

UCSF

UC San Francisco Previously Published Works

Title

The Incidence of Pulmonary Embolism and Associated FDG-PET Findings in IV Contrast-Enhanced PET/CT

Permalink

<https://escholarship.org/uc/item/0bq0k160>

Journal

Academic Radiology, 21(6)

ISSN

1076-6332

Authors

Flavell, Robert R
Behr, Spencer C
Brunsing, Ryan L
et al.

Publication Date

2014-06-01

DOI

10.1016/j.acra.2014.02.013

Peer reviewed

The Incidence of Pulmonary Embolism and Associated FDG-PET Findings in IV Contrast-Enhanced PET/CT

Robert R. Flavell, MD, PhD, Spencer C. Behr, MD, Ryan L. Brunsing, MD, PhD, David M. Naeger, MD, Miguel Hernandez Pampaloni, MD, PhD

Rationale and Objectives: Most fluorine-18 fluorodeoxyglucose (FDG)-positron emission tomography with computed tomography (PET/CT) studies are performed on cancer patients. These patients are at increased risk of pulmonary embolism (PE). In this retrospective review, we determined the rate of PE, and the prevalence of associated FDG-PET findings on intravenous (IV) contrast-enhanced PET/CT.

Materials and Methods: We identified all PET/CT studies performed at our institution with a reported finding of PE between January 2005 and October 2012. The medical record was reviewed for symptoms, which were identified after the diagnosis of PE, and whether the patients received treatment. The prevalence of associated FDG-PET findings was determined.

Results: A total of 65 total cases of PE (of 182,72 total PET/CT examinations) were identified of which 59 were previously unknown. This gives an incidental PE (IPE) rate of 0.32%. Of the patients where sufficient clinical information was available, 34 of 36 (94%) were treated either with therapeutic anticoagulation or inferior vena cava filter, and 30 of 36 (83%) were asymptomatic in retrospect. Of the patients with IPE, we found nine (15.2%) with associated focal pulmonary artery hypermetabolism, three (5.1%) with hypermetabolic pulmonary infarction, and one with increased isolated right ventricular FDG uptake (1.7%). One case of chronic PE demonstrated a focal hypometabolic filling defect in a pulmonary artery on PET.

Conclusions: We found IPE in 0.32% of PET/CT scans. Focal pulmonary artery hypermetabolism or hypometabolism, and hypermetabolic pulmonary artery infarction with the “rim sign” were uncommonly associated with PE. These findings could raise the possibility of IPE in non-IV contrast-enhanced PET/CT studies.

Key Words: Pulmonary embolism; PET/CT; thromboembolism; pulmonary infarction; pulmonary artery hypermetabolism.

©AUR, 2014

Acute pulmonary embolism (PE) represents a potential life threatening complication of venous thrombosis and is notoriously variable in presentation. Oncology patients have a higher risk of PE (1,2) and often present without the typical clinical manifestations (3). These so-called asymptomatic PEs are clinically significant in oncology patients. They serve as a marker for future symptomatic venous thromboembolism (VTE), a term that encompasses both deep venous thrombosis (DVT) and PE (4), and are

associated with decreased survival (5). The current consensus is for therapeutic intervention in these patients despite the absence of symptoms (1).

Several previous studies have investigated the rates of incidental PE (IPE) on contrast-enhanced computed tomography (CT) studies in the oncologic population, with reported rates ranging from 0.58% to 4.0% (2,4,6–13). Since its approval by the Food and Drug Administration in 2000, fluorine-18 fluorodeoxyglucose (FDG)-positron emission tomography fused with concurrent computed tomography (PET/CT) has rapidly evolved into a cornerstone imaging modality in oncology. At our institution, approximately 95% of PET/CT studies are performed in patients with known or suspected malignancy, and the routine imaging protocol includes intravenous (IV) contrast. In general, most centers perform low-dose, noncontrast-enhanced CT as part of their routine FDG-PET/CT protocol. At our institution, there is a consensus between referring clinicians and the radiology department that patients undergoing PET/CT have a contrast-enhanced diagnostic quality CT unless there is a

Acad Radiol 2014; 21:718–725

From the Department of Radiology and Biomedical Imaging, University of California, San Francisco, 505 Parnassus Avenue, M-396, San Francisco, CA, 94143 (R.R.F., S.C.B., D.M.N., M.H.P.); Biomedical Sciences Graduate Program, University of New Mexico School of Medicine, Albuquerque, New Mexico (R.L.B.); and Department of Medicine, Newton-Wellesley Hospital, Newton, Massachusetts (R.L.B.). Received December 7, 2013; accepted February 17, 2014. **Address correspondence to:** M.H.P. e-mail: Miguel.Pampaloni@ucsf.edu

©AUR, 2014

<http://dx.doi.org/10.1016/j.acra.2014.02.013>

contraindication such as renal failure or contrast allergy. One previous study found IPE in 13 of 2216 patients (0.59%) who had a contrast-enhanced PET/CT (13). The aim of our study was to evaluate the incidence of IPE in patients referred for FDG-PET/CT studies in a much larger cohort. An additional goal was to identify and characterize associated PET findings and their relative frequency.

MATERIALS AND METHODS

Patient Selection

This is a retrospective study based on the analysis of all FDG-PET/CT studies performed at our institution from January 1, 2005 to October 31, 2012. We searched our database of all PET/CT performed at our institution for the presence of any one of the terms “embolus,” “emboli,” “embolism,” “PE,” or “thromboembolism.” Only FDG-PET/CT studies performed at our institution were included in this study. Thus, outside studies submitted for internal review were excluded from further analysis. This was because of the potential for inhomogeneity in preparation and imaging acquisition protocols. The study was approved by our institutional review board.

Scan Technique

FDG-PET/CT examinations were performed on either a Biograph 16 (Hi-Rez) PET/CT scanner (Siemens AG, Erlangen, Germany) with an integrated PET and 16-MDCT scanner or a Discovery VCT PET/CT scanner (GE Medical Systems, Milwaukee, WI) with an integrated PET and 64-MDCT scanner. All patients fasted with hydration for at least 6 hours. Patients had blood glucose levels <200 mg/dL. Fluorine-18 FDG (0.45 ± 0.09 GBq) was injected intravenously followed by a 10-mL normal saline flush. Patients rested for 60 ± 15 minutes and voided before being positioned supine on the scanner table.

CT examinations were performed in neutral breath hold after the injection of 150 mL of iohexol (Omnipaque 350; GE Healthcare) at 3 mL/second. Acquisition was performed at kilovoltage peak of 120 with auto-milliampere second. Images were reconstructed as contiguous 5-mm slices. Additional lung reformats were generated with contiguous 2-mm slices. PET was performed immediately after CT, without patient repositioning. PET images were obtained in a three-dimensional mode at 7–10 bed positions per patient, with an acquisition time of 3–4 minutes per station, from the skull vertex through the mid thigh, except for patients with clinical indication to scan to the toes such as melanoma and myeloma. The CT, PET, and fused PET/CT images were displayed in orthogonal planes on an Advantage Workstation (GE Healthcare). Maximum standardized uptake value (SUV_{max}) was based on a total body weight and determined on the Advantage Workstation. All scans were initially interpreted and reported by members of both the nuclear medicine section and the thoracic, abdominal, or neuroradiology sections.

Imaging and Chart Review

The medical charts of patients included in the study were reviewed for demographic data, initial diagnosis, metastatic disease, and the presence and location of PE.

All cases were reviewed independently for the presence of PE by three radiologists (R.F., a radiology resident with 2 years of experience; S.B., attending radiologist with fellowship training in body CT and nuclear medicine; and D.N., attending radiologist with fellowship training in chest CT and nuclear medicine). The presence of PE was determined using commonly accepted methodology (14). Only cases in which all three radiologists independently agreed with the diagnosis of PE were included. The largest affected vessel (main, lobar, segmental, or subsegmental) was noted. The presence or absence of PE on previous chest CT was reviewed. The PET scan associated with each CT was reviewed for cardiac or pulmonary uptake, which was not attributable to metastatic disease. The electronic medical record was reviewed for the presence of symptoms attributable to PE or DVT. The medical record was also reviewed to see if patients were treated with anticoagulation or inferior vena cava (IVC) filter placement.

To estimate the total number of IV contrast-enhanced FDG-PET/CT studies performed over the relevant time frame, we used the search engine to identify all studies done in the month of August for each year from 2005 through 2012. All studies were independently reviewed to determine whether IV contrast was injected. The number of IV contrast-enhanced and noncontrast-enhanced FDG-PET/CT performed in the month of August from a given year was used to determine an estimated number of total for that year in its entirety. Numbers were adjusted from 2008 (leap year) to 2012 (leap year and only days up through October) to reflect the difference in total number of days included in the period of our search. A search of all reports for the year 2011, using the search term “Omnipaque” (the type of IV contrast used at our institution), demonstrated that the rate of IV contrast-enhanced FDG-PET/CT including this term in the report was consistent throughout the year at approximately 8.02 scans per day with a standard deviation of 0.88. Although this term was not included in all reports, it suggests a rate of FDG-PET/CT studies that did not vary significantly over months, which validated our approach. A similar approach was used to determine the total number of PET/CT studies done in breast cancer and melanoma patients.

RESULTS

Total Number of Cases Reviewed

Because of the very large number of PET/CT examinations performed it was not possible to individually review all reports for the administration of contrast. Using the method outlined previously, we estimate a total of 18,272 IV contrast-enhanced PET/CT scans performed during the study period (Supplemental Fig. 1A). Our data show that there has been

a steady increase in the number of FDG-PET/CT studies performed at our hospital from 2005 to 2011, and the percentage of FDG-PET/CT studies performed with IV contrast enhancement has steadily declined at a rate of about 1.25% per year (Supplemental Fig. 1B). This likely reflects a variety of factors, including advanced age of the patients being referred for imaging leading to an increased prevalence of renal failure, and perhaps trends in patient refusal of IV contrast administration.

Incidence of PE

Between January 2005 and October 2012, 72 FDG-PET/CT studies were identified which reported a PE. On review, seven studies were excluded because of insufficient findings for the diagnosis of PE by any of the three readers. An additional six cases had been seen on previous contrast-enhanced CT, with the amount of time from initial detection to the PET/CT ranging from 17 to 1702 days. None of the studies were done as a result of clinical concern for a PE. In total, 59 FDG-PET/CT studies containing a finding of an IPE were included in our cohort. This represents an estimated 0.32% incidence of IPE on FDG-PET/CT imaging at our institution (Table 1).

The demographics of patients with IPE are included as Table 2. The mean age was 55.8 years old. There were nearly equal numbers of males ($n = 29$) and females ($n = 30$). The vast majority ($n = 57$; 96.6%) had a primary diagnosis of malignancy, reflecting the prevalence of cancer in the population of patients who undergo PET/CT scanning. Of those patients diagnosed with a solid tumor, 34 of 51 (66.7%) had metastatic disease. Information about treatment was available in 47 of 59 patients with IPE. Of those 47 patients, 30 were undergoing chemotherapy (63.8%), 13 had prior surgical resection of their neoplasm (27.6%), 7 had undergone radiation (14.9%), and 19 had not yet received treatment.

The primary diagnosis of all cases in our cohort was determined by chart review. Two patients did not have diagnosis of cancer, although all were imaged because of concern for possible malignant disease. In all, 57 of 59 cases (96.6%) had a primary malignant disease at the time of IPE. This result is not surprising, because of the prevalence of malignancy in the patient population undergoing PET/CT. The most prevalent diagnoses are summarized in Table 3. This distribution is different from previously reported cohorts of IPE (4,10), likely reflecting institution-specific referral patterns in addition to the proven utility of PET/CT in these cancers.

Previous studies have shown that there is large variability in the incidence of VTE according to the type of primary malignancy, with lower rates in breast cancer compared to other primary sites like pancreatic cancer, lung cancer, and tumors of gastrointestinal origin (4,10,12,15,16). Given that the largest number of IPE cases in our cohort were found in breast cancer and melanoma patients, we next determined the rate of IPE detection per tumor type to delineate if our incidence reflected a higher number of these patients at our

TABLE 1. Pulmonary Embolism Characteristics

Total IV contrast-enhanced PET/CT scans*	18,272
Number of cases with PE	65
Number of cases of incidental PE	59 (0.32%)
Number of cases with chronic PE	6
Location in pulmonary artery	
Main PA	9/59 (15%)
Lobar PA	15/59 (25%)
Segmental PA	27/59 (46%)
Subsegmental PA	8/59 (14%)

IV, intravenous; PA, pulmonary artery; PE, pulmonary embolism; PET/CT, positron emission tomography with computed tomography.

*Estimate.

TABLE 2. Demographics of Patients with IPE

Age	
Mean	55.8
Standard deviation	15
Range	23–84
Malignant neoplasm*	57/59 (97%)
Metastases	34/51 (67%)
Female	30/59 (51%)
Male	29/59 (49%)
Treatment†	
Chemotherapy	30/47 (64%)
Surgery	13/47 (28%)
Radiation	7/47 (15%)
None	19/47 (21%)

IPE, incidental pulmonary embolism.

*Solid tumors only.

†In patients where sufficient clinical information was available. Twelve patients did not have sufficient data to determine treatment course.

TABLE 3. Characteristics of Tumors

Type	# IPE	Total*	Incidence*
Breast	13	2563	0.51%
Melanoma	10	2839	0.35%
Lymphoma	6		
Lung	5		
Esophageal	4		
Colon	3		
Anorectal	2		
Ewing's sarcoma	2		
Multiple myeloma	2		
Other	10		
No malignancy	2		

IPE, incidental pulmonary embolism.

*Estimated.

institution, or a true deviation from previously reported IPE patterns. Our data show that within our cohort of patients the overall incidence of IPE on IV contrast-enhanced FDG-PET/CT studies is 0.51% in breast cancer patients and

0.35% in melanoma patients (Table 3), which is consistent with a previous report (15).

Pulmonary Emboli Characteristics

In nearly half of the 59 cases ($n = 27$; 45.8%) a segmental branch of the pulmonary artery was the largest vessel involved, whereas in 24 of 59 cases (40.7%) either a lobar or main branch of the pulmonary artery was involved (Table 1). This distribution is similar to a previous retrospective study in a cohort of cancer patients, where 58.8% of IPE were segmental and 21.6% were in major vessels, namely main or lobar branches (4). Sample images from several cases of variable severity are included as Figures 1–5 and Supplemental Figures 2–4.

Clinical Symptoms and Treatment with Anticoagulation

Of the 59 patients with IPE, 23 did not have sufficient information in the electronic medical record to determine if they were treated with anticoagulation or if they had any clinical symptoms. Of the 36 remaining patients, 2 (6%) were not treated. One patient was not treated because of the presence of hemorrhagic brain metastasis and very poor prognosis precluding IVC filter placement. A second patient was not treated because of the presence of only isolated subsegmental PE, brain metastases, and a negative lower extremity ultrasound study for DVT. Of the remaining patients, 31 (86%) were treated with therapeutic anticoagulation, and 3 (8%) had an IVC filter placed. All the patients were treated on the basis of the PET/CT findings, without confirmatory CT angiogram. In the 36 patients where sufficient clinical information was available, 30 patients were asymptomatic, even in retrospect (83%), whereas two had lower extremity swelling, suggesting DVT (6%), one had fever (3%), and three had dyspnea (8%).

Review of Associated FDG-PET Findings

We reviewed the PET images associated with all the studies that were positive for PE for additional findings. These results are summarized in Table 4. We found a qualitatively discernible focal increase in FDG uptake within the pulmonary artery at or adjacent to the PE, in comparison to other pulmonary arteries in the same patient, in 9 of 59 cases of IPE. A sample case is demonstrated in Figure 1 and Supplemental Figure 2. The mean SUV_{max} of these lesions was 2.2 ± 0.7 (SD). A single case of documented chronic PE in the right main pulmonary artery, which had been seen on multiple CT scans over a period of 1702 days, demonstrated an associated hypometabolic filling defect (Fig. 2 and Supplemental Fig. 3).

We also reviewed all cases for the associated finding of a pulmonary infarction. We used commonly accepted criteria for pulmonary infarction as a peripheral, triangular consolidation with central lucency, located in the expected vascular

TABLE 4. PET Findings Associated with IPE

High SUV in right ventricle	1/59 (2%)
Hypermetabolic pulmonary infarction	3/59 (5%)
Hypermetabolic pulmonary artery	9/59 (15%)
Mean SUV	2.2 ± 0.7
Max SUV	3.2
Main PA	1/9
Lobar PA	2/9
Segmental PA	4/9
Subsegmental PA	2/9

IPE, incidental pulmonary embolism; PA, pulmonary artery; SUV, standardized uptake value.

distribution of the associated embolism (17). We found pulmonary infarction in only three cases of PE (Figs. 3 and 4 and Supplemental Fig. 4). All these cases demonstrated associated hypermetabolism. One case demonstrated a complete rim of FDG avidity (Fig. 3), whereas the other two did not (Fig. 4 and Supplemental Fig. 4).

Given the potential for right heart strain as a result of increased vascular resistance in the setting of PE, we also reviewed all cases for the presence of an abnormal pattern of cardiac uptake. One case demonstrated markedly increased uptake in the right ventricular wall (Fig. 5). This case was associated with saddle PE and bowing of the interventricular septum seen on CT.

DISCUSSION

Cancer patients are known to be prone to the development of VTE because of an underlying hypercoagulable state (18,19). Thromboembolism is the second leading cause of death among cancer patients undergoing chemotherapy (20). In fact, 15% of patients with malignant disease develop a clinically significant VTE at some point during the course of their disease (19), a number that does not include asymptomatic PE.

We estimate that between January 2005 and October 2012 there were 20,815 FDG-PET/CT scan done at our institution, 18,272 (87.8%) of which were done with IV contrast (Supplemental Fig. 1). From the cohort of IV contrast-enhanced images, we identified 59 cases of IPE, representing an estimated 0.32% of studies. The rate of IPE in cancer patients, when reported on a per CT study basis has varied significantly across previous studies, ranging from 0.58% to 4.0% (7–9,15). Our rate of 0.33% is lower than these reported values, likely because of multiple factors that have previously been reported to affect the rate of IPE (21). Firstly, our study includes only outpatients, as the PET/CT scanners at our institution are located at separate facility from inpatients. One meta-analysis found an overall rate of 1.2% of IPE in outpatients, similar but still greater than the rate determined in our study (21). Secondly, our study was a retrospective review from reports, so the reported incidence is likely less than the true incidence because of search error

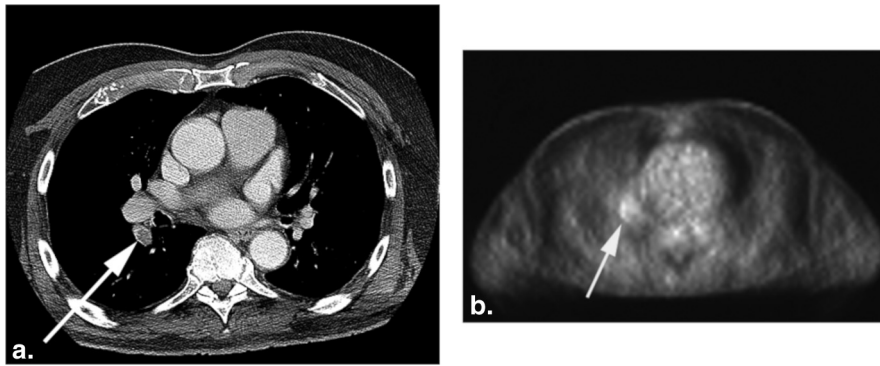


Figure 1. Example of pulmonary artery hypermetabolism. A 75-year-old woman with history of melanoma. **(a)** Incidental right lower lobe segmental pulmonary embolism seen on computed tomography. **(b)** The associated PET demonstrates mild focal pulmonary arterial hypermetabolism, with an SUV_{max} of 2.0. PET, positron emission tomography; SUV, standardized uptake value.

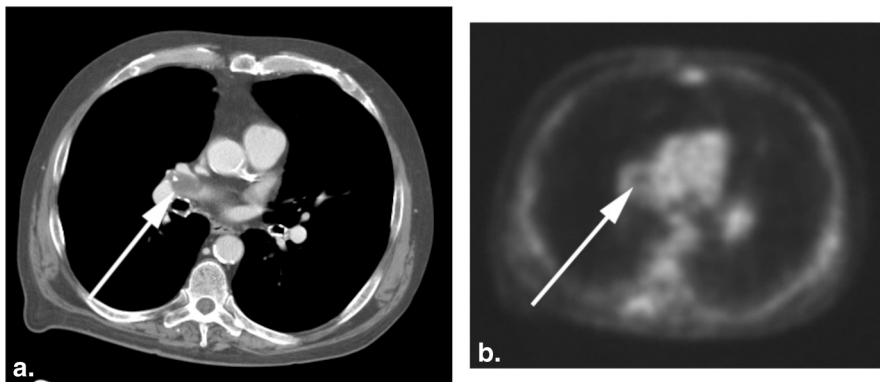


Figure 2. Example showing hypometabolism in chronic pulmonary embolism. A 66-year-old man with history of non-Hodgkin's lymphoma. **(a)** Contrast-enhanced CT demonstrates a calcified, eccentrically located filling defect in the right main pulmonary artery. This defect had been present on previous CTs and is consistent with a chronic pulmonary embolism. **(b)** The associated PET demonstrates a hypometabolic pulmonary arterial filling defect. CT, computed tomography; PET, positron emission tomography.

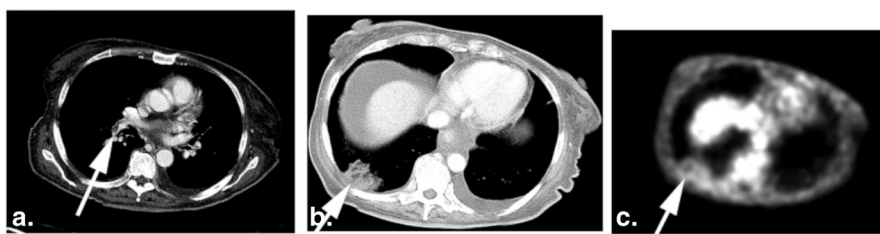


Figure 3. - Example of peripheral pulmonary infarction with associated complete rim of FDG avidity. A 66-year-old woman with a history of gastric cancer. Contrast-enhanced computed tomography demonstrates a pulmonary embolism involving the interlobar pulmonary artery **(a)** and a right lower lobe pulmonary infarction **(b)**. The associated PET demonstrates a rim of FDG avidity at the borders of the infarction **(c)**. FDG, fluorodeoxyglucose; PET, positron emission tomography.

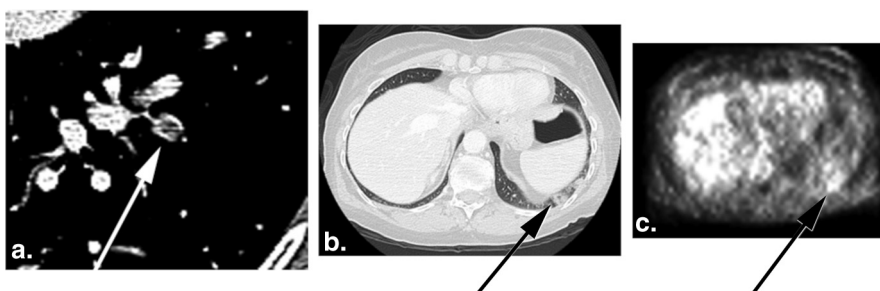


Figure 4. Example of peripheral pulmonary infarction with associated FDG avidity. A 61-year-old woman with history of breast cancer. Contrast-enhanced computed tomography demonstrates a segmental left lower lobe pulmonary embolism **(a)** and a small left lower lobe pulmonary infarction **(b)**. The associated PET demonstrates an incomplete rim of FDG avidity surrounding the area of the infarction **(c)**. FDG, fluorodeoxyglucose; PET, positron emission tomography.

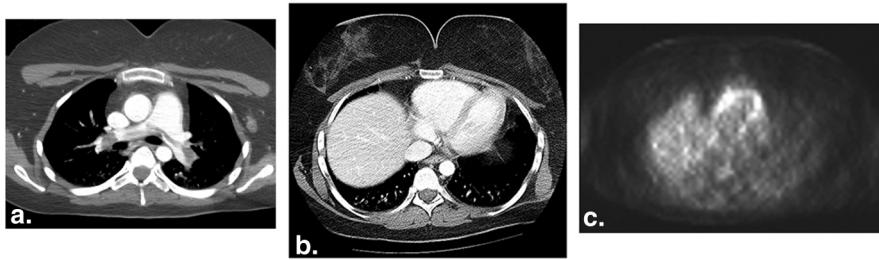


Figure 5. Example showing increased right ventricular uptake. A 29-year-old woman with metastatic melanoma. **(a)** Saddle pulmonary embolus was seen on contrast-enhanced CT. **(b)** CT demonstrating bowing of the interventricular septum. **(c)** PET demonstrating near-circumferential increased FDG uptake in the right ventricular wall. CT, computed tomography; FDG, fluorodeoxyglucose; PET, positron emission tomography.

and prospectively underdetecting IPE. Thirdly, the contrast bolus timing in our contrast-enhanced studies is optimized for tissue enhancement rather than for detection of PE. It is likely that a substantially greater fraction of examinations would demonstrate PE in these patients if they had undergone an appropriate protocol and interpreted CT angiogram. Finally, it is likely that the referral base for PET/CT represents a different patient cohort than those receiving standard chest CT. For example, our cohort includes a large number of patients with breast cancer and melanoma, but relatively few patients with gastrointestinal malignancies. Accordingly, when comparing disease-specific incidence of IPE in breast cancer and melanoma, we found rates of 0.51% and 0.35%, respectively, similar in comparison to a previous report (15). One previous study reviewed a smaller cohort of cancer patients undergoing contrast-enhanced PET/CT and found the rate of PE to be 0.59% (13). These results are similar to our own, likely because of similar patient cohort as in our study. Taken together, these factors of CT protocol, interpretation, and patient selection explain the relatively low rate of IPE in our study.

In the 36 cases of IPE where sufficient clinical information was available, most cases (34, 94%) were treated either with therapeutic anticoagulation or an IVC filter. These data are similar to previously published series and congruent with the clinical consensus that incidental pulmonary embolus be treated with anticoagulation (1,4,6,12,15). Most patients (30/36, 83%) were asymptomatic, even in retrospect. Two patients had retrospectively identified symptoms of DVT (6%), and four (11%) had either fever or pulmonary symptoms.

Multiple PET findings have been described in association with pulmonary emboli, including pulmonary artery hypermetabolism (13,22–24), four-chamber cardiac uptake (25), and the presence of a rim of hypermetabolism surrounding a pulmonary infarction (26,27). However, the relative frequency of these findings has not been previously described. One additional novel sign of PE identified in this study was the presence of a hypometabolic filling defect in the pulmonary artery tree. This finding was associated with a single case of a large chronic right main pulmonary artery embolism (Fig. 2 and Supplemental Fig. 3). This finding was only identified in one of six cases of previously identified PE and is therefore unlikely to be present in most cases of chronic PE. However, as no mimics of this sign have been pre-

viously reported, we propose that the finding of a hypometabolic pulmonary arterial filling defect should prompt search for underlying embolism.

We found qualitatively increased focal or curvilinear FDG uptake in the pulmonary arterial in 9 of 59 cases of IPE, with a mean SUV_{max} of 2.2 (Fig. 1, Supplemental Fig. 2, and Table 4). This value is similar to previously published reports of 1.65 (13), 2.3 (23), and 1.7 (24). We found that focal pulmonary arterial hypermetabolism could only be qualitatively detected in 15.2% of cases, indicating that this is an insensitive secondary sign of PE. The differential diagnosis for the finding of pulmonary arterial hypermetabolism includes pulmonary artery sarcoma, pulmonary artery metastasis, and FDG microembolism secondary to paravenous injection (23,24,28). These should be readily distinguishable because of the much higher SUV reported for pulmonary artery malignancies, and because of the presence of extravasation of FDG at the injection site in the case of microembolism.

The physiology underlying the variability in pulmonary artery uptake associated with PE is unclear. The resolution of PET/CT is insufficient to determine if the uptake is associated with the thrombus itself, or with the wall of the vessel. It is possible that the hypermetabolism may reflect vessel wall inflammation, as previously suggested (13). Pathologic analysis of pulmonary thrombectomy specimens commonly demonstrates mild inflammation, but may demonstrate moderate or severe inflammation in a minority of cases (13.4% and 1.3% of cases, respectively) (29). The fact that we observed increased pulmonary artery hypermetabolism in 15.2% of cases is similar to the incidence of inflammation in previous pathologic series, suggesting that these findings may be correlated. However, this remains a possibility for further study.

We found pulmonary infarction in only 3 of 59 cases of PE (Figs. 3 and 4 and Supplemental Fig. 4). All these cases demonstrated some focal hypermetabolism associated with the infarction, as has been previously described (13,30). One demonstrated a complete rim of FDG avidity (Fig. 3), the previously described “rim sign” (27). Although only seen in a single case in our series, the rim sign may represent a relatively specific secondary sign of PE. A more homogeneous pattern of uptake, such as seen in other cases of pulmonary infarction (Fig. 4 and Supplemental Fig. 4), is not specific and could be seen in pneumonia or atelectasis.

Finally, we assessed for an abnormal pattern of cardiac uptake. We did not identify any cases of four-chamber increase in cardiac uptake, as has been previously described (25). We identified a single case that demonstrated markedly increased right ventricular myocardial uptake in a patient with a saddle PE (Fig. 5). CT also demonstrated marked dilation of the right ventricle in proportion to the left ventricle and bowing of the interventricular septum, a known secondary sign of elevated right heart pressure (31). An increase in right ventricular myocardial FDG uptake has been described in association with pulmonary arterial hypertension (32–34). Therefore, the finding of increased right ventricular wall uptake in this patient likely reflects acutely increased pulmonary arterial and right heart pressure because of the massive PE. The finding of markedly increased right ventricular myocardial uptake is unlikely to represent a specific sign for PE, as there are numerous causes of pulmonary arterial hypertension.

Our study has several limitations. As a single institution study, our patient population undergoing FDG-PET/CT may not be the same as other institutions. Patient populations at other institutions will reflect the referral pattern unique to that institution. The patient population undergoing PET/CT imaging at our institution may have a different prevalence of metastatic disease, undergo chemotherapy at a different rate, and have a different proportion of outpatients than those being imaged at other facilities. As a tertiary care hospital with a large associated cancer center, we are probably witnessing the evolving role of FDG-PET/CT from a predominantly first line imaging modality for staging to the modality of choice for the assessment of response to therapy. Finally, because of the very large number of studies analyzed, it was not possible to estimate the rate of PE on a per-patient basis, limiting direct comparison against some previous studies.

CONCLUSIONS

Our findings indicate that IPE are found in 0.32% of IV contrast-enhanced FDG-PET/CT scans, and that a large percentage of these lesions are found in the lobar or main branches of the pulmonary artery. We found focal pulmonary artery hypermetabolism in 15.2% of the cases of IPE and focal pulmonary artery hypometabolism in a single case of chronic PE. Furthermore, we found three cases of hypermetabolic pulmonary infarction (5.1% of cases) and one case of pulmonary infarction with a complete rim of FDG avidity. A single case (1.7%) demonstrated isolated increased right ventricular uptake, associated with a saddle pulmonary embolus and CT evidence of elevated right heart pressure. Although these secondary signs are present in a minority of cases, we propose that the findings of focal pulmonary artery hypermetabolism or hypometabolism, and hypermetabolic pulmonary infarction with rim sign are likely to represent specific secondary signs for PE. Therefore, the presence of these findings on noncontrast PET/CT should alert the radiologist to the possibility of underlying PE.

SUPPLEMENTARY DATA

Supplementary data related to this article can be found at <http://dx.doi.org/10.1016/j.acra.2014.02.013>.

REFERENCES

- den Exter PL, Jimenez D, Kroft LJ, et al. Outcome of incidentally diagnosed pulmonary embolism in patients with malignancy. *Curr Opin Pulm Med* 2012; 18:399–405.
- Heit JA, Silverstein MD, Mohr DN, et al. Risk factors for deep vein thrombosis and pulmonary embolism: a population-based case-control study. *Arch Intern Med* 2000; 160:809–815.
- Fulkerson WJ, Coleman RE, Ravin CE, et al. Diagnosis of pulmonary embolism. *Arch Intern Med* 1986; 146:961–967.
- den Exter PL, Hooijer J, Dekkers OM, et al. Risk of recurrent venous thromboembolism and mortality in patients with cancer incidentally diagnosed with pulmonary embolism: a comparison with symptomatic patients. *J Clin Oncol* 2011; 29:2405–2409.
- Sorensen HT, Mellemejaer L, Olsen JH, et al. Prognosis of cancers associated with venous thromboembolism. *N Engl J Med* 2000; 343:1846–1850.
- Gladish GW, Choe DH, Marom EM, et al. Incidental pulmonary emboli in oncology patients: prevalence, CT evaluation, and natural history. *Radiology* 2006; 240:246–255.
- Sebastian AJ, Paddon AJ. Clinically unsuspected pulmonary embolism—an important secondary finding in oncology CT. *Clin Radiol* 2006; 61:81–85.
- Cronin CG, Lohan DG, Keane M, et al. Prevalence and significance of asymptomatic venous thromboembolic disease found on oncologic staging CT. *AJR Am J Roentgenol* 2007; 189:162–170.
- Rita Larici A, Calandriello L, Maggi F, et al. Prevalence of incidental pulmonary emboli in oncology patients. *Radiology* 2007; 245:921–922. author reply—2.
- Browne AM, Cronin CG, English C, et al. Unsuspected pulmonary emboli in oncology patients undergoing routine computed tomography imaging. *J Thorac Oncol* 2010; 5:798–803.
- Gladish GW, Erasmus JJ. Unsuspected pulmonary emboli in oncology patients undergoing routine computed tomography imaging. *J Thorac Oncol* 2010; 5:759–760.
- Abdel-Razeq HN, Mansour AH, Ismael YM. Incidental pulmonary embolism in cancer patients: clinical characteristics and outcome—a comprehensive cancer center experience. *Vasc Health Risk Manag* 2011; 7:153–158.
- Wittram C, Scott JA. 18F-FDG PET of pulmonary embolism. *AJR Am J Roentgenol* 2007; 189:171–176.
- Wittram C, Maher MM, Yoo AJ, et al. CT angiography of pulmonary embolism: diagnostic criteria and causes of misdiagnosis. *Radiographics* 2004; 24:1219–1238.
- Di Nisio M, Ferrante N, De Tursi M, et al. Incidental venous thromboembolism in ambulatory cancer patients receiving chemotherapy. *Thromb Haemost* 2010; 104:1049–1054.
- Menapace LA, Peterson DR, Berry A, et al. Symptomatic and incidental thromboembolism are both associated with mortality in pancreatic cancer. *Thromb Haemost* 2010; 106:371–378.
- Revel MP, Triki R, Chatellier G, et al. Is it possible to recognize pulmonary infarction on multisection CT images? *Radiology* 2007; 244:875–882.
- Blom JW, Doggen CJ, Osanto S, et al. Malignancies, prothrombotic mutations, and the risk of venous thrombosis. *JAMA* 2005; 293:715–722.
- Caine GJ, Stonelake PS, Lip GY, et al. The hypercoagulable state of malignancy: pathogenesis and current debate. *Neoplasia (New York, NY)* 2002; 4:465–473.
- Khorana AA, Francis CW, Culakova E, et al. Thromboembolism is a leading cause of death in cancer patients receiving outpatient chemotherapy. *J Thromb Haemost* 2007; 5:632–634.
- Dentali F, Ageno W, Becattini C, et al. Prevalence and clinical history of incidental, asymptomatic pulmonary embolism: a meta-analysis. *Thromb Res* 2010; 125:518–522.
- Goethals I, Smeets P, De Winter O, et al. Focally enhanced f-18 fluorodeoxyglucose (FDG) uptake in incidentally detected pulmonary embolism on PET/CT scanning. *Clin Nucl Med* 2006; 31:497–498.

23. Ito K, Kubota K, Morooka M, et al. Diagnostic usefulness of 18F-FDG PET/CT in the differentiation of pulmonary artery sarcoma and pulmonary embolism. *Ann Nucl Med* 2009; 23:671–676.
24. Lee EJ, Moon SH, Choi JY, et al. Usefulness of fluorodeoxyglucose positron emission tomography in malignancy of pulmonary artery mimicking pulmonary embolism. *ANZ J Surg* 2013; 83:342–347.
25. Franceschi AM, Matthews R, Mankes S, et al. Four chamber FDG uptake in the heart: an indirect sign of pulmonary embolism. *Clin Nucl Med* 2012; 37: 687–691.
26. Badr A, Joyce JM, Durick J. Rim of FDG uptake around a pulmonary infarct on PET/CT in a patient with unsuspected pulmonary embolism. *Clin Nucl Med* 2009; 34:285–286.
27. Soussan M, Rust E, Pop G, et al. The rim sign: FDG-PET/CT pattern of pulmonary infarction. *Insights Imaging* 2012; 3:629–633.
28. Farsad M, Ambrosini V, Nanni C, et al. Focal lung uptake of 18F-fluorodeoxyglucose (18F-FDG) without computed tomography findings. *Nucl Med Commun* 2005; 26:827–830.
29. Bernard J, Yi ES. Pulmonary thromboendarterectomy: a clinicopathologic study of 200 consecutive pulmonary thromboendarterectomy cases in one institution. *Hum Pathol* 2007; 38:871–877.
30. Kamel EM, McKee TA, Calcagni ML, et al. Occult lung infarction may induce false interpretation of 18F-FDG PET in primary staging of pulmonary malignancies. *Eur J Nucl Med Mol Imaging* 2005; 32: 641–646.
31. Reid JH, Murchison JT. Acute right ventricular dilatation: a new helical CT sign of massive pulmonary embolism. *Clin Radiol* 1998; 53: 694–698.
32. Oikawa M, Kagaya Y, Otani H, et al. Increased [18F]fluorodeoxyglucose accumulation in right ventricular free wall in patients with pulmonary hypertension and the effect of epoprostenol. *J Am Coll Cardiol* 2005; 45: 1849–1855.
33. Basu S, Alzeair S, Li G, et al. Etiopathologies associated with intercostal muscle hypermetabolism and prominent right ventricle visualization on 2-deoxy-2[F-18]fluoro-D-glucose-positron emission tomography: significance of an incidental finding and in the setting of a known pulmonary disease. *Mol Imaging Biol* 2007; 9:333–339.
34. Bokhari S, Raina A, Rosenweig EB, et al. PET imaging may provide a novel biomarker and understanding of right ventricular dysfunction in patients with idiopathic pulmonary arterial hypertension. *Circulation* 2011; 4: 641–647.