# Recommendations for reducing radiation exposure in myocardial perfusion imaging

Manuel D. Cerqueira, MD,<sup>a</sup> Kevin C. Allman, MBBS,<sup>b</sup> Edward P. Ficaro, PhD,<sup>c</sup> Christopher L. Hansen, MD,<sup>d</sup> Kenneth J. Nichols, PhD,<sup>e</sup> Randall C. Thompson, MD,<sup>f</sup> William A. Van Decker, MD,<sup>g</sup> and Marko Yakovlevitch, MD<sup>h</sup>

#### INTRODUCTION

Radionuclide myocardial perfusion imaging (MPI) using single photon emission computed tomography (SPECT) or positron emission tomography (PET) for the detection of ischemia in patients with known or suspected coronary artery disease (CAD) has widespread clinical utilization and has been shown to have high accuracy and incremental prognostic value.<sup>1-3</sup> Amidst the recent publicity regarding the increasing use of all types of ionizing radiation in the United States, patients and medical professionals are scrutinizing the need for diagnostic testing and how radiation exposure can be reduced.<sup>4,5</sup> There are three critical questions that physicians must consider and answer with regard to radiation exposure and performing MPI in a particular patient:

- Is MPI testing appropriate and necessary in this patient?
- How can the MPI protocol be optimized to give the lowest possible radiation dose while maintaining diagnostic accuracy?
- How can new technologies be utilized to provide the lowest possible radiation dose while maintaining diagnostic accuracy?

Lowering the radiation dose while maintaining or improving image quality should be considered an improvement in quality of care; the lower the radiation and the higher the image quality, the greater the improvement in the quality of patient care. In general, all MPI studies should be performed in appropriate patients using relatively short-lived radionuclides, and using all possible measures to minimize radiation exposure. Under such circumstances, the benefits of the diagnostic and prognostic information outweigh the risks of radiation exposure.

This document identifies the best practice methods to optimize the benefits of MPI testing by obtaining the highest quality diagnostic images while minimizing radiation exposure. The focus will be on the appropriate selection of patients, the use of protocols that lessen total radiation exposure, and the use of equipment and processing methods that achieve the best image quality at the lowest possible radiation dose.

# **APPROPRIATE PATIENT SELECTION**

Diagnostic radionuclide tracers are distributed over the entire body and are not associated with deterministic radiation effects which include skin burns, cataracts, and permanent sterility. They are related directly to the total dose received, have a latency of weeks to months, and there is a threshold below which effects will not occur. Theoretically, radionuclide tracers add a very small risk due to the stochastic effects of radiation, which are based on probability of chromosomal damage, independent of the dose; there is no threshold for the occurrence of cancer or genetic effects that may take years to decades to develop. Alternative diagnostic techniques such as computed tomography (CT) coronary angiography may expose the patient to a comparable or lower total body radiation dose, but the exposure to critical organs, such as the breast in female patients, is much higher. Since the currently favored model for stochastic effects does not acknowledge a lower radiation threshold, performance of MPI should be governed by the principle of ALARA (As Low As Reasonably Achievable).<sup>6</sup> Patient selection is the initial and most important component of managing radiation exposure

From the Cleveland Clinic,<sup>a</sup> Cleveland, OH; Royal Prince Alfred Hospital,<sup>b</sup> Camperdown, New South Wales, Australia; University of Michigan,<sup>c</sup> Ann Arbor, MI; Jefferson Heart Institute,<sup>d</sup> Philadelphia, PA; Long Island Jewish Medical Center,<sup>c</sup> New Hyde Park, NY; Mid America Heart Institute,<sup>f</sup> Leawood, KS; Temple University Hospital,<sup>g</sup> Philadelphia, PA; and Summit Cardiology,<sup>h</sup> Seattle, WA.

Unless reaffirmed, retired, or amended by express action of the Board of Directors of the American Society of Nuclear Cardiology, this Information Statement shall expire as of July 2015.

Reprint requests: Manuel D. Cerqueira, MD, Cleveland Clinic (Jb3), 9500 Euclid Ave., Cleveland, OH 44195.

J Nucl Cardiol

<sup>1071-3581/\$34.00</sup> 

Copyright @ 2010 by the American Society of Nuclear Cardiology. doi:10.1007/s12350-010-9244-0

from MPI, and is aided by the availability of appropriate use criteria (AUC).<sup>7</sup> When MPI is clinically appropriate, the benefits of accurate diagnosis and management are orders of magnitude greater than the potential risk from the radiation exposure.<sup>8</sup> Therefore, it would increase overall patient risk to avoid MPI or perform less optimal diagnostic testing in order to limit radiation exposure. When MPI is inappropriate, there is no amount of radiation exposure that can be considered acceptable. Inappropriate indications for cardiac radionuclide imaging are listed in Table 1.<sup>7</sup> MPI should be chosen for its ability to meaningfully advance clinical decisionmaking and care.

Once MPI is deemed appropriate, choosing the best diagnostic testing strategy for a patient requires evaluating the characteristics of both the patient and the test. With regard to radiation, the risk is greatest in the youngest patients. The latency period between radiation exposure and any measurable increase in cancer risk is believed to be 10-20 years for most cancers.<sup>9</sup> Accordingly, radiation exposure in younger patients results in a greater lifetime risk of potential radiation-induced cancer than in older patients. Older patients also are less sensitive to the oncologic effects of radiation than are children and young adults. Estimates of lifetime risk of cancer from a specific radiologic examination are four times greater for a 20-year-old woman than for a 50-year-old man.<sup>10</sup> The additional risk of malignancy from exposure to a 10 mSv dose is estimated to be approximately 1 in 2000, for the age distribution of the US population. This estimate includes children and young adults. Since most cardiac patients are older than the average assumed in this calculation, the estimated risk would be much smaller in the MPI population. The mean age of patients undergoing MPI in the United States is in the range of 60-65 years and the risks from their underlying suspected or known CAD are far greater than the theoretical risk from an appropriately ordered MPI study.

The greatest incremental value of diagnostic testing is obtained in the intermediate likelihood patient, and it is for this group of patients that MPI has the greatest utility.<sup>1,2</sup> MPI can also be used to obtain important management and prognostic information in patients, even when the diagnosis is established and the coronary anatomy is known.<sup>3</sup> In addition to establishing a diagnosis and providing prognostic information, MPI can also be of value in demonstrating the absence of disease in certain low-likelihood patients with persistent, unexplained symptoms that may indicate the presence of a life-threatening disease. Clinical judgment, therefore, must be patient specific. However, general principles of utility and appropriateness of diagnostic testing can be applied to clinical scenarios based on evidence. These scenarios have been described<sup>7</sup> and reviewed.<sup>11</sup>

Layered or serial testing should be avoided. Every effort should be made to select the best test based on the unique features of the patient and the specific clinical presentation in order to make a diagnosis or guide management without the need for multiple subsequent "layered" tests. Requests for serial testing in asymptomatic revascularized patients should be carefully scrutinized in light of appropriate use criteria and the likelihood of changing management. When appropriate repeat studies are performed in patients without interval myocardial infarction, repeating the rest study may not be necessary when a prior rest study is available.

#### **Recommendations:**

- Apply appropriate use criteria.
- Consider alternative modalities with comparable diagnostic accuracy without radiation in younger patients.
- Consider utilization in the following patients in whom MPI has the most clinical utility: intermediate CAD risk, those requiring prognostic or management information, and those with persistent and unexplained symptoms.
- Layered or serial testing should be avoided.

# PROTOCOLS, RADIOTRACERS, AND IMAGING SYSTEMS

SPECT and PET MPI can be performed using several different protocols and radionuclide tracers.<sup>12-14</sup> Patient radiation exposure depends on the type and dose of injected radiotracer. Table 2 lists the effective dose estimates using tissue dose coefficients,  $E_1$ , or effective dose coefficients,  $E_2$ , for standard myocardial perfusion imaging protocols. Administered radiation dose may vary considerably based on patient weight and characteristics of the imaging system. There are a number of approaches available to significantly decrease patient exposure while maintaining diagnostic study quality.

# SPECT

In general, Tc-99m-based SPECT protocols (sestamibi and tetrofosmin) offer lower patient radiation exposure than Tl-201 (stress/redistribution and stress/ reinjection) or dual-isotope (Tl-201 rest/Tc-99m stress) protocols.<sup>10</sup> Patient radiation exposure per mCi is similar for the available Tc-99m-based tracers. For the diagnosis of ischemia, Tc-99m-based protocols are preferred.

**Stress-only SPECT imaging.** The use of stressonly imaging to exclude significant myocardial ischemia with Tc-99m-based tracers in appropriately selected

# Table 1. Inappropriate indications (median score 1-3)

Indication	Appropriateness score (1-9)
Detection of CAD: Symptomatic evaluation of ischemic equivalent (non-acute)	
1 Low pretest probability of CAD	I(3)
ECG interpretable AND able to exercise	(-)
Detection of CAD: Symptomatic acute chest pain	
10 Define ACS*	I(1)
Detection of CAD/risk assessment without ischemic equivalent: Asumptomatic	
12 Low CHD Risk (ATP III Risk Criteria)	I(1)
13 Intermediate CHD risk (ATP III Risk Criteria)	I(3)
ECG interpretable	(- )
Detection of CAD/risk assessment without ischemic equivalent: Syncope	
20 Low CHD risk (ATP III Risk Criteria)	I(3)
Risk assessment with prior test results and/or known chronic stable CAD, asymptomatic OR stable	e sumptoms. normal prior
stress imaging study	<b>. . . . . . . . . .</b>
23 Low CHD risk (ATP III Risk Criteria)	I(1)
Last stress imaging study done less than 2 years ago	-(-/
24 Intermediate to high CHD risk (ATP III Risk Criteria)	I(3)
Last stress imaging study done less than 2 years ago	
25 Low CHD risk (ATP III Risk Criteria)	I(3)
Last stress imaging study done more than or equal to 2 years ago	(- )
Risk assessment with prior test results and/or known chronic stable CAD, asumptomatic OR sta	ble sumptoms. abnormal
coronaru anaioaraphu OR abnormal prior stress imaging studu, no prior revascularization	
27 Known CAD on coronary angiography OR prior abnormal stress imaging study	I(3)
Last stress imaging study done less than 2 years ago	-(-)
Risk assessment with prior test results and/or known chronic stable CAD, asymptomatic, prior co	oronaru calcium Aaatston
score	i ontang ballotan in igatoton
33 Agatston score less than 100	I(2)
Risk assessment with prior test results and/or known chronic stable CAD. Duke treadmill score	
37 Low-risk Duke treadmill score	I(2)
Risk assessment: Preoperative evaluation for non-cardiac surgery without active cardiac condition	ns. low-risk suraeru
40 Preoperative evaluation for non-cardiac surgery risk assessment	I(1)
Risk assessment: Preoperative evaluation for non-cardiac suraery without active cardiac condition	ns. intermediate-risk
surgeru	,
41 Moderate to good functional capacity (greater than or equal to 4 METs)	I(3)
42 No clinical risk factors	I(2)
44 Asymptomatic up to 1 year post normal catheterization, non-invasive test, or previous revascularization	l(2)
Risk assessment: Preoperative evaluation for non-cardiac surgery without active cardiac condition	ns. vascular suraeru
45 Moderate to good functional capacity (greater than or equal to 4 METs)	I(3)
46 No clinical risk factors	I(2)
48 Asymptomatic up to 1 year post normal catheterization non-invasive test or previous	I(2)
revascularization	1(2)
KISK assessment: Within 3 months of an acute coronary syndrome, STEMI	1/2)
49 Primary PCI with complete revascularization	I(2)
51 Hemodynamically unstable, signs of cardiogenic shock, or mechanical complications	I(1)
<i>Risk assessment: Within 3 months of an acute coronary syndrome, ACS-asymptomatic post-rev</i> <i>CABG</i> )	ascularization (PCI or
53 Evaluation prior to hospital discharge	I(1)

# Table 1. continued

Indication	Appropriateness score (1-9)
Risk assessment: Within 3 months of an acute coronary syndrome, cardiac rehabilitation	
54 Prior to initiation of cardiac rehabilitation (as a stand-alone indication)	I(3)
Risk assessment: Post-revascularization (PCI or CABG), asymptomatic	
59 Less than 2 years after PCI	I(3)
Risk assessment: Post-revascularization (PCI or CABG), cardiac rehabilitation	
61 Prior to initiation of cardiac rehabilitation (as a stand-alone indication)	I(3)
Evaluation of ventricular function, evaluation of left ventricular function	
65 Routine use of stress FP RNA in conjunction with rest/stress gated SPECT N	1PI I(3)

Reproduced with permission from 2009 Appropriate Use Criteria for Cardiac Radionuclide Imaging.<sup>7</sup>

patients provides the lowest radiation dose for SPECT.<sup>15-18</sup> The accuracy of stress-only studies can be enhanced using attenuation correction. The prognostic value of an unequivocally normal stress-only gated Tc-99m SPECT study performed with attenuation correction and employing careful quality control and experienced readers has been demonstrated to be equivalent to that obtained from a conventional rest/stress study in a large single-center study of 16,854 patients.<sup>19</sup>

Performing stress first, reviewing the images, and if the stress portion is normal, canceling the rest portion, minimizes radiation exposure for one-day studies. Reststress sequences are acceptable. In obese patients requiring two-day studies, the stress study should be performed first, using attenuation correction if available, and the rest study canceled when the stress portion is normal.

**Dual-isotope protocol.** A dual-isotope protocol is most useful when myocardial viability is an overriding clinical consideration in patients with advanced CAD, when there is substantially impaired left ventricular systolic function, and when PET is not available. The patient dose for the detection of myocardial ischemia using this protocol involves a higher patient radiation exposure than is necessary to obtain the diagnostic information by other protocols. The use of PET with a flow tracer and fluorodeoxyglucose (FDG) for viability can achieve a lower patient exposure than existing thallium protocols. However, in patients in whom viability is an important clinical issue, the risk of shortterm cardiovascular mortality is much higher and should be the predominant focus of clinical decision making.

**Dose adjustment for patient weight.** Injected doses for SPECT studies are typically adjusted for patient weight in order to maintain acquired count statistics. In obese patients, the consideration of performing PET rather than SPECT MPI may allow for lower patient radiation exposure with maintained (or even improved) diagnostic performance.<sup>20</sup> In lighterweight patients, reducing the injected dose can still allow adequate counts statistics while reducing exposure.

**Reduction of injected activity with new SPECT camera technologies.** SPECT studies performed with higher sensitivity solid-state cameras with high sensitivity crystal material and utilizing advanced reconstruction algorithms permit lower doses of injected activity for diagnostic studies compared with Anger gamma cameras using sodium iodide crystals.<sup>21</sup> Dose reductions on the order of 50% compared with conventional doses appear to be feasible.

# PET

For PET MPI, the tracers currently in use are Rb-82 and N-13 ammonia, in decreasing frequency of clinical utilization. Protocols most commonly involve both rest and stress imaging on a single day. Injected dose and resulting patient radiation dose depends in part on the system sensitivity and mode of operation (two-dimensional [2D] vs three-dimensional [3D]). In general, the newer lutetium oxyothosilicate (LSO) systems will require less injected activity than older bismuth germinate (BGO)-based systems. Systems operating in 3D acquisition mode require lower injected activity than those acquiring in 2D mode.

Because of their short half lives, PET studies potentially offer a lower patient radiation dose compared to SPECT, and may result in more favorable patient dosimetry, especially in obese patients. The issue of stress-only PET MPI has not been widely investigated but is worthy of consideration.

**Attenuation correction.** For both PET and SPECT MPI, the use of either radionuclide or CT-based transmission scans for attenuation correction contributes a negligible component of the radiation exposure received by the patient when the dose from the injected tracer is considered. The use of attenuation correction is

			<b>Effective doses (mSv)</b>			
	Injected activity (mCi)		From ICRP tables		From manufacturers' PIs	
Protocol	Rest	Stress	<b>E</b> <sub>1</sub>	E <sub>2</sub>	<b>E</b> <sub>1</sub>	E <sub>2</sub>
<sup>99m</sup> Tc Sestamibi rest-stress	10.0	27.5	11.3	11.4	14.6	NR
<sup>99m</sup> Tc Sestamibi stress only	0.0	27.5	7.9	8.0	10.0	NR
<sup>99m</sup> Tc Sestamibi two day	25.0	25.0	15.7	15.6	20.6	NR
<sup>99m</sup> Tc Tetrofosmin rest-stress	10.0	27.5	9.3	9.9	9.7	12.9
99mTc Tetrofosmin stress only	0.0	27.5	6.6	7.1	6.7	8.8
<sup>99m</sup> Tc Tetrofosmin two day	25.0	25.0	12.8	13.5	13.7	18.3
<sup>201</sup> Tl stress-redistribution	0.0	3.5	22.0	22.0	28.7, 9.3, 28.4	46.6, NR, 46.6
<sup>201</sup> Tl stress-reinjection	1.5	3.0	31.4	31.5	43, 14.0, 42.6	69.9, NR, 69.9
Dual isotope <sup>201</sup> Tl- <sup>99m</sup> Tc Sestamibi	3.5	25.0	29.2	29.3	37.8, 18.4, 37.5	NR, NR, NR
<sup>99m</sup> Tc labeled erythrocytes	22.5	0.0	5.7	5.8	2.3	NR
<sup>82</sup> Rb	50.0	50.0	13.5	12.6	3.0	NR
<sup>13</sup> N-ammonia	15.0	15.0	2.4	2.2	n/a	n/a
<sup>15</sup> O-water*	29.7	29.7	2.5	2.4	n/a	n/a
<sup>18</sup> F-FDG	10.0	0.0	7.0	7.0	n/a	n/a

# Table 2. Estimates of effective doses of standard myocardial perfusion imaging protocols

\* American Society of Nuclear Cardiology guidelines do not prescribe a recommended dose. Stress and rest doses of 1100 MBq (29.7 mCi) used, as per European Association of Nuclear Medicine/European Society of Cardiology guidelines. *PI*, Package Insert (or product information).

 $E_1$  = Effective dose estimated from tissue dose coefficients, using ICRP Publication 60 tissue weighting factors. Calculations performed using the "splitting rule," arithmetic averaging rather than mass averaging of individual remainder organ dose contributions, and upper large intestine rather than extrathoracic airways as a remainder organ, as was originally specified in ICRP Publication 60. If dose to the colon was not specified in a data source, then the average of the upper large intestine and lower large intestine doses was substituted.

 $E_2$  = Effective dose estimated from whole-body dose coefficients, using ICRP Publication 60 tissue weighting factors.

*NR*, Not Reported in PI (Total body dose provided rather than effective dose); n/a, Not available for cyclotron-produced tracers. Reproduced with permission from Estimates of Effective Doses of Standard Myocardial Perfusion Imaging Protocols.<sup>10</sup>

desirable, especially when stress-only imaging is performed.<sup>18,22</sup>

#### **Recommendations:**

The clinical indications and physical stature of each patient should be reviewed and the best combination of radiotracers and protocols selected using the following guidelines:

- Use radionuclides with shorter half-life such as Tc-99m and PET tracers.
- Perform stress-only testing.
- Use weight-based dosing.

#### TECHNOLOGY

Once the appropriate patient, radiotracer, and protocol have been selected, technological considerations can further reduce radiation exposure. The various hardware and software options are discussed below, and the recommendations that can be made based on the existing data are listed in Table 3.

#### **Image Reconstruction**

Filtered back projection (FBP) has been the standard method for reconstructing SPECT MPI. FBP has problems with low-count SPECT images due to the inevitable amplification of image noise, and mandates higher radiation doses or longer acquisition times. It is unable to correct for photon attenuation and scatter.<sup>23,24</sup>

Iterative reconstruction (IR) refers to a broad category of SPECT and PET reconstruction techniques that estimate the distribution of the radioactivity being imaged by mathematically generating projections that such a distribution would produce.<sup>23,24</sup> The generated projections are compared to the actual acquired projections and the difference is used to improve the estimate of the estimated distribution. If the algorithm is designed correctly, the estimate approaches the actual distribution being imaged and the difference is minimized. Using IR allows the injection of a lower radiation dose and correction for the Poisson nature of the noise as well as attenuation and scatter.

Feature	Potential for dose reduction	Recommendation
Patient selection	Significant	Apply appropriate use criteria. Consider alternative modalities with comparable diagnostic accuracy without radiation in younger patients. Consider utilization in the following patients in whom MPI has the most clinical utility: intermediate CAD risk, those requiring prognostic or management information, and those with persistent and unexplained symptoms. Layered or serial testing should be avoided
Protocols, radiotracers and imaging systems	Significant	The clinical indications and physical stature of each patient should be reviewed and the best combination of radiotracers and protocols selected using the following guidelines: use radionuclides with shorter half-life such as Tc-99m and PET tracers, perform stress-only testing, and use weight-based dosing
Reconstruction-FBP	Standard	No recommendation
Reconstruction- iterative	Potential for significant	Strongly recommend
Multi-detector systems	Significant	Strongly recommend minimum of two detectors
New camera geometries	Significant (same effect as multi-detector systems)	Use when available. Consider for new equipment purchase
Solid-state detector systems	Minor unless part of a multi- detector or new geometry system.	No recommendation
Collimators-custom	Unproven, probably minor	Further exploration and research
Energy settings	Probably minor	Further exploration and research
Step and shoot	Minor	No recommendation
Count consistency	Minor	No recommendation

#### Table 3. Recommendations for achieving MPI radiation reduction

Clinical acceptance of IR had been limited due to the computationally intensive nature of the algorithms and resultant slow reconstruction times. With the increased computation speed currently available, IR is now practical for clinical use. Studies have been performed to identify the potential lower bounds of the required count density that is needed with IR techniques compared to the established protocols using FBP. The results have shown that with IR techniques, image quality comparable to FBP can be obtained with as little as 50% to 75% of the counts; the degree of reduction is dependent on the software vendor's implementation of the algorithm.<sup>18,25-27</sup> Since the detected count density for a SPECT study is dependent on the injected radioactivity. IR techniques hold the promise of significantly reducing administered dose and patient radiation exposure. PET protocols routinely use IR with attenuation and scatter correction.

IR techniques are very dependent on the manufacturer, class of algorithm, the geometry of the imaging system, and other models incorporated into the reconstruction (e.g., photon attenuation). As a result, not all algorithms will provide the same results for the same acquisition protocol. It is advised to follow the manufacturer's recommendations and review the results and protocols that clinically validated the technology.

#### **Recommendation:**

• IR should be used for SPECT and PET MPI processing.

# **Imaging Systems**

The majority of imaging laboratories perform SPECT using conventional Anger cameras with NaI(Tl) crystals

and parallel-hole collimators with protocols optimized to acquire high-quality SPECT MPI with the minimum radiation dose.<sup>28</sup> The following suggestions minimize injected radiation dose, and therefore deliver less radiation to the patient, while maintaining high image quality.

Multiple detector systems. Multi-detector imaging systems are able to obtain high-quality diagnostic images using lower radionuclide doses than a single-detector system. Two-detector systems are widely available and enable a routine reduction by half of the injected dose while imaging for the same time as a onedetector system. Since 180-degree acquisition is preferable with FBP and comparable results are obtained with IR using 180-degree and 360-degree acquisitions (with correction for patient attenuation, detector response characteristics, and radiation scatter), 180degree acquisition is recommended.<sup>29</sup> Some of the newer, unconventional radiation detector designs, especially those that employ arrays of small solid state detectors,<sup>30</sup> have the equivalent of many more "heads" than two-detector or three-detector gamma cameras. The potential for these newer SPECT systems to reduce radiation dose are discussed below.

**Solid-state detector-based SPECT systems.** New technology based on innovative detector designs using semiconductors have greater count sensitivity and the potential for reducing radiation dose relative to conventional anger cameras.<sup>30</sup>

Improved sensitivity. New imaging geometries using solid-state detectors offer increased sensitivity compared to conventional parallel collimated Anger cameras. While conventional Anger cameras have count sensitivities on the order of 0.5-0.7 kcps, new SPECT system geometries using cadmium-zinc-telluride (CZT) have sensitivities of 2.2-4.7 kcps.<sup>31,32</sup> Some detector designs have reported a 10 fold greater count sensitivity than Anger cameras, without a reduction in energy or spatial resolution.<sup>33</sup> Anger cameras have 9-10% Tc-99m energy resolution while CZT-based SPECT cameras have energy resolution of 5.7%. Similarly, Anger cameras have system spatial resolution on the order of 9-10 mm, while CZT SPECT systems have system spatial resolution of 4-5 mm. Consequently, it is feasible to acquire high-quality MPI images with solid-state detector-based SPECT systems using greatly reduced injected radioactivity.31,34,35

*Overall image quality*. Initial reports indicated the image quality achieved with CZT cameras in 5-6 min is comparable to or better than conventional parallel hole collimated Anger camera acquisitions of 14-15 min.<sup>36,37</sup> A subsequent analysis of 168 patients showed that CZT studies with 4-minute rest and 2-minute stress CZT acquisitions produced image quality comparable to that of 14-minute rest and 12-minute stress Anger camera

studies.<sup>38</sup> CZT imaging devices can acquire images of similar quality as Anger cameras in 12-14 minutes studies using only 15%-30% of the current standard injected isotope dose. In clinical trials the newer detectors produce images that are at least comparable to those obtained with conventional SPECT cameras,<sup>21,37,39,40</sup> so that significant radiation dose reduction is feasible.

**Collimation.** Conventional collimators. Most cardiac Anger cameras use low-energy high resolution parallel-hole collimators with a collimator resolution of about 7 mm at 10 cm from the collimator surface. "General purpose" collimators typically have  $2\times$  higher sensitivity, but a reduced collimator spatial resolution of 9 mm. "High sensitivity" collimators have  $4\times$  higher sensitivity than "high resolution" collimators, but a collimator spatial resolution of only 13 mm. Incorporating the intrinsic camera resolution, going from a "high resolution" to a "general purpose" collimator acquisition would allow a reduction of the injected dose by 50% while achieving the same number of counts, but the system spatial resolution would decline from 8-8.5 to 9.5-10 mm.<sup>28</sup>

Custom-designed collimators. An alternative approach is to use specially designed focusing collimators for cardiac imaging. The geometry of the collimator holes and septa are not parallel, but are variably aligned across the face of the collimator to achieve higher spatial resolution and higher sensitivity in the area of the heart. Special reconstruction software is required. Because of their higher extrinsic sensitivity compared to conventional parallel-hole collimators, the injected dose can be reduced and quality images obtained using standard imaging times. Pin-hole collimators have the greatest spatial resolution, and a solid-state based detector has been constructed that uses multiple pin-hole collimators, as described below. Other specialized collimator designs include multiple simultaneous pin-hole collimators that are used in conjunction with standard Anger camera technology, which also require special reconstruction software.

Considerable development over the past several years has been devoted to creating sophisticated IR algorithms that incorporate physical details of the collimators. These algorithms are designed to account for their different spatial resolution with depth.<sup>34,41</sup>

#### **Recommendations:**

- Anger camera MPI studies should be performed using a minimum of 2 detectors with 180-degree acquisition.
- High sensitivity SPECT imaging geometries should be used whenever available and considered at the time of new equipment purchase.
- High sensitivity and custom designed collimators have the potential for reducing injected radiation

dose .but require further validation before implementation into clinical practice.

# **Imaging System Acquisition Parameters**

**Energy settings, energy resolution.** One method for increasing the total number of counts from a lower injected radiation dose is to widen the energy window. This results in a larger total number of acquired counts, but also increases the percentage of scattered gamma rays, which will reduce spatial resolution and reduce image contrast. For institutions that routinely employ scatter corrections as part of IR,<sup>42,43</sup> this may have a less deleterious effect than for those that use FBP. If the attempt is made to obtain acceptable myocardial perfusion studies by widening the energy window, institutions that have more than one Anger camera should use the one with the best intrinsic energy resolution, in order to minimize the blurring effect of including a larger percentage of scattered radiation.

When using Tl-201, acquiring the 167 keV peak increases counts by about 5%, which can be used to reduce the injected activity. The higher energy Tl-201 gamma rays improve the spatial resolution compared to the lower energy Tl-201 photons.

**Continuous acquisition.** Continuous acquisition is an alternative to conventionally used step-andshoot protocols where the detectors are not acquiring counts while moving from one projection to the next. Continuous acquisition protocols acquire additional data for only a few seconds per projection.<sup>44</sup> Step-and-shoot protocols are used instead of continuous acquisition, because reconstruction algorithms are designed for input data that are assumed to be a sequence of images acquired at discrete angles.<sup>45</sup> However, using reconstruction software designed to accommodate true continuous acquisitions allows a reduced injection dose while obtaining the same target count rate. The reduction is on the order of 5%.

**Count consistency.** One means to reduce total radiation dose for sites that use multiple injections of the same isotope involves obtaining consistent counts for the rest and stress portions of the study. By performing a one-minute anterior-view myocardial image for the initial injection, it is possible to adjust the amount of time per projection in order to obtain a target information density.<sup>46</sup> This approach could be extended to enable acquiring the initial tomogram for a standardized initial total number of myocardial counts and then, for the second injection, to reduce the injected activity so as to acquire the second tomogram for a longer period of time that is still tolerable for the patient. This would be most feasible for two-day protocols, and would be more likely useful in imaging smaller patients rather than larger patients.

#### **Recommendations:**

- Widening the energy window requires further validation before implementation into clinical practice.
- Continuous acquisition, and count consistency methods require further validation before implementation into clinical practice.



Figure 1. Proposed algorithm for maximal reduction in patient radiation exposure.

# SUMMARY

Figure 1 illustrates the patient flow resulting from the above discussion. It is consistent with the concept of lowering the radiation dose while maintaining or improving image quality which should be considered an improvement in quality of care. The lower the radiation and the higher the image quality the greater the improvement in the quality of patient care. This approach combines the radiation reduction factors, which if implemented correctly, will provide the highest quality MPI studies at the lowest possible radiation dose. It must be recognized that such an approach requires moving away from the comfortable, familiar, time-tested, and efficient protocols adapted in most nuclear cardiology laboratories. It requires patient assessment at the time of scheduling, flexibility in patient flow as each patient will potentially go through a different protocol, and ultimately modifying existing imaging systems or purchasing new equipment.

Based on these recommendations, we expect that for the population of patients referred for SPECT or PET MPI, on average a total radiation exposure of  $\leq 9$  mSv can be achieved in 50% of studies by 2014.

# Acknowledgments

Dr. Nichols serves as a consultant for IEAE and receives royalties from Syntermed, Inc. The authors have no conflicts of interest to disclose except as noted above.

#### References

- Berman DS, Hachamovitch R, Kiat H, et al. Incremental value of prognostic testing in patients with known or suspected ischemic heart disease: A basis for optimal utilization of exercise technetium-99m sestamibi myocardial perfusion single-photon emission computed tomography. J Am Coll Cardiol 1995;26: 639-47.
- Giri S, Shaw LJ, Murthy DR, et al. Impact of diabetes on the risk stratification using stress single-photon emission computed tomography myocardial perfusion imaging in patients with symptoms suggestive of coronary artery disease. Circulation 2002;105:32-40.
- Iskandrian AS, Chae SC, Heo J, et al. Independent and incremental prognostic value of exercise single-photon emission computed tomographic (SPECT) thallium imaging in coronary artery disease. J Am Coll Cardiol 1993;22:665-70.
- Beller GA. Importance of consideration of radiation doses from cardiac imaging procedures and risks of cancer. J Nucl Cardiol 2010;17:1-3.
- Cohen MC. Radiation reduction: How long can you go? J Nucl Cardiol 2010;17. doi:10.1007/s12350-010-9212-8.
- United States Nuclear Regulatory Commission. Definitions. 10 CFR § 20.1003. http://www.nrc.gov/reading-rm/doc-collections/cfr/ part020/part020-1003.html. Revised December 1, 2009. Effective May 21, 1991. Accessed April 1, 2010.

- Hendel RC, Berman DS, Di Carli MF, et al. 2009 appopriate use criteria for cardiac radionuclide imaging. J Am Coll Cardiol 2009;53:2201-29.
- Committee to Assess Health Risks from Exposure to Low Levels of Ionizing Radiation, National Research Council. Health Risks from Exposure to Low Levels of Ionizing Radiation: BEIR VII Phase 2. Washington, DC: National Academies Press; 2006.
- National Council on Radiation Protection and Measurement. Comparative carcinogenicity of ionizing radiation and chemicals. Report No. 096. Washington, DC: National Council on Radiation Protection and Measurement, 1989.
- Einstein AJ, Moser KW, Thompson RC, Cerqueira MD, Henzlova MJ. Radiation dose to patients from cardiac diagnostic imaging. Circulation 2007;116:1290-305.
- Ward RP, Al-Mallah MH, Grossman GB. American Society of Nuclear Cardiology review of the ACCF/ASNC appropriateness criteria for single-photon emission computed tomography myocardial perfusion imaging (SPECT MPI). J Nucl Cardiol 2007;14: e26-38.
- Henzlova MJ, Cerqueira MD, Hansen CL, Taillefer R, Yao S. Imaging guidelines for nuclear cardiology procedures: Stress protocols and tracers. J Nucl Cardiol 2009;16. doi:10.1007/s12350-009-9062-4.
- Hansen CL, Goldstein RA, Akinboboye OO, et al. Imaging guidelines for nuclear cardiology procedures: Myocardial perfusion and function: Single photon emission computed tomography. J Nucl Cardiol 2007;14:e39-60.
- Dilsizian V, Bacharach SL, Beanlands RS, Bergmann SR, Delbeke D, Gropler RJ, et al. Imaging guidelines for nuclear cardiology procedures: PET myocardial perfusion and metabolism clinical imaging. J Nucl Cardiol 2009;16. doi:10.1007/s12350-009-9094-9.
- 15. Gibson PB, Demus D, Noto R, Hudson W, Johnson LL. Low event rate for stress-only perfusion imaging in patients evaluated for chest pain. J Am Coll Cardiol 2002;39:999-1004.
- Santana CA, Garcia EV, Vansant JP, et al. Gated stress-only 99mTc myocardial perfusion SPECT imaging accurately assesses coronary artery disease. Nucl Med Commun 2003;24:241-9.
- Heller GV, Bateman TM, Johnson LL, et al. Clinical value of attenuation correction in stress-only Tc-99m sestamibi SPECT imaging. J Nucl Cardiol 2004;11:273-81.
- Bateman T, Heller GV, McGhie AI, et al. Multicenter investigation comparing a highly efficient half-time stress-only attenuation correction approach against standard rest-stress Tc-99m SPECT imaging. J Nucl Cardiol 2009;16:726-35.
- Chang SM, Nabi F, Xu J, Raza U, Mahmarian JJ. Normal stressonly versus standard stress/rest myocardial perfusion imaging: Similar patient mortality with reduced radiation exposure. J Am Coll Cardiol 2010;55:221-30.
- Bateman TM, Friedman JD, Heller GV, et al. Diagnostic accuracy of rest/stress ECG-gated Rb-82 myocardial perfusion PET: Comparison with ECG-gated Tc-99m sestamibi SPECT. J Nucl Cardiol 2006;13:24-33.
- Berman DS, Kang X, Tamarappoo B, et al. Stress thallium-201/ rest technetium-99m sequential dual isotope high-speed myocardial perfusion imaging. J Am Coll Cardiol Imaging 2009;2:273-82.
- 22. Chang H, George RT, Schuleri KH, et al. Prospective electrocardiogram-gated delayed enhanced multidetector computed tomography accurately quantifies infarct size and reduces radiation exposure. J Am Coll Cardiol Imaging 2009;2:412-20.
- Hansen C. Digital image processing for clinicians, part III: SPECT reconstruction. J Nucl Cardiol 2002;9:542-9.
- Kak A, Slaney M. Algorithms for reconstruction with nondiffracting sources. In: Kak A, Slaney M, editors. Principals of computerized tomographic imaging. New York: IEEE; 1999.

- 25. DePuey EG, Gadiraju R, Clark J, et al. Ordered subset expectation maximization and wide beam reconstruction "half-time" gated myocardial perfusion SPECT functional imaging: A comparison to "full-time" filtered backprojection. J Nucl Cardiol 2008;14:547-63.
- DePuey EG, Bommireddipalli S, Clark J, Thompson L, Srour Y. Wide beam reconstruction "quarter-time" gated myocardial perfusion SPECT functional imaging: A comparison to "full-time" ordered subset expectation maximum. J Nucl Cardiol 2009;16: 736-52.
- 27. Venero CV, Heller GV, Bateman TM, et al. A multicenter evaluation of a new postprocessing method with depth-dependent collimator resolution applied to full-time and half-time acquisitions without and with simultaneously acquired attenuation correction. J Nucl Cardiol 2009;16:714-25.
- Cherry SR, Sorenson JA, Phelps ME. Physics in nuclear medicine. 3rd ed. Philadelphia, PA: Elsevier Science; 2003.
- He X, Links JM, Gilland KL, Tsui BMW, Frey EC. Comparison of 180° and 360° acquisition for myocardial perfusion SPECT with compensation for attenuation, detector response, and scatter: Monte Carlo and mathematical observer results. J Nucl Cardiol 2006;13:345-53.
- Slomka PJ, Patton JA, Berman DS, Germano G. Advances in technical aspects of myocardial perfusion SPECT imaging. J Nucl Cardiol 2009;16:255-76.
- Garcia EV, Tsukerman L, Keidar Z. A new solid state, ultra fast cardiac multi-detector SPECT system [abstract]. J Nucl Cardiol 2008;15:S3.
- Kennedy JA, Yosilevsky G, Przewloka K, Israel O, Frenkel A. 3D spatial resolution map and sensitivity characterization of a dedicated cardiac CZT SPECT camera [abstract]. J Nucl Med 2009; 50:107P.
- Gambhir SS, Berman DS, Ziffer JA, et al. A novel high-sensitivity rapid-acquisition single-photon cardiac imaging camera. J Nucl Med 2009;50:635-43.
- Marie PY, Djaballah W, Franken PR, et al. OSEM reconstruction, associated with temporal Fourier and depth-dependant resolution recovery filtering, enhances results from sestamibi and 201Tl 16-interval gated SPECT. J Nucl Med 2005;46:1789-95.
- 35. Sharir T, Slomka PJ, Hayes S, et al. Multicenter trial of high-speed versus conventional single-photon emission computed tomography

imaging: Quantitative results of myocardial perfusion and left ventricular function. J Am Coll Cardiol 2010;55:1965-74.

- Ben-Haim S, Van Gramberg D, Bomanji J, et al. Thallium 201 (Tl) myocardial perfusion imaging (MPI) with a novel fast cardiac camera versus conventional camera (SPECT) [abstract]. J Nucl Med 2009;50:124P.
- Keidar Z, Kagna O, Frenkel A, Israel O. A novel ultrafast cardiac scanner for myocardial perfusion imaging (MPI): Comparison with a standard dual-head camera [abstract]. J Nucl Med 2009;50:125P.
- Esteves FP, Raggi P, Folks RD, et al. Novel solid-state-detector dedicated cardiac camera for fast myocardial perfusion imaging: Multicenter comparison with standard dual detector cameras. J Nucl Cardiol 2009;16:927-34.
- Sharir T, Ben-Haim S, Merzon K, et al. High-speed myocardial perfusion imaging initial clinical comparison with conventional dual detector anger camera imaging. J Am Coll Cardiol Imaging 2008;1:156-63.
- Maddahi J, Mendez R, Mahmarian JJ, et al. Prospective multicenter evaluation of rapid, gated SPECT myocardial perfusion upright imaging. J Nucl Cardiol 2009;16:351-7.
- Daou D, Pointurier I, Coaguila C, et al. Performance of OSEM and depth-dependent resolution recovery algorithms for the evaluation of global left ventricular function in 201Tl gated myocardial perfusion SPECT. J Nucl Med 2003;44:155-62.
- Bowsher JE, Floyd CE. Treatment of Compton scattering in maximum likelihood, expectation-maximization reconstructions of SPECT images. J Nucl Med 1991;32:1285-91.
- Xaio J, de Wit TC, Staelen SG, Beekman FJ. Evaluation of 3D Monte Carlo-based scatter correction for 99mTc cardiac perfusion SPECT. J Nucl Med 2006;47:1662-9.
- Cao ZJ, Maunouroy C, Chen CC, et al. Comparison of continuous step-and-shoot versus step-and-shoot acquisition SPECT. J Nucl Med 1996;37:2037-40.
- Bieszk JA, Hawman EG. Evaluation of SPECT angular sampling effects: Continuous versus step-and-shoot acquisition. J Nucl Med 1987;28:1308-14.
- 46. Case J, Bateman T, Cullom S. Obtaining optimum and consistent SPEW myocardial counts using an anterior planar view to determine SPECT acquisition times [abstract]. J Am Coll Cardiol 1998;31:82A.