberrations and malignant disease among A-bomb survivors. *Amer J Pub Health* **60**: 641-644, 1970.
26. Vaughan J. Bone disease induced by radiation. *Int Rev Exp Path* **1**: 243-396, 1962.
27. Fischer P, Golob E, Kunze-Mühl E, Ham AB, et al. Chromosome aberrations in peripheral blood cells in man following chronic irradiation from internal deposits of Thorotrast. *Radiat. Res.* **29**: 505-515, 1968.
28. Garza-Chana R, Leah CH, Alvarez M, and Sanchez FI. Chromosome analysis in males occupationally exposed to lead. Abstract 325. In *Abstracts V International Congress of Human Genetics*, S. Armentrout and R. Usker, eds. Excerpta Medica, Amsterdam, 1976.
29. Sampson RJ, Key CR, Buncher CR, and Iijima A. Thyroid carcinoma in Hiroshima and Nagasaki. I. Prevalence of thyroid carcinoma at autopsy, Hiroshima 1957-68, Nagasaki 1951-67. *ABCC Technical Report* 25-68, 1968.
30. Muller J, David A, Rejskova M, and Brezikova D. Chronic occupational exposure to strontium-90 and radium-225. *Lancet* **2**: 129-131, 1961.
31. Fischer P, Golob E, Kunze-Mühl E, and Müllner T. Chromosomal aberrations in thorium dioxide patients. *Ann NY Acad Sci* **145**: 769-786, 1967.

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