

cine CT images eliminates artifactual defects in PET  $^{82}\text{Rb}$  images. Our concern with the study is that it contrasts false-positive findings from cine CT with software alignment and false-positive findings from helical CT without software alignment. Because most manufacturers of PET/CT scanners have a software alignment tool to be used in conjunction with helical CT, we suggest that it is appropriate and important for Gould et al. to compare false-positive findings from software-aligned cine CT and false-positive findings from software-aligned helical CT (e.g., their slow helical CT scan). Table 5 of Gould et al. lists an artifact frequency for unshifted slow helical CT studies (27%, or 39/145) similar to that found for “conventional” PET  $^{82}\text{Rb}$  studies (21%, or 252/1,177) in an earlier publication by Dr. Gould’s group (2), in which a  $^{68}\text{Ge}$  rod source was used for the attenuation correction. After visual checking of the PET  $^{82}\text{Rb}$  and attenuation images for misregistration, the misaligned conventional rod source studies were manually shifted using computer software (2). Moreover, even with cine CT, Gould et al. reported that 19% (22/114) of patient datasets were misaligned with PET and required software alignment (1). That is, the new paper (1) combined with the earlier publication (2) supports the conclusion that free-breathing helical CT and cine CT have nearly the same frequency of artifacts as does conventional PET. Significantly, all techniques required a software alignment solution.

In our institution, we have used a somewhat different approach on our 64-slice Biograph PET/CT scanner (Siemens). We acquire 3 very fast (2.7-s) helical CT scans during free breathing, 1 immediately before and 2 immediately after acquiring the stress PET  $^{82}\text{Rb}$  images (3). The exposure,  $\text{CTDI}_{\text{vol}}$ , is 0.7 mGy in each scan. We have found that this protocol increases the probability of alignment between a PET  $^{82}\text{Rb}$  image and an acquired CT image. Our protocol also includes 3 CT scans at rest for correction of the rest PET  $^{82}\text{Rb}$  scan. Like Gould et al. (1), we have estimated visually the degree of PET/CT misalignment with this procedure using the PET/CT 3-dimensional fusion software of the manufacturer (3). We found no apparent misalignment between PET and at least 1 CT scan in 85% of studies at stress and 89% of studies at rest. The best-case misalignment was small, and appropriate for PET attenuation correction, in an additional 14% of the studies at stress and 11% of the studies at rest (3). In only a few cases (<1%) did we observe a large or severe PET misalignment with all 3 of the CT scans that then required computer software alignment. We have acquired 1,400 rest/stress PET/CT  $^{82}\text{Rb}$  clinical studies with this protocol. The total  $\text{CTDI}_{\text{vol}}$  is 4.2 mGy with our 6-CT scan protocol. Gould et al. quoted a radiation exposure of 5.7 mGy for their helical CT scan and a radiation dose of 10 mGy for cine CT. We are studying techniques to reduce the dose even further. These steps include reducing the x-ray voltage from 120 to 100 kVp and even to 80 kVp in very thin patients and reducing the number of CT scans, thus requiring a greater reliance on software alignment.

In summary, the slow helical non-breath-hold CT approach originally proposed by Brunken et al. (4) produces a frequency of misalignment-related artifacts that is similar to the frequency reported for cine CT (1) and conventional PET (2). Gould et al. (1) did not provide the false-positive rate for software-shifted non-breath-hold helical CT, and this omission represents a major limitation of the paper. PET and CT alignment can be achieved with a fast helical multi-CT scan protocol that limits the need for software alignment tools to a small percentage of studies, while using an even lower radiation dose (3).

## REFERENCES

1. Gould KL, Pan T, Loghin C, Johnson NP, Guha A, Sdringola S. Frequent diagnostic errors in cardiac PET/CT due to misregistration of CT attenuation and emission PET images: a definitive analysis of causes, consequences, and corrections. *J Nucl Med.* 2007;48:1112–1121.
2. Loghin C, Sdringola S, Gould KL. Common artifacts in PET myocardial perfusion images due to attenuation-emission misregistration: clinical significance, causes, and solutions. *J Nucl Med.* 2004;45:1029–1039.
3. Streeter J, Eisner R, Hamill J, Nelson M, Patterson R. Attenuation correction of stress PET Rb82 with ultrafast CT images [abstract]. *J Nucl Med.* 2007;48(suppl 2):446P.
4. Brunken RC, DiFilippo FP, Bybel B, Neumann DR, Kaczur T, White RD. Clinical evaluation of cardiac PET attenuation correction using “fast” and “slow” CT images [abstract]. *J Nucl Med.* 2004;45(suppl):120P.

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**REPLY:** In their letter to the editor, Eisner and Patterson make 3 criticisms that they call “major limitations” to our report on attenuation–emission misregistration in cardiac PET/CT (1), as follows: First, the frequency of attenuation–emission artifacts in PET/CT is similar using slow helical CT during breathing, using a rotating rod during breathing, and using cine CT during breathing, all without manual shifting for final optimal coregistration. In our report, this “baseline” frequency of misregistration was corrected by manual coregistration of attenuation and emission scans for the rotating rod (2) and cine CT attenuation data (1). Second, in their protocol, 1 of 3 sequential fast helical CT scans were acquired during breathing without shifting to achieve coregistration. Artifacts were small or, in 85% of cases, absent, and only more severe artifacts were corrected by shift software. Third, the radiation dose for PET/CT is too high and needs to be reduced.

Our paper in *The Journal of Nuclear Medicine* validated a PET/CT protocol that eliminates all misregistration artifacts, thereby providing a definitive, quantitative, standalone noninvasive guide for the management of coronary artery disease. To our knowledge, the paper was the first large, systematic clinical report defining and solving this problem in PET/CT and having significant implications.

The vehemence of their terming their criticisms as “major limitations” is puzzling for several reasons. Basically, Eisner and Patterson agree that attenuation–emission misregistration is a real problem in cardiac PET, a problem not widely addressed clinically until our first reports in *The Journal of Nuclear Medicine* (1,2). Contrary to the emphasis in their letter, the frequency of misregistration artifacts with cine CT attenuation correction (1) without manual shifting should be and is similar to that with the rotating rod (2) because both acquire attenuation data that are averaged over the breathing cycle. The data would be inconsistent otherwise. However, breathing during slow helical CT distorts the attenuation data such that manual shifting to achieve coregistration fails to eliminate the corresponding artifacts, as our data show (1). Despite averaging of attenuation correction during breathing using either a rotating rod or cine CT, misregistration still occurs, requiring manual shifting to optimize coregistration in all patients.

A careful paper from Bacharach’s laboratory (3) on quantitative PET demonstrates that the degree of attenuation varies substantially with respiration even when cardiac borders are coregistered.

Therefore, structures of time-changing attenuation vary in intensity with breathing during emission scans, and this variation can be reproduced only by averaging the attenuation measurements obtained during breathing. Two recent papers (4,5) and a thoughtful editorial (6) published after submission of our reports (1,2) also confirmed the importance of performing attenuation correction on PET/CT to account for varying attenuation with breathing during emission scans.

It is not apparent how multiple-CT fast helical scans acquired during breathing, as proposed by Eisner and Patterson, address this documented variation in attenuation during breathing even if 1 of 3 fast helical CT scans coregisters with the emission images. Paradoxically, their statement indicates that 2 of the 3 fast helical CT scans during breathing cause misregistration, consistent with our data, but that a random 1 of 3 fast CT scans provides correct coregistration in 85% of cases but not all patients. In our cine CT approach, we acquire data at any projection angle 20 times during 2–3 respiratory cycles for effective averaging of time-changing attenuation for all CT slice locations. However, the multiple fast helical CT scans may not produce a predictable result because they can sample moving objects only in a snapshot of time that fails to account for time-changing attenuation.

Two references are quoted in support of their criticisms of our paper and in support of their approach: an abstract from 2004 (7) and another from 2007 (8). Neither of these abstracts was published as a peer-reviewed paper, and no quantitative data have been reported. Finally, their technique of acquiring 3 fast helical CT scans in hopes that one will fit the emission data—a technique that by their own admission fails in 15% of cases—does not seem a good solution from our viewpoint of using cardiac PET as a definitive, standalone guide to the management of coronary artery disease, particularly in the absence of published quantitative clinical data.

Some CT scanners are limited in performing cine CT, as reported by Low et al. (9). In an example cine scan obtained on a 4-slice Siemens VZ, each 0.5-s acquisition is followed by 0.25 s of dead time; a maximum of 7 couch positions (1 cm per couch position) can be programmed; and reprogramming for additional coverage requires approximately 2 min. If the scanner used by Eisner and Patterson is similarly limited in doing cine CT, they need to develop other approaches that address the problem effectively with quantitative published data rather than attacking a rational good solution that works.

We notice an inconsistency between the 0.7-mGy absorbed dose in their letter and the 0.7-mSv effective dose reported in an abstract (8) by the same group. Eisner and Patterson should address this discrepancy before comparing radiation doses between different CT techniques. However, we agree that radiation exposure is excessive in PET/CT, particularly compared with the relatively

negligible exposure from rotating rod attenuation. The estimated risk of cancer from excessive use of CT angiograms causes substantial concern (10), particularly with its limited resolution and technical factors that fail to separate 23% diameter stenosis from 77% diameter stenosis of coronary arteries (11). Cardiac PET needs to avoid this radiation risk. Therefore, rotating rod attenuation for cardiac PET remains an excellent option of diagnostic value comparable to PET/CT. We have also developed protocols for substantially reducing radiation exposure from cine PET/CT. These protocols have been applied to large numbers of patients, and objective quantitative data prove their quantitative accuracy. We anticipate that Eisner and Patterson will provide a definitive clinical study with quantitative data on their proposed approach to attenuation correction in PET/CT that is not available to date.

## REFERENCES

1. Gould KL, Pan T, Loghin C, Johnson NP, Guha A, Sdringola S. Frequent diagnostic errors in cardiac PET/CT due to misregistration of CT attenuation and emission PET images: a definitive analysis of causes, consequences, and corrections. *J Nucl Med.* 2007;48:1112–1121.
2. Loghin C, Sdringola S, Gould KL. Common artifacts in PET myocardial perfusion images due to attenuation-emission misregistration: clinical significance, causes and solutions. *J Nucl Med.* 2004;45:1029–1039.
3. Le Meunier L, Maass-Moreno R, Carrasquillo JA, Diekmann W, Bacharach SL. PET/CT imaging: effect of respiratory motion on apparent myocardial uptake. *J Nucl Cardiol.* 2006;13:821–830.
4. Alessio AM, Kohlmyer S, Branch K, Chen G, Caldwell J, Kinahan P. Cine CT for attenuation correction in cardiac PET/CT. *J Nucl Med.* 2007;48:794–801.
5. Cook RA, Carnes G, Lee Ty, Wells RG. Respiration-averaged CT for attenuation correction in canine cardiac PET/CT. *J Nucl Med.* 2007;48:811–818.
6. Bacharach SL. PET/CT attenuation correction: breathing lessons. *J Nucl Med.* 2007;48:677–679.
7. Brunken RC, DiFilippo FP, Bybel B, Neumann DR, Kaczur T, White RD. Clinical evaluation of cardiac PET attenuation correction using “fast” and “slow” CT images [abstract]. *J Nucl Med.* 2004;45(suppl):120P.
8. Streeter J, Eisner R, Hamill J, Nelson M, Patterson R. Attenuation correction of stress PET Rb-82 with ultrafast CT images [abstract]. *J Nucl Med.* 2007; 48(suppl 2):446P.
9. Low D, Nystrom M, Kalinin P, et al. A method for the reconstruction of four-dimensional synchronized CT scans acquired during free-breathing. *Med Phys.* 2003;30:1254–1263.
10. Einstein AJ, Henzlova MJ, Rajagopalan SR. Estimating risk of cancer associated with radiation exposure from 64-slice computed tomography coronary angiography. *JAMA.* 2007;298:317–323.
11. Gould KL. Assessing progression or regression of CAD: the role of perfusion imaging. *J Nucl Cardiol.* 2005;12:625–638.

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