Tumor characteristics of ductal carcinoma in situ of breast visualized on [F-18] fluorodeoxyglucose-positron emission tomography/computed tomography: Results from a retrospective study

Tomoyuki Fujioka, Kazunori Kubota, Akira Toriihara, Youichi Machida, Kaori Okazawa, Tsuyoshi Nakagawa, Yukihisa Saida, Ukihide Tateishi

AIM
To clarify clinicopathological features of ductal carcinoma in situ (DCIS) visualized on [F-18] fluorodeoxyglucose-positron emission tomography/computed tomography (FDG-PET/CT).

METHODS
This study retrospectively reviewed 52 consecutive tumors in 50 patients with pathologically proven pure DCIS who underwent [F-18] FDG-PET/CT before surgery. [F-18] FDG-PET/CT was performed after biopsy in all patients. The mean interval from biopsy to [F-18] FDG-PET/CT was 29.2 d. [F-18] FDG uptake by visual analysis and maximum standardized uptake value (SUVmax) was compared with clinicopathological characteristics.
RESULTS

[F-18] FDG uptake was visualized in 28 lesions (53.8%) and the mean and standard deviation of SUVmax was 1.63 and 0.90. On univariate analysis, visual analysis and the SUVmax were associated with symptomatic presentation ($P = 0.012$ and 0.002, respectively), palpability ($P = 0.030$ and 0.024, respectively), use of core-needle biopsy (CNB) ($P = 0.023$ and 0.012, respectively), ultrasound-guided biopsy ($P = 0.040$ and 0.006, respectively), enhancing lesion $\geq 20$ mm on magnetic resonance imaging (MRI) ($P = 0.001$ and 0.010, respectively), tumor size $\geq 20$ mm on histopathology ($P = 0.002$ and 0.008, respectively). However, [F-18] FDG uptake parameters were not significantly associated with age, presence of calcification on mammography, mass formation on MRI, presence of comedo necrosis, hormone status (estrogen receptor, progesterone receptor and human epidermal growth factor receptor-2), and nuclear grade. The factors significantly associated with visual analysis and SUVmax were symptomatic presentation ($P = 0.019$ and 0.001, respectively), use of CNB ($P = 0.001$ and 0.031, respectively), and enhancing lesion $\geq 20$ mm on MRI ($P = 0.001$ and 0.049, respectively) on multivariate analysis.

CONCLUSION

Although DCIS of breast is generally non-avid tumor, symptomatic and large tumors ($\geq 20$ mm) tend to be visualized on [F-18] FDG-PET/CT.

Key words: Ductal carcinoma in situ; Positron emission tomography; Breast cancer; [F-18] fluorodeoxyglucose-positron emission tomography/computed tomography

© The Author(s) 2016. Published by Baishideng Publishing Group Inc. All rights reserved.

Core tip: Symptomatic tumor or large ductal carcinoma in situ (DCIS) ($\geq 20$ mm) is often visualized on [F-18] fluorodeoxyglucose-positron emission tomography/computed tomography (FDG-PET/CT). This evidence suggests that large DCIS ($\geq 20$ mm) has possibility to be selected as target lesion on [F-18] FDG-PET/CT prior to neoadjuvant chemotherapy.

Fujioka T, Kubota K, Torihara A, Machida Y, Okazawa K, Nakagawa T, Saida Y, Tateishi U. Tumor characteristics of ductal carcinoma in situ of breast visualized on [F-18] fluorodeoxyglucose-positron emission tomography/computed tomography: Results from a retrospective study. World J Radiol 2016; 8(8): 743-749. Available from: URL: http://www.wjgnet.com/1949-8470/full/v8/i8/743.htm DOI: http://dx.doi.org/10.4329/wjr.v8.i8.743

INTRODUCTION

Since widespread of and technical improvements to screening mammography, the frequency of ductal carcinoma in situ (DCIS) has increased substantially. Mammography is able to detect even small DCIS if they have suspicious microcalcifications[1]. With the increasing use of ultrasound and magnetic resonance imaging (MRI), occult DCIS that cannot be identified by mammography, such as DCIS without microcalcification and in patients with dense breasts, have been occasionally detected[2-5]. Nowadays, maximally 20%-25% of new breast cancer cases consist of DCIS[6-8].

[F-18] fluorodeoxyglucose-positron emission tomography/computed tomography (FDG-PET/CT) has been recognized as an essential modality for detecting hypermetabolic activity in primary breast tumor, diagnosing and staging local and distant sites, and evaluating the response to therapy[9-13]. It has been reported that the 25%-77% sensitivity of DCIS is lower than that of invasive ductal carcinoma (IDC) on [F-18] FDG-PET/CT[10,11]. However, DCIS often can be detected on [F-18] FDG-PET/CT. The purpose of this study was to clarify clinicopathological features of DCIS visualized on [F-18] FDG-PET/CT with special reference to pathologic specimens obtained by surgery.

MATERIALS AND METHODS

Patients

This retrospective study was officially approved by the Ethical Commission of our institution (No. 1987). The inclusion criteria for patients with breast cancer were as follows: (1) those who underwent [F-18] FDG-PET/CT and operation at our hospital between March 2008 and May 2014; and (2) those with a pathologically proven pure DCIS, which means DCIS without microinvasion. The decision to perform [F-18] FDG-PET/CT was left to the discretion of the surgeon. In this study, after searching the database of radiology reports and clinical records at our institute, we identified 52 consecutive lesions in 50 patients with a mean age of 56.3 years (range, 33-85 years). All of the patients were not pregnant or breast feeding. [F-18] FDG-PET/CT was performed after biopsy in all patients. The mean interval from biopsy to [F-18] FDG-PET/CT was 29.2 d (range, 43-133 d) and from biopsy to operation was 78.8 d (range, 34-224 d). The treatment comprised of total mastectomy in 14 lesions, breast conserving surgery in 27, and skin sparing mastectomy in 11.

Clinical examination

All patients had physical examinations, mammography, ultrasound, and MRI. Mammography examination (craniocaudal and mediolateral oblique views) was performed using Lorad Selenia (Hologic Inc, Bedford, Massachusetts). EUB-7500 scanner with a EUP-L54MA 9.75-MHz linear probe (Hitachi Medical Systems, Tokyo) or Aplio XG scanner with a PLT-805AT 8.0-MHz linear probe (Toshiba Medical Systems, Tochigi) was used for the ultrasound examinations. MRI examination was performed with 1.5-T system...
Magnetom Vision, Siemens, Erlangen) for 7 patients and 3.0-T system (Signa HDxt, General Electric Medical Systems, Milwaukee, Wisconsin) for 43 patients using a breast coil in the prone position using a breast coil. To evaluate the MRI imaging, the early phase of a contrast enhancement study within 1 and 2 min after intravenous bolus injection of Gd-DTPA (0.2 mL/kg) was acquired. Unilateral coronal T1 weighted sequence (repetition time (TR)/echo time (TE) = 170/4.7, flip angle = 40°, 4 mm thick section, 256 × 256 matrix, 210 mm field of view) using 1.5-T system and bilateral axial fat suppressed T1 weighted sequence (TR/TE = 6.5/2.4, flip angle = 10°, 2 mm thick section, 512 × 512 matrix, 360 mm field of view) using 3.0-T system were employed.

Percutaneous needle biopsy was done with either ultrasound-guided core-needle biopsy (CNB): 14-gauge Biopsy System (C.R. Bard, Covington, Georgia), ultrasound-guided or stereotactic vacuum-assisted biopsy (VAB); 8- or 11-gauge MammoToome (Ethicon Endo-Surgery, Cincinnati, Ohio) or 11- or 14-gauge Vacora (CR Bard, Murray Hill, New Jersey).

One radiologist had 5 years of experience in breast imaging recorded the reason for presentation (screening-detected or symptomatic lesion), presence of a palpable lesion, use of biopsy device, and image guidance, then evaluated presence of clustered micro calcifications within the tumor on mammography, presence of mass formation, and size (largest diameter) of enhancing lesion on MRI by the American College of Radiology Breast Imaging Reporting and Data System without access to FDG-PET/CT or pathological data[14] (Figure 1A and 2A).

FDG-PET/CT protocol
After at least a 4-h fasting period, the patients received an intravenous injection of 3.7 MBq/kg (0.1 mCi/kg) [F-18] FDG. Images were obtained by whole-body mode with a PET/CT system (Aquiduo; Toshiba Medical Systems, Tokyo). CT images were performed by the following parameters: Pitch 0.938; 0.5 s gantry rotation time; 30 mm/s table time; 120 KVP; auto-exposure control (SD 20); and 2.0-mm slice thickness. Contrast agents were not used during our study. Approximately 60 min after the FDG injection, whole-body emission PET scan was obtained with the following parameters: 7-8 bed positions; 2-min emission time per bed position; 3.375-mm slice thickness; and 128 × 128 matrix.

Data analysis of FDG-PET/CT
[F-18] FDG-PET/CT images were reviewed by two nuclear medicine physicians (with 5 and 17 years of experience) in consensus; although they knew that the patients had DCIS, they were blind to clinical information including menopausal status and phase of the menstrual cycles, and presence of fibrocystic changes in patients. They performed visual analysis without defining a cutoff point. Lesions showing [F-18] FDG uptake higher than the surrounding background of normal breast tissue were defined as FDG positive. Region of interest (ROI) was placed on the PET images for measuring the maximum standardized uptake values (SUVmax) of the tumor. If the tumor was FDG negative, ROI was drawn on the background breast tissue area, and the SUVmax was established (Figures 1B, 1C, 2B and 2C).

Pathological evaluation
Specimens were cut into 5-10 mm contiguous sections and then stained using hematoxylin and eosin staining. Additional immunohistochemistry of markers for estrogen receptor (ER), progesterone receptor (PgR), and human epidermal growth factor receptor (HER)-2 was used to...
evaluate the hormone receptor status. All cases were diagnosed by more than two pathologists, and the following histological features were recorded: Presence of comedo necrosis, nuclear grade (1, 2 or 3), hormone receptor status (ER, PgR and HER-2), and tumor size (largest diameter).

**Statistical analysis**

The patient population showed normal distribution with the Smirnov-Kolmogorov analysis in this study. The patient characteristics and findings of [F-18] FDG-PET/CT (results of visual analysis and the SUVmax) were compared in each group using univariate analysis of Fisher’s exact test and Mann-Whitney U test. Further, we also performed multiple logistic regression analyses. Statistical calculations were performed with IBM SPSS statistics 22. P values of < 0.05 were regarded as significant.

**RESULTS**

**PET/CT findings**

Visual analysis revealed that [F-18] FDG uptake was seen in 28 lesions (53.8%, Figure 1). All values are provided as mean ± SD. The SUVmax of all lesions was 1.63 ± 0.90 (range, 0.62-5.49), of [F-18] FDG-positive lesions were 2.18 ± 1.16 (range, 1.16-5.49), and of [F-18] FDG-negative lesions were 0.99 ± 0.19 (range, 0.62-1.29, Figure 2).

**Clinicopathological characteristics**

Clinically, 39 lesions (75.0%) were symptomatic, 13 (25.0%) were detected on screening, and 15 (28.8%) were palpable (Table 1). Percutaneous biopsy was performed using ultrasound-guided CNB in 20 lesions (38.5%), ultrasound-guided VAB in 25 (48.1%), and stereotactic VAB in 7 (13.5%). Moreover, calcification on mammography was found in 27 lesions (52.0%). Eight patients (15.4%) had mass formation; 42 lesions (80.8%) did not have mass formation and 2 (3.8%) were undetected at MRI. The mean size of enhancing lesions at MRI was 31.7 ± 30.0 mm (range, 0-80 mm), of [F-18] FDG-negative lesions was 22.8 ± 18.5 mm (range, 0-60 mm), and of [F-18] FDG-positive lesions was 39.3 ± 18.2 mm (range, 7-80 mm). In 34 lesions (65.4%), sizes of enhancing lesions were ≥ 20 mm.

Microscopic observation revealed that 22 lesions (42.3%) had comedo necrosis, 42 (80.8%) were ER positive, 38 (73.1%) were PgR positive, and 22 (42.3%) were HER-2 positive. Further, 42 lesions (80.8%) were nuclear grade 1, 8 (15.4%) were nuclear grade 2, and 2 (3.8%) were nuclear grade 3. The mean pathological size of tumors was 41.4 ± 32.2 mm (range, 1-150 mm), of [F-18] FDG-negative lesions was 34.5 ± 39.4 mm (range, 1-150 mm), and of [F-18] FDG-positive lesions was 47.3 ± 24.0 mm (range, 2-85 mm); 37 lesions (71.1%) were ≥ 20 mm in size.

**Comparison between clinicopathological characteristics and PET/CT findings**

On univariate analysis (Table 2), visual analysis and significant association was found between the SUVmax and symptomatic presentation (P = 0.012 and 0.002, respectively), palpability (P = 0.030 and 0.024, respectively), use of CNB (P = 0.023 and 0.012, respectively), ultrasound-guided biopsy (P = 0.040 and 0.006, respectively), large size (≥ 20 mm) of enhancing lesion on MRI (P = 0.001 and 0.010, respectively), and large tumor size (≥ 20 mm) on histopathology (P = 0.002.
and 0.008, respectively). However, visual analysis and SUVmax failed to show association with age, presence of calcification on mammography, mass formation on MRI, presence of comedo necrosis, hormone receptor status (ER, PgR), HER-2 expression, and nuclear grade.

On multivariate analysis (Table 2), the factors significantly associated with visual analysis and SUVmax were symptomatic presentation ($P = 0.019$ and 0.001, respectively), use of CNB ($P = 0.001$ and 0.031, respectively), and large size ($\geq 20$ mm) of enhancing lesion on MRI ($P = 0.001$ and 0.049, respectively).

**DISCUSSION**

This study demonstrated that symptomatic and large DCIS ($\geq 20$ mm) often can be visualized on [F-18] FDG-PET/CT. The results mirrored those of the previous study showing sensitivity of 25%-77% \cite{10,11}. Although [F-18]
FDG uptake depends on cell density in patients with predominant and pure DCIS, the association between [F-18] FDG and clinicopathological features of DCIS has not been fully elucidated. To the best of our knowledge, our study has the largest sample number among studies examining clinicopathological features of DCIS visualized on [F-18] FDG-PET/CT. Our hypothesis is that large DCIS (≥ 20 mm) has possibility to be selected as target lesion on [F-18] FDG-PET/CT prior to neoadjuvant chemotherapy. In our study, FDG uptake was highly seen in enhancing lesions of ≥ 20 mm in size on MRI (24/34, 70.6%) and in tumors histopathologically measuring ≥ 20 mm on (25/37, 67.6%). These results show that large tumors tend to have [F-18] FDG uptake in DCIS, similar to a previous study that reported that sensitivity of [F-18] FDG-PET and PET/CT is associated to a large tumor size in IDC[10,11].

In this study, FDG uptake was highly detected in symptomatic (11/13, 84.6%) and palpable lesions (12/15, 80.0%); [F-18] FDG-PET/CT findings were markedly associated with symptomatic presentation and palpability. It has been reported that a large tumor tend to be symptomatic and palpable and that most tumors do not become palpable until a size > 10 mm in diameter is reached[10]. These factors may reflect the association between symptomatic presentation, palpability, and [F-18] FDG-PET/CT findings. In contrast, [F-18] FDG uptake was not highly visualized in screening-detected lesions (17/39, 43.6%) and non-palpable lesions (16/37, 43.2%). These results indicate that FDG-PET/CT was not appropriate for the screening of DCIS.

In our study, FDG uptake of symptomatic (11/13, 84.6%) and palpable lesions (12/15, 80.0%); [F-18] FDG-PET/CT findings were markedly associated with symptomatic presentation and palpability. It has been reported that a large tumor tend to be symptomatic and palpable and that most tumors do not become palpable until a size > 10 mm in diameter is reached[10]. These factors may reflect the association between symptomatic presentation, palpability, and [F-18] FDG-PET/CT findings. In contrast, [F-18] FDG uptake was not highly visualized in screening-detected lesions (17/39, 43.6%) and non-palpable lesions (16/37, 43.2%). These results indicate that FDG-PET/CT was not appropriate for the screening of DCIS.

We presented clinicopathological features of DCIS visualized on [F-18] FDG-PET/CT with the largest sample number and hypothesized that large DCIS (≥ 20 mm) might be selected as target lesion on [F-18] FDG-PET/CT prior to neoadjuvant chemotherapy. However, this study had several limitations. First, our study was conducted retrospectively. Second, our study could not correlate the findings with menopausal status and phase of the menstrual cycles, and presence of fibrocystic changes in patients, which can influence normal breast parenchymal enhancement at MRI and [F-18] FDG uptake on PET/CT[19,20]. Third, invasive intervention might affect [F-18] FDG uptake of the lesion in our study, because all patients had biopsy prior to [F-18] FDG-PET/CT study and the mean interval from biopsy to [F-18] FDG-PET/CT was 29.2 d (range, 43-133 d). And finally, PET/CT has limitation in DCIS less than 20 mm; however, MRI might be more sensitive in lesion less than 20 mm.

In conclusion, although most DCIS are non-avid on [F-18] FDG-PET/CT, [F-18] FDG uptake of symptomatic tumors or tumors greater than or equal to 20 mm often can be visualized.

COMMENTS

Background

Nowadays, the incidence of ductal carcinoma in situ (DCIS) has increased by prevailing mammography, ultrasound, and magnetic resonance imaging and over 20% of newly identified breast cancer consists of DCIS. However, the sensitivity of [F-18] fluorodeoxyglucose-positron emission tomography/ computed tomography (FDG PET/CT) for DCIS remains obscure.

Research frontiers

[F-18] FDG PET/CT has been an essential modality for detecting metabolic activity in primary breast tumor, diagnosing and staging local and distant sites, and evaluating the response to therapy. Wide range of the sensitivity to detect primary tumor in patients with DCIS exists.

Innovations and breakthroughs

DCIS often can be detected on [F-18] FDG-PET/CT in daily practice. The authors assessed clinicopathological features of tumor visualized on [F-18] FDG-PET/CT using pathologic specimens obtained by surgery.
Applications
This study revealed that symptomatic tumor and large DCIS (≥ 20 mm) is often visualized on [F-18] FDG-PET/CT. The results suggest that large DCIS (≥ 20 mm) has possibility to be selected as target lesion on [F-18] FDG-PET/CT prior to neoadjuvant chemotherapy. Devulking after neoadjuvant chemotherapy may affect surgical approach.

Terminology
DCIS is a group of non-invasive malignant epithelial tumors characterized by non-invasion of adjacent tissues and mostly adenocarcinoma derived from the mammary parenchymal epithelium. [F-18] FDG-PET/CT is one of the hybrid non-invasive imaging modalities which provides activity of glucose metabolism.

Peer-review
This article mentioned that DCIS of breast with diameter > 20 mm can be visualized on [F-18] FDG-PET/CT. This result is very useful in clinical diagnosis.

REFERENCES
1 Barreau B, de Mascarel I, Feuga C, MacGrogan G, Dilhady MH, Picot V, Dilhady JM, de Lara CT, Bussières E, Schreer I. Mammography of ductal carcinoma in situ of the breast: review of 909 cases with radiographic-pathologic correlations. Eur J Radiol 2005; 54: 55-61 [PMID: 15797293 DOI: 10.1016/j.ejrad.2004.11.019]
2 Berg WA, Gutierrez L, NessAiver MS, Carter WB, Bhargavan M, Lewis RS, Iffie OB. Diagnostic accuracy of mammography, clinical examination, US, and MR imaging in preoperative assessment of breast cancer. Radiology 2004; 233: 830-849 [PMID: 15486214 DOI: 10.1148/radiol.233301484]
3 Schouten van der Velden AP, Schlooz-Vries MS, Boetes C, Wobbes T. Magnetic resonance imaging of ductal carcinoma in situ: what is its clinical application? A review. Am J Surg 2009; 198: 262-269 [PMID: 19375068 DOI: 10.1016/j.amjsurg.2009.01.010]
4 Baur A, Bahrs SD, Speck S, Wietek BM, Krämer B, Vogel U, Claussen CD, Siegmann-Luz KC. Breast MRI of pure ductal carcinoma in situ: sensitivity of diagnosis and influence of lesion characteristics. Eur J Radiol 2013; 82: 1731-1737 [PMID: 23743052 DOI: 10.1016/j.ejrad.2013.05.002]
5 Jin ZQ, Lin MY, Hao WQ, Jiang HT, Zhang L, Hu WH, Zhang M. Diagnostic evaluation of ductal carcinoma in situ of the breast: ultrasonographic, mammographic and histopathologic correlations. Ultraschall Med 2015; 41: 47-55 [PMID: 25479813 DOI: 10.1055/s-0040-1714023]
6 Jemal A, Siegel R, Ward E, Murray T, Xu J, Thun MJ. Cancer statistics, 2007. CA Cancer J Clin 2007; 57: 43-66 [PMID: 17237035 DOI: 10.3322/canjclin.57.1.43]
7 Li CI, Daling JR, Malone KE. Age-specific incidence rates of in situ breast carcinomas by histologic type, 1980 to 2001. Cancer Epidemiol Biomarkers Prev 2005; 14: 1008-1011 [PMID: 15824180 DOI: 10.1158/1055-9156.EPI-04-0849]
8 Rosen EL, Eubank WB, Mankoff DA. FDG PET, PET/CT, and breast cancer imaging. Radiographics 2007; 27 Suppl 1: S215-S229 [PMID: 18180228 DOI: 10.1148/rg.27s075517]
9 Rostom AY, Powe J, Kandil A, Ezzat A, Bakheet S, el-KhWSky F, el-Hussainy G, Sobriss R, Sjöklint O. Positron emission tomography in breast cancer: a clinicopathological correlation of results. Br J Radiol 1999; 72: 1064-1069 [PMID: 10709822 DOI: 10.1259/bjr.72.863.10700822]
10 Avril N, Rosé CA, Schelling M, Dose J, Kuhn W, Bensse S, Weber W, Ziegler S, Graeff H, Schwager M. Breast imaging with positron emission tomography and fluorine-18 fluorodeoxyglucose: use and limitations. J Clin Oncol 2000; 18: 3495-3502 [PMID: 11032590]
11 Mavi A, Urban M, Yu JQ, Zhuang H, Houseni M, Cermik TF, Thiruvenkatasamy D, Czernecki B, Schnall M, Alavi A. Dual time point 18F-FDG PET imaging detects breast cancer with high sensitivity and correlates well with histologic subtypes. J Nucl Med 2006; 47: 1440-1446 [PMID: 16954551]
12 Groheux D, Giacchetti S, Delord M, Hindé E, Vercellino L, Cuvier C, Toutbert ME, Merlet P, Hennequin C, Espié M. 18F-FDG PET/CT in staging patients with locally advanced or inflammatory breast cancer: comparison to conventional staging. J Nucl Med 2013; 54: 5-11 [PMID: 23213197 DOI: 10.2967/jnumed.112.108684]
13 Wang Y, Zhang C, Liu J, Huang G. Is 18F-FDG PET accurate to predict neoadjuvant therapy response in breast cancer? A meta-analysis. Breast Cancer Res Treat 2012; 131: 357-369 [PMID: 21906110 DOI: 10.1007/s10549-011-1780-z]
14 D’Orsi CJ, Sickles EA, Mendelson EB, Morris EA. Breast Imaging Reporting and Data System: BI-RADS Atlas. 5th ed. American College of Radiology, Reston, 2013
15 Wang SY, Shamiyanan T, Viening BA, Kane R. Tumor characteristics as predictors of local recurrence after treatment of ductal carcinoma in situ: a meta-analysis. Breast Cancer Res Treat 2011; 127: 1-14 [PMID: 21327465 DOI: 10.1007/s10549-011-1387-4]
16 Groheux D, Espié M, Giacchetti S, Hindé E. Performance of FDG PET/CT in the clinical management of breast cancer. Radiology 2013; 266: 388-405 [PMID: 23220901 DOI: 10.1148/radiol.12110853]
17 Brennan ME, Turner RM, Giatto S, Marinovich ML, French JR, Macaskill P, Houssami N. Ductal carcinoma in situ at core-needle biopsy: meta-analysis of underestimation and predictors of invasive breast cancer. Radiology 2011; 260: 119-128 [PMID: 21493791 DOI: 10.1148/radiol.11102368]
18 Shigematsu H, Kadoya T, Masumoto N, Matsuura K, Ema A, Kajitani K, Amioka A, Okada M. Role of FDG-PET/CT in prediction of underestimation of invasive breast cancer in cases of ductal carcinoma in situ diagnosed at needle biopsy. Clin Breast Cancer 2014; 14: 358-364 [PMID: 24962555 DOI: 10.1016/j.clb.2014.04.006]
19 Kang SS, Ko EY, Han BK, Shin JH, Hahn SY, Ko ES. Background parenchymal enhancement on breast MRI: influence of menstrual cycle and breast composition. J Magn Reson Imaging 2014; 39: 526-534 [PMID: 23633296 DOI: 10.1002/jmri.24185]
20 Vranješević D, Schiepers C, Silverman DH, Quon A, Villalpando J, Dahlbom M, Phelps ME, Czernin J. Relationship between 18F-FDG uptake and breast density in women with normal breast tissue. J Nucl Med 2003; 44: 1238-1242 [PMID: 12902413]

Fujikawa T et al., Ductal carcinoma in situ of breast

P-Reviewer: Chu JP, Tsikouras PPT, Zafrakas M
S-Editor: Gong XM
L-Editor: A
E-Editor: Zhang FF
