

# COMPARISON OF FREE-BREATHING AND BREATH-HELD CINE DENSE STRAIN IN BOYS WITH DUCHENNE MUSCULAR DYSTROPHY

UCLA

Patrick Magrath<sup>1,2</sup>, Nyasha Maforo<sup>2,3</sup>, Michael Loecher<sup>4</sup>, Holden Wu<sup>1-3</sup>, Nancy Halnon<sup>2</sup>, Pierangelo Renella<sup>2,</sup> Daniel B. Ennis<sup>4</sup>

¹Department of Bioengineering, University of California, Los Angeles, ²Department of Radiology, University of California, Los Angeles

³Physics and Biology in Medicine Interdepartmental Program, University of California, Los Angeles, ⁴Department of Radiology, Stanford University

MAGNETIC RESONANCE Research Labs

## Introduction

Duchenne Muscular Dystrophy is a fatal, genetic muscle wasting disorder impacting 1:3800 boys [1]. The most common cause of mortality is cardiac failure and/or respiratory failure [2]. In order to assess the progression of cardiac disease and response to current and novel therapeutics, peak mid-wall circumferential strain (Ecc) has been proposed as a biomarker [3].

CINE Displacement Encoding with Stimulated Echoes (CINE DENSE)

- permits rapid, navigator-gated assessment of  $E_{cc}$  in the heart
- validated against MRI tagging
- has not been reported in a DMD cohort.

Breath holding can impact measures of cardiac function [2], but no comparison of free-breathing and breath-held Ecc measurements has been conducted in boys with DMD using cine DENSE encoding.

## **OBJECTIVES**

Our *objectives* were:

- 1) To characterize  $E_{cc}$  in a DMD cohort using CINE DENSE
- 2) To compare the  $E_{cc}$  measurements derived from free-breathing and breath-held CINE DENSE.

#### METHODS

Study Population: Boys with DMD (N = 15) were prospectively enrolled in this IRB approved, HIPAA compliant study after obtaining informed consent. DMD boys had an age of (13  $\pm$  2.1 years) and a Left Ventricular Ejection Fraction (LVEF) of (50.35  $\pm$  10.78)

Acquisition: Single slice, short-axis, mid-ventricular data was acquired on a 3T Siemens Skyra using two protocols:

- 1. Navigator-gated, free-breathing CINE DENSE examination (2.5x2.5x8mm, TE/ $T_{Res}$ =1.2/15 ms, Denc = .08 cycles/mm, spirals = 10, Navg = 3, scan time = 2.5 minutes)
- 2. Breath-held CINE DENSE (3.0x3.0x8mm, TE/ $T_{Res}=1.2/15$  ms, Denc = .08 cycles/mm, spirals = 6, Navg = 1, scan time = 25 seconds)

#### Post processing:

- •A custom post-processing tool (Matlab) was used to segment the data and transform the x and y displacement encoded phase information into Lagrangian displacements and  $E_{cc}$ .
- The mid-wall wall was identified and global  $E_{cc}$  values were computed for both acquisitions.

#### Statistics:

 $\cdot$  E<sub>cc</sub> values were summarized as mean  $\pm$  standard deviation for each subject.

# METHODS (CONTINUED)

- •Bland-Altman analysis summarized mean difference and bias between the breath-held and free-breathing acquisition.
- A paired two-tailed t-test was used to compare differences in E<sub>cc</sub> values for the free-breathing and breath-held acquisitions. A p-val≤0.05 indicated statistical significance.

#### RESTITES

Strain Values: Mid-wall  $E_{cc}$  in DMD patients was -14.9  $\pm$  4.4% and -15.2  $\pm$  5.0% for the breath held and free breathing technique respectively (P=0.53).

Further Analysis: Figure 1 shows Bland-Altman analysis between the breath-held and free-breathing acquisitions ( $E_{cc}$  Bias = - 0.05%, Limits of agreement [-5.50%, 4.50%]) and Figure 2 demonstrates a paired box-plot comparing the free breathing and breath held technique with individual pairs of data mapped to one another.

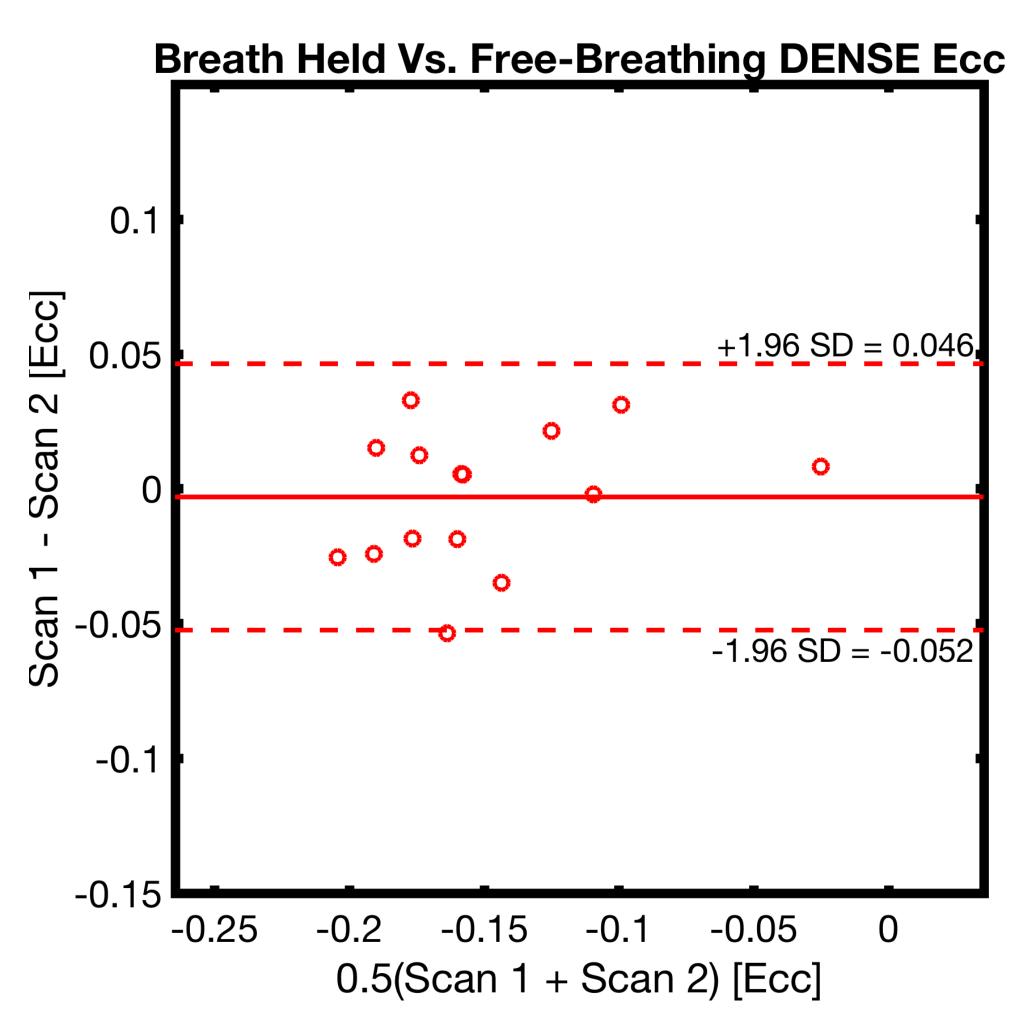


Figure 1. Bland-Altman analysis of peak mid-wall E<sub>cc</sub> measured using breath held and free breathing CINE DENSE.

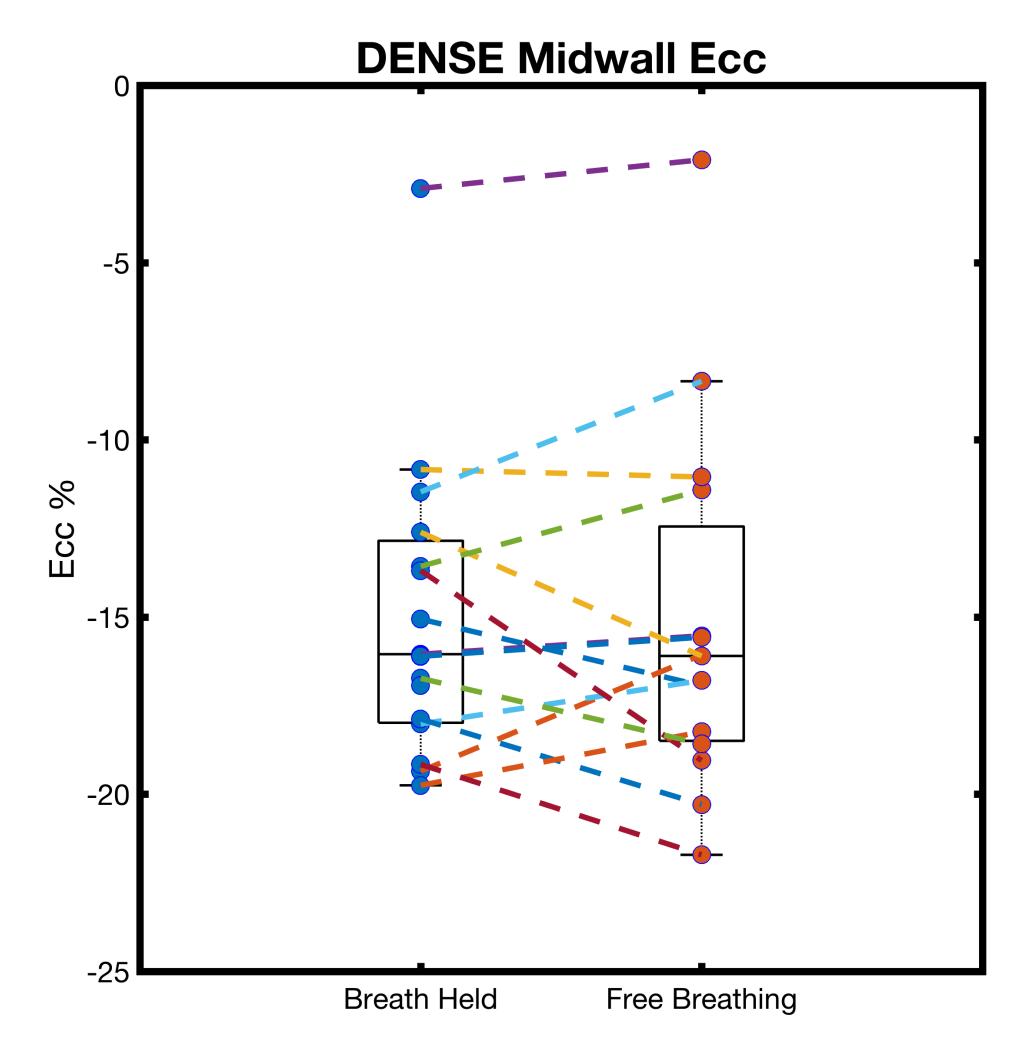


Figure 2. Paired box-plot comparing  $E_{cc}$  results for breath held and free breathing CINE DENSE. There is no significant difference between the two groups. Each individual's scans are connected with dashed lines for reference.

## RESULTS (CONTINUED)

**Figure 3** shows a side by-side comparison of breath-held and free-breathing cine DENSE at peak systole in a representative subject.

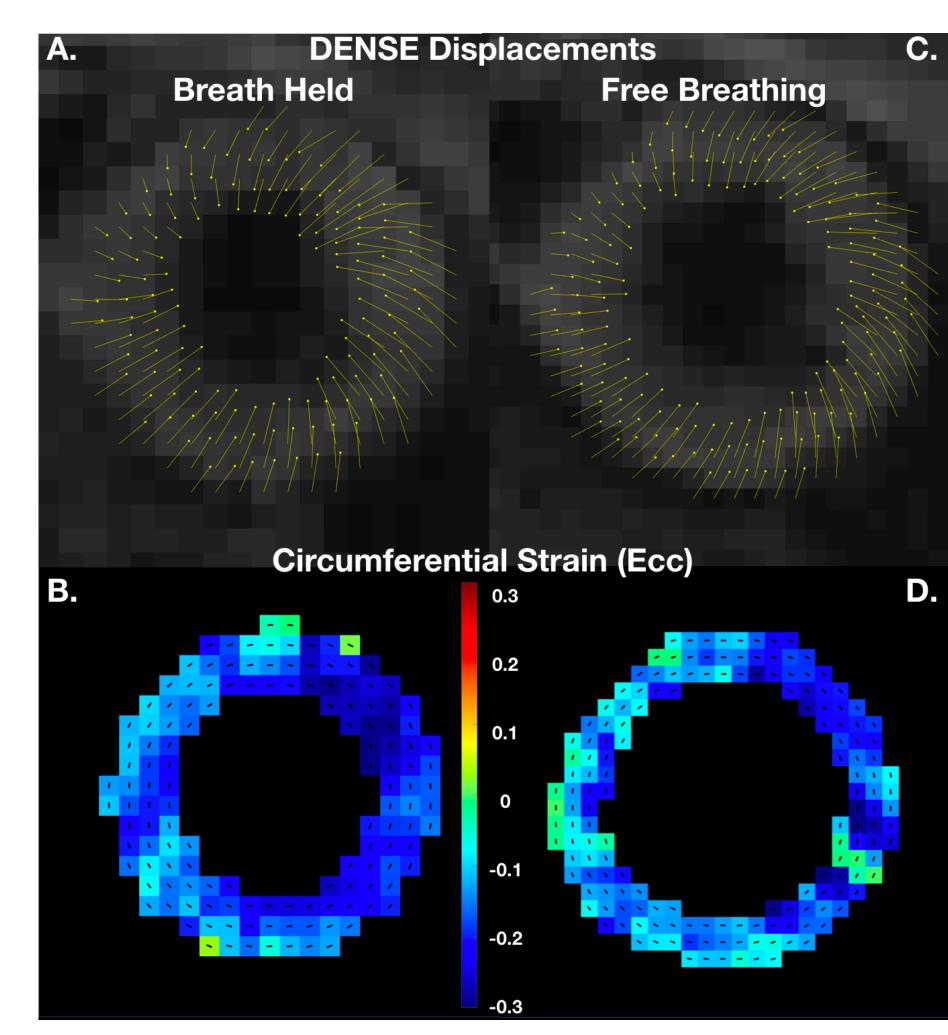


Figure 3. Representative image of DENSE displacements and  $E_{cc}$  in one DMD patient for the breath held acquisition (A and B) and the free breathing acquisition (C and D).

## DISCUSSION

- This report is the first time that normative values for peak mid-ventricular mid-wall  $E_{cc}$  have been reported using CINE DENSE in a DMD cohort.
- The results show good agreement with previous  $E_{cc}$  reports in boys with DMD using MRI tagging (-12.4% and-15%  $E_{cc}$ , [3,4]).
- The use of a breath-held or free breathing technique did not significantly alter the resultant strain values.
- However, because of the difficulty DMD patients frequently have with breath-holding, capitalizing on DENSE's ability to acquire strain information without breath holding is a compelling way to improve clinical workflow and patient compliance, especially for more severely effected boys.
- Anecdotally, image quality is better with free breathing data and is more easily processed.

### CONCLUSION

Free breathing CINE DENSE strain estimates are an excellent technique for characterizing myocardial function in boys with DMD.

#### ACKNOWLEDGEMENTS

This work was supported by NIH K25HL135408 & NIH NHLBI R01 HL131975.

#### REFERENCES

- [1] McNally EM, Kaltman JR, Benson DW *et al.* Contemporary cardiac issues in
- Duchenne muscular dystrophy. *Circulation*, 131(18), 1590-1598 (2015).

  [2] Hor KN, Baumann R, Pedrizzetti G *et al.* Magnetic Resonance Derived Myocardial Strain Assessment Using Feature Tracking. *Journal of Visualized Experiments : JoVE*, (48), 2356 (2011)
- [3]Hor KN, Kissoon N, Mazur W *et al.* Regional circumferential strain is a biomarker for disease severity in duchenne muscular dystrophy heart disease: a cross-sectional study. *Pediatric cardiology*, 36(1), 111-119 (2015).