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Reference # 31

DIFFUSE PLEURA ASBESTOS NORTH WEST

J C WAGNER C A

From the Pathology Division, Pneumo-
Industrial Research, Johannesburg
Department of Thoracic Surgery, Cancer

(RECEIVED 16)

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from elsewhere in the Union. During this period
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This is a preliminary publication and the problem
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the eastern portion of the magisterial district of
Postmasburg and finally, to the western area of the

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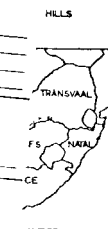
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again in the Kuruman dis-
trict in 1918 a large number of
This northward trend has
since 1950 mining started at
malind border (Table 2).
quarried in numerous small
his was followed gradually

TABLE 1
CROCIDOLITE AND AMONITE
mines (1952)

mine	total tonnage	amonite
Adams	41.94	49.47
Adams	0.20	0.63
Adams	18.64	4.15
Adams	19.16	35.63
Adams	1.17	6.57
Adams	0.18	0.11
Adams	6.07	0.02
Adams	8.10	0.20
Adams	0.11	0.07
Adams	2.48	2.33
Adams	—	0.35
Adams	—	0.61
Adams	—	0.01
Total	100.73	100.46

and here whites may
vicinity of the mines
The type of asbestos mined throughout this area
is crocidolite, better known as Cape Blue Asbestos
The chemical analysis of the fibre is given in Table 1
Crocidolite is the fibrous form of riebeckite and all
stages of transformation from a massive riebeckite
rock through lamellar riebeckite to asbestiform
crocidolite occur in this region. Magnetite is
frequently associated with the crocidolite (Vermaas,
1952).
Mining of asbestos first began in the Prieska
district in 1893 and gradually spread northwards

DIFFUSE PLEURAL MESOTHELIOMA AND ASBESTOS EXPOSURE IN THE NORTH WESTERN CAPE PROVINCE

BY

J. C. WAGNER, C. A. SLEGGES, and PAUL MARCHAND

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(RECEIVED FOR PUBLICATION APRIL 24, 1960)

Primary malignant tumours of the pleura are uncommon. Thirty-three cases (22 males, 11 females, ages 31 to 68) of diffuse pleural mesothelioma are described, all but one have a probable exposure to crocidolite asbestos (Cape blue). In a majority this exposure was in the Asbestos Hills which lie to the west of Kimberley in the north west of Cape Province. The tumour is rarely seen elsewhere in South Africa.

Mesothelioma of the pleura is regarded as an uncommon tumour. In the last four years we have seen 33 histologically proven cases. 28 of these had some association with the Cape asbestos field and four cases had been exposed to asbestos in industry.

The tumour is rarely encountered elsewhere in South Africa. During the past five years, with the exception of the present series, no neoplasm of this nature has been diagnosed amongst 10 000 lungs examined at the Pneumonomus Bureau in Johannesburg, or in the Pathology Department of the South African Institute for Medical Research (Higginson and Oxley (1957) did not observe a single case in their survey of malignant tumours occurring in the Bantu and Cape Coloured population of Johannesburg and the North Eastern Transvaal).

Our first necropsy specimen of pleural mesothelioma with asbestos was examined at the Pneumonomus Research Unit in February, 1956 (Case 1). During the early months of that year, one of us (C.A.S.) in the Northern Cape, treated six patients with gross pleural thickening. Pleural biopsies from two of them showed the features of mesothelioma. In the ensuing two years eight further cases were found from this region and five from elsewhere in the Union. During this period C.A.S. had become perturbed at the number of these unusual tumours occurring amongst his patients, and stimulated an investigation. At this stage there were two reasons to suggest that asbestos might be implicated. First, asbestos was found in

the lungs of the first case (Case 1) and secondly 10 of the cases came from a hospital to which suspected cases of tuberculosis were referred from a large asbestos mining area. This hypothesis could not be supported if once from the original histories obtained from the patients, for they included housewives, domestic servants, cattle herders, farmers, a water buff, an insurance agent and an accountant none of whom were working on the asbestos mines at the time. We therefore undertook a detailed investigation of their past occupation and place of residence and the association with asbestos exposure was discovered. The cases are summarized in Table 3. Previous biopsy specimens were re-examined and new cases diagnosed, including asbestos miners. In only one case do the relatives deny that the patient either visited the asbestos mines or was exposed to asbestos.

This is a preliminary publication and the problem is being intensively investigated.

The Asbestos Area of the North-West Cape

According to Hill (1930) asbestos was discovered by Lichenstein near Prieska during his travels between 1893 and 1906. Since then it has been established that the asbestos deposits extend from 20 miles south of Prieska, northwards through the western part of the magisterial district of Hay, to the eastern portion of the magisterial district of Postmasburg and finally, to the western area of the

DIFFUSE PLEURAL MESOTHELIOMA

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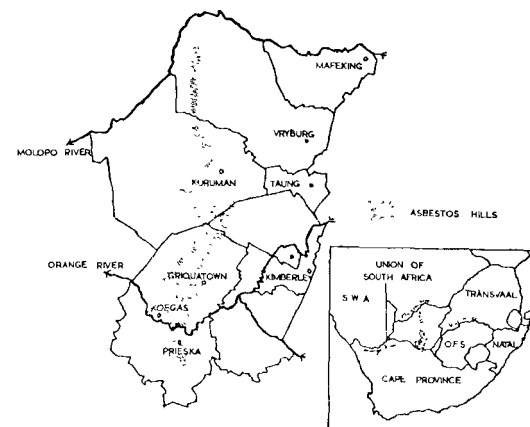


FIG. 1. Map of Grouped West asbestos fields.

district of Kuruman (Fig. 1). These deposits occur in the slopes of a range of hills covering an area of approximately 8 000 square miles. Known as the "Asbestos Mountains", these hills extend more or less longitudinally between 22° 30' and 23° E. The whole area is semi-arid, sparsely populated and, apart from mining, cattle ranching is the only important occupation. In the Kuruman area several large tracts of land in or abutting on these mountains have been reserved for the aboriginal inhabitants and here whites may only reside in the immediate vicinity of the mines.

The type of asbestos mined throughout this area is crocidolite, better known as Cape Blue Asbestos. The chemical analysis of the fibre is given in Table 1. Crocidolite is the fibrous form of riebeckite and all stages of transformation from a massive riebeckite rock through luncular riebeckite to asbestos-form crocidolite occur in this region. Magnetite is frequently associated with the crocidolite (Vermaas, 1952).

Mining of asbestos first began in the Prieska district in 1893 and gradually spread northwards

In 1908 production had begun in the Kuruman district. Between 1916 and 1918 a large number of claims were taken up. This northward trend has continued and in about 1950 mining started at Poffert near the Bechuanaland border (Table 2). Initially the ore was quarried in numerous small open cast workings. This was followed gradually

TABLE 1
CHEMICAL ANALYSIS OF CROCIDOLITE AND ANOMITE
(From Vermaas, 1952)

	Crocidolite	Anomite
SiO ₂	51.04	49.43
Al ₂ O ₃	0.20	0.63
FeO	18.84	4.15
CaO	19.39	35.63
MgO	0.97	0.52
CaO	0.19	0.52
Na ₂ O	6.07	0.02
K ₂ O	0.04	0.20
H ₂ O	0.31	0.01
H ₂ O	2.18	2.13
Totals	—	0.25
Loss	—	0.61
F	—	0.01
Total	100.73	100.46

TABLE 2
POPULATION OF THE MAGISTERIAL DISTRICTS IN WHICH THE ASBESTOS MINES ARE SITUATED
(taken from Union Census 1921 and 1951)

District	Population 1921			Population 1951		
	Whites	Non-whites	Total	Whites	Non-whites	Total
Prickia	1410	4879	6289	1361	10987	12348
Kuruman	4713	16608	21321	4818	21779	26597
Hay	4389	8814	13203	2757	17108	19865
Prinsburg		Included in the Hay District		4987	21019	25906

by a type of shallow mining, inclined shafting became more common after 1930.

According to Froid (1915) the quarrying was a family affair undertaken by the local inhabitants. The men quarried the rock, which was sorted and then hand cobbled by the women and children. Hand cobbling consists of separating the fibre from the banded transverse by striking the rock cob with a small hammer. As the market was selective, a rotary sieve was sometimes used to grade the fibre and eliminate the small particles of dust. After grading, it is banded and weighed. The asbestos was then transported by donkey wagon to the nearest railway. Before the establishment of the railway in this area between 1923 and 1930, such a journey could last as long as 10 days. This transport was generally undertaken by white youths (farmers' sons) under contract. The white inhabitants played very little part in actual recovery of the fibre, being nearly always employed in the role of managers or overseers, issuing stores and supervising the grading and weighing of the fibres. Mining activities were and are still carried out round the mines.

Since the 1939-1945 war, the demand for asbestos has enormously increased. The lucrative claims have been bought by registered companies; others have been abandoned. As deeper and richer deposits have been found, vertical shafts are being sunk. However, the mines with large shallow deposits are still using the inclined shaft and tunnel and quarrying is still used by the few remaining outcrop producers. With the building of more mills, hand cobbling has diminished. In 1915 the first crushing mill was established at Koppies in the south. This was followed by a large mill at Kuruman, operated between 1926 and 1931, where it was situated within 300 yards of the main street, close to which cobbling was also done for a few years (1927-1930). This was followed by a mill at Prickia in 1930, which was completely rebuilt in 1957. Georgetown had a small mill in 1928. The practice today is for one mill to serve several mines in the immediate vicinity.

In the early days the manager and labourers lived within a few yards of their place of work, and even

today the non-white prefers to live as close as possible, and the children play on the dumps from the mine and mill.

Case Histories

The following eight case histories illustrate various aspects of the disease and the different types of exposure to asbestos dust.

Case 1. B.P., a Bantu male, 36 years of age (born 1920), was a mine labourer, and was the first case diagnosed as a mesothelioma with evidence of asbestosis. He was born in the Kuruman district but it is not known whether he worked in the asbestos mines. He was employed on the Witwatersrand gold mines underground for two years, and in the change rooms for a further 11 years.

A radiograph taken at a mine hospital on August 18, 1955, showed a massive right-sided pleural effusion, and 3,000 ml. of fluid was withdrawn. He was admitted to the Witwatersrand Native Labour Association Hospital on August 24, 1955, and two days later aspiration yielded 1,000 ml. of thick, gelatinous pus, which could be pulled out in threads. He was treated with frequent aspirations and instillation of vasodose but without improvement, and he died on February 15, 1956 (Manning, 1956).

At autopsy the right thoracic cavity was occupied by a large gelatinous tumour which displaced the mediastinum and compressed the left lung. The tumour had infiltrated the pericardium. The right lung was completely compressed by neoplastic tissue (Fig. 2) but the right bronchial tree did not show any evidence of a primary bronchogenic carcinoma. Histological sections of the pleural growth showed a papillary mesothelioma (Fig. 3). There was evidence of asbestosis in both lungs (Fig. 4).

Case 4. K.H., a white female, 56 years of age (born 1898), was a social worker, who could only have had a short exposure to asbestos as a child and probably a further short exposure as a young woman. She may have had several short visits to the mines with her husband at a later period.

She was born in Georgetown where she lived until she was 5 years of age. Her family then moved to Kimberley where from 1916 to 1922 she worked as a clerk in an asbestos warehouse. Her husband owned an asbestos mine from 1913-1940.



FIG. 2

Fig. 2. Right lung of Case 1, showing a tumour compressing the lung substance.

Fig. 3. Cellular structure of tumour of Case 1. (H&E, 40x magnification and stain).

Fig. 4. Sections of lung from Case 1 showing asbestos (1 day 40 H&E, 40x magnification and stain).



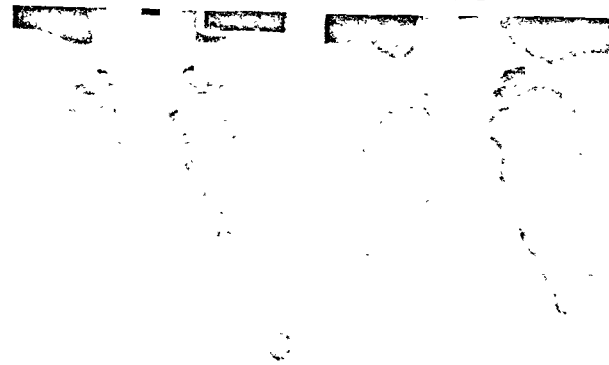


Fig. 5. On July 21, 1954.

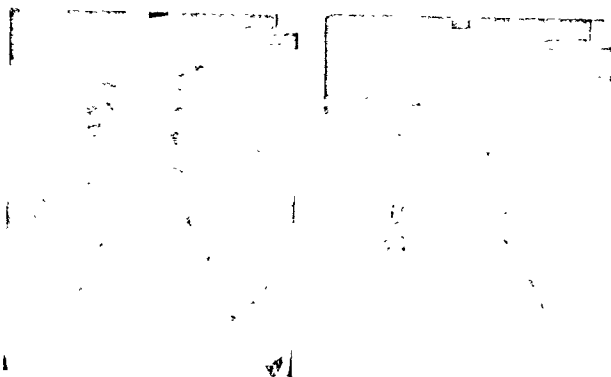


Fig. 6. On July 2, 1955.

Fig. 7. On March 3, 1956.

Fig. 8. On July 3, 1956.

Figs 5, 6, 7, and 8. Radiographs showing development of tumour in Case 4.



Fig. 9. Right lung of Case 14.



Fig. 10. Histological sections of tumour from Case 14. Hematoxylin and eosin.

She was referred to the West End Hospital, Kimberley, on July 19, 1954, because of dyspnoea and right chest discomfort of acute onset. The chest radiograph showed a localized pleural effusion in the right base. Straw-coloured fluid was aspirated, which on examination was found to be negative for *M. tuberculosis* but contained mesothelial cells arranged in acini. At thoracoscopy in Johannesburg on August 11, 1954, tumour nodules were seen on the parietal pleura. Histological examination of the biopsy specimen showed a mesothelioma. Deep therapy and radio-active gold implantation in September, 1955, caused initial regression of both the pleural fluid and nodules but did not arrest the progress of the disease. She received a further course of *oxy* therapy without improvement and died in January, 1957. The liver was enlarged and nodular before death but no necropsy was done. (Figs 5, 6, 7, and 8 show the radiological development of the tumour in this case.)

Case 14. S.S., a Bantu male, 40 years of age (born 1918), was an asbestos miner. He was born in the Kuruman district and as a child often played on the asbestos dump near an asbestos mine. He subsequently worked with asbestos for several years, being employed at an asbestos mine weighing fibre from 1938-41. In 1942 he worked on a Witwatersrand gold mine but

returned to the Kuruman district in 1943 where he worked as a farm labourer until his final illness.

He was admitted to the West End Hospital on March 3, 1959. He had been well until July, 1958, when he became aware of pains in the right side of his chest. In addition he had a slightly productive cough and shortness of breath on exertion. Chest radiograph on March 9, 1959, showed obliteration of the right lung field. Asbestos bodies were found in the sputum. Needle biopsy showed histological features consistent with a mesothelioma.

At autopsy. The right chest was occupied by a large whitish tumour which infiltrated the mediastinum, extending through the diaphragm onto the superior surface of the liver. A straw-coloured effusion was present in the left pleural space but the left lung was not affected. The mediastinal, cervical, and para-aortic abdominal lymph glands to the level of the renal arteries were infiltrated with tumour.

In the right lung it was found that the tumour mass had infiltrated along the interlobar fissures. The lobes of the lung were compressed into three small segments (Fig. 9). On histological section a papillary type of pleural mesothelioma was seen (Fig. 10) which had infiltrated into the lung parenchyma in some areas. There was marked lymphatic spread throughout the

lung substance. Numerous asbestos bodies were present in the air spaces and occasional bodies and fibres were seen in the interstitial tissue. The lung itself was markedly atelectatic. Evidence of asbestosis was observed in the left lung.

Case 15. S.S.V., a white female, 42 years of age (born 1916), is a housewife, who lived on a mine near Johannesburg and was treated for tuberculous pleurisy for six months. It was only after the histological diagnosis of a mesothelioma had been made that her previous history was obtained. In addition, it was subsequently learnt that her father (Case 22) had been born on the asbestos belt and had a long mining history.

The patient was born in the Griquatown district and went to school at Kuruman. She spent most of her youth in the asbestos belt. She has lived near Johannesburg for 20 years.

In March 1958 this patient developed right chest pain and rapidly increasing shortness of breath. She was admitted to hospital and 2 500 ml of slightly blood-stained fluid was removed from the right chest. A thoracoscopic examination was carried out on July 27 when a small biopsy was taken but this consisted of fibrous tissue. Bronchoscopy was normal and no neoplastic cells were observed in the bronchial washings. *M. tuberculosis* was not cultured from numerous specimens of sputum, pleural fluid, nor from the bronchial washings. She was admitted to a Johannesburg nursing home on October 16, 1958.

Radiographs then showed massive thickening of the right pleura with fluid in the right chest. At thoracotomy (Mr. Dennis Laffert) the pleural fluid was 2 cm thick, hard as rock, and stripped with great difficulty from the chest wall. The visceral pleura stripped fairly easily from the upper lobe but separation became more difficult over the lower lobe where the peel was thicker. During the dissection a necrotic mass was found in the periphery of the left lower lobe. At the base, confluent with the diaphragm and perhaps the mediastinum, was a hard orange mass which could not be removed. Histological sections of the specimen revealed an advanced form of tubular mesothelioma. Subsequently three specimens of sputum were examined for asbestos bodies but none were found.

The patient is still alive (April, 1960) but her condition is deteriorating.

Case 22. D.A.B. a white male, 68 years of age, and a farmer was the father of the previous patient. Although a diagnosis of pleural mesothelioma was made in January, 1957 it was not until July, 1959 that details of his exposure to asbestos were obtained.

He was born in Griquatown and went to school there. He worked in Kuruman, Pommersburg and Griquatown until the age of 62. Between 1918 and 1932 he worked as an asbestos miner underground and was mostly in charge of blasting. Between 1932 and 1934 he worked in the district but returned to asbestos mining from 1945 to 1952.

In January, 1957 the patient consulted his doctor because of left-sided chest pain with cough and slight

haemoptysis. He was referred to the Johannesburg General Hospital for investigation. The chest radiograph showed gross pleural thickening with a basal pleural effusion on the left side. He was admitted to hospital and on June 10, 1957 a thoracoscopic pleural biopsy was taken. This showed the features of a mesothelioma. On June 24, 1957 in attempt at decortication was made. The left upper lobe was stripped easily but the grossly thickened pleura was so adherent to the lower lobe that the attempt was abandoned. Sections of the pleura confirmed the diagnosis of mesothelioma. He died on December 13, 1957.

Case 16. A.S.V. a white male, 50 years of age (born 1909) is a farmer who was born at Griquatown and went to a village school near an asbestos mine. From 1914 to 1916 he transported asbestos in donkey wagons from Illovo to Pekaia. During these journeys he often slept on asbestos bags. In 1911 he worked in an asbestos mine for a month.

The patient became ill in December, 1955 with vague subcostal pain and dyspnoea. A left-sided pleural effusion was diagnosed and he was referred to Mr. J. K. Brimmer in Pretoria. There was no evidence of malignancy in bronchial and sclerotic biopsies. Some upper pleural thickening and a dense homogeneous opacity at the base of the left lung was seen in the radiograph of August 3, 1956. Thoracotomy (Mr. Brimmer) in February, 1957 revealed a thickened nodular pleura with three masses on the left diaphragm. A mesothelioma was seen on histological section.

In April, 1959 he was examined by Mr. J. D. Visser at Kimberley who found similar nodules in the subcutaneous tissue over the neck and chest. A large firm mass was found in the thoracotomy wound and a mass was present in the left hypochondrium. There was generalized lymphadenopathy and numerous nodules were present on the femora. Chest radiographs showed the left pleural cavity to be obliterated. It was one of the few cases to develop widespread metastases.

A biopsy of one of the subcutaneous chest nodules showed a similar type of mass to those seen in the original specimen from the pleura. The patient died on May 22, 1959. No necropsy was performed.

Case 24. D.G., a white male, 32 years of age (born 1927) by profession an accountant, was born in England but came to South Africa as an infant. He spent his early childhood in Kuruman and left at the age of 7. He later qualified as a chartered accountant in Kimberley in 1953 at the age of 22. He had been discharged medically unfit from the army because of a left pleural effusion. Neither then nor on many subsequent occasions were acid-fast bacilli found either in sputum or pleural fluid. His only subsequent contact with asbestos was the making of the books of an asbestos mine in the Transvaal.

In September, 1953 he complained of left-sided chest pains and shortness of breath. A radiograph of the chest showed flattening of the diaphragm on the left side with obliteration of the costophrenic angle and thickening of the oblique fissure. He was seen on numerous occasions

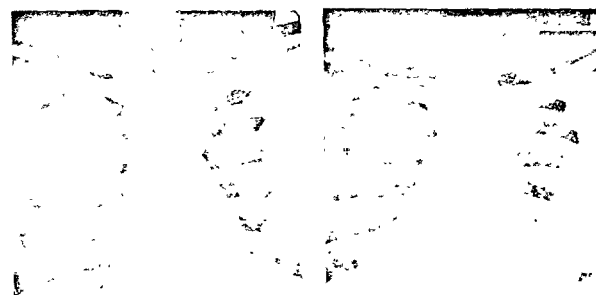


FIG. 11. On September 11, 1956.

FIG. 12. On August 23, 1957.

FIGS. 11 and 12. Radiographs showing development of tumour in Case 24.

from 1953 to 1957. A radiograph taken early in 1956 showed the development of a left superior mediastinal shadow which enlarged progressively (Fig. 11) and later the development of diffuse nodular pleural thickening on the left. During 1957 similar but less marked changes developed on the right side (Fig. 12). He consulted to Swaps in June, 1957 and at operation, a diffuse pleural thickening was present.

Sections of the parietal pleura only were removed, and the histological appearances were those of a mesothelioma. He died on October 6, 1957. A necropsy was not performed.

Case 30. G.F., a white male, 50 years of age (born 1909) is a stockkeeper with a history of industrial exposure. He has never been near the Cape Asbestos Fields.

He was born in South West Africa and lived there until the age of 21. From the age of 21 to 26 he worked in a factory in the Cape as a book-maker and later. He volunteered the information that he had worked continuously with asbestos, lagging pipes and boilers. He remarked that the quality of abestos lagging from old pipes created a very dusty atmosphere and that he would cough a lot at such work. Between 26 and 32 years of age he worked in a factory in Johannesburg in a similar capacity but was not exposed to such heavy dust concentrations. For the past 20 years he had not been in contact with asbestos. Enquiries were made from the factory in the Cape as to the nature of the asbestos used

and these have revealed that during the relevant periods Cape crocidolite was exclusively in use.

He was first seen on March 14, 1959 complaining of cough, pain in the right chest and recent onset of breathlessness. Three pints of fluid had been aspirated from the pleura and cytological examination showed the presence of malignant cells. The radiograph showed marked pleural thickening and hydro pneumothorax.

On March 17, 1959, a thoracoscopy was done. The pleura felt thickened and the lung was collapsed and immobile. Several biopsies were taken. These showed the presence of a mesothelioma. The underlying lung showed the presence of asbestosis. He was treated with nitrogen mustard intrapleurally and his condition temporarily improved. When last examined he complained of increased breathlessness.

Discussion

In 1924 Robertson denied the existence of primary malignant tumours of the pleura and considered them to be secondary in origin. Since then, on the one hand, Willis (1948, 1953) and Smart and Hirsman (1957) have supported Robertson's view, while on the other hand, primary neoplasms of this nature have been described by many authors in recent years. These include Tobiansen (1955) in Sweden, Bilimori and Bovo (1957) in Italy, Godwin (1957) in the United States, and McCaughey (1959)

in Britain. Evidence in support of the mesothelial origin of these tumours can be found in the tissue culture experiments of Stout and Murray (1942) and Sano, Weiss, and Gault (1950).

The variegated histological pattern of tumours arising from the mesothelium of the pleura was remarked on by Klempner and Rabin (1931). Such variation is appreciated when the multipotentiality of the cells lining embryonic cavities are considered. Maximow first demonstrated this feature in 1927. Novák (1931) stated that the mucosa of all parts of the Mullerian canal and the germinal epithelium were derived from these cells. Keasbey (1947) showed that in the embryo the mesonephros, metanephros, Wolffian body, genital ridge and all dependent mesodermal structures are derived from the mesothelium.

Campbell (1950) considered the presence of both epithelial and mesenchymal elements a major diagnostic feature. In describing the histology of 11 of these tumours McLaughley (1958) demonstrated that either the epithelial or the mesenchymal element might predominate. He classified his cases into the following four groups: tumours of epithelial character, tumours of mesenchymal type, tumours of mixed type, and tumours of anaplastic type.

Using McLaughley's classification on our series of tumours, the majority are, as in Campbell's series, of the mixed type. Of the remainder a few showed the papillary tubular structure of the tumours of epithelial character, and there were several of the anaplastic type. One case had the appearance of a tumour of mesenchymal type, but even in this case primitive tubular structures were seen.

Apart from the original case all the histological diagnoses were made on biopsy material. Three of these biopsy findings were confirmed at necropsy. In the remainder we have had to rely on clinical and radiological examination to exclude other primary sites of malignancy.

Following the work of Meyer and Chaffee (1939, 1940), the possibility of demonstrating hyaluronic acid in these tumours both chemically and histochemically has been considered as a diagnostic aid. Both these investigations are still in an early stage (Harrington, 1959).

Preliminary results of the histochemical experiments show that there is metachromatic substance in both the stroma and glandular structures of the tumours. This metachromasia can be reversed by incubation with testicular hyaluronidase. This material stains strongly with Hales's (1946) colloidal iron method. The periodic acid Schiff technique has given variable reactions. These results are not specific for hyaluronic acid but are strongly suggestive of its presence.

The amount of metachromatic substance in these tumours has varied considerably. This we think is partly due to the fact that until recently we had not appreciated the solubility of hyaluronic acid in aqueous media and the majority of these tumours were fixed in 10% formal-saline. Further, Liso (1951) states that the metachromasia of hyaluronic acid is optimal at a concentration of 1/10,000 and gradually decreases at higher levels. When the high concentrations that Meyer and Chaffee (1939, 1940) found in their tumours are considered this variation in metachromatic properties seems to have been partially explained. In addition with the great differences in the histology of the various lesions would be logical to expect certain tumours with marked glandular appearance to secrete more than those with an essentially non-glandular structure.

Three of the five autopsies performed showed evidence of peritoneal metastases. In the other two intra-thoracic spread was observed: in one case to the other lung and chest wall, and in the second to the mediastinal lymph glands and pericardium. Biopsy evidence of metastases has been obtained in two cases: one from the omentum and the other from the subcutaneous tissue of the chest wall, while a third patient developed an implantation nodule in the thoracotomy scar which showed a similar appearance to that of the previous biopsy specimen.

The first recorded case of carcinoma of the lung associated with asbestos was described by Lyns and Smith (1935). By 1955, according to Doll, a total of 61 cases had been reported. Included in these cases was one mesothelioma. Carter (1955) mentioned two cases of diffuse mesothelioma from Canadian chrysotile mines. A further three cases were described by van der Schoot (1958). Unfortunately no indication is given in the literature regarding the type of asbestos to which the majority of recorded cases of carcinoma were exposed. However, discussion with management and medical officers of two of the factories in which the majority of the cases reported in Britain were employed suggests that most of these workers were handling chrysotile asbestos (Wagner, 1958). The possibility that some of these people may also have been exposed to crocidolite dust cannot be excluded.

Attempts to produce tumours in experiment animals by exposing them to asbestos dust have been made. Novak and Karr (1938) were unsuccessful. Lynch, Melver, and Cain (1957) succeeded in producing tumours in mice, but lung neoplasms in the control animals were far too numerous for the results to be considered significant. On the other hand, Schwab (1958) working in Drucker's laboratory has been able to produce sarcomas in rats. This has occurred after subcutaneous and

intra-peritoneal inoculation of both asbestos fibres and dust. He states that "mineral asbestos" was used but does not name the variety.

In our series of mesothelioma histological evidence of asbestos has been observed in eight of the 10 cases in which lung parenchyma was included in the specimen examined. No lung tissue was present in the biopsies from the remaining 23 cases. It was only in the four cases that came to necropsy that large sections of the lung tissue were available for examination. In all of these specimens evidence of asbestosis was found.

At first it was thought that the presence of numerous asbestos bodies (fragments of fibre and dust immediately below the pleural elastic laminae in these specimens) might have been significant in the pathogenesis of the tumours. However, it is more probable that this distribution is a result of the marked atelectasis. Similar features have been observed in sticklefin lungs of asbestos miners in which no mesothelioma have been observed.

Pleural fibrosis has been a common finding in cases of asbestosis and in some cases large pleural plaques measuring up to 1.0 cm in thickness have been seen. These plaques which were first described by Clayton (1933) have been observed following exposure to both amosite and crocidolite asbestos dusts. In these cases of benign pleural thickening no evidence of stromal metachromasia has been observed.

In all the histological sections of pleura examined in the cases of these mesotheliomas and in more than 100 cases of asbestosis, no asbestos bodies (fragments of fibres or dust) have been observed beyond the pleural elastic laminae.

Three patients from the Kuruman district, with clinical and radiological features consistent with those of diffuse pleural mesothelioma had markedly abnormal cells in the pleural fluid. These cells showed no specific features to distinguish them from cells originating in secondary malignant pleural deposits nor from the grossly atypical cells sometimes seen in non-malignant pleural effusions. As no biopsy or autopsy examinations were obtained in these cases they have been excluded from the series.

The pathological evidence for associating these tumours with asbestos exposure is not conclusive. As previously stated, only in eight of the 33 cases has evidence of asbestos been demonstrated. Of these six had a definite mining history and one had been exposed to asbestos while lagging steam pipes. The other case was born in the Kuruman district and nothing else is known of him, until his arrival at a Witwatersrand gold-mine at the age of 23. In the remaining 25 cases we can only present circumstantial evidence of exposure to asbestos dust.

(Table 3). Eighteen of these 25 cases were born in the vicinity of the mines and two arrived in the district as infants. Of these 18 people 11 admit definite childhood exposure to the dust and two others were exposed industrially in later life. In addition two patients with childhood exposure later worked in the asbestos mines. Three cases arrived in the region at an older age but were employed either on the mines or in transporting asbestos. A further three of these 25 cases have had industrial exposure, and in only one case do the relatives deny any exposure to asbestos dust.

The four industrial cases are significant. Two of the patients were lagging locomotive boilers and one was lagging steam pipes. A man, who was an upholsterer by trade, was employed in making fire proof clothing from 1939-1945. As far as can be ascertained these people were never in the Griqualand district. These findings tend to add support to asbestos being the common factor in the development of these tumours, and to counter the suggestion that there may be some other environmental cause in the region of Griqualand West.

If asbestos dust is a factor in the occurrence of these tumours similar cases might have been expected from the neighbourhood of the Transvaal asbestos mines in the Pietersburg and Lydenburg districts where crocidolite and amosite asbestos is mined. According to Vermaas (1952), crocidolite and amosite occur in the same seams in the Pietersburg district. As can be seen from Table 1, amosite is similar in composition to crocidolite. In the past four years the lungs of 24 cases of asbestosis from the Lydenburg district have been examined. In this material two cases of adenocarcinoma have been observed. The one was in a white miner with 19 years' service, who had an adenocarcinoma arising from a bronchus. The other case was a Bantu miner who had a peripheral tumour. No service record was obtainable in this case. Only one case of asbestosis has been received from the Pietersburg area. All of these men were actually employed on the mines at the time of death, the majority having had a relatively short service. Our findings suggest that mesothelioma occurs 20 to 40 years or more after exposure to dust. Until comparatively recently the mining in the Transvaal has been on a small scale, and there were no settlements in the vicinity of the mines.

The lungs of 20 asbestos miners from the Cape Asbestos Field have been examined in whom no mesothelioma was observed. One autopsy specimen consisting of three fragments of lung showed asbestosis and an adenocarcinoma. Radiological features of asbestosis have been observed in many miners from this region, without any suggestion of

TABLE 3
DIFFUSE PLEURAL MESOTHELIOMA: ASSOCIATION WITH ASBESTOS

(1)	(2)	(3)	(4)	(5)	(6)	(7)	(8)	(9)	(10)	(11)
Case No.	Year of Birth	Age at Diagnosis	Race	Sex	Born on Asbestos Fields	Asbestos Exposure	Diagnosed on Biopsy	Necropsy	Histological Evidence of Asbestos	Surv. Time (yr)
1	1920	36	B	M		Other history unknown until came to the Witwatersrand at the age of 23.				
2	1913	42	MND	M		Mined asbestos from 1930-33, left area at the age of 27.				
3	1902	53	B	F		Lived while life in a location near an asbestos mill.				
4	1896	58	W	F		Lived on asbestos fields until the age of 5, worked in asbestos warehouse 1916-20.				
5	1925	34	W	M		Spent all his working life in the vicinity of mines.				
6	1903	61	W	F		Lived all his life in the vicinity of mines.				
7	1920	46	W	M		Lived all his life in the vicinity of mines, worked at a mine.				
8	1894	63	MND	F		From the age of 24 lived in a village serving local mines, often visited mines, watched cobbing outside houses.				
9	1905	52	MND	M		Whole life spent near mines, digging with a shovel, from the age of 7-17 years, lived on dumps and in mine as a boy, returned to work from age 21-25.				
10	1909	49	W	M		Lived at the mine from age 7-17 years, lived on dumps and in mine as a boy, returned to work from age 21-25.				
11	1898	60	MND	M		Whole life spent near mines, miner 1911-33.				
12	1910	48	B	M		Lived in mines until the age of 17, miner 1922.				
13	1909	50	W	F		Lived at mine until the age of 21, played with there as a child.				
14	1918	40	B	M		Whole life spent near mines, miner 1938-41.				
15	1916	42	W	F		Placed in dumps as a child.				
16	1896	60	W	M		Placed in dumps at the age of 17, until age of 20, spent in school near cobbing sheds.				
17	1913	48	MND	F		Went to school near cobbing sheds.				
18	1920	38	B	M		Lived on dumps, spent some time at the age of 15, worked in mines.				
19	1922	37	MND	M		Whole life in the vicinity of the mines, miner 1915-58.				
20	1906	53	W	F		Lived at mine, miner 1928-1939.				
21	1912	44	W	M		Spent whole life in village on wagon route to KwaZulu.				
22	1889	48	W	M		Lived in the vicinity of mines until the age of 16, then on dumps as a child.				
23	1895	63	W	F		Whole life in the vicinity of the mines, miner 1913-15, and 1945-52.				
24	1922	35	W	M		Lived in the vicinity of the mines until the age of 40.				
25	1899	50	W	F		Lived in the vicinity of the mines until the age of 17, placed in dumps as a child.				
26	1895	49	W	M		Lived on a wagon route from the age of 10 to 18 years, then 1918 spent whole life in some mines.				
27	1904	52	W	M		Mined and transported asbestos from 1920-34.				
28	1899	60	W	M		Worked in mines, transported asbestos from 1920-34.				
29	1913	44	W	M		Lived whole life in a farm in mining area from 1913-15, then in a farm near asbestos 1916-21.				
30	1900	50	W	M		Spent whole life in a farm near asbestos 1916-21.				
31	1913	44	W	M		Mining, steam pumps in explosive factories 1916-20.				
32	1908	49	W	M		Worked as a fitter on railways, maintaining locomotives, broken down in 1945.				
33	1840	69	W	F		Mining, a Diggers' Hunker for the Air Force, 1929-45.				
						No history of exposure to asbestos.				

Column 4: Race: W, White; MND, Mixed (Coloured); B, Bantu.

Column 5: Sex: M, Male; F, Female.

Column 6: Born on asbestos fields: 1, Born on asbestos fields; 0, Not born on asbestos fields.

Column 7: Asbestos exposure: 1, Asbestos exposure; 0, No asbestos exposure.

Column 8: Diagnosed on biopsy: 1, Diagnosed on biopsy; 0, Not diagnosed on biopsy.

Column 9: Necropsy: 1, Necropsy done; 0, Necropsy not done.

Column 10: Histological evidence of asbestos: 1, Histological evidence of asbestos; 0, No histological evidence of asbestos.

Column 11: Survival time (yr): 1, Survival time; 0, No survival time.

tumour formation. However, pleural involvement has been a common finding in these cases (Hartwitz, 1959). It is possible that there has not been a sufficient lapse of time after exposure for tumour development in these cases. Further, the factor of individual susceptibility must also be considered. We wish to record the great assistance that we have received in this investigation from the medical practitioners of the Grigoland West District and the

DIFFUSE PLEURAL MESOTHELIOMA

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Addendum

By the end of June 1960, a total of 47 cases of mesothelioma had been identified. In 45 of these a possible association with exposure to crocidolite has been established. In one case a mesothelioma of the peritoneum was present.

THE JULY (1960) ISSUE

The July (1960) issue contains the following papers:

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- Phosgene Poisoning Caused by the Use of Chlorine Paint Removers Containing Methylene Chloride in Ill-ventilated Rooms Heated by Kerosene Stoves. By W. B. Gerritsen and C. H. Buschmann.
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- The Measurement of Occupational Mortality. I. D. K. Liddell.
- Observations on the Occupational Life History of the Coal Face Worker at Two Collieries. By O. P. Edmonds and D. S. Kerr.
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