

# Comparing Left Atrial Strain in Sinus versus Atrial Fibrillation Patients

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## Abstract

**Background:** Assessment of left atrial (LA) function and its variability among patients with sinus rhythm and atrial fibrillation (AF) has been the subject of research recently. **Aim:** This research investigated whether LA function parameters by means of tissue Doppler imaging (TDI) were comparable in atrial fibrillation patients and patients with normal sinus rhythm. **Subjects and methods:** This observational study was done in Benha University Hospital, from March 2022 to January 2023, included 200 patients to study difference in LA parameters between the patients with sinus rhythm and AF patients. All subjects underwent history taking, clinical examination, Electrocardiogram and Echocardiography with TDI values. **Results:** We included (I)100 patients with sinus rhythm and (II)100 atrial fibrillation patients in the study (mean age  $57.7 \pm 13.6$  years, mean body mass index  $25.21 \pm 4.12$  kg/m<sup>2</sup>, 44% males). Patients of group II had significantly higher LA volumes and LA Volume index (P values=.0001 each), patients of group I had higher LA emptying fraction (LAEF) (P value=.006). LA  $\Sigma$  Strain and Strain rate were lower among patients of group II (P value=.0001 each). LAEF was positively correlated with LA  $\Sigma$  strain rate (r =0.226) (P value .001). LA volume index was negatively correlated with LA  $\Sigma$  strain and strain rate (r =-0.35, -0.207), (P values .0001, .003) respectively. **Conclusion:** LA function parameters were significantly lower among patients with AF. LAEF was positively correlated with LA  $\Sigma$  strain rate. LA volume index was negatively correlated with LA  $\Sigma$  strain and strain rate.

**Keywords:** left atrium, strain, atrial fibrillation, tissue Doppler imaging.

## Introduction

Left atrial function has been the topic of research for many years<sup>(1)</sup>. The left atrium has five main functions. First, the left atrium serves as a blood-receiving reservoir chamber. Second, it is a contractile chamber that by presystolic

kick helps complete left ventricular filling. Third, the left atrium works as a conduit that empties its contents into the left ventricle down the pressure gradient after the mitral valve opens. Fourth, it is a sensor of blood volume in the heart and

releases atrial natriuretic peptide (ANP) in response to stretch so that ANP- induced diuresis can help keep blood volume normal. Fifth, the left atrium contains receptors for the afferent nerves of various reflexes<sup>(2,3)</sup>.

Studies suggest that tissue Doppler-derived parameters of LA function may provide clinically helpful data. tissue Doppler has allowed the study of LA strain and strain rate<sup>(2,4)</sup>.

Tissue Doppler echocardiography-derived analysis of LA performance (namely LA strain) provides a window on all phases of LA function (reservoir, conduit, and booster pump) and has shown prognostic significance in different pathological conditions, and the results obtained with this method are not directly comparable to those obtained by volumetric approaches<sup>(5)</sup>.

If LA strain is to become accepted for evaluation of LA performance, then a standardized methodology, common reference values, and the impact of different diseases on LA function need to be precisely described<sup>(6)</sup>.

The clinical value of end-systolic LA volume assessed in 2- and 4-chamber view (normally  $\leq 34$  mL/m<sup>2</sup>) has been highlighted as a key prognostic marker<sup>(7,8)</sup>. and is a component of the parameters for the evaluation of LV diastolic function in current guidelines<sup>(9)</sup>.

The parameters described before<sup>(10)</sup> have been studied in the research context by many but are not widely used in the clinical setting. Those researchers<sup>(10)</sup> are to be congratulated for their effort to demonstrate the robustness and the feasibility of LA strain indices in many subjects.

Their results are consistent with a recent Task Force from the American Society of

Echocardiography that collaborated on standardization for measuring LA strain. The role of LA strain has also been underscored in a recent European Association of Cardio-Vascular Imaging recommendation paper on the exploitation of imaging in atrial fibrillation<sup>(11)</sup>.

This endorsement is supported by the growing number of studies, which have looked at LA strain for the evaluation of LA remodeling, thromboembolic risk, and prognosis and management in patients with atrial fibrillation. Nevertheless, these studies are often hindered by the small sample size and by several technical and methodological drawbacks<sup>(11,12)</sup>.

There are several important limitations regarding left atrial strain-derived assessment of left atrial function that should be highlighted:

- left atrial strain is load dependent and influenced by LV function. These limits have been highlighted, and it has been proposed that instead of focusing on the reservoir function, researchers should look more carefully at the booster pump function. This approach has not been widely accepted but worthy of consideration in upcoming studies<sup>(14)</sup>.
- The value of left atrial strain in the field of supraventricular arrhythmias. Left atrial strain is a predictor of atrial fibrillation incidence and recurrence, and its value seems to be in collaboration with thromboembolic risk.

Larger randomized trials are needed to confirm the association with thromboembolic risk and to assess whether LA strain can be used to risk-stratify population<sup>(15,16)</sup>.

- The potential value of LA strain in the evaluation of LV diastolic function and filling pressures <sup>(17)</sup>. The reduction in LV filling pressures decreases but scarcely normalizes LA volumes. There seems to be a strong relation between reduction in LV filling pressure and improvement in LA function as indicated by the improvement in LA strain, and this may be of use not only clinically, but also from a research perspective.

## Patients and methods

This observational study was done in Benha University Hospital in the period from March 2022 to January 2023, and included 200 patients divided into group I of patients with normal sinus rhythm and group II of AF patients.

The Ethics Committee of Benha Faculty of Medicine approved the study protocol {MS. 37.1.2022}. All individuals gave their consent to participate in this research.

### Inclusion criteria:

- Consecutive adult male and female patients.
- Patients with Ejection Fraction (EF) more than 50%.

### Exclusion criteria

- Patients with documented ischemic heart disease.
- Patients with prior PCI or CABG.
- Patients with rheumatic heart disease.
- Patients with significant (more than moderate) degenerative valvular heart disease.

### The following data were collected:

A. *Patients Characteristics:* age, Sex and Body mass index {BMI= Weight (Kg) / (Height in meters) 2} and risk factors

including HTN, DM, dyslipidemia, drugs used and smoking.

B. *Investigations including ECG:* for rhythm, whether sinus or AF, presence of ischemic changes and chamber enlargement signs and laboratory investigations including complete blood count, blood sugar, serum creatinine, urea, lipid profile (cholesterol, triglycerides, LDL & HDL).

C. *Conventional transthoracic 2D Echocardiography:*

Patients were examined in the left lateral decubitus with a commercially available ultrasound system (Philips, Epic 7, equipped with a 5.5 X transducer).

- Left ventricular ejection fraction (LVEF), that was measured using M-Mode in left para sternal long axis view <sup>(18)</sup>.
- Pulsed wave Doppler across the mitral valve in the apical 4-chamber was used to assess peak (E), and late (A) velocities. The E/e' ratio was then calculated <sup>(19)</sup>.
- Volumetric Measurements of LA:
- LA maximal volume (Vmax) at end-systole, just before the mitral valve opening; LA minimal volume (Vmin) at end-diastole, after mitral valve closure <sup>(20)</sup>.
- LA emptying fraction was calculated according to the following formula =  $\{(LA \text{ maximum volume} - LA \text{ minimum volume}) / LA \text{ maximum volume} \times 100\%$
- The LA size was represented by LA volume index {indexed by body surface area} by the following formula =  $(0.85 \times A1 \times A2) / (L1-L2 / 2)$  <sup>(21)</sup>.
- Where:
- A1 = planimetered LA area in apical 4-chamber view

- A2 = Max. planimetered LA area in apical 2-chamber view
- L1 = the LA long-axis length determined in apical 4-chamber view
- L2 = the LA long-axis length determined in apical 2-chamber view
- Measurements were repeated 3 times in each individual, and the average was used for analysis.
- We defined LA enlargement as a LA volume index of  $\geq 32 \text{ mL/m}^2$  (22).
- The accuracy of this echocardiographic technique has been validated previously against the computed tomography and magnetic resonance imaging methods (23).

#### D. Tissue Doppler imaging:

- Tissue Doppler imaging (frame rate range: 75-150/s) was obtained in apical 4- and 2-chamber views mid and basal during end-expiration. The Doppler tissue signal angle was less than 20 degrees. Digitally stored loops of Doppler tissue imaging were used for offline derivation of tissue tracking images and strain rate recordings (24).
- Strain was defined as a dimensionless quantity produced by the application of stress, and it represents the fractional or percentage change from the original unstressed dimension. Strain rate is equal with the temporal derivative of strain, and the total strain can, therefore, be determined by combining the strain rate values for a given time interval (25).
- The strain rate is calculated as the difference between myocardial velocities from 2 points divided by the distance separating the points (26,27).
- **Statistical analysis:**
- Statistical analysis was conducted using statistical pack of social studies

(SPSS) 25<sup>th</sup> edition, quantitative variables were presented in mean and standard deviation, it was compared between study groups using Mann Whitney U test. Categorical variables were presented in frequency and percentages and compared using the Chi<sup>2</sup> test. The receiver curve of characteristic curve was constructed to visualize the area under the curve for predicted outcomes. All P values were two sided. P values less than .05 were considered significant (28).

## Results

- The mean age was  $57.7 \pm 13.6$  years and mean BMI  $25.21 \pm 4.12 \text{ kg/m}^2$ . 88 patients (44%) were males. Dyslipidemia was the most common comorbidity reported in the current study accounting for 114 patients (57%), followed by Diabetes reported in 96 patients (48%). 39 patients (19.5%) were smokers while 81 patients (40.5%) were hypertensive, 43 patients (21.5%) had positive family history for cardiac diseases (**table 1**).
- Among the included patients, ejection fraction showed a mean of 60.8%, mean LVEDD 4.6 cm, mean LVESD 2.7 cm, while mean LA Vmax was 76.6 mL, LA Vmin was 44.8 mL, LA volume index is  $30.1 \text{ mL/m}^2$ , and mean LA emptying fraction is 30.0 (**table 2**).
- Patients of group II had significantly lower EF than those of group I (P value=.0001) and had significantly higher LA Vmax, LA Vmin and LA Volume index (P values=.0001 each), on the other hand patients of group I had higher LA emptying fraction (P value =.006) (**table 3**).

- LA  $\Sigma$  Strain and LA  $\Sigma$  Strain rate were significantly lower among patients of group II than patients of group I (P value 0.0001 each) (**table 4**)
- Correlation matrix showed that LA emptying fraction was positively correlated with LA  $\Sigma$  strain rate with (r = 0.226) and (P value 0.001). While LA volume index was negatively correlated with LA  $\Sigma$  strain and  $\Sigma$  strain rate with (r = -0.35 and -0.207), (P values 0.0001 and 0.003) respectively (**table 5**).

**Table 1:** comparison of demographics and clinical characteristics between study groups.

		<i>Group I (Sinus rhythm)</i>		<i>Group II (AF)</i>		<i>P value</i>
		<i>Mean ± SD / Count</i>	<i>Range / %</i>	<i>Mean± SD / Count</i>	<i>Range / %</i>	
<i>Age</i>	<i>Years</i>	57.2 ±12.7	26 -94	58.1 ±14.5	26 -94	0.867
<i>BMI</i>	<i>Kg/m<sup>2</sup></i>	25.24 ±4.04	17.2 -34.5	25.19 ±4.22	16.5 -35.7	0.866
<i>Sex</i>	<i>Male</i>	46	46%	42	42%	0.569
	<i>Female</i>	54	54%	58	58%	
<i>Diabetes</i>	<i>Yes</i>	49	49%	47	47%	0.777
<i>Smoking</i>	<i>Yes</i>	18	18%	21	21%	0.801
<i>Dyslipidemia</i>	<i>Yes</i>	56	56%	58	58%	0.775
<i>Hypertension</i>	<i>Yes</i>	40	40%	41	41%	0.823
<i>Family History</i>	<i>Positive</i>	22	22%	21	21%	0.863

**Table 2:** 2D Echocardiography findings among the included patients (n=200).

	<b>Mean</b>	<b>SD</b>	<b>Min</b>	<b>Max</b>
<b>Ejection fraction (%)</b>	60.8	3.8	55.2	73.4
<b>LVEDD (cm)</b>	4.6	0.4	4.0	5.5
<b>LVESD (cm)</b>	2.7	0.3	2.0	3.5
<b>LVPWDD (cm)</b>	1.1	0.1	0.7	1.4
<b>IVSDD (cm)</b>	1.1	0.1	0.7	1.4
<b>LA Vmax (mL)</b>	76.6	8.6	61.1	92
<b>LA Vmin (mL)</b>	44.8	6.4	33.1	57.8
<b>LA Volume index (mL/m<sup>2</sup>)</b>	30.1	5.4	20	42.8
<b>LA emptying fraction</b>	30.0	5.4	16.8	42.8

**Table 3:** comparison of 2D echocardiography findings between study groups.

	Group I (Sinus rhythm)				Group II (AF)				P value
	Mean	SD	Min	Max	Mean	SD	Min	Max	
Ejection fraction (%)	62.9	3.8	56.7	73.4	58.7	2.4	55.2	69.5	.0001
LVEDD (cm)	4.5	0.3	4.0	5.0	4.8	0.4	4.2	5.5	.0001
LVEDS (cm)	2.5	0.3	2.0	3.0	2.8	0.4	2.2	3.5	.0001
LVPWDD (cm)	1.0	0.1	0.8	1.3	0.9	0.1	0.7	1.2	0.0016
IVSDD (cm)	1.0	0.1	0.8	1.3	0.9	0.1	0.7	1.2	0.0015
LA Vmax (mL)	73.3	8.2	61.1	87.2	80	7.5	67	92	.0001
LA Vmin (mL)	41.5	4.9	33.1	50	48.2	6	38.1	57.8	.0001
LA Volume index (mL/m <sup>2</sup> )	27.4	4.6	20	38	32.8	4.7	20	42.8	.0001
LA emptying fraction	43.2	9.2	22.6	69.9	39.1	10.0	17.4	57.6	.006

Table 4: comparison of LA strain by TDI between study groups.

	Group I (Sinus rhythm)				Group II (AF)				P value
	Mean	SD	Min	Max	Mean	SD	Min	Max	
TDI LA $\Sigma$ Strain (%)	24.3	3.4	18.6	30	19.7	2.7	15.7	25.1	.0001
TDI LA $\Sigma$ Strain rate (sec <sup>-1</sup> )	1.9	0.3	1.4	2.3	1.7	0.3	1.3	2.1	.0001

Table 5: correlation matrix between 2D echo parameters and LA strain and strain rate.

		<i>LA <math>\Sigma</math> Strain</i>	<i>LA <math>\Sigma</math> Strain rate</i>
<b>LA emptying fraction</b>	r	.082	.226**
	P value	.246	.001
<b>LA Volume index</b>	r		-.207**
	P value	.0001	.003

## Discussion

Two dimension (2D) echocardiography remains to be the most used imaging modality for evaluation of LA parameters and its primary variables<sup>(29)</sup>. Tissue Doppler imaging (TDI) is a useful echocardiographic technique; which is readily available on most commercially echocardiographic systems; to evaluate global and regional myocardial systolic as well as diastolic function<sup>(30)</sup>.

The aim of our study was to assess left atrial strain in patients with sinus rhythm and atrial fibrillation using tissue Doppler

imaging. Patients in this study were divided into two groups:

- Group (I): 100 patients with normal sinus rhythm.
- Group (II): 100 patients with atrial fibrillation (Table 1).

In our study patients of group II, have significantly lower LVEF than those of group I, and have significantly higher LVEDD, LVESD (P values .0001 each), which was consistent with a similar study in 2019<sup>(31)</sup>. Additionally, a group of scientists later the same year<sup>(32)</sup> reported that patients with AF had significant

higher LVESD than patients without AF (Table 2).

Moreover, another trial<sup>(33)</sup> which studied difference in LVEF, LVEDD, and LVESD between patients with AF and patients who restored sinus rhythm successfully came to a same conclusion (table 2,3).

In the current study, patients of group II have significantly higher LA Vmax, LA Vmin and LA Volume index (P values .0001 each), which was similar to the results concluded by a group of scientists in 2018<sup>(34)</sup> with similar demographic characters. This was also similar to a single-center study in 2020<sup>(35)</sup> which reported a significant reduction in LA Volume index in patients who successfully restored sinus rhythm after ablation, and another larger trial in 2020<sup>(36)</sup> which reported that LA dilatation is a strong, independent predictor of worse outcome (table 2,3).

It was reported by two independent groups of researchers<sup>(37, 38)</sup> that patients with AF had lower LA emptying fraction than patients without AF with P value .013, .0051 respectively, which was similar to what we found in our study with P value .006 (table 2, 3).

To our knowledge, there is no study that has reported a correlation between LA emptying fraction and LA volume index with LA  $\Sigma$  strain and  $\Sigma$  strain rate.

## Conclusion

LA parameters including strain and strain rate, assessed by TDI, were significantly lower among patients with AF than patients with sinus rhythm. LA emptying fraction was positively correlated with LA  $\Sigma$  strain rate. LA volume index was negatively correlated with LA  $\Sigma$  strain and strain rate.

Notably, it was concluded in 2020<sup>(39)</sup> that patient who developed AF had significantly larger LA volumes and lower LAEF than participants free of AF which was like what we found in our study. Also, a similar trial<sup>(40)</sup> found similar result as patients with AF had lower LAEF compared with patients without AF (table 2, 3).

In the current study, LA  $\Sigma$  Strain and LA  $\Sigma$  Strain rate were significantly lower among patients of group II than patients of group I (P value 0.0001 each), this was similar to the results clarified in 2015 by a group of Japanese researchers<sup>(41)</sup> who found that patients with AF had significantly lower LA  $\Sigma$  Strain and LA  $\Sigma$  Strain rate than patients without AF (table 4).

Notably, it was attributed in 2016<sup>(5)</sup> that patient with AF had significantly lower LA  $\Sigma$  Strain than people with sinus rhythm, this was concordant to our study where LA  $\Sigma$  Strain rate was significantly lower among patients of group II than patients of group I (P value .0001). Also, it was recorded in 2017<sup>(42)</sup> that LA  $\Sigma$  Strain was lower among patients with persistent AF than patients with paroxysmal AF.

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