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1	20	21	40	41	60	61	62
Trop. Infect. Dis. P. Goulet L						11	
						12	
						13	

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1	2	61	62
		21	
Birth Defects in a Community Located Near		22	
a Uranium Chemical Plant - - HUMAN, CANADA		23	
Epidemiology		24	

Source (Journal, Vol., Number, Pages, Date)

1	2	61	62
Report by the Dept. of Social and Family		31	
Medicine, Laval University, Quebec		32	

Brief Summary

1	2	10	61	62
SUMMARY:			61	
			62	
			63	
			64	

R&S 110847

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BIRTH DEFECTS IN A COMMUNITY
LOCATED NEAR A VINYL CHLORIDE PLANT

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R&S 110848

BIRTH DEFECTS IN A COMMUNITY
LOCATED NEAR A VINYL CHLORIDE PLANT

The discovery of the carcinogenic action of vinyl chloride monomer in man has swiftly raised the question of the teratogenic properties of this chemical substance.

The mutagenic capacity of vinyl chloride monomer and its metabolites has been confirmed in vitro within recent months.⁽¹⁻⁴⁾ The presence of chromosomal aberrations at abnormally high levels in the lymphocytes of vinyl chloride monomer workers has also been reported⁽⁵⁻⁷⁾.

Studies showing the presence of birth defects in humans following exposure to vinyl chloride monomer are however much less convincing. The rate of fetal mortality and abortion has been noted to be higher than expected in the wives of vinyl chloride monomer workers.⁽⁸⁾ However, this rate was established using questionnaires filled out by the workers themselves thus attenuating the strength of any conclusion.

A recent study⁽⁹⁾ concerning the oncogenic and mutagenic risks of residents in communities where there are vinyl chloride producing plants, suggests the existence of an association between the presence of such plants in a population and elevated rates of malformations in the population. Using data from Ohio State's Birth Registry (congenital malformations have been reported since 1968), the author compared the rate of birth defects in three cities where vinyl chloride polymerization plants are located with rates observed in other cities of the same State. The study shows that the

rate of birth defects is higher in these three cities than in the rest of the State. The difference was particularly marked for malformations of the central nervous system, upper digestive system and the feet (club-feet).

Using data from a different source (Center for Disease Control's hospital-based birth defects monitoring program) other authors confirm that the rate of defects of the central nervous system is higher in Painesville Ohio where a vinyl chloride plant⁽¹⁰⁾ is located. They show however that the mothers of the malformed children were less exposed to vinyl chloride than the control mothers and thus conclude that the high rate of defects is not related to exposure to vinyl chloride. Moreover, in another city where a vinyl chloride plant was located, the rate of birth defects was not higher than predicted.

The aim of the study is to confirm the presence of an increase in the frequency of congenital malformations in a community living next to a vinyl chloride polymerisation plant by comparing birth defects in such a community (Shawinigan) with those of another one (Drummondville) chosen as a control.

METHOD

There has been a vinyl chloride polymerisation plant in Shawinigan since 1943 which hired 250 men on average over the years. Production has fluctuated with time. At the moment, the amount of vinyl chloride monomer released in the atmosphere is estimated at approximately 200 ppm. No data is available to estimate the quantity of vinyl chloride monomer released in the air in the past; it is doubtful it was any lower than today. Ten workers of the plant were found having died of angiosarcoma of the liver.⁽¹¹⁾ The plant is surrounded by other heavy industries such as an aluminum refinery plant, a carbide plant, a chemical factory and a pulpe and paper mill. As the city is located in a hollow between hills and predominant

winds blow from the industries toward the residential zone, the community has been exposed to relatively high concentrations of atmospheric pollutants.

Drummondville is also an industrialised city. Its population (31,000) shows an age and sex structure similar to Shawinigan (29,000). Its major industries comprise textile factories one of which handling synthetic fibers, a lamp factory and many small businesses. It is located fifty miles south west of Shawinigan.

DATA SOURCES

Data concerning birth defects were obtained from the files of hospitals where the women residing in the two cities give birth. For geographic reasons, Shawinigan data were obtained from six regional hospitals while Drummondville data could be found at the city only hospital. In both cases, more than 96% of all records could be retrieved.

Malformed children were ascertained through the following sources: the mother's file, the newborn's file, the obstetric room's daybook and the autopsy reports.

DEFINITION OF A BIRTH DEFECT

Birth defects included in this study were those either apparent at birth, or discovered by an X-Ray or a laboratory examination (this excludes heart defects) or described in autopsy reports.

The following abnormalities were not included as such in the study:

a) locomotor apparatus

- metatarsus varus
- calcaneo valgus and varus
- misimplanted toes
- internal rotation of the foot
- flat-foot

b) genitals

cryptorchidism
hydrocele
undescended testes
phimosis

c) cutaneous-mucous membrane

nevus vasculosus, nevus venosus
mongolian spots
cavernous hemangiomas
pilonidal sinus
benign teratoma

d) others

pectus excavatum
stenosis of external urethral meatus
thymus hypertrophia
small cleft palate
eyelid ptosis
minor deformity of the thorax
short string of the tongue

It was further required, for a child to be considered malformed that defects be diagnosed at the time of the discharge from hospital.

Still-births comprised only children still-born at or after twenty weeks of pregnancy.

The annual number of live births whose parents resided in each city was obtained from the Quebec birth registration file.

RESULTS

Table I gives the distribution of malformed children by parents' place of residence. The Chi square test indicates that the difference is significant between the two cities.

R&S 110852

Table II shows that the ratio of malformed children per 1,000 live births were higher in the city where a vinyl chloride polymerization plant is located. This is true for all types of defects except those of the endocrine system. It may also be noted that the mean age of the mother at delivery in Shawinigan is often lower than in Drummondville.

Table III shows the distribution of malformed children according to number of defects per year and city of residence. Among malformed children, the proportion with multiple malformations is similar in both cities (Shawinigan 28/124, Drummondville 28/116). It would seem therefore that the higher ratio of malformed children in the city where the vinyl chloride plant (RR. 1.56) is located does not extend to multiple malformations.

Except for the years 1971-72, the increase in birth defects in Shawinigan is constant over the nine year period of observation. (Figure I) No reason could explain this difference in the ratio of defects each year.

Table IV shows that the ratio of still-births per 1,000 live births is similar in both cities. However, proportion of malformed children among stillborns is somewhat higher in Shawinigan (21% versus 18%). This is concurrent with the finding of a greater number of children with birth defects in Shawinigan.

DISCUSSION

The present study is in agreement with the studies carried out in Ohio, where a higher birth defect rate was observed in a city with a vinyl chloride polymerization plant. However, the presence of other atmospheric pollutants in the environment of Shawinigan should be noted. The present study does not permit linking the observed phenomenon to a particular pollutant be it vinyl chloride or any other teratogenical agent.

The congenital defects observed hereby in Shawinigan as well as in Drummondville were higher than those observed in Ohio. Differences could be accounted for by the data collection method. The present study gathered data from hospital records while the Ohio study used birth registers. It could be expected that certain cases be not reported to a state register.

It is puzzling to observe that the increase of birth defects is in the same order of magnitude in several organ systems. Usually, any particular chemical would produce a specific effect rather than a broad increase in birth defects generally. The explanation to such an observation could be twofold:

- 1 - The chemical in the Shawinigan environment produces by itself or in combinaison with other air pollutants a broad spectrum of developmental defects. This observation disagrees with previous studies showing a possible association between vinyl chloride in the environment and central nervous system defects.⁽¹⁰⁾ The problems faced with in Shawinigan could be of a different order of magnitude than those seen elsewhere.
- 2 - Even if careful attention was given to the collection of data and even if more than 96% of all births were reviewed in both cities, the apparent difference between Shawinigan and Drummondville can result from an artifact such as ignoring defects diagnosed after the hospital discharge, different diagnosis or different reporting criteria used in the seven hospitals contacted or some inherent lack of comparability between the study and control population.

Further studies are needed to assert that vinyl chloride causes birth defects in the community living near a polymerization plant. Should this be the case, stringent emission standards will have to be devised.

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ABSTRACT

Birth defect ratios observed in a community where a vinyl chloride polymerization plant is located since 1943 (and where ten cases of angiosarcoma of the liver have been diagnosed among vinyl chloride monomer workers) are compared with the birth defect ratios observed in another city chosen as control. The frequency of congenital defects is higher in the exposed community. This phenomenon is observed in seven out of the nine years covered by the study. Women who gave birth to malformed children were on average younger in the exposed community. Both sexes are affected equally. These findings are partly in agreement with other already published previous studies and should lead to further researches.

R&S 110855

REFERENCES

1. RANNUG V., JOHANSSON A., RAMEL C. and WACHMEISTER C.A.: "The mutagenicity of vinyl chloride after metabolic activation" *Ambio* 3, 174, 1974.
2. BARTSCH H., MALAVIELLE C., MONTESAND R.: "Human rat and mouse liver-mediated mutagenicity of vinyl chloride in *S. Typhimurium* strains" *Int. J. Cancer* 15, 429, 1975.
3. HUBERMAN E., BARTSCH H. and SACHS L.: "Mutation induction in chinese Hamster V 79 cells by two vinyl chloride metabolites, chloroethylene oxide and 2-chloroacetaldehyde" *Int. J. Cancer* 16, 639, 1975.
4. ELMORE J.D., WONG J.L., LAUNBACH A.P. and STREIPS U.N.: "Vinyl chloride mutagenicity via the metabolism chloroxirane and chloroacetaldehyde monomer hydrate" *Biochim. Biophys. Acta* 442 (3): 405, 1976.
5. DUCATMAN E., HIRSCHORN K. and SELIKOFF I.: "Vinyl chloride exposure and human chromosome aberrations". *J. Mutation Res.* 31, 163, 1975.
6. FUNES-CARIOTO F., LAMBERT B., LINDSTEN J., EHRENBURG L., NATARAJAN A.T. and OSTERMAN-GOLKAR S.: "Chromosome aberrations in workers exposed to vinyl chloride". *Lancet*, i., 459, 1975.
7. FOMENKO V.N., KATOSOVA L.D. and PAVLENKO G.: "Cytogenetic analysis of the peripheral blood in workers exposed to vinyl chloride polymerization". *Gigiena Truda i Professional'nye Zabolevania* Vol. 9, Sept. 1976, 48-50.

8. INFANTE P.F., WAGONER J.K., McMICHAEL A.J., WAXWEILLER R.J. and FALK H.: "Genetic risks of vinyl chloride" Lancet 734, April 3, 1976.
9. INFANTE P.F.: "Oncogenic and mutagenic risks in communities with polyvinyl chloride production facilities". Annals N.Y. Aca. of Sciences, 271, 49, 1976.
10. EDMONDS L.D., FALK H. and NISSIM J.E.: "Congenital malformations and vinyl chloride". Lancet, Novembre 29, 1098, 1975.
11. DELORME F. and THERIAULT G.P.: "Ten cases of angiosarcoma of the liver in Shawinigan, Quebec" to the press.

R&S 110857

TABLE I - Distribution of malformed children / 1,000 live births by parents' place of residence 1966-1974.

	Shawinigan	Drummondville
Number of malformed children	124	115
Ratio malformed children per 1,000 live births	30.8	19.7

$$\chi^2 = 12.359, \text{ d.f. } 1, p < 0.001$$

Vinyl chloride and birth defects

TABLE II

Distribution of malformed children/1,000 live births, relative risk and mean age of mother at delivery time, by type of malformations and city, 1966-1974.

	Ratio for both sexes		Relative Risk Sh./Dr.	Mean age of mother	
	Shawinigan	Drummondville		Shawinigan	Drummondville
	(4,022)*	(5,875)*			
Central Nervous System	(21)** 5.2	(24) 4.1	1.27	24.6	28.7
Eye, Ear, Face	(13) 3.2	(8) 1.4	2.29	22.7	23.0
Cardio-vascular	(28) 7.0	(26) 4.4	1.59	26.7	28.8
Respiratory	(9) 2.2	(3) 0.5	4.40	24.2	23.8
Digestive	(12) 3.0	(10) 1.7	1.76	24.7	29.7
Genital	(15) 3.7	(18) 3.1	1.19	25.6	24.6
Urinary	(8) 2.0	(6) 1.0	2.0	22.8	21.7
Locomotor	(49) 12.2	(43) 7.3	1.67	27.0	29.2
Chromosomal	(6) 1.5	(8) 1.4	1.07	28.5	36.2
Endocrine	(0) -	(2) 0.3	0	-	26.4

* Number of live-births

** Number of children with that defect.

TABLE III

Distribution of malformed children by year and parent's city of residence according to number of defects (1966-1974)*.

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of defects	1966		1967		1968		1969		1970		1971		1972		1973		1974		T O T A L	
	S (610) **	D (761)	S (510)	D (689)	S (454)	D (681)	S (466)	D (677)	S (438)	D (648)	S (396)	D (636)	S (372)	D (583)	S (368)	D (540)	S (408)	D (660)	Shawinigan (4022)	Drummondville (5875)
1	14	8	10	7	9	7	6	7	16	10	3	9	6	11	18	13	14	16	96	88
2	2	2	2	1	2	2			1	1	1	4	2	5	1	2	2	4	13	21
3	3		2		1				1		1	1		1		1			8	3
4		1					1				1								2	1
5									1				1					1	2	1
6			1																1	
7																				
8			1					1											1	1
9																				
10																	1		1	
11				1																1
Total	19 (31.1) ***	11 (14.5)	16 (31.4)	9 (13.1)	12 (26.4)	9 (13.2)	7 (15.0)	8 (11.8)	19 (43.4)	11 (17.0)	6 (15.2)	14 (22.8)	9 (24.2)	17 (29.2)	19 (51.6)	16 (29.6)	17 (41.7)	21 (31.8)	124 (30.8)	116 (19.7)

* Still born defectives included

** Number of live births

*** Ratio malformed children/1,000 live-births

S = Shawinigan

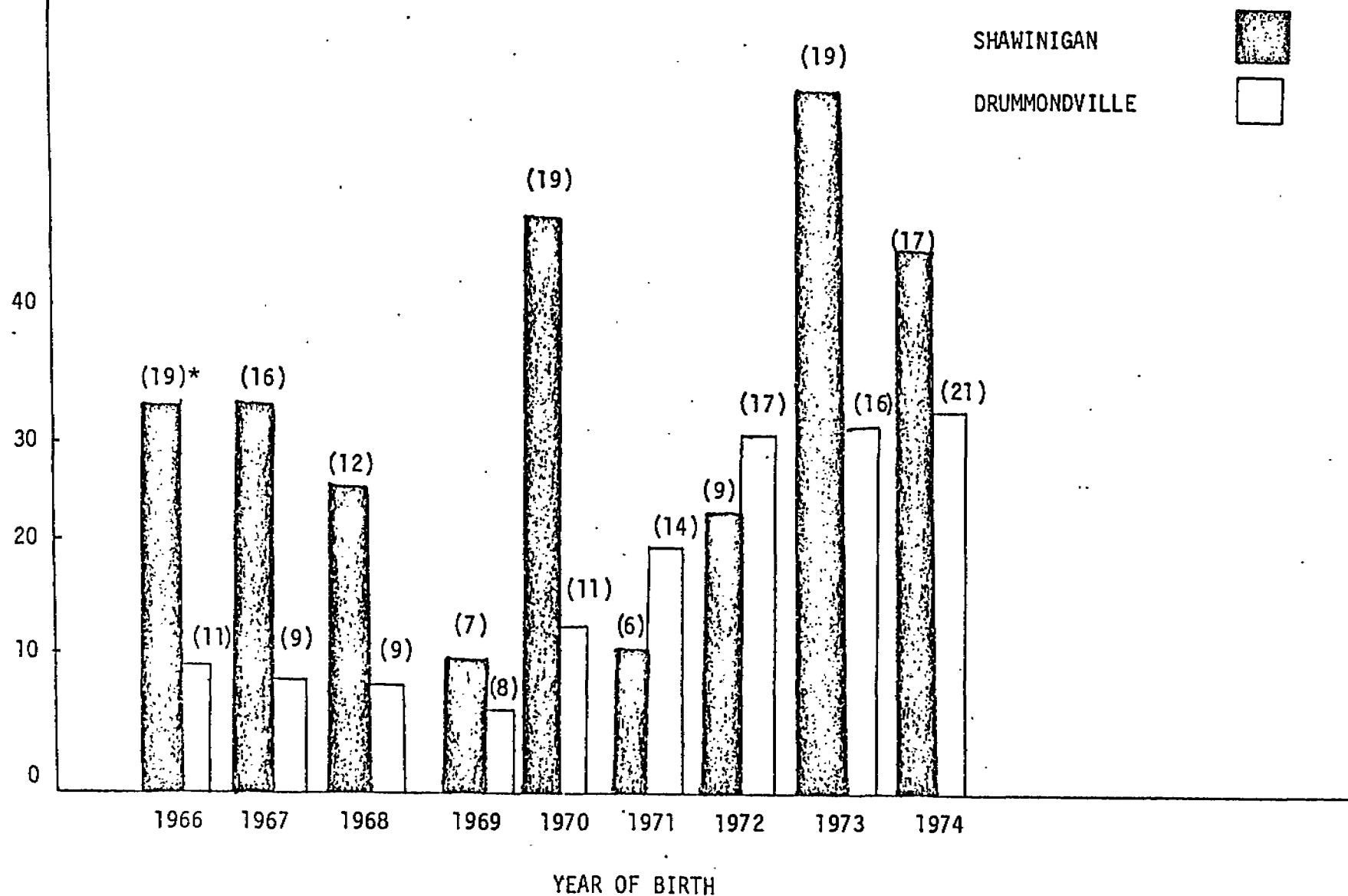
D = Drummondville

Vinyl chloride and birth defects

FIGURE 1

Annual distribution of malformed children / 1,000 live-births by parents city of residence for the years 1966-1974

Malformed children / 1,000 live-births



* Number of children

R&S 110861

TABLE IV

Distribution of still births/1,000 live births by parents place of residence, category of still births and mean age of the mother 1966-1974

	SHAWINIGAN	DRUMMONDVILLE	MEAN AGE OF THE MOTHER	
			Shawinigan	Drummondville
Still birth without malformation	(37)* 9.2	(64) 10.89	28.3	28.8
Still birth with malformation	(10) 2.49	(14) 2.38	26.1	29.3
TOTAL still-births ratio per 1,000 live births	(47) 11.69	(78) 13.27	27.2	29.1

* number of cases.