Xeromammography: a Study of 59 Cancers and 204 Non-Cancerous Lesions of the Breast*⁹

Masayuki NISHIKI, Motoi YAMANE, Kuniki AMANO,

Tsuneo OKUMICHI, Toshiya MATSUYAMA and Haruo EZAKI

The Second Department of Surgery, Hiroshima University School of Medicine, 1–2–3, Kasumi, Minami-ku, Hiroshima 734, Japan

(Received March 8, 1984)

Key words: Xeromammography, Breast cancer, Risk factor

ABSTRACT

Of female patients who were subjected to mammary gland xeromammography at the 2nd Dept. of Surgery, Hiroshima University School of Medicine for a period from 1979 to 1983, 59 patients with palpable breast cancer and 204 non-cancerous patients confirmed pathologically were studied mainly in regard to direct signs.

The incidence of cancer was relatively high in P2 and DY (so-called high-risk) parenchymal pattern groups, indicating necessity of minute interpretation of mammograms in these two groups. As for direct signs of breast cancer, the incidence of tumor shadow was 77.9% (46/59) and that of calcification including microcalcification (21 cases) was 42.9% (25/59) in cancer patients.

The false negative rate of xerommography was 13.6% (8/59), attributable to misinterpretation in 4 cases and to mammograms themselves which were completely normal in the remaining 4 cases. The mean age of the false negative cases seemed to be younger than that of whole cancer cases (59 cases) while neither histopathological findings nor parenchymal patterns significantly correlated with the results.

INTRODUCTION

The mammography is considered as a standard technique of supplementary diagnosis of breast cancer. Equipment, films, and roentgenographic techniques have been improved greatly for the last 20 years, and especially the xeromammography which applies electlic treatment to contrast and record but not chemical treatment as in the conventional film mammography, has been used more widely since 1970s. The most remarkable characteristic of xeromammography is edge enhancement of the margin of calcification and tumor shadow, which facilitates reading of mammograms.

MATERIAL & METHOD

Confirmatory pathological diagnosis could be made for 263 cases of total female patients who underwent mammary gland xeromammography at the 2nd Dept. of Surgery, Hiroshima University School of Medicine for a period from June 1979 to June 1983. Fifty-nine patients had cancers and 204 did not. Either aspiration biopsy, needle biopsy, or open biopsy was applied to pathologically fonfirm the mammographic data. The 263 patients were studied in terms of direct signs, that is, tumor shadow and calcification. No cases of occult breast cancer nor under 30-year old were included in the study. The classification of mammographic parenchymal patterns followed that of Wolfe.

RESULTS

The incidence of breast cancer according to mammographic parenchymal pattern was significantly higher in the combination P2 and DY (so-called high-risk) parencymal patterns, compared with the combined N1 and P1 patterns (Table 1): 31.4% (38/121) in the former; and

江崎治夫:ゼロマンモグラー:59例の乳癌および204例の非

^{*&}lt;sup>1</sup> 西亀正之,山根 基,天野国幹,奥道恒夫,松山敏哉, 癌性乳腺疾患の検討

Parenchymal Pattern	No. of Cases with Cancer	No. of Total Cases**	%
N1 & P1	21	117	17.9*
P2 & DY	38	121	31.4^{st}

Table 1. Incidence of Breast Cancer: According to Parenchymal Pattern

* Significant difference (p < 0.05)

** Excluding cases of 30 years old and under

Dethalast 1 D	0	Tumor Shadow		Calcification	
Pathological Diagnosis	Case	clear	unclear	fine	coarse
Mastopathy	93	8	4	2	7
Fibroadenoma	47	22	6	0	2
Cyst	16	5	1	0	0
Fat necrosis	11	1	0	0	1
Papilloma	7	2	1	0	0
Normal gland	26	0	0	0	2
Others	4	1	0	0	0
Total	204	39	12	2	12

Table 2. Incidence of Tumor Shadow and Calcification in Non-Cancerous Cases

17.9% (21/117) in the latter. Tumor shadow and calcification were seen in 25% (51/204) and 6.9% (14/204) of non-cancerous cases, respectively (Table 2).

In mastopathy cases, the incidence of tumor shadow was 12.6% (12/93), not different from that of other benigh lesion groups, but that of calcification was 9.8% (9/93), significantly higher compared to other groups (Table 2).

On the other hand, tumor shadow appeared in 46 of 59 cancer cases (77.9%): 16 cases (27.1%) with both tumor shadow and calcification; and 30 cases (50.8%) with only tumor shadow. Calcification was seen in 25 of 59 cases (25/59), including 21 patients with microcalcification. Four of 59 cancer patients (6.8%)who had neither tumor shadow nor calcification showed completely normal mammograms (Table 3).

Table 3. Direct Signs in 59 Breast Cancer Cases

	T + C +	T+C-	T-C-	T-C-
No. of cases	16	30	9	4
%: Cases/total cases	27.1	50.8	15.3	6.8
T : tumor shadow + : positive	W C : calcification — ; negative			

Table 4 shows incidence of direct signs in 44 patients with infiltrating carcinomas, classified as histologically common types. The direct signs appeared in all patients with papillotubular type carcinoma, compared with in 87.5% of medullotubular cases. Scirrhous cases showed a relatively low incidence of calcification.

 Table 4. Direct Signs in 44 Infiltrating Carcinoma (Common Type)

Pathological	No. of Cases			
Classification*	T+C+	T+C-	T-C+	T-C-
pap. tub.	5	6	1	0
med. tub.	6	10	5	3
scirr.	1	6	1	0
T : tumor shadowC : calcification+ : positive- : nagative				
+ : positive * Pathological clas		Ŭ		eneral

Rule for Clinical and Pathological Record of Mammary Cancer 1982, The 6th edition, edited by Japan Mammary Cancer Society, Tokyo (in Japanese)

As for the incidence of direct signs according to mammographic parenchymal pattern, tumor shadow was seen in 63.1% (12/19) of the DY pattern group, significantly lower compared with 100% (9/9) of P1 and 94.7% (18/19) of P2

Parenchymal Pattern	No. of Cases with Tumor Shadow	No. of Total Cases	%
N1	8	12	66.7 ^a
P1	9	9	100 ^b
P2	18	19	94.7°
DY	12	19	63.1ª

 Table 5. Incidence of Tumor Shadow in Breast

 Cancer: According to Parenchymal Pattern

a-b, a-c, a-d, and b-c : no significant difference

b-d, and c-d : significant difference (p < 0.05)

pattern groups (Table 5). The incidence of calcification by parenchymal patterfi showed no significant difference between DY and N1, P1 or P2 pattern groups (Table 6).

 Table 6. Incidence of Calcification in Breast

 Cancer: According to Parenchymal Pattern

Parenchymal pattern	No. of Cases with Calcification	No. of Total Cases	%
N1	7	12	58.3ª
P1	3	9	33.3 ^b
P2	5	19	26.3°
DY	9	19	47.6ª

No significant difference among a, b, c, and d.

Table 7 shows the accuracy of our data obtained from xeromammography which revealed cancers in 59 out of 59 cancer patients (true positive rate: 86.4%, 51/59), and failed to reveal cancers in the remaining 8 cases (false negative rate: 13.6%, 8/59). The faults were resulted from misinterpretation in 4 cases and mammograms themselves which were completely normal in other 4 cases. The false positive rate in xeromammography was 1.5%(3/204), acceptably low.

Table 7. Accuracy in Xeromammography

Pathology	М	M	В	В
Xeromammography	Μ	В	Μ	В
No of Cases	51	8	3	201

M: diagnosis as Malignant lesion

B : diagnosis as Benign lesion

Table 8 shows the incidence of false negative case according to mammographic parenchymal pattern: the incidence was 21.1% (4/19) in DY pattern group, slightly higher but not significantly different from that in N1, P1, or P2 pattern group,

 Table 8. Incidence of False Negative Case in

 Breast Cancer: According to parenchymal Pattern

Parenchymal Pattern	No. of Cases of False Negative	No. of Total Cases	%
N1	1	12	8.3ª
P1	1	9	11.1 ^b
P2	2	19	10.5°
DY	4	19	21.1^{d}

No significant difference among a, b, c, and d.

DISCUSSION

The following advantage of xeromammography over conventional film mammography were pointed out by other researchers^{11, 24, 25, 33)}: (1) enhancement of margins of lesions; (2) highlightening of microcalcification and (3) ability to penetrate dense dysplastic breasts. However, a number of researchers^{14, 27, 28)} demonstrated that with the aid of microfocal spot roentgenography, the conventional film mammography could reveal lesions as delicate as those discovered by xeromammography.

Wolfe classified mammograms into four groups (N1, P1, P2, and DY) according to the roentgenographic appearance of breast parenchyma, that is, prominent duct and dystrophy, demonstrated a significantly greater cancer incidence in the P2 and DY pattern groups compared with N1 or P1 groups^{35,36)}. Since his reports, considerable retrospective and prospective studies were made to confirm the correlation between parenchymal patterns and risk factor for present or future breast cancer.

Doyle⁶⁾, Moskowitz²²⁾, Tabar³¹⁾, Weich³²⁾, etc found no correlation between them. Egan⁷⁾ stated that the mammographic parenchymal patterns were not reliable indicators for initial cancer or developing cancer and that the apparently high incidence of developing cancer in dense fibroglandular cases might be attributable to overlook small cancers at the first examination.

Boyd²⁾, however, contradicted Egan's statement⁷⁾, demonstrating particularly close correlation between dysplasia and breast cancer in his age-matched case control study. He also suggested that benign lesion cases included in cotrol groups might lead to negative data regarding parencymal pattern and breast cancer. Others^{8,4,6,12)} were also in favor of Wolfe's hypothesis.

Our study in 59 cancer cases and 204 benign

lesion cases including 26 cases with normal mammary glands showed the cancer incidence in the P2 and DY group (32.4%, 38/121) was higher than that in the N1 and P1 group (17.9%, 21/117). As pointed out by Parsons²⁴⁾ and Egan⁷⁾, it is difficult to interpret mammo-grams of the P2 and Dy groups, suggesting that a minute interpretation particularly on delicate lesions is necessary in these groups.

Two major signs of breast cancer on mammograms are tumor shadow and microcalcification. The tumor shadow generally occurs as an indeterminate shape with irregular margin, but it is known that patients with colloid type and medullary type breast cancers rarely have tumor shadow with completely smooth border^{11,} ²⁶⁾. Microcalcification is also a sign indictating the development of serious cancer. Radiography of cancer specimens is known to detect microcalcification at the incidence of 60–80%^{8,} ²¹⁾, compared with 30–50% on mammograms^{1, 34)}.

Our data of xeromammography, that is, 13.6 % (8/59) of false negative rate and 1.5% (3/204) of false positive rate, were almost consistent with other reports^{5, 9, 18, 23, 29, 30)}. Xeromammograms of 8 false negative cases consisted of 4 completely normal ones and 4 misinterpreted ones, including 2 cases with tumor shadow, 1 with microcalcification, and 1 with skin change revealed by retrospective study.

Martin¹⁹⁾ demonstrated an analysis result of 48 false negative cases that no abnormal radiography finding was seen in 37% (16/48), clear abnormal findings were overlooked in 29% (14/ 48), and subttle indirect signs were overlooked in 38% (18/48), and pointed out that interpretation of subttle indirect signs is important to improve diagnositic rate of mamography. Lesnick¹⁷⁾ stated that 63% of 52 breast cancer cases of 45 years old and under showed no abnormality on their mammograms. In general, it seems that cancers in patient showing mammary dysplastic changes or dense breasts (often seen in younger patients) are liable to be overlooked^{7,24}).

Our analysis of the 8 false negative cases revealed a trend that their mean age was slightly younger than that of the total cancer cases $(45.9\pm5.7 \text{ years old vs } 49.6\pm10.4 \text{ years old})$ but found no significant correlation with histopathological findings or parenchymal patterns.

We make it a rule to perform aspiration

cytology in all patients with palpable mass or abnormal findings in xeromammography. Our study including 59 palpable cancer cases found 13.6% of false negative cases in xeromammography, supporting the view of Niloff²³⁾ and Mann²⁰⁾ that xeromammography should be positively performed concomitantly with aspiration cytology or open biopsy in patients showing normomammograms but clinically suspected sings of cancer.

REFERENCE

- 1. Black, J. W. and Young, B. 1964. A radiological and pathological study of the incidence of calcification in disease of the breast and neoplasms of other tissues Br. J. Radio. 38: 596-598.
- Boyd, N.F., O'sullivan, B., Compbell, J.E., Fishell, E., Simor, I., Cooke, G. and Germanson, T. 1982. Mammographic signs as risk factors for breast cancer. Br. J. Cancer 45: 185-193.
- Boyd, N.F., o'sullivan, B., Campbell, J.E., Fishell, E., Simor, I., Cook, G. and Germanson, T. 1982. Mamographic patterns and bias in breast cancer detection. Radiology 143:671-674.
- Brisson, J., Merletti, F., Sadowsky, N., Twaddle, J. A., Morrison, A. S. and Cole, P. Mammographic features of breast and breast cancer risk. Am. J. Epidemiol. 115: 428-437.
- Cohn, H. E. 1972. Mammography in its proper perspective. Surg. Gynecol. Obstet. 134:97-98.
- Doyler, P. J., Blamey, R. W., Chatte, A. and Roeduck, E. 1979. Rate of breast cancer related to parenchymal pattern of mammogram. Clin. Oncol. 5: 390-391.
- Egan, R. L. and McSweenney, M. B. 1979. Mammographic parenchymal patterns and risk of breast cancer. Radiology 135: 65-70.
- 8. Fischer, E. R., Gregorio, R. M. and Fischer, B. With the assistance of Redmond, C., Vellios, F., Sommers, S. C. and Cooperative Investigators 1975. The pathology of invasive breast cancer. A syllabus derived from findings of the National Surgical Adjuvant Breast Project (Protocol No 4) Cancer 36: 1-85.
- Frankl, G. and Ackerman, M. 1983. Xeromammography and 1200 Breast Cancers. Symposium of Mammography, p. 81-91, *In* Wolfe J. N. (ed.), Radiologic Clinics of North America. Vol. 21, No. 1, W. B. Saunders Company Ltd, London
- Gershon-Cohen, J., Berger, S. M. and curcio, B. M. 1966. Breast cancer with microcalcification, diagnostic difficulties, Radiology 87: 613-622.
- Gravelle, I. H. 1982. Diagnostic Image in Breast Cancer, p. 795-820. In Baum M. (ed.), Clinics in Oncology. Vol. 1, No. 3, W. B. Saunders Company

Ltd. London.

- Hainline, S., Myers, L., Mclelland, R., Newell, J., Grufferman, S. and Shingleton, W. 1978. Mammographic pattern and risk of breast cancer. Am. J. Roentganol. 130: 1157-1158.
- Hassler, O. 1966. Microradiographic investigations of calcifications of the female breast. Cancer 23: 1103-1109.
- Hans, A. G., Paulus, D. D., Dodd, G. D., Cowart, R. W. and Bencomo, J. 1979. Magnification mammography: evaluation of screen-film and xeroradiographic techniques. Radiology 133: 233– 226.
- Koehl, R. H., Snyder, R. E., Hutler, R. V. and Foote, F. W. 1970. The incidence and sifnificance of calcifications within operative breast specimens, Am. J. Clin. Pathol. 53: 3-14.
- Krook, P. M., Carlile, T., Bush, W. and Hall, M. H. 1978. Mammographic parenchymal patterns as a risk indicator for prevalent and incident cancer. Cancer 41: 1093-1097.
- 17. Lesnick, G. J. 1977. Detection of breast cancer in young women. J. A. M. A. 237 : 967-969.
- Lewis, J. D., Milbrath, J. R., Shaffer, K. A., Davin, J. C. and DeCosse, J. J. 1976. Which breast to biopsy; an expanding dilemma. Ann. Surg. 184: 253-257.
- Martin, J. E., Moskowitz, M. and Milbrath, J. R. 1979. Breast cancer missed by mammography. A. J. R. 132: 737-739.
- Mann, B.D., Giuliano, A.E., Bassett, L.W., Barber, M.S., Hallaner, W. and Morton, D. L. 1983. Delayed diagnosis of breast cancer as a result of normal mammograms. Arch. Surg. 118: 23-24.
- Millis, R. R., Davis, R. and Stacey, A. J. 1975. The detection and significance of calcifications in the breast: a radiological and pathological study. Br. J. Radiol. 49: 12-26.
- Moskowitz, M., Cartside, P. and McLaughlin, C. 1980. Mammographic pattern as markers for high-risk benign breast disease and incident cancers. Radiology 134 : 293-295.
- Niloff, P.H. and Sheiner, N.M. 1981. Falsenegative mammograms in patients with breast cancer. Canadian J. of Surgery 24: 50-52.

- Parsons, C. A. 1979. The role of xeromammography in the detection of breast cancer. Invest. Cell, Pathol. 2: 163-170.
- Ruzicka, F. F., JR., Kaufman, L., Shapiro, G., Perez, J. V. and Grossi, C. E. 1965. Xerommamography and film mammography. A comparative study. Radiology 85: 260-269.
- Sadowsky, N. and Kopans, D. B. 1983. Breast Cancer. Symposium of Mammography, p. 81-91, *In* Wolfe J. N. (ed.) Radiologic Clinics of North America Vol. 21, No. 1, W. B. Saunders Company Ltd. London.
- 27. Sickles, E.A. 1980. Further experience with microfocal spot magnification mammography in the assessment of clustered breast microcalcification. Radiology 137: 9-14.
- Sickles, E. A., Doi, K. and Genant, H. K. 1977. Magnification film mammography: image quality and clinical studies. Radiology 125: 69-76.
- Sokol, E. S., Walker, B., Terz, J. J. and Lawrence, W. JR. 1970. Role of mammography with palpable breast lesions. Surgery 67: 748-753.
- 30. Stark, A.M. and Way, S. 1974. The screening of well women for the early detection of breast cancer using clinical examination with thermography and mammography. Cancer 33:1671-1679.
- Tabar, L. and Dean, P.B. 1982. Mammographic parenchymal patterns. Risk indicator for breast cancer? J. A. M. A. 247 : 185-189.
- Weicj, J. and Adler, O. B. 1981. Breast pattern as a risk for development of cancer. Israel J. Med. Sci. 17: 9-10.
- 33. Wolfe, J. N., Dooley, R. P. and Harkins, L. E. 1971. Xeroradiography of the breast: A comparative study with conventional film mammography. Cancer 28: 1569-1574.
- Wolfe, J. N. 1974. Analysis of 462 breast carcinomas A. J. R. 121 : 864-853.
- Wolfe, J. N. 1976. Risk for breast cancer development determined by mammographic parenchymal pattern Cancer 37: 2486-2492.
- Wolfe, J. N. 1979. Breast parenchymal patterns: Prevalent and incident. Carcinoma. Radiology 131: 267-268.