

CT perfusion angiography; beware of artifacts!

E. E. van der Wall · J. D. Schuijf · J. J. Bax ·
J. W. Jukema · M. J. Schalij

Received: 1 December 2009 / Accepted: 4 December 2009 / Published online: 24 December 2009
© The Author(s) 2009. This article is published with open access at Springerlink.com

Over the past years, myocardial perfusion imaging has been established as the reference standard for prognosis and clinical decision making of patients with coronary artery disease (CAD). Myocardial perfusion has predominantly been assessed by single photon emission tomography (SPECT) [1–7] and, more recently, by positron emission tomography (PET) and magnetic resonance imaging (MRI) [8–16].

Amongst the advanced cardiac imaging modalities, multi detector computed tomography (CT) angiography has emerged as a reliable non-invasive method for the assessment of coronary anatomy, CAD, and cardiac function [17–22]. Multiple studies involving over several thousands of patients have established that CT angiography is highly accurate for delineation of the presence and severity of coronary atherosclerosis [23–30]. CT angiography may also reveal the total plaque burden, i.e. both calcified and non-calcified components, for individual patients with coronary atherosclerosis [31–40].

The advent of prospectively gated acquisition techniques for 64-slice CT angiography has allowed

a significant reduction in dose exposure. Consequently, a combined approach of CT coronary angiography and myocardial perfusion imaging with CT angiography might potentially become feasible at a total radiation dose of less than 10 mSv, particularly for the assessment of patients with established CAD, who are likely to have diffuse calcification [40–47].

Since myocardial perfusion by CT angiography is based on myocardial signal density, it is crucial to determine the normal values of myocardial signal density and to identify potential mechanisms of misinterpretation of perfusion defects. In routine CT angiography acquisitions, there might be a considerable signal density drop at the posterobasal wall resembling perfusion defects possibly being attributed to beam hardening artifacts.

In the current issue of the *International Journal of Cardiovascular Imaging* Rodríguez-Granillo et al. [48] investigated normal myocardial signal density levels during CT angiography and evaluated the impact of artifacts due to beam hardening. A group of 36 consecutive asymptomatic patients with a low probability of CAD were referred for CT angiography because of inconclusive or discordant functional tests. Perfusion defects were defined as a myocardial segment having a signal density two standard deviations below the average myocardial signal density for the 16 left ventricular American Heart Association (AHA) segments. Signal density was evaluated in 576 American Heart Association (AHA) segments

Editorial comment on the article of Rodríguez-Granillo et al.
(doi: 10.1007/s10554-009-9531-5).

E. E. van der Wall (✉) · J. D. Schuijf ·
J. J. Bax · J. W. Jukema · M. J. Schalij
Department of Cardiology, Leiden University Medical
Center, P.O. Box 9600, Leiden, The Netherlands
e-mail: e.e.van_der_wall@lumc.nl

and 36 posterobasal segments. The mean myocardial signal density at the posterobasal segment was 53.5 ± 35.1 Hounsfield Units, whereas the mean myocardial signal density at the basal, mid and apical myocardium was 97.4 ± 17.3 Hounsfield Units, with significant differences between posterobasal and all AHA segments. Posterobasal perfusion defects were identified in 26 (72%) patients. The only variable associated to the presence of posterobasal perfusion defect was heart rate, whereas body mass index, blood signal density of the left and right ventricles, contrast-to-noise ratio, and the extent of atherosclerosis were not related to the presence of perfusion defects. The main findings of the study were that (1) beam hardening artifacts are a common finding at CT angiography of asymptomatic patients affecting predominantly the posterobasal wall, (2) perfusion defects at the short axis plane including the mitral valve and the left ventricular outflow tract are not related to technical issues whereas heart rate may be associated with this finding, (3) perfusion defects can be also identified at the inferior and anteroapical segments but this occurs less often.

Occurrence of attenuation artifacts during radionuclide SPECT perfusion imaging has been considered an important limitation of the technique [49–51]. Apical thinning due to the overlying diaphragm and the occurrence of anteroapical defects as a result of breast attenuation are very common causes for unwanted perfusion deficits, leading to image misinterpretation and potentially a wrong diagnosis. The present study [48] indicates that perfusion artifacts also occur at CT perfusion angiography and could therefore be a reason for misinterpretation. These perfusion defects can largely be ascribed to beam hardening artifacts most likely from the spine for posterobasal segments and from the sternum for anteroapical segments.

In radionuclide myocardial perfusion SPECT imaging, successful attenuation correction programs have been developed in order to discriminate between true and false perfusion defects [52–55]. Similarly, correction algorithms are currently being designed for CT angiography with the same purpose. Recently, So et al. [56] designed phantoms to simulate the beam hardening artifacts encountered in cardiac CT perfusion studies of humans and animals. These phantoms were used to investigate whether beam hardening artifacts could be reduced with this approach and to

determine the optimal settings of the correction algorithm for patient and animal studies, which depend upon the anatomy of the scanned subject. The correction algorithm was also applied to correct beam hardening in a clinical study to further demonstrate the effectiveness of this technique.

To summarize, the study by Rodríguez-Granillo et al. [48] convincingly shows that in an asymptomatic population with no history of CAD, who undergo CT perfusion angiography, artifacts in the posterobasal wall are a common finding in more than two-thirds of patients. This phenomenon of pseudo-perfusion defects may considerably affect proper image interpretation and should be taken into account in the judgment of CT perfusion images. In the near future, correction algorithms for CT perfusion angiography will assist in identifying the true nature of the defects in order to establish the correct diagnosis.

Open Access This article is distributed under the terms of the Creative Commons Attribution Noncommercial License which permits any noncommercial use, distribution, and reproduction in any medium, provided the original author(s) and source are credited.

References

1. Bavelaar-Croon CD, Pauwels EK, van der Wall EE (2001) Gated single-photon emission computed tomographic myocardial imaging: a new tool in clinical cardiology. *Am Heart J* 141:383–390
2. van der Wall EE, Heidendal GA, den Hollander W, Westera G, Roos JP (1980) I-123 labeled hexadecenoic acid in comparison with thallium-201 for myocardial imaging in coronary heart disease. A preliminary study. *Eur J Nucl Med* 5:401–405
3. Bax JJ, Lamb H, Dibbets P, Pelikan H et al (2000) Comparison of gated single-photon emission computed tomography with magnetic resonance imaging for evaluation of left ventricular function in ischemic cardiomyopathy. *Am J Cardiol* 86:1299–1305
4. Chamuleau SA, van Eck-Smit BL, Meuwissen M et al (2007) Long-term prognostic value of CFVR and FFR versus perfusion scintigraphy in patients with multivessel disease. *Neth Heart J* 15:369–374
5. Bavelaar-Croon CD, Kayser HW, van der Wall EE et al (2000) Left ventricular function: correlation of quantitative gated SPECT and MR imaging over a wide range of values. *Radiology* 217:572–575
6. van der Wall EE, den Hollander W, Heidendal GA, Westera G, Majid PA, Roos JP (1981) Dynamic myocardial scintigraphy with I-123I-labeled free fatty acids in patients with myocardial infarction. *Eur J Nucl Med* 6:383–389

7. Kurvers MJ, Braam RL, Verzijlbergen JF, Heestermaas AA, Ten Berg JM (2007) Myocardial salvage in STEMI patients treated with primary coronary angioplasty as demonstrated by myocardial SPECT. *Neth Heart J* 15:422–423
8. van der Wall EE, van Dijkman PR, de Roos A et al (1990) Diagnostic significance of gadolinium-DTPA (diethylenetriamine penta-acetic acid) enhanced magnetic resonance imaging in thrombolytic treatment for acute myocardial infarction: its potential in assessing reperfusion. *Br Heart J* 63:12–17
9. van Dijkman PR, van der Wall EE, de Roos A et al (1991) Acute, subacute, and chronic myocardial infarction: quantitative analysis of gadolinium-enhanced MR images. *Radiology* 180:147–151
10. de Roos A, Matheijssen NA, Doornbos J, van Dijkman PR, van Voorthuisen AE, van der Wall EE (1990) Myocardial infarct size after reperfusion therapy: assessment with Gd-DTPA-enhanced MR imaging. *Radiology* 176:517–521
11. van de Wal RM, van Werkum JW, le Cocq d'Armandville MC et al (2007) Giant aneurysm of an aortocoronary venous bypass graft compressing the right ventricle. *Neth Heart J* 15:252–254
12. Niezen RA, Helbing WA, van der Wall EE, van der Geest RJ, Rebergen SA, de Roos A (1996) Biventricular systolic function and mass studied with MR imaging in children with pulmonary regurgitation after repair for tetralogy of Fallot. *Radiology* 201:135–140
13. Pluim BM, Lamb HJ, Kayser HW et al (1998) Functional and metabolic evaluation of the athlete's heart by magnetic resonance imaging and dobutamine stress magnetic resonance spectroscopy. *Circulation* 97:666–672
14. Vliegen HW, Doornbos J, de Roos A, Jukema JW, Bekendam MA, van der Wall EE (1997) Value of fast gradient echo magnetic resonance angiography as an adjunct to coronary arteriography in detecting and confirming the course of clinically significant coronary artery anomalies. *Am J Cardiol* 79:773–776
15. Hoogendoorn LI, Pattynama PM, Buis B, van der Geest RJ, van der Wall EE, de Roos A (1995) Noninvasive evaluation of aortocoronary bypass grafts with magnetic resonance flow mapping. *Am J Cardiol* 75:845–848
16. Holman ER, Buller VG, de Roos A et al (1997) Detection and quantification of dysfunctional myocardium by magnetic resonance imaging. A new three-dimensional method for quantitative wall-thickening analysis. *Circulation* 95:924–931
17. Schuijff JD, Bax JJ, van der Wall EE (2007) Anatomical and functional imaging techniques: basically similar or fundamentally different? *Neth Heart J* 15:43–44
18. van Werkhoven JM, Schuijff JD, Jukema JW et al (2008) Anatomic correlates of a normal perfusion scan using 64-slice computed tomographic coronary angiography. *Am J Cardiol* 101:40–45
19. Scholte AJ, Bax JJ, Wackers FJ (2006) Screening of asymptomatic patients with type 2 diabetes mellitus for silent coronary artery disease: combined use of stress myocardial perfusion imaging and coronary calcium scoring. *J Nucl Cardiol* 13:11–18
20. Wijpkema JS, Dorgelo J, Willems TP et al (2007) Discordance between anatomical and functional coronary stenosis severity. *Neth Heart J* 15:5–11
21. Molhoek SG, Bax JJ, Bleeker GB et al (2004) Comparison of response to cardiac resynchronization therapy in patients with sinus rhythm versus chronic atrial fibrillation. *Am J Cardiol* 94:1506–1509
22. Thygesen K, Alpert JS, White HD (2007) Universal definition of myocardial infarction; Joint ESC/ACCF/AHA/WHF Task Force for the Redefinition of Myocardial Infarction. *Eur Heart J* 28:2525–2538
23. van Lennep JE, Westerveld HT, van Lennep HW, Zwinderman AH, Erkelens DW, van der Wall EE (2000) Apolipoprotein concentrations during treatment and recurrent coronary artery disease events. *Arterioscler Thromb Vasc Biol* 20:2408–2413
24. Chamuleau SA, Vrijsen KR, Rokosh DG, Tang XL, Piek JJ, Bolli R (2009) Cell therapy for ischaemic heart disease: focus on the role of resident cardiac stem cells. *Neth Heart J* 17:199–207
25. de Leeuw JG, Wardeh A, Sramek A, van der Wall EE (2007) Pseudo-aortic dissection after primary PCI. *Neth Heart J* 15:265–266
26. Braun S, van der Wall EE, Emanuelsson S, Kobrin I (1996) Effects of a new calcium antagonist, mibefradil (Ro 40–5967), on silent ischemia in patients with stable chronic angina pectoris: a multicenter placebo-controlled study. The mibefradil international study group. *J Am Coll Cardiol* 27:317–322
27. Portegies MC, Schmitt R, Kraaij CJ et al (1991) Lack of negative inotropic effects of the new calcium antagonist Ro 40–5967 in patients with stable angina pectoris. *J Cardiovasc Pharmacol* 18:746–751
28. ten Kate GJ, Wuestink AC, de Feyter PJ (2008) Coronary artery anomalies detected by MSCT-angiography in the adult. *Neth Heart J* 16:369–375
29. Schuijff JD, Jukema JW, van der Wall EE, Bax JJ (2007) Multi-slice computed tomography in the evaluation of patients with acute chest pain. *Acute Card Care* 9: 214–221
30. Groen JM, Greuter MJ, Vliegenthart R et al (2008) Calcium scoring using 64-slice MDCT, dual source CT and EBT: a comparative phantom study. *Int J Cardiovasc Imaging* 24:547–556
31. Bax AL, van der Wall EE, Braun S, Emanuelsson H, Brusckhe AV, Kobrin I (1995) Effects of the new calcium antagonist mibefradil (Ro 40–5967) on exercise duration in patients with chronic stable angina pectoris: a multicenter, placebo-controlled study. *Ro 40–5967 International Study Group. Am Heart J* 130:748–757
32. Schuijff JD, Pundziute G, Jukema JW et al (2006) Diagnostic accuracy of 64-slice multislice computed tomography in the noninvasive evaluation of significant coronary artery disease. *Am J Cardiol* 98:145–148
33. Jongbloed MR, Lamb HJ, Bax JJ et al (2005) Noninvasive visualization of the cardiac venous system using multislice computed tomography. *J Am Coll Cardiol* 45:749–753
34. Schuijff JD, Wijns W, Jukema JW et al (2006) Relationship between noninvasive coronary angiography with multislice computed tomography and myocardial perfusion imaging. *J Am Coll Cardiol* 48:2508–2514
35. Pundziute G, Schuijff JD, Jukema JW et al (2007) Prognostic value of multislice computed tomography coronary

- angiography in patients with known or suspected coronary artery disease. *J Am Coll Cardiol* 49:62–70
36. Henneman MM, Schuijf JD, Pundziute G et al (2008) Noninvasive evaluation with multislice computed tomography in suspected acute coronary syndrome: plaque morphology on multislice computed tomography versus coronary calcium score. *J Am Coll Cardiol* 52:216–222
 37. de Nooijer R, Verkleij CJ, von der Thüsen JH et al (2006) Lesional overexpression of matrix metalloproteinase-9 promotes intraplaque hemorrhage in advanced lesions but not at earlier stages of atherogenesis. *Arterioscler Thromb Vasc Biol* 26:340–346
 38. van der Laarse A, Kerkhof PL, Vermeer F et al (1988) Relation between infarct size and left ventricular performance assessed in patients with first acute myocardial infarction randomized to intracoronary thrombolytic therapy or to conventional treatment. *Am J Cardiol* 61:1–7
 39. van der Hoeven BL, Pires NM, Warda HM et al (2005) Drug-eluting stents: results, promises and problems. *Int J Cardiol* 99:9–17
 40. Ertaş G, van Beusekom HM, van der Giessen WJ (2009) Late stent thrombosis, endothelialisation and drug-eluting stents. *Neth Heart J* 17:177–180
 41. Scholte AJ, Schuijf JD, Kharagjitsingh AV et al (2008) Different manifestations of coronary artery disease by stress SPECT myocardial perfusion imaging, coronary calcium scoring, and multislice CT coronary angiography in asymptomatic patients with type 2 diabetes mellitus. *J Nucl Cardiol* 15:503–509
 42. Juwana YB, Wirianta J, Suryapranata H, de Boer MJ (2007) Left main coronary artery stenosis undetected by 64-slice computed tomography: a word of caution. *Neth Heart J* 15:255–256
 43. Scholte AJ, Schuijf JD, Kharagjitsingh AV et al (2008) Prevalence of coronary artery disease and plaque morphology assessed by multi-slice computed tomography coronary angiography and calcium scoring in asymptomatic patients with type 2 diabetes. *Heart* 94:290–295
 44. Akram K, Voros S (2008) Absolute coronary artery calcium scores are superior to MESA percentile rank in predicting obstructive coronary artery disease. *Int J Cardiovasc Imaging* 24:743–749
 45. Tops LF, Bax JJ, Zeppenfeld K et al (2005) Fusion of multislice computed tomography imaging with three-dimensional electroanatomic mapping to guide radiofrequency catheter ablation procedures. *Heart Rhythm* 2:1076–1081
 46. Sirineni GK, Raggi P, Shaw LJ, Stillman AE (2008) Calculation of coronary age using calcium scores in multiple ethnicities. *Int J Cardiovasc Imaging* 24:107–111
 47. Marques KM, Westerhof N (2008) Characteristics of the flow velocity-pressure gradient relation in the assessment of stenoses: an in vitro study. *Neth Heart J* 16:156–162
 48. Rodríguez-Granillo GA, Rosales MA, Degrossi E, Rodríguez AE (2009) Signal density of left ventricular myocardial segments and impact of beam hardening artifact: implications for myocardial perfusion assessment by multidetector CT coronary angiography. *Int J Cardiovasc Imaging*. [Epub ahead of print]
 49. Morita K, Tsukamoto E, Tamaki N (2002) Perfusion-BMIPP mismatch: specific finding or artifact? *Int J Cardiovasc Imaging* 18:279–282
 50. America YG, Bax JJ, Dibbets-Schneider P, Pauwels EK, Van der Wall EE (2005) Evaluation of the Quantitative Gated SPECT (QGS) software program in the presence of large perfusion defects. *Int J Cardiovasc Imaging* 21: 519–529
 51. Purser NJ, Armstrong IS, Williams HA, Tonge CM, Lawson RS (2008) Apical thinning: real or artefact? *Nucl Med Commun* 29:382–389
 52. Stinis CT, Lizotte PE, Movahed MR (2006) Impaired myocardial SPECT imaging secondary to silicon- and saline-containing breast implants. *Int J Cardiovasc Imaging* 22:449–455
 53. Verburg FA, Romijn RL, Nekolla S, Verzijlbergen JF (2009) A phantom assessment of cold stomach-related artifacts in myocardial perfusion imaging. *Nucl Med Commun* 30:569–573
 54. Kovalski G, Keidar Z, Frenkel A, Israel O, Azhari H (2009) Correction for respiration artefacts in myocardial perfusion SPECT is more effective when reconstructions supporting collimator detector response compensation are applied. *J Nucl Cardiol* 16:949–955
 55. Ali I, Ruddy TD, Almgrahi A, Anstett FG, Wells RG (2009) Half-time SPECT myocardial perfusion imaging with attenuation correction. *J Nucl Med* 50:554–562
 56. So A, Hsieh J, Li JY, Lee TY (2009) Beam hardening correction in CT myocardial perfusion measurement. *Phys Med Biol* 21:3031–3050