

Specialty Area: Cardiac MRI: Function, Perfusion & Viability

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Highlights:

- The fundamental basis for estimating cardiac perfusion with dynamic contrast-enhanced MRI is signal changes that occur during first passage of contrast agent; normally perfused tissues appear relatively bright, whereas hypoperfused tissues appear relatively dark
- Image acquisition must be fast enough to track the bolus of contrast agent passing through multiple planes of the heart; the most commonly used pulse sequence is saturation-recovery TurboFLASH
- Dynamic contrast-enhanced images can be analyzed qualitatively, semi-quantitatively, or quantitatively
- Major confounders include “dark rim” artifacts, nonlinear relationship between signal and [Gd], and cardiac and respiratory motion

Title: Technical Foundations: Physics of Perfusion Imaging

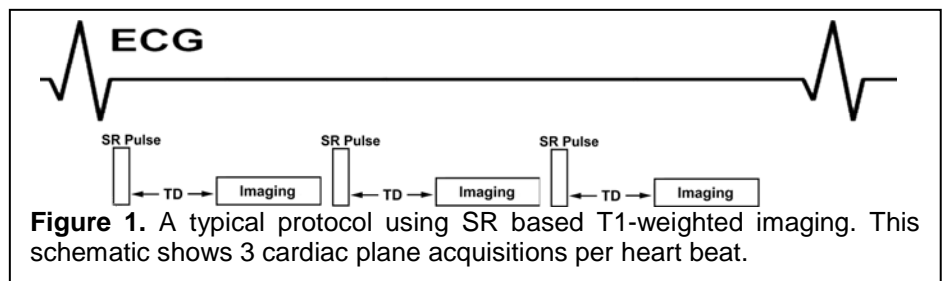
Target audience: Basic science and clinical imagers involved with cardiovascular MRI

Objectives: Upon completion of this educational presentation, the participants should be able to:

- Understand the fundamental basis for assessment of myocardial perfusion with dynamic contrast-enhanced MRI
- Understand the basics of image acquisition and analysis
- Understand the pitfalls of cardiac perfusion MRI and how to address them, to the extent that it is possible
- Appreciate the benefits of k-t acceleration methods to increase the spatio-temporal resolution and/or spatial coverage

Purpose/Introduction: First-pass cardiac perfusion MRI (1-4) has been an active area of investigation for over 2 decades. Recent clinical trials have shown that cardiac perfusion MRI is as accurate as SPECT for coronary artery disease (CAD) detection (5-7). Cardiac perfusion MRI features higher spatial resolution than SPECT; this advantage may be beneficial for detection of subendocardial perfusion defects. This presentation will cover the basics of cardiac perfusion MRI, including image acquisition and image analysis. For more details on key factors that are critical for successful cardiac perfusion MRI, please see (8, 9).

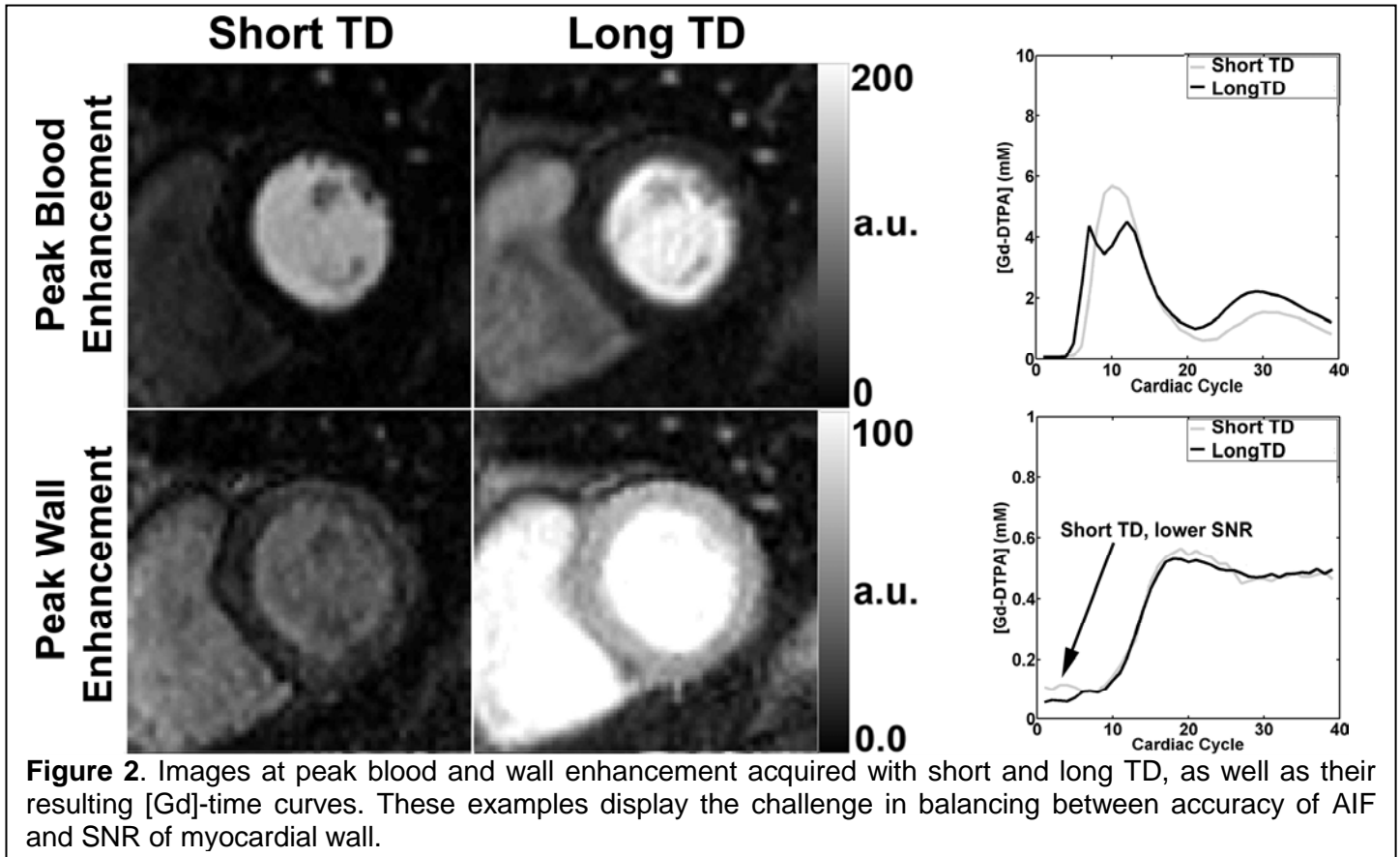
Methods: (Image Acquisition) First-pass perfusion MRI is the one of most challenging cardiovascular MRI techniques; it needs to be performed rapidly to acquire multiple (3-4) planes of the heart with high temporal resolution (~150 ms), and repeated for about 1 min to characterize the bolus kinetics (see Figure 1). Consequently, the intrinsic SNR is relatively low; the dynamic range of signal over time is wide, which makes it challenging to optimize the pulse sequence parameters. The most commonly used pulse sequence is saturation-recovery (SR) TurboFLASH. Other possible readouts include segmented EPI (10, 11) and b-SSFP (12-14), but they are more susceptible to image artifacts than TurboFLASH (13).



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(Signal-to-[Gd] Conversion) For quantitative evaluation of myocardial perfusion, one must convert the signal to concentration of contrast agent [Gd]. When choosing a SR delay (TD), the user must consider multiple factors, which include: myocardial enhancement (longer TD, higher SNR), arterial input function (shorter TD, less clipping of peak signal), spatial coverage (longer TD, less time to acquire multiple planes). Figure 2 shows examples that compares different results with varying TD. With shorter TD, the images suffer from lower SNR but avoids signal clipping for the AIF (i.e., accurate). With longer TD, the images have higher SNR but suffers from signal clipping for AIF (i.e., underestimation). This conundrum can be avoid by acquiring the AIF

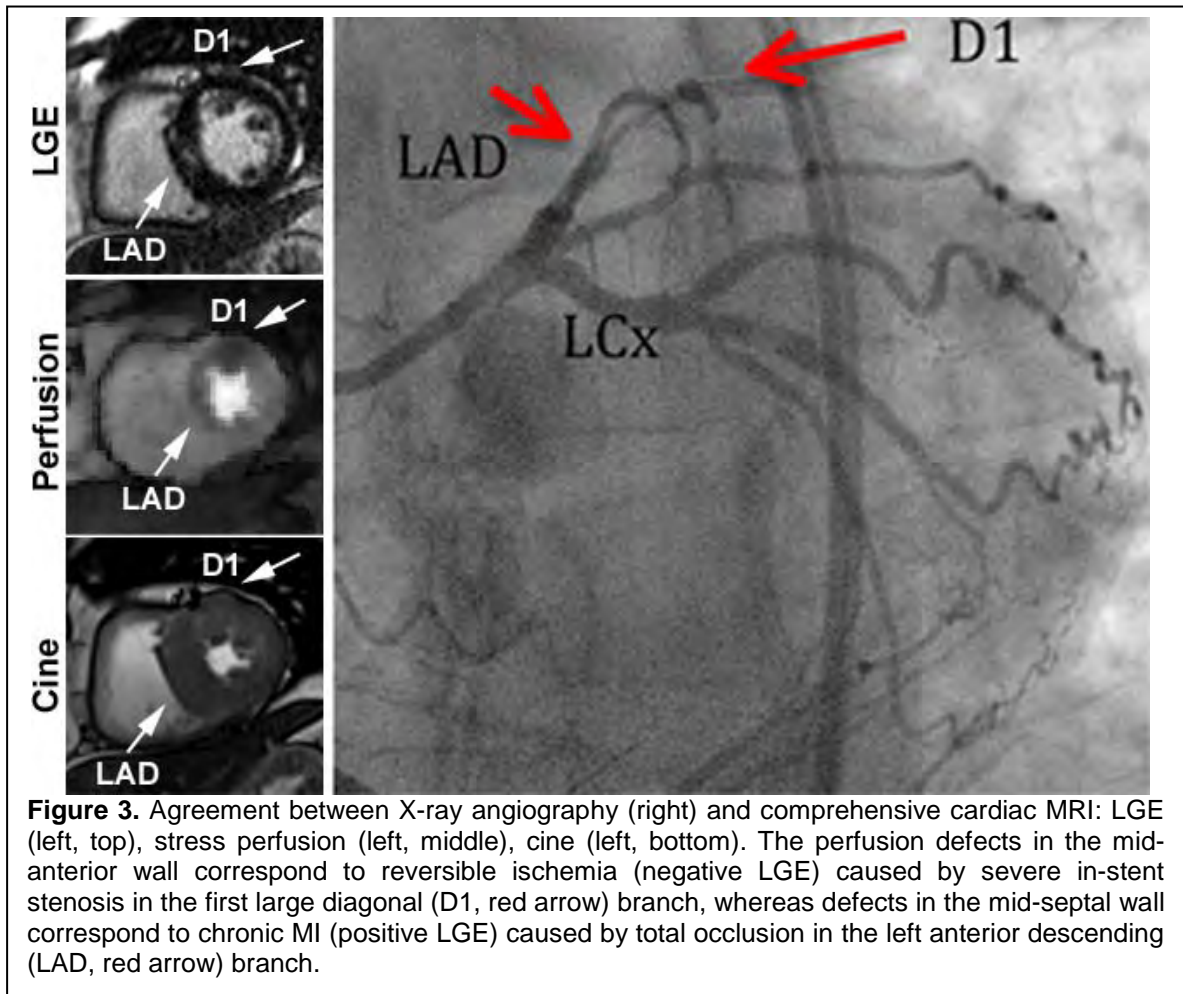
acquisition with short TD and myocardial wall acquisitions with long TD (15-17), or with a dual-bolus approach (18).



(Perfusion Estimation) The method of choice to estimate myocardial perfusion will depend largely on the experience level of the imaging center. Qualitative evaluation is the most common in clinical practice. Semi-quantitative evaluation methods (myocardial perfusion reserve index) are also available (19, 20). Quantitative evaluation methods can improve accuracy and help detect multi-vessel disease (21, 22), but require extensive expertise and experience to produce reliable results.

Results: Figure 3 shows a perfusion image with high resolution (2 mm x 2 mm x 8 mm) and temporal resolution (45 ms) that was used to diagnose CAD (23). The perfusion defects in the mid-anterior wall correspond to reversible ischemia (negative LGE) caused by severe in-stent stenosis in the first large diagonal branch, whereas defects in the mid-septal wall correspond to chronic MI (positive LGE) caused by total occlusion in the left anterior descending branch. These MRI images are consistent with X-ray angiography as shown. Note that this rapid perfusion image acquisition with radial k-space undersampling and constrained reconstruction (24, 25) suppresses dark rim artifacts, owing to the high spatio-temporal resolution (26).

Discussion: Cardiac perfusion MRI is a clinically useful method for CAD detection. Technically, cardiac perfusion MRI is one of the most challenging cardiovascular MRI methods; it needs to be performed rapidly to acquire multiple (3-4) planes of the heart with high temporal resolution (~150 ms), and repeated for about 1 min to characterize the bolus kinetics. Recent advances in acceleration methods provide a means to extend spatial coverage and/or increase spatio-temporal resolution. These methods include non-Cartesian acquisitions (27), dynamic parallel imaging (28), and k-t acceleration methods (29-31). These promising methods, while encouraging, will need to be tested rigorously in clinical settings to ensure they yield high accuracy for CAD detection. Finally, for clinical translation, more robust and easy to use analysis methods are needed to achieve reliable estimates of myocardial perfusion.



Bibliography & References Cited

1. van Rugge FP, Boreel JJ, van der Wall EE, et al. Cardiac first-pass and myocardial perfusion in normal subjects assessed by sub-second Gd-DTPA enhanced MR imaging. *J Comput Assist Tomogr.* 1991;15(6):959-65.
2. Cullen JH, Horsfield MA, Reek CR, et al. A myocardial perfusion reserve index in humans using first-pass contrast-enhanced magnetic resonance imaging. *J Am Coll Cardiol.* 1999;33(5):1386-94.
3. Nagel E, Underwood R, Pennell D, et al. New developments in non-invasive cardiac imaging: critical assessment of the clinical role of cardiac magnetic resonance imaging. *Eur Heart J.* 1998;19(9):1286-93.
4. Atkinson DJ, Burstein D, Edelman RR. First-pass cardiac perfusion: evaluation with ultrafast MR imaging. *Radiology.* 1990;174(3 Pt 1):757-62.
5. Schwitter J, Wacker CM, van Rossum AC, et al. MR-IMPACT: comparison of perfusion-cardiac magnetic resonance with single-photon emission computed tomography for the detection of coronary artery disease in a multicentre, multivendor, randomized trial. *Eur Heart J.* 2008;29(4):480-9.
6. Schwitter J, Wacker CM, Wilke N, et al. MR-IMPACT II: Magnetic Resonance Imaging for Myocardial Perfusion Assessment in Coronary artery disease Trial: perfusion-cardiac magnetic resonance vs. single-photon emission computed tomography for the detection of coronary artery disease: a comparative multicentre, multivendor trial. *Eur Heart J.* 2013;34(10):775-81.
7. Schwitter J, Wacker CM, Wilke N, et al. Superior diagnostic performance of perfusion-cardiovascular magnetic resonance versus SPECT to detect coronary artery disease: The secondary endpoints of the multicenter multivendor MR-IMPACT II (Magnetic Resonance Imaging for Myocardial Perfusion Assessment in Coronary Artery Disease Trial). *J Cardiovasc Magn Reson.* 2012;14:61.
8. Gerber BL, Raman SV, Nayak K, et al. Myocardial first-pass perfusion cardiovascular magnetic resonance: history, theory, and current state of the art. *J Cardiovasc Magn Reson.* 2008;10:18.
9. Kellman P, Arai AE. Imaging sequences for first pass perfusion --a review. *J Cardiovasc Magn Reson.* 2007;9(3):525-37.
10. Debatin JF, McKinnon GC, von Schulthess GK. Technical note--approach to myocardial perfusion with echo planar imaging. *MAGMA.* 1996;4(1):7-11.
11. Ding S, Wolff SD, Epstein FH. Improved coverage in dynamic contrast-enhanced cardiac MRI using interleaved gradient-echo EPI. *Magn Reson Med.* 1998;39(4):514-9.
12. Fenchel M, Helber U, Simonetti OP, et al. Multislice first-pass myocardial perfusion imaging: Comparison of saturation recovery (SR)-TrueFISP-two-dimensional (2D) and SR-TurboFLASH-2D pulse sequences. *J Magn Reson Imaging.* 2004;19(5):555-63.
13. Wang Y, Moin K, Akinboboye O, Reichek N. Myocardial first pass perfusion: steady-state free precession versus spoiled gradient echo and segmented echo planar imaging. *Magn Reson Med.* 2005;54(5):1123-9.
14. Schreiber WG, Schmitt M, Kalden P, et al. Dynamic contrast-enhanced myocardial perfusion imaging using saturation-prepared TrueFISP. *J Magn Reson Imaging.* 2002;16(6):641-52.
15. Gatehouse PD, Elkinington AG, Ablitt NA, et al. Accurate assessment of the arterial input function during high-dose myocardial perfusion cardiovascular magnetic resonance. *J Magn Reson Imaging.* 2004;20(1):39-45.
16. Kim D. Influence of the k-space trajectory on the dynamic T1-weighted signal in quantitative first-pass cardiac perfusion MRI at 3T. *Magn Reson Med.* 2008;59(1):202-8.
17. Kim D, Axel L. Multislice, dual-imaging sequence for increasing the dynamic range of the contrast-enhanced blood signal and CNR of myocardial enhancement at 3T. *J Magn Reson Imaging.* 2006;23(1):81-6.
18. Christian TF, Rettmann DW, Aletras AH, et al. Absolute myocardial perfusion in canines measured by using dual-bolus first-pass MR imaging. *Radiology.* 2004;232(3):677-84.
19. Al-Saadi N, Nagel E, Gross M, et al. Noninvasive detection of myocardial ischemia from perfusion reserve based on cardiovascular magnetic resonance. *Circulation.* 2000;101(12):1379-83.
20. Nagel E, Klein C, Paetsch I, et al. Magnetic resonance perfusion measurements for the noninvasive detection of coronary artery disease. *Circulation.* 2003;108(4):432-7.
21. Mordini FE, Haddad T, Hsu LY, et al. Diagnostic Accuracy of Stress Perfusion CMR in Comparison With Quantitative Coronary Angiography: Fully Quantitative, Semiquantitative, and Qualitative Assessment. *JACC Cardiovasc Imaging.* 2014;7(1):14-22.

22. Patel AR, Antkowiak PF, Nandalur KR, et al. Assessment of advanced coronary artery disease: advantages of quantitative cardiac magnetic resonance perfusion analysis. *J Am Coll Cardiol*. 2010;56(7):561-9.
23. Harrison A, Adluru G, Damal K, et al. Rapid ungated myocardial perfusion cardiovascular magnetic resonance: preliminary diagnostic accuracy. *J Cardiovasc Magn Reson*. 2013;15(1):26.
24. Adluru G, McGann C, Speier P, et al. Acquisition and reconstruction of undersampled radial data for myocardial perfusion magnetic resonance imaging. *J Magn Reson Imaging*. 2009;29(2):466-73.
25. Adluru G, Whitaker RT, DiBella EVR. Spatio-Temporal Constrained Reconstruction of sparse dynamic contrast enhanced radial MRI data. *IEEE Int Symp Biomed Imaging*, 2007. Arlington, VA 109-12.
26. Sharif B, Dharmakumar R, LaBounty T, et al. Eliminating dark-rim artifacts in first-pass myocardial perfusion imaging. *J Cardiovasc Magn Reson*. 2013;15(Suppl 1):O3.
27. Salerno M, Sica C, Kramer CM, Meyer CH. Improved first-pass spiral myocardial perfusion imaging with variable density trajectories. *Magn Reson Med*. 2013;70(5):1369-79.
28. Kellman P, Derbyshire JA, Agyeman KO, et al. Extended coverage first-pass perfusion imaging using slice-interleaved TSENSE. *Magn Reson Med*. 2004;51(1):200-4.
29. Plein S, Kozerke S, Suerder D, et al. High spatial resolution myocardial perfusion cardiac magnetic resonance for the detection of coronary artery disease. *Eur Heart J*. 2008;29(17):2148-55.
30. Plein S, Ryf S, Schwitter J, et al. Dynamic contrast-enhanced myocardial perfusion MRI accelerated with k-t sense. *Magn Reson Med*. 2007;58(4):777-85.
31. Otazo R, Kim D, Axel L, Sodickson DK. Combination of compressed sensing and parallel imaging for highly accelerated first-pass cardiac perfusion MRI. *Magn Reson Med*. 2010;64(3):767-76.