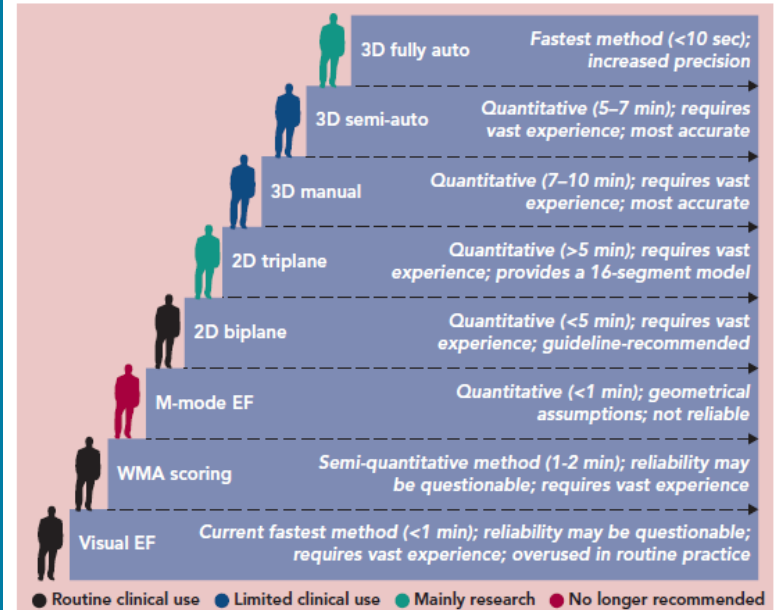


Assessing LV Systolic Function: when to use contrast or 3D echo

Michael H. Picard, MD, FASE
No disclosures

Figure 3: Stairway of Echocardiographic Methods for the Assessment of Left Ventricular Ejection Fraction



EF = ejection fraction; M-mode = mono-dimensional; WMA = wall motion abnormalities.
Spitzer et al. Cardiac Failure Review 2017;3:97-101

LV function in clinical practice: role of echo

- **Diagnosis** – systolic and diastolic dysfunction
 - Etiology for symptoms
- Assessing **response** to treatment
- Assessing risk and **prognosis**
 - Need for interventions
 - Defibrillators, valve surgery, meds, CRT
 - Timing of interventions



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Assessing function of the LV pump

Myocardial deformation

- Vcf
- Isovolumic acceleration
- dP/dt
- Tissue Doppler
- Strain
- Strain rate
- Torsion
- Twist

Volume change

- Shortening fraction
- Stroke volume
- LVEF
- Stroke work
- Elastance
 - End systolic pressure-volume relation



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Quantitation of global LV systolic function: *how?*

Isovolumic indices

- dP / dt
- E_{max}

Ejection phase indices

- Area change
- Myocardial Performance Index (MPI, Tei index)
- Fractional shortening
- Velocity of circumferential fiber shortening (Vcf)
- LV EF



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Assessing global systolic function by LV EF

- Has flaws but most commonly used measure
- Qualitative
 - Internal check
- Single dimension
 - Obsolete
- Volumetric
 - **Simpson's Rule Method / Method of Discs**
 - 2/3 Area length
 - When apical endocardium can't be traced
 - **Three dimensional**



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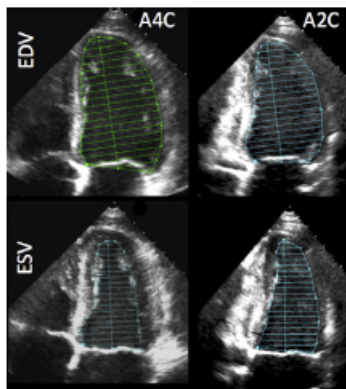
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ASE Best Practice: LVEF should be calculated from LV volumes

Volumes.

Volume measurements are usually based on tracings of the blood-tissue interface in the apical four- and two-chamber views. At the mitral valve level, the contour is closed by connecting the two opposite sections of the mitral ring with a straight line. LV length is defined as the distance between the middle of this line and the most distant point of the LV contour.

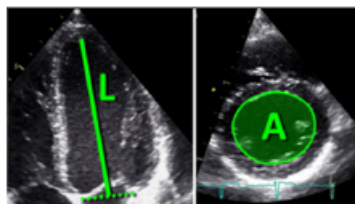
Biplane disk summation



- Corrects for shape distortions
- Less geometrical assumptions compared with linear dimensions
- Apex frequently foreshortened
- Endocardial dropout
- Blind to shape distortions not visualized in the apical two- and four-chamber planes

Single plane ok if no WMA

Area-length



- Partial correction for shape distortion
- Apex frequently foreshortened
- Heavily based on geometrical assumptions
- Limited published data on normal population

MOD – trace interface between compacted and noncompacted myocardium



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LV function guideline recommendations

Lang et al, JASE 2015;28:1-39

- LVEF from 2D volumes
 - < 52% for men abnormal
 - < 54% women abnormal



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Pitfalls of 2D LVEF

- 2D LVEF from biplane method of discs (MOD) or area length (AL) is a reasonable screen for relative degrees of dysfunction
 - AL assumes a geometric shape of the LV
 - MOD assumes elliptical shape of each disc
 - MOD requires accurate delineation of endocardial borders
 - MOD over-weights the size and motion of the LV from 2 apical views
- 2D LVEF may have reduced accuracy in remodeled LV, states with abnormal septal motion, focal RWMA, foreshortened LV
- Solutions
 - Ultrasound Enhancing Agents (aka Contrast)
 - 3D LV echo

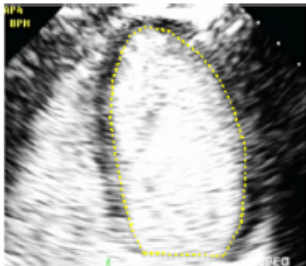


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Enhancements to 2D LV volume measurement

- Contrast agents (aka ultrasound enhancing agents UEA)

| Parameter and method | Technique | Advantages | Limitations |
|----------------------|---|--|---|
| | Endocardial border enhancement  | <ul style="list-style-type: none">• Helpful in patients with suboptimal acoustic window• Provides volumes that are closer to those measured with cardiac magnetic resonance | <ul style="list-style-type: none">• Same limitations as the above non-contrast 2D techniques• Acoustic shadowing in LV basal segments with excess contrast |

Lang et al, JASE 2015;28:1-39

- Many studies have documented beneficial effect of UEA on early outcomes in critically ill patients and cost effectiveness in those with suboptimal windows
- Appropriate use criteria
 - 2 or more contiguous segments not seen on noncontrast images



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- Very low mechanical index imaging ($MI < 0.2$) RECOMMENDED
 - Non linear acoustic signal differentiated from tissue
 - High spatial and good temporal resolution
 - LVEF and RWMA
- Low MI (<0.3)
 - Harmonic techniques
- Intermediate MI (0.3-0.5)
 - Harmonic techniques
 - More destruction of microbubbles and swirling artifacts
 - Same problem with high MI (>0.5)



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Value of contrast – 2018 ASE guidelines

Porter et al, JASE 2018;31:241-74

- Details on agents
- Details on settings for various machines
- Details on indications
- Practical tips

Table 4 Location and description of VLMI imaging software on commercially available echocardiographic scanners

| Manufacturer | Platform and portability* | Location and name of enhanced imaging software on front end | High-MI “flash” impulse location on front end | Specific pulse sequence scheme used (dominant nonlinear activity detected) | Frequency/MI recommended for VLMI imaging |
|--------------|----------------------------|---|---|---|--|
| Philips | iE33 Not portable | Contrast key On/off LVO and low-MI choices | Touch screen/ flash label | Amplitude modulation and pulse inversion (fundamental and harmonic) | <2.0 MHz/MI < 0.2 (GEN or PEN setting) |
| Philips | Epiq Not portable | Contrast key On/off Low-MI and LVO choices | Touch screen/ flash label | Amplitude modulation and pulse inversion (fundamental and harmonic) | <2.0 MHz/MI < 0.2 (GEN or PEN setting) |
| Philips | CX50 Portable | Contrast key On/off LVO choice | Control panel | Amplitude modulation (harmonic) | <2.0 MHz/MI < 0.3 |
| GE | Vivid E95 Not portable | Advanced contrast option | Touch screen/ flash label | Pulse inversion 1.5/3.0 and 1.6/3.2 MHz and 1.7/3.4 MHz (harmonic) Amplitude modulation 2.1 and 2.4 MHz (fundamental and harmonic) | 1.5–1.7 MHz/MI < 0.2 2.1–2.4 MHz/MI < 0.2 |
| Siemens | SC2000 Not portable | | Not available; need to use “color Doppler” knob | Pulse inversion and alternating polarity/amplitude (fundamental and harmonic) | 2.0 MHz/MI < 0.2 |
| Toshiba | Aplio i900 Not portable | Touch screen/ CHI label | Control panel | Pulse subtraction (amplitude modulation; harmonic) | h3.5/MI < 0.2 (PEN setting) |
| Toshiba | Aplio 500 Not portable | Touch screen/ low label | Touch screen/ flash label | Pulse subtraction (amplitude modulation; harmonic) | h2.8–h3.6/MI < 0.2 |
| Esaote | MyLabEight Not portable | Contrast key On/off LVO choice | Touch screen/ flash label | Phase cancellation | PEN frequency/MI < 0.2 |
| Esaote | MyLabSeven Not portable | Contrast key On/off LVO choice | Touch screen/ flash label | Phase cancellation | 1.5 MHz/MI < 0.2 |
| Esaote | MyLabAlpha Portable | Contrast key On/off LVO choice | Touch screen/ flash label | Contrast tuned imaging | 1.5 MHz/MI < 0.2 |

Where to end tracing; where is the MV annulus ?

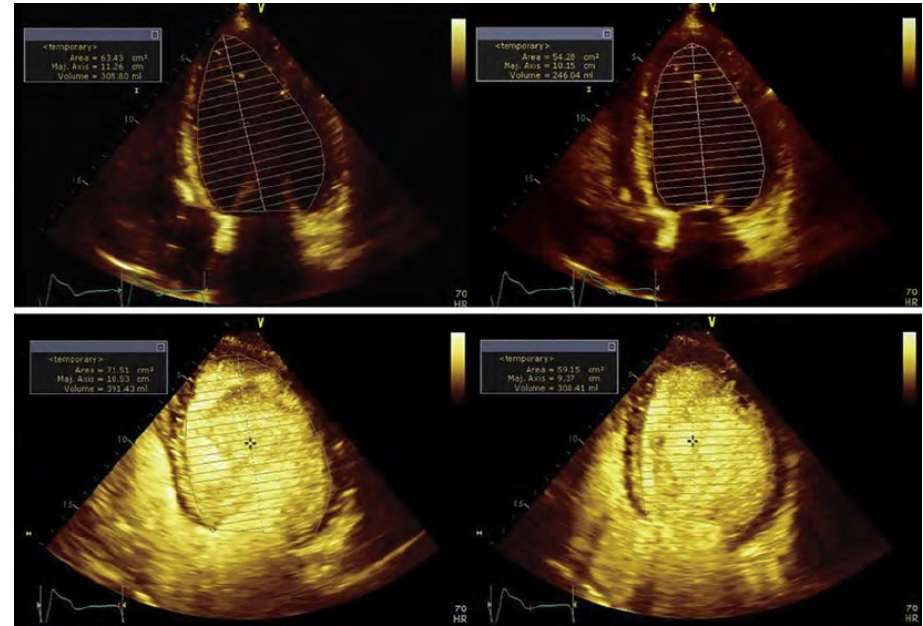


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Contrast for LV systolic function - volumes

- Contrast enhanced LV volumes will be larger than unenhanced
 - Less foreshortening
 - Better delineation of the border between non-compacted and compacted myocardium
- Larger ULN for LV volume
 - EDV
 - Women 81 ml/sq m
 - Men 98 ml/sq m
 - Better agreement with CMR



Porter et al, JASE 2018;31:241-74

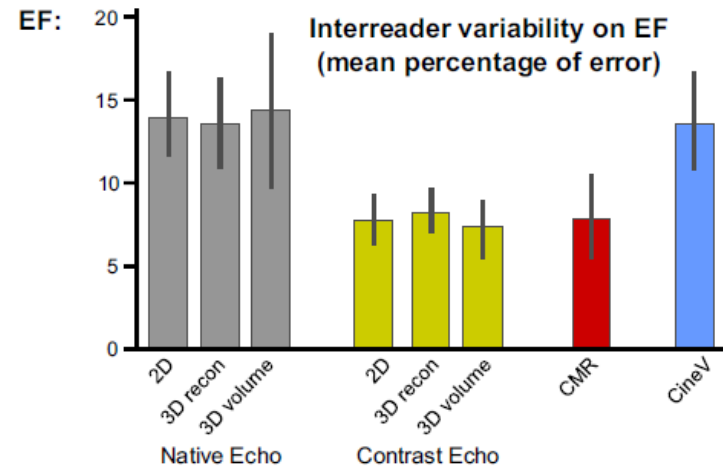


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Contrast for LVEF

- Improved accuracy (CMR gold standard)
- Reduced interobserver variability
- Therefore, value when precise quantification of LVEF required
 - Defibrillator, CRT, Chemotherapy follow up, valve disease intervention timing



Hoffmann et al, JASE 2014;27:292-301



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Routine use of contrast on admission TTE for heart failure reduces rate of repeat echoes

Lee et al JASE 2021;34:1253-61

- 9,115 HF admissions over 4 year period
 - 5,600 UEA on 1st TTE, 3,515 no UEA
 - 104 repeat TTEs during hospital stay considered unjustified
 - 77% were in the no contrast 1st TTE
 - As rate of contrast increased over 4 year period
 - » Unjustified TTE rate decreased
 - Use of contrast associated with reduced LOS



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Overcoming barriers to use

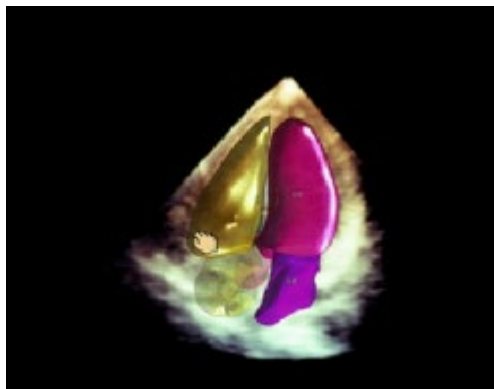
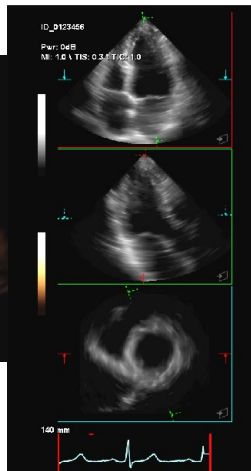
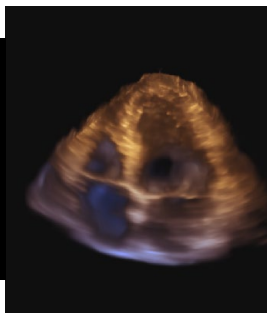
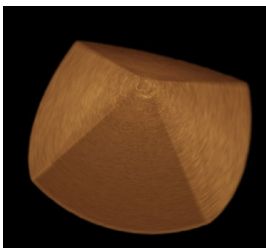
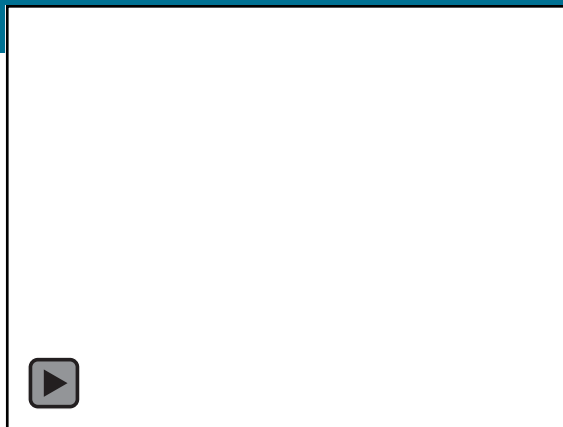
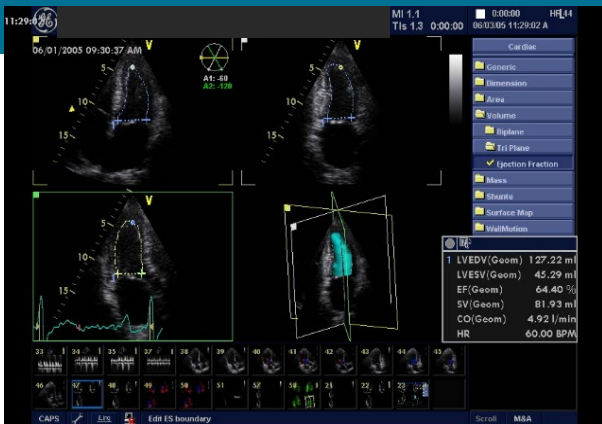
- IV placement
 - Various models
 - Sonographers place IV
 - Echo lab nurse for IV
- Should we use contrast for all LVEF assessments even when image quality is optimal ?
 - Benefit Study



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3D echo for volume and EF

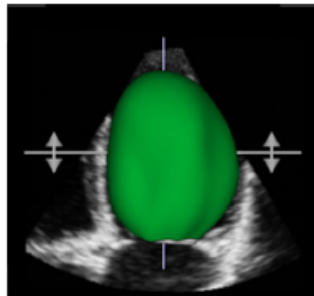


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Enhancements to LV volume measurement

3D data sets



- No geometrical assumption
- Unaffected by foreshortening
- More accurate and reproducible compared to other imaging modalities
- Lower temporal resolution
- Less published data on normal values
- Image quality dependent



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3D normal values

- Volumes larger than 2D
- EF range different

Table 3 Normal values for LV parameters obtained with 3DE

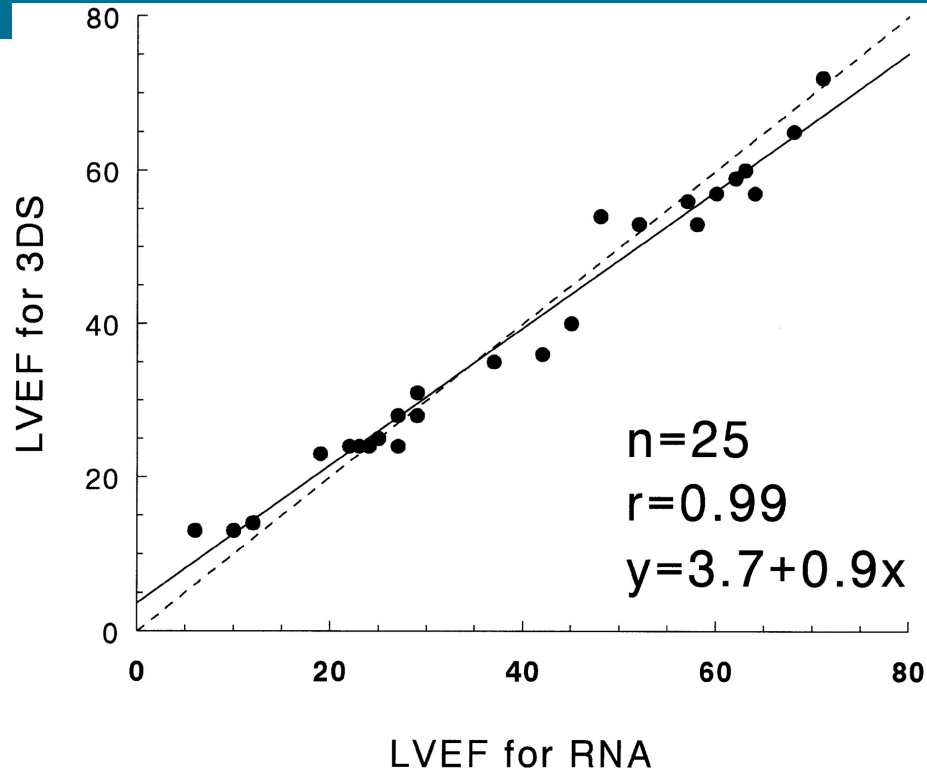
| | Aune <i>et al.</i> (2010) | Fukuda <i>et al.</i> (2012) | Chahal <i>et al.</i> (2012) | Muraru <i>et al.</i> (2013) |
|-----------------------------|---------------------------|-----------------------------|---|-----------------------------|
| Number of subjects | 166 | 410 | 978 | 226 |
| Ethnic makeup of population | Scandinavian | Japanese | 51% European white, 49% Asian Indian | White European |
| EDVi (mL/m ²) | | | | |
| Men, mean (LLN, ULN) | 66 (46, 86) | 50 (26, 74) | White: 49 (31, 67); Indian: 41 (23, 59) | 63 (41, 85) |
| Women, mean (LLN, ULN) | 58 (42, 74) | 46 (28, 64) | White: 42 (26, 58); Indian: 39 (23, 55) | 56 (40, 78) |
| ESVi (mL/m ²) | | | | |
| Men, mean (LLN, ULN) | 29 (17, 41) | 19 (9, 29) | White: 19 (9, 29); Indian: 16 (6, 26) | 24 (14, 34) |
| Women, mean (LLN, ULN) | 23 (13, 33) | 17 (9, 25) | White: 16 (8, 24); Indian: 15 (7, 23) | 20 (12, 28) |
| EF (%) | | | | |
| Men, mean (LLN, ULN) | 57 (49, 65) | 61 (53, 69) | White: 61 (49, 73); Indian: 62 (52, 72) | 62 (54, 70) |
| Women, mean (LLN, ULN) | 61 (49, 73) | 63 (55, 71) | White: 62 (52, 72); Indian: 62 (52, 72) | 65 (57, 73) |



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Linear regression of LVEF in all patients, measured by 3D echocardiography by Simpson's method (3DS) vs radionuclide angiography (RNA)



Nosir, Y. F.M. et al. *Circulation* 1996;94:460-466



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Accuracy of 3D echo for LV volume

- function of image quality:
 - the number of elements in the matrix array transducer
 - the voxel size
 - the spatial resolution of the image
 - Temporal resolution



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Challenges to using 3D LVEF in clinical practice

- Since normal range is different than 2D how do you deal with teaching your referring clinicians how they should interpret the different tests
 - An LVEF of 50% on 3D is normal but if the MD is used to the 2D range they may think of it as abnormal

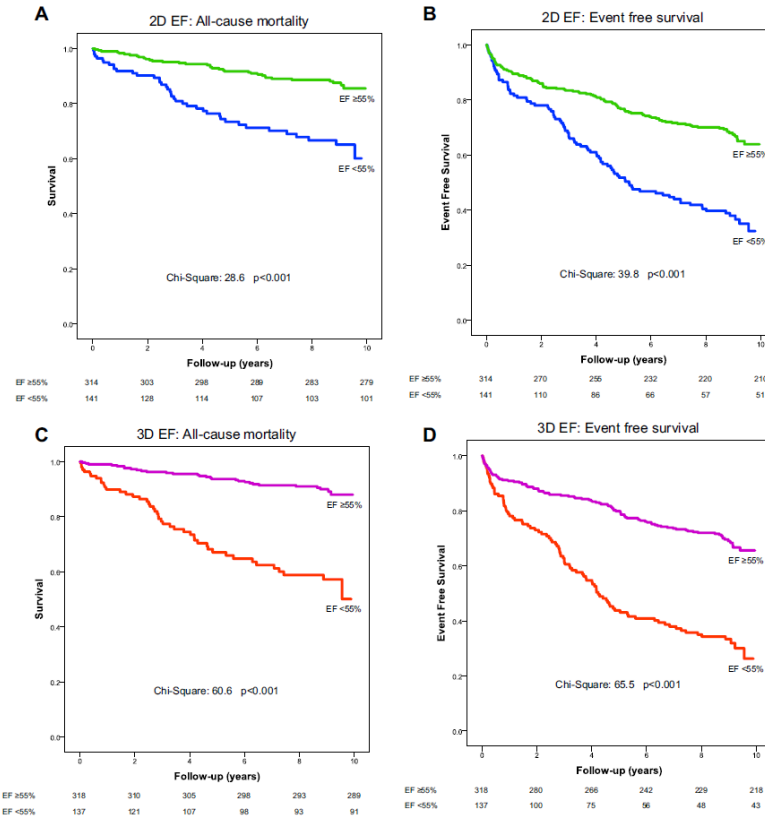


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3D EF stronger association with outcomes than those from 2D

Stanton et al, JASE 2014;27:65-73



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Automated 3D echo LVEF performance

Spitzer et al. *Cardiac Failure Review* 2017;3:97-101

Table 1: Comparisons Among Fully Automated 3D Methods and Either Cardiac Magnetic Resonance or Manual Echocardiography

| Authors | Reference | Software | n | Feasibility | LVEDV bias (ml) | LVESV bias (ml) | LVEF bias (%) |
|--|------------|---------------------------------|-----|-------------|-----------------|-----------------|---------------|
| Thavandiranathan, et al., 2012 ¹⁵ (I) | CMR | eSie LVA™ (Siemens Healthcare) | 101 | 66 % | -18 ± 54 | -10 ± 36 | -0 ± 6 |
| Thavandiranathan, et al., 2012 ¹⁵ (II)* | 2D Simpson | eSie LVA | 27 | 89 % | 2 ± 16 | 4 ± 13 | -2 ± 4 |
| Ren, et al., 2014 ¹⁹ | Manual 3D | eSie LVA | 48 | 85 % | -3 ± 23 | -2 ± 14 | -0 ± 9 |
| Otani, et al., 2016 ^{21,*} | 2D Simpson | HeartModel (Philips Healthcare) | 10 | 100 % | -3 ± 26 | -1 ± 17 | -0 ± 10 |
| Tsang, et al., 2016 ³⁰ (I) | CMR | HeartModel | 69 | 94 % | 2 ± 40 | 10 ± 40 | -6 ± 16 |
| Tsang, et al., 2016 ³⁰ (II) | Manual 3D | HeartModel | 104 | 90 % | -24 ± 50 | -13 ± 58 | -2 ± 18 |
| Spitzer, et al., 2017 ²² | Manual 3D | HeartModel | 72 | 93 % | -6 ± 39 | -2 ± 39 | -1 ± 15 |
| Levy, et al., 2017 ³³ | CMR | HeartModel | 63 | 86 % | -22 ± 34 | -13 ± 33 | -1 ± 7 |
| Medvedofsky, et al., 2017 ^{34,†} | Manual 3D | HeartModel | 180 | 100 % | -14 ± 20 | -6 ± 16 | -2 ± 7 |
| Medvedofsky, et al., 2017 ^{34,‡} | Manual 3D | HeartModel | 300 | 66 % | -3 ± 22 | 1 ± 16 | 0 ± 10 |

*Atrial fibrillation; †Including patients with arrhythmias; ‡Consecutive patients. I and II describe two reference modalities used in a single report. CMR = cardiac magnetic resonance; LVA = left ventricle analysis; LVEDV = left ventricle end-diastolic volume; LVEF = left ventricle ejection fraction; LVESV = left ventricle end-systolic volume.



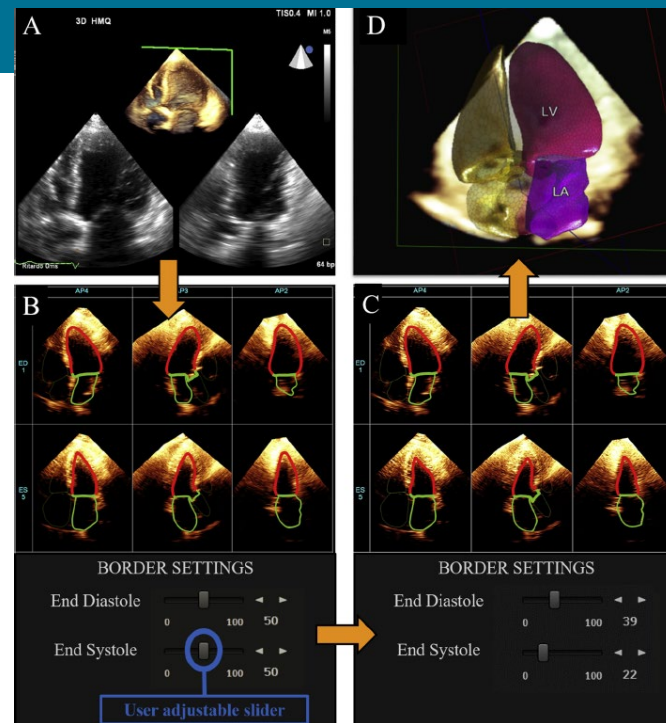
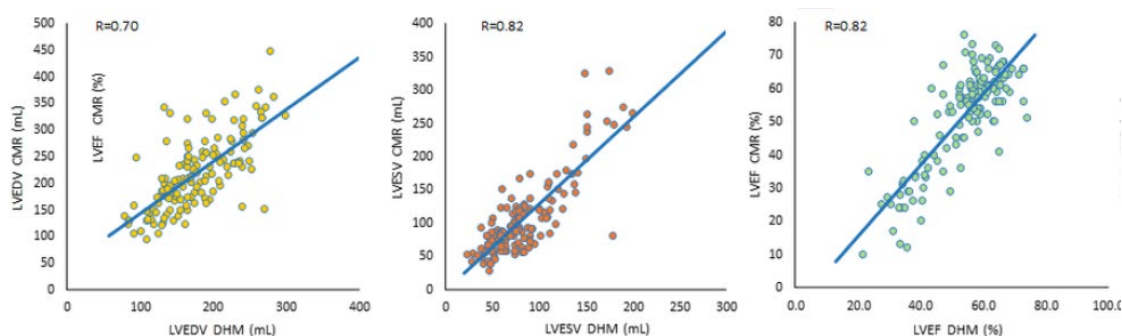
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Automated machine learning-based 3D quantification of LV volume and function

Italiano et al, DOI [10.21203/rs.3.rs-355587/v1](https://doi.org/10.21203/rs.3.rs-355587/v1)

- Automated LV border detection and surface rendering
 - Based on library of shape descriptions of the LV optimally place the contours
- 600 unselected patients (12% AF)
 - 140 with CMR
 - 88% feasibility
 - 64% accurate borders (9% major border corrections required)
 - Small LVs, distorted shapes



Tamborini et al JASE 2017;30:1049-58



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Contrast for 2D LVEF

- When endocardial border not delineated on 2 or more contiguous segments
- LV volumes will be larger than unenhanced LV images
 - EFs are equivalent
- Patience and practice required to identify mitral annular plane for biplane method of disc tracings
- Of particular benefit when precise LV volumes or EF are needed for serial assessment and clinical decision making

3D for LV systolic function:

- Limitations
 - Image quality
 - Rhythm (use single beat capture)
 - Lower temporal resolution than 2D
 - Less published data for normal values
- Advantages
 - Better precision than 2D especially in asymmetric LVs
 - Can pick up subtle differences on serial studies
 - Follow course of a disease
- **BENEFITS OUTWEIGH LIMITATIONS**



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A photograph of the Massachusetts General Hospital building at dusk. The building is a large, multi-story structure with many windows, some of which are illuminated from within, creating a warm glow against the dark blue sky. The building is situated on a hillside overlooking a body of water. In the foreground, there are trees and a road with some cars. The overall scene is a mix of urban architecture and natural elements.

Thank you



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