

PET imaging of sympathetic innervation with [^{18}F]Flurobenguan vs [^{11}C]mHED in a patient with ischemic cardiomyopathy

Jason G. E. Zelt, MSc,^{a,b} Lisa M. Mielniczuk, MSc, MD,^{a,b} Cesare Orlandi, MD,^c Simon Robinson, PhD,^c Tayebah Hadizad, PhD,^a Olga Walter, RN,^a Linda Garrard, RN,^a Rob S. B. Beanlands, MD,^{a,b} and Robert A. deKemp, PhD^a

^a Division of Cardiology, Department of Medicine, Molecular Function and Imaging Program, The National Cardiac PET Centre, University of Ottawa Heart Institute and University of Ottawa, Ottawa, ON, Canada

^b Department of Cellular and Molecular Medicine, Faculty of Medicine, University of Ottawa, Ottawa, Canada

^c Lantheus Medical Imaging, North Billerica, MA

Received Nov 6, 2018; accepted Nov 7, 2018
doi:10.1007/s12350-018-01527-5

INTRODUCTION

Although reduced ejection fraction defines a high risk subset, most current risk assessment approaches fail to identify the majority of patients with ischemic cardiomyopathy at risk of sudden arrhythmic death (SAD).¹ Nuclear imaging of the cardiac sympathetic nervous system offers the potential for improving the prediction of SAD in post-MI subjects and guiding therapy to improve survival. [^{123}I]meta-iodobenzylguanidine (MIBG) for planar and SPECT imaging and [^{11}C]meta-hydroxyephedrine (mHED) for PET have been the most widely used to characterize sympathetic nervous system (SNS) function in humans. Their utility in prognostication and risk stratification has been demonstrated in prospective trials.^{2,3} They have also been used as endpoints to evaluate effectiveness of some therapies in small randomized studies.⁴ However, widespread clinical use of PET SNS imaging is limited by the short half-life (20min) of ^{11}C -labeled tracers. A new ^{18}F -labeled tracer, such as Flurobenguan (N-[3-bromo-

4-(3-[^{18}F]fluoro-propoxy)-benzyl]-guanidine), would take advantage of the high resolution and quantification advantages of PET, due to its longer half-life (110 min), allowing for wide distribution to maximize patient and clinical impact.

CASE SUMMARY

We report the first comparison of Flurobenguan (or LMI-1195) vs mHED PET images in a 74-year-old man with a previous anterior MI and ischemic cardiomyopathy. The patient has NYHA class 3 dyspnea, with impaired LV function (EF: 34%) and had received primary prevention ICD therapy. mHED and Flurobenguan PET/CT imaging was performed on two separate imaging visits within one week. A focal defect was identified on mHED revealing a zone of denervation in the area of previous infarct within the anteroseptal wall (Figure 1A). The size and severity of regional denervation were closely recapitulated using the new tracer Flurobenguan (Figure 1B). The definitive assessment of whether Flurobenguan yields a similar estimate of cardiac sympathetic innervation as measured by mHED is the subject of our ongoing trial.

Acknowledgments

Jason Zelt is an MD/PhD student supported in part by the Vanier Canada Graduate Scholarship, The University of Ottawa, and by a government/industry Grant from the Ontario Research Fund (ORF RE07-021) (industry partner:

Reprint requests: Robert A. deKemp, PhD, Division of Cardiology, Department of Medicine, Molecular Function and Imaging Program, The National Cardiac PET Centre, University of Ottawa Heart Institute and University of Ottawa, 40 Ruskin Street, Ottawa, Ontario, K1Y 4W7, Canada; RAdKemp@ottawaheart.ca

J Nucl Cardiol 2019;26:2151–3.

1071-3581/\$34.00

Copyright © 2018 American Society of Nuclear Cardiology.

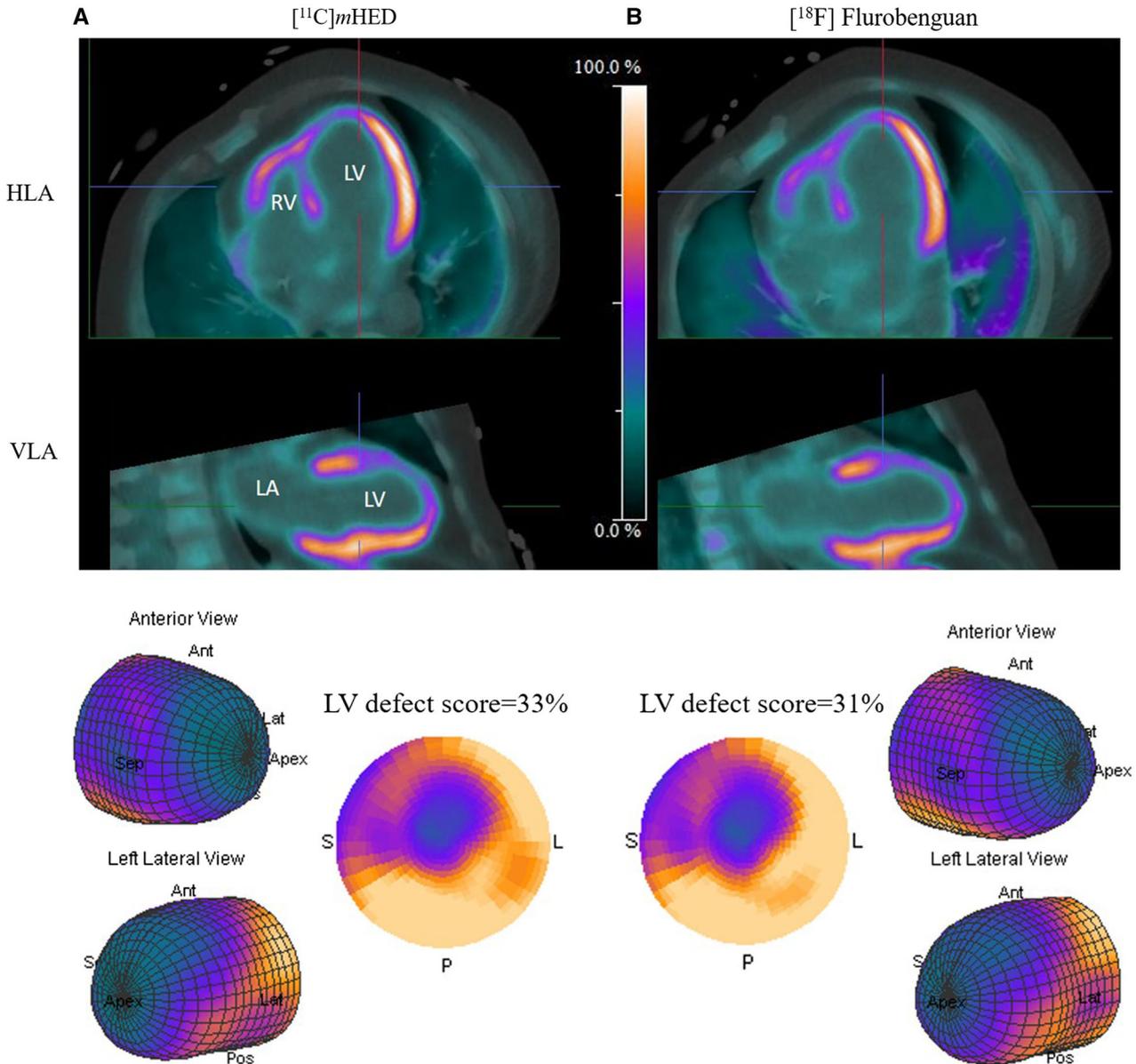


Figure 1. Representative mHED (A) and Fluorobenguan (B) PET images in a patient with ischemic cardiomyopathy and previous anterior STEMI. The size and severity of the regional denervation were similar when assessed with both mHED and Fluorobenguan.

Lantheus Medical Imaging). The latter also supported this work. Rob Beanlands is a Career Investigator supported by the Heart and Stroke Foundation of Ontario (HSFO), a Tier 1 Chair in Cardiac Imaging Research at the University of Ottawa and Vered Chair in Cardiology at the University of Ottawa Heart Institute. Lisa Mielniczuk is a Mid-career Clinician Scientist supported by HSFO and Tier 2 Chair in HF Research at the University of Ottawa.

Disclosure

Rob S. B. Beanlands and Robert A. deKemp received unrestricted grant funding from Lantheus Medical Imaging (LMI) for this study. Rob S. B. Beanlands is a consultant for LMI. Cesare Orlandi and Simon Robinson are employees of LMI. Jason G. E. Zelt, Lisa M. Mielniczuk, Tayebah Hadizad, Olga Walter, and Linda Garrard declare they have no conflict of interest related to this study.

References

1. Stecker EC, Vickers C, Waltz J, et al. Population-based analysis of sudden cardiac death with and without left ventricular systolic dysfunction. *J Am Coll Cardiol* 2006;47:1161-6. <https://doi.org/10.1016/j.jacc.2005.11.045>.
2. Jacobson AF, Senior R, Cerqueira MD, et al. Myocardial iodine-123 meta-iodobenzylguanidine imaging and cardiac events in heart failure. *J Am Coll Cardiol* 2010;55:2212-21. <https://doi.org/10.1016/j.jacc.2010.01.014>.
3. Fallavollita JA, Heavey BM, Luisi AJ, et al. Regional myocardial sympathetic denervation predicts the risk of sudden cardiac arrest in ischemic cardiomyopathy. *J Am Coll Cardiol* 2014;63:141-9. <https://doi.org/10.1016/j.jacc.2013.07.096>.
4. Hall AB, Ziadi MC, Leech JA, et al. Effects of short-term continuous positive airway pressure on myocardial sympathetic nerve function and energetics in patients with heart failure and obstructive sleep apnea: a randomized study. *Circulation* 2014;130:892-901. <https://doi.org/10.1161/CIRCULATIONAHA.113.005893>.