



CARDIOIMAGING NEL DIABETE MELLITO ASINTOMATICO

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Disclosures

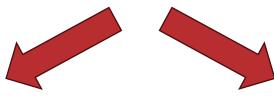
SOBI Srl (July 2021)

BACKGROUND

DIABETES



<u>High prevalence</u>: 1/10 worldwide (undiagnosed in > 1/3) <u>Increased risk</u> (2-5 fold) of **Major CV events** (HF, AF/stroke, CAD)

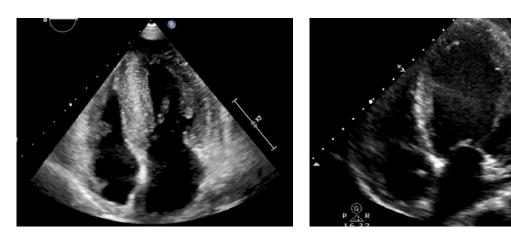


Cardiac dysfunction

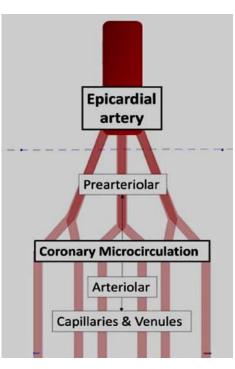
54% Diastolic Dysfunction (RR 3.74 for death)
17% Diabetic Cardiomyopathy (RR 5.06 for death) (overt systolic dysfunction or at least moderate diastolic dysfunction)

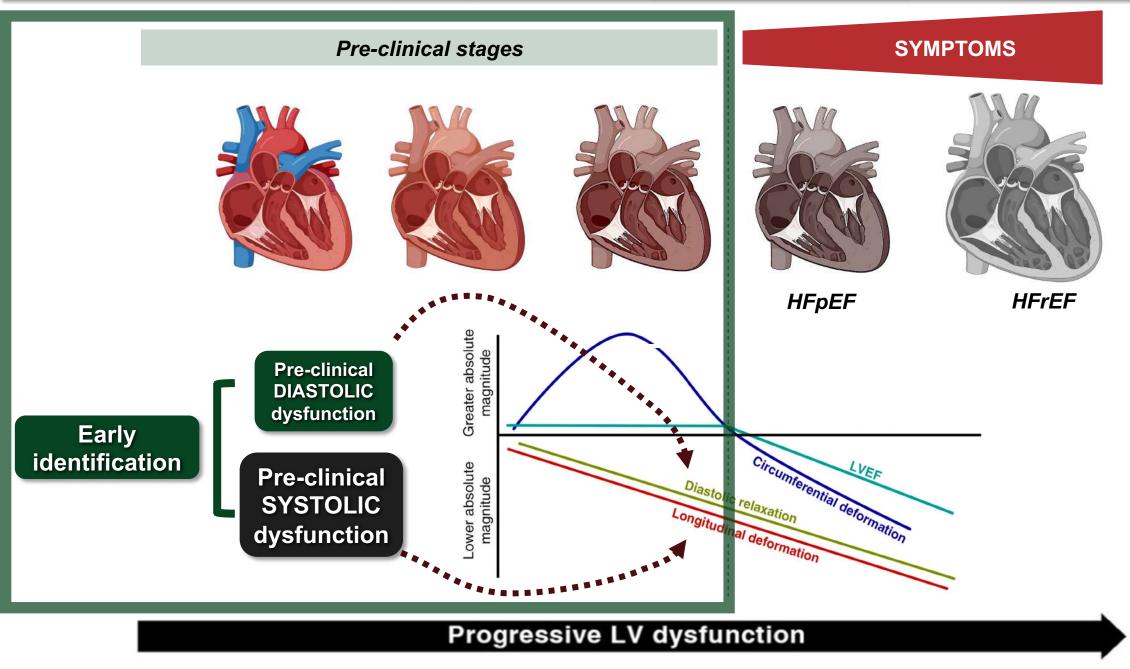
Myocardial ischemia (CAD / INOCA)

23% >moderate (>50%) coronary artery stenosis20-58% perfusion defects on SPECT



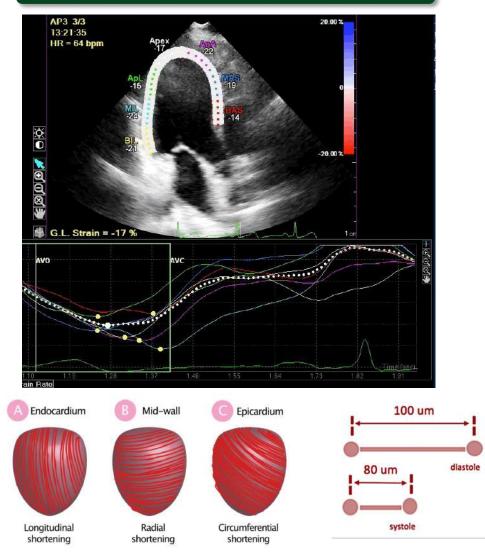
What's the role of CARDIAC IMAGING in <u>asymptomatic</u> DM patients ?

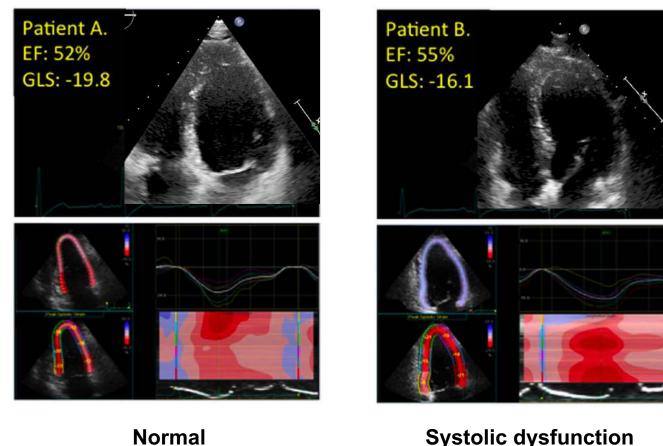




Voigt et al, JACC 2019; Cikes et al, EHJ 2016

Speckle Tracking Echocardiography





Systolic dysfunction

Global Longitudinal Strain (GLS) vs. Ejection Fraction (EF)

- Higher **reproducibility** than Echocardiography (r=0.93 vs. cardiac magnetic resonance) ٠
- Higher **prognostic value** for major CV events (AUC 0.82 vs. 0.72) ٠

Table 1 Clinical characteristics

	Cardiac risk factors and prevention
ORIGINAL ARTICLE	
Subclinical LV dysfunction an in type 2 diabetes mellitus	d 10-year outcomes
David J Holland, ^{1,2,3} Thomas H Marwick, ⁴ Bria	n A Haluska, ¹ Rodel Leano, ¹
(mean 66.3±6.9) assessed w echocardiography	atients with normal EF vith <u>speckle tracking</u>

	Normal (n=126)	LVD (n=104)	p Value
Clinical variables			
Age (years)	56±10	56±10	0.834
Male sex	56 (44%)	70 (67%)	0.001
Body mass index (kg/m ²)	31.7±6.0	31.9±5.5	0.844
Waist-to-hip ratio	0.94±0.08	0.97±0.09	0.004
Systolic BP (mm Hg)	130±13	132±14	0.228
Diastolic BP (mm Hg)	79±8	81±7	0.019
Aerobic capacity (METs)	6.4±2.0	6.3±1.9	0.943
Biochemical variables			
Total cholesterol (mmol/L)	4.82±1.09	4.88±0.95	0.663
eGFR (mL/min/1.73m ²)	114±36	120±45	0.316
Fasting glucose (mmol/L)	8.36±3.08	8.65±3.28	0.490
HbA _{1c} (%)	7.5±1.5	7.7±1.6	0.291
Echocardiography			
EF (%)	66.7±6.7	65.9±7.0	0.337
EDV (mL)	66.1±16.7	67.7±20.7	0.528
ESV (mL)	22.3±7.6	23.7±10.3	0.236
Left atrial area (cm ²)	18.5±3.6	18.9±4.0	0.503
LV mass index (g/m ^{2.7})	42±14	44±13	0.396
E (cm/s)	68.4±14.7	66.9±15.2	0.481
A (cm/s)	68.0±14.7	66.9±17.7	0.338
E/A	1.04±0.26	1.02±0.37	0.674
Sm (cm/s)	6.9±1.4	6.7±1.1	0.184
Em (cm/s)	6.5±1.6	6.0±1.7	0.049
E/Em	11.0±2.9	11.8±3.6	0.099
GLS (%)	-20.8±1.5	-16.6±1.9	<0.001
Strain rate (s ⁻¹)	-1.12±0.13	-0.98±0.13	<0.001
Medication use			
Use of metformin	72 (57%)	69 (66%)	0.191
Use of ACE inhibitors	40 (32%)	26 (25%)	0.231
Use of β-blockers	5 (4%)	4 (4%)	0.945

Data are mean±SD or n (%). A, late diastolic transmitral flow; BP, blood pressure; E, early diastolic transmitral flow; EDV, end diastolic volume; eGFR, estimated glomerular filtration rate; Em, early diastolic tissue velocity; ESV, end systolic volume; GLS, global longitudinal strain; HbA_{1c}, glycosylated haemoglobin; LVD, LV dysfunction; METs, metabolic equivalents; Sm, systolic tissue velocity.

Table 2 Statistical predictors of the primary endpoint

ORIGINAL ARTICLE

Subclinical LV dysfunction and 10-year outcomes in type 2 diabetes mellitus

Cardiac risk factors and prevention

David J Holland, ^{1,2,3} Thomas H Marwick,⁴ Brian A Haluska,¹ Rodel Leano,¹

236 asymptomatic DMT2 patients with normal EF (mean 66.3±6.9) assessed with speckle tracking echocardiography

Pre-clinical left ventricular dysfunction in 45% (GLS < -18.9%)

...increased risk of all-cause mortality and hospitalization

		Univa	riate anal	ysis					Multivari	ate analys	is	
		HR (9	5% CI)			p Valu	2		HR (95%	CI)		p Val
linical												
Age		1.04 (1.01 to 1.0	7)		0.004	*		1.04 (1.00) to 1.07)		0.025
Male sex		0.90 (0.56 to 1.4	6)		0.678						
Height (m)		0.98 (0.97 to 1.0	2)		0.824						
Weight (kg)		1.01 (1.00 to 1.0	2)		0.148						
Body mass index (kg/m ²)		1.03 (1.00 to 1.0	7)		0.083						
Systolic BP (mm Hg)												
Exercise capacity (METs)	Ĩ											
liochemical	1.0-	S		-								
Total cholesterol (mmol/L)												
eGFR (mL/min/1.73m ²)				1	· · · · ·	_						
Fasting glucose (mmol/L)					7		~~					
HbA _{1c} (%)	0.8-				14-1	٦.	~	<u> </u>			Normal	
chocardiography	0.0					-۱.	1	-		~_	Normai	
EF (%)†	_											
GLS (%)	No.						(19)					
*Multivariate analysis performe	<u>.</u> <u>S</u>										-	
tEF was not a significant indep	Un 0.6-										I	
BP, blood pressure; eGFR, estim	0											
	ee										LVD	
	Event-free Survival											
	t 0.4-											
	ş					Log ra	ank χ²=	4.73				
	ш											
						р	=0.030)				
	0.2-											
		100	405	100	447		105	100	00	07	06	Narmal
	0.0-	126	125	122	117	111	105	100	99	97	96	Normal
	0.0	104	103	102	93	82	77	72	70	68	66	
		ò	1	2	3	4	5	6	ł	8	9	10
						Follo	11 un (years)				

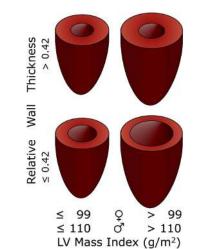
Longitudinal Myocardial Strain Alteration Is Associated with Left Ventricular Remodeling in Asymptomatic Patients with Type 2 Diabetes Mellitus

Laura Ernande, MD, PhD, Cyrille Bergerot, MD, Nicolas Girerd, MD, Hélène Thibault, MD, PhD,

172 asymptomatic DMT2 patients with normal EF (mean 70±7) assessed with <u>speckle tracking</u> <u>echocardiography</u>

Pre-clinical left ventricular dysfunction in **23%** (GLS < -18%)

...associated with higher relative wall thickness and end-systolic volume



Variable	Total population $(n = 154)$	Normal strain (ε _L ≥ 18%) (<i>n</i> = 118)	Altered strain ($ \varepsilon_L < 18\%$) (n = 36)	P*
Clinical characteristics				
Age (y)	58 ± 8	58 ± 8	57 ± 8	.51
Men	88 (57%)	60 (51%)	28 (78%)	.003
BMI (kg/m ²)	29.5 ± 4.4	29.1 ± 4.3	30.6 ± 4.9	.09
Diabetes duration (y)	13 ± 8	13 ± 8	13 ± 7	.84
Treated hypertension	80 (52%)	58 (49%)	22 (61%)	.11
Dyslipidemia	90 (58%)	71 (60%)	19 (53%)	.30
Current smokers	26 (17%)	18 (15%)	8 (22%)	.21
Peripheral artery disease	51 (33%)	39 (33%)	12 (33%)	.37
Retinopathy	33 (21%)	25 (21%)	8 (22%)	.48
Systolic blood pressure (mm Hg)	132 ± 16	131 ± 16	135 ± 18	.20
Heart rate (beats/min)	75 ± 11	74 ± 12	78 ± 10	.07
Medications	10 - 11		10 = 10	
Metformin	110 (71%)	87 (74%)	23 (64%)	.21
Sulfonylureas	64 (42%)	55 (47%)	9 (25%)	.02
Glitazones	33 (21%)	28 (24%)	5 (14%)	.02
Insulin	70 (45%)		and the second	.07
ACE inhibitors or ARBs		50 (42%)	20 (56%)	.07
	89 (58%)	61 (52%)	28 (78%)	
Statins	89 (58%)	69 (58%)	20 (56%)	.52
Antiplatelet agents	45 (29%)	32 (27%)	13 (36%)	.18
Biologic characteristics		70.40		
HbA _{1c} (%)	7.7 ± 1.3	7.8 ± 1.3	7.4 ± 1.4	.17
Triglycerides (mmol/L)	1.8 ± 1.4	1.8 ± 1.4	1.8 ± 1.4	.93
Total cholesterol (mmol/L)	4.7 ± 1.1	4.8 ± 1.1	4.5 ± 1.0	.17
eGFR (mL/min/1.73 m ²)	83 ± 19	83 ± 22	83 ± 19	.96
Microalbuminuria (mg/L)	24 (11–69)	18 (10–60)	42 (15–148)	.045
Echocardiographic characteristics				
LV dimensions				
Total LV wall thickness (mm)	20 ± 3	20 ± 3	21 ± 2	.12
LV mass index (g/m ²)	93 ± 18	92 ± 19	95 ± 17	.48
RWT	0.41 ± 0.07	0.40 ± 0.07	0.44 ± 0.06	.008
2D/speckle-tracking imaging				
LVEDV (mL)	79 ± 22	77 ± 20	85 ± 24	.08
LVEDVi (mL/m ²)	42 ± 10	79 ± 22	79 ± 22	.50
LVESV (mL)	24 ± 9	23 ± 9	28 ± 11	<.001
LVESVi (mL/m ²)	13 ± 5	12 ± 4	14 ± 5	.01
LVEF (%)	70 ± 7	71 ± 7	67 ± 7	.002
ε _L (%)	-19.8 ± 2.4	-20.8 ± 1.8	-17.7 ± 1.2	-
LV diastolic function				
E/A ratio	1.0 ± 0.2	1.0 ± 0.3	1.0 ± 0.2	.63
mDT (msec)	240 ± 54	240 ± 51	239 ± 61	.00
E/e' ratio	9.7 ± 2.7	9.8 ± 2.8	9.3 ± 2.6	.32
	46 ± 13	9.6 ± 2.8 45 ± 13	9.3 ± 2.0	.32
LA volume (mL)	40 ± 13	40 ± 13	47 王 14	.42

Ernande, ASE 2014

Longitudinal Myocardial Strain Alteration Is Associated with Left Ventricular Remodeling in Asymptomatic Patients with Type 2 Diabetes Mellitus

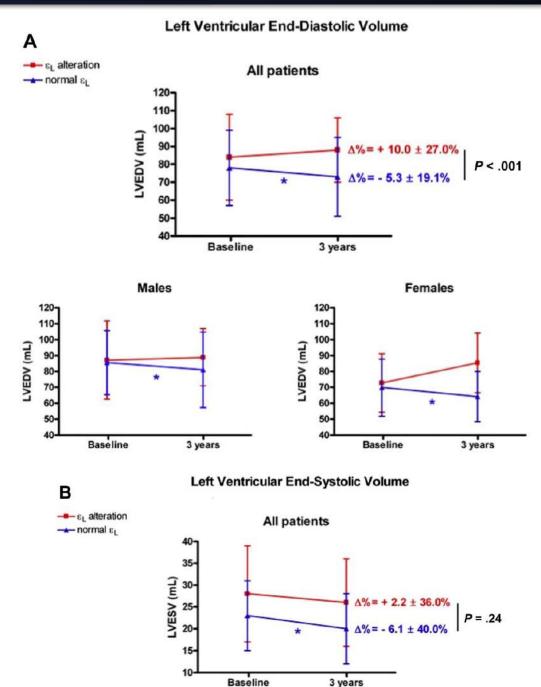
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172 asymptomatic DMT2 patients with normal EF (mean 70±7) assessed with <u>speckle tracking</u> <u>echocardiography</u>

Pre-clinical left ventricular dysfunction in **23%** (GLS < -18%)

...associated with higher relative wall thickness and end-systolic volume

...associated with adverse LV remodeling at 3 years



The International Journal of Cardiovascular Imaging (2023) 39:977–989 https://doi.org/10.1007/s10554-023-02810-4

ORIGINAL PAPER

Subclinical systolic dysfunction detected by 2D speckle tracking echocardiography in adults with diabetes mellitus: systematic review and meta-analysis of 6668 individuals with diabetes mellitus and 7218 controls

Seyed-Mohammad Ghoreyshi-Hefzabad¹ · Prajith Jeyaprakash^{1,2} · Ha Q. Vo^{2,3} · Alpa Gupta¹ · Koya Ozawa¹ · Faraz Pathan^{1,3} · Kazuaki Negishi^{1,2,3,4}

6668 asymptomatic DM patients compared with 7218 controls, assessed with speckle tracking echocardiography

...lower LV strain, LA reservoir strain, RV strain

Strain variable	Studies (n)	DM (n)	Control (n)	Mean [95% CI] in DM	Mean [95% CI] in Control	MD [95% CI] Ran- dom Effects model
LVGLS	32	6114	6729	17.9 [17.1, 18.4]	19.8 [19.1, 20.5]	- 1.98 [- 2.46, - 1.51]
LVGCS	14	1626	3549	20.3 [18.6, 21.9]	21.3 [19.6, 22.9]	- 0.96 [- 1.48, - 0.45]
LVGRS	9	529	400	42.7 [39.7, 45.6]	47.0 [43.1, 50.9]	- 4.0 [- 5.50, - 2.52]
LVSR	13	1029	924	1.0 [0.9, 1.1]	1.1 [1.0, 1.2]	- 0.07[- 0.13, - 0.02]
LA reservoir strain	7	543	428	28.0 [24.4, 31.6]	36.5 [34.0, 39.0]	- 8.42[- 11.6, 5.25]
RVGLS	7	341	311	23.8 [20.1, 27.4]	26.0 [23.4, 28.6]	- 2.38 [- 4.67, - 0.09]

Ghoreyshi-Hefzabad, Intl Journal Cardiovasc Imaging 2023

Study		Experimental Mean SD	Total Me	Control an SD	Mean	Difference	MD	95%-CI	Weight (fixed)	Weight (random)
Nakai,2009	60	17.60 2.6000	25 20.	80 1.8000		<u> </u>	-3.20	[-4.16; -2.24]	0.6%	2.4%
NG.2009		18.30 2.2000		90 1.9000				[-2.41; -0.79]	0.8%	2.6%
Ernande,2011		19.00 3.0000		00 2.0000				[-3.69; -2.31]	1.1%	2.8%
D'Andrea.2012		15.40 3.3000		50 3.5000		_		[-6.61; -3.59]	0.2%	1.8%
Cognet 2013	28	13.90 3.7000	35 15.	00 4.5000			1.10	[-3.13; 0.93]	0.1%	1.4%
Conte-1,2013		20.30 2.6000		90 1.3000				[-1.53; 0.33]	0.6%	2.5%
Conte-2,2013	27	19.00 2.0000	24 20.	90 1.3000				[-2.82; -0.98]	0.6%	2.5%
Zoroufian,2014	39	15.10 2.9000	37 18.	30 1.7000				[-4.26; -2.14]	0.5%	2.3%
Jensen,2015	1065	18.30 2.6000	198 18.	80 2.5000				[-0.88; -0.12]	3.6%	3.0%
Karagov,2015	82	17.90 2.7000	90 21.	10 3.2000			-3.20	[-4.08; -2.32]	0.7%	2.5%
Skali,2015	1322	17.80 2.4000	1742 18.	50 2.3000			-0.70	[-0.87; -0.53]	18.5%	3.1%
Tadic-1,2015	57	19.00 1.6000	54 21.	00 2.0000			-2.00	[-2.68; -1.32]	1.1%	2.8%
Tadic-2,2015	50	20.90 1.7000	50 21.	70 1.8000		<u> </u>	-0.80	[-1.49; -0.11]	1.1%	2.8%
Abdel-Salem,2016	30	17.70 2.5000	30 21.	20 1.7000	-		-3.50	[-4.58; -2.42]	0.4%	2.3%
Bakhum,2016	60	16.87 3.4100	30 19.	20 2.0200			-2.33	[-3.46; -1.20]	0.4%	2.3%
Jedrzejewska 2016	50	20.30 2.0000	50 22.	20 1.8000		-}+	-1.90	[-2.65; -1.15]	0.9%	2.7%
Jorgensen,2016	770	14.10 2.4000	234 15.	00 2.0000			-0.90	[-1.21; -0.59]	5.6%	3.1%
Loncarevic,2016	70	17.36 1.8000	80 18.	71 1.8600				[-1.94; -0.76]	1.5%	2.9%
Mochizuki-1,2016	137	19.20 2.6000	69 20.	70 2.1000		<u>++</u>	-1.50	[-2.16; -0.84]	1.2%	2.8%
Mochizuki-2,2016	61	17.00 2.7000	69 20.	70 2.1000	-		-3.70	[-4.54; -2.86]	0.7%	2.6%
Kishi-1,2017	368	14.10 2.5000	1485 15.	50 2.4000		*	-1.40	[-1.68; -1.12]	6.5%	3.1%
Kishi-2,2017	85	13.50 2.9000	1485 15.	50 2.4000				[-2.63; -1.37]	1.3%	2.8%
Suto,2017	145	18.00 2.6000	90 20.	30 2.0000		-++	-2.30	[-2.89; -1.71]	1.5%	2.9%
Vukomanovic,2017	50	18.50 2.0000	40 21.	10 2.8000			-2.60	[-3.63; -1.57]	0.5%	2.4%
Jorgensen-1,2018	57	15.30 2.4000	80 15.	90 2.0000			-0.60	[-1.36; 0.16]	0.9%	2.7%
Jorgensen-2,2018	349	14.60 2.8000	80 15.	90 2.0000			-1.30	[-1.83; -0.77]	1.9%	2.9%
Jorgensen-3,2018	345	14.00 2.8000	80 15.	90 2.0000		+	-1.90	[-2.43; -1.37]	1.9%	2.9%
Lin,2018	505	19.00 1.8000	1416 20.	50 1.9000			-1.50	[-1.69; -1.31]	15.2%	3.1%
NG,2018	337	17.60 2.3000	316 18.	90 2.4000			-1.30	[-1.66; -0.94]	4.0%	3.0%
Philouze,2018	44	20.20 2.7000	35 20.	80 2.3000		÷++	-0.60	[-1.70; 0.50]	0.4%	2.3%
Ringle,2018	66	18.90 2.0000	26 20.	50 2.0000			-1.60	[-2.51; -0.69]	0.6%	2.5%
Stevanovic,2018	121	17.50 2.1000	41 24.	40 2.4000	<u> </u>		-6.90	[-7.72; -6.08]	0.8%	2.6%
Tadic,2018	48	18.80 2.0000	44 20.	80 2.4000			-2.00	[-2.91; -1.09]	0.6%	2.5%
Bogdanovic-1,2019	67	19.60 0.4000	20 21.	90 0.4000		E28	-2.30	[-2.50; -2.10]	13.1%	3.1%
Bogdanovic-2,2019	20	21.30 0.4000	20 21.	90 0.4000			-0.60	[-0.85; -0.35]	8.5%	3.1%
Cameli,2019	52	16.70 3.7000	60 17.	90 3.7000			-1.20	[-2.57; 0.17]	0.3%	2.0%
Haley,2020	151	15.10 3.1000	146 18.	00 2.8000			-2.90	[-3.57; -2.23]	1.2%	2.8%
Roberts,2020	34	18.60 2.2000	17 19.	70 2.3000			-1.10	[-2.42, 0.22]	0.3%	2.0%
Fixed effect model	7002		8498			0	-1.45	[-1.52; -1.38]	100.0%	-
Random effects model						\$	-1.96	[-2.27; -1.64]		100.0%
Heterogeneity: $I^2 = 94\%$, τ	2 = 0.81	44, p < 0.01			0 6	1 1 1				

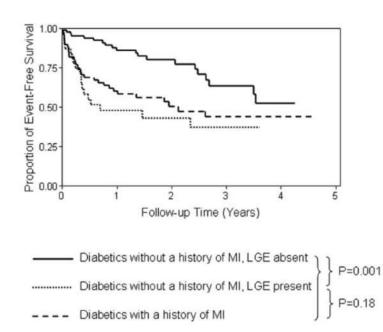
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Cardiac Magnetic Resonance

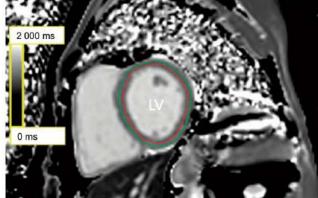
Focal fibrosis (ischemic LGE scar) - observed in 28% asymptomatic DM patients without clinical evidence of MI (normal ECG)

- associated with increased risk of MACE (HR 4.13) comparable with those with history of MI

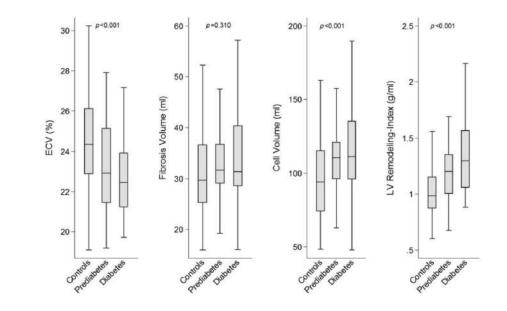


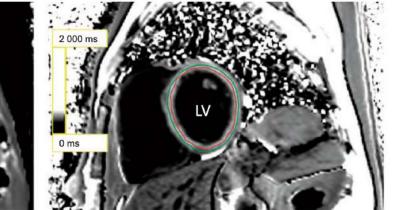


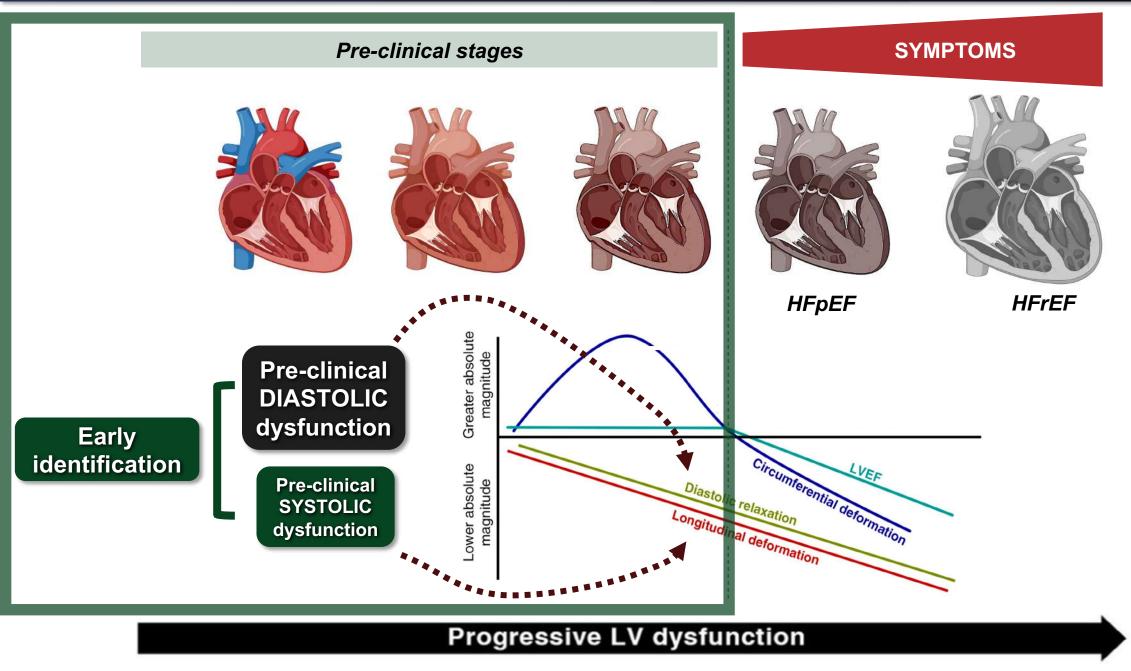




<u>Diffuse fibrosis (T1 mapping, ECV)</u> <u>decreased ECV in asymptomatic DM patients</u>, higher cell volume (consistent with early LV remodelling)







Voigt et al, JACC 2019; Cikes et al, EHJ 2016

Stress-Echocardiography

European Society of Cardiology doi:10.1093/ehjci/jeaa070

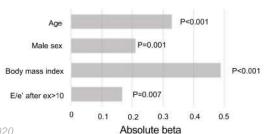
Incremental value of diastolic stress test in identifying subclinical heart failure in patients with diabetes mellitus

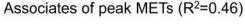
Tomoko Nishi^{1,2†}, Yukari Kobayashi 💿 ^{1,2†}*, Jeffrey W. Christle^{1,3}, Nicholas

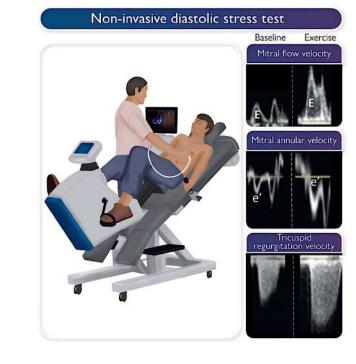
180 asymptomatic DMT2 patients assessed with stressechocardiography

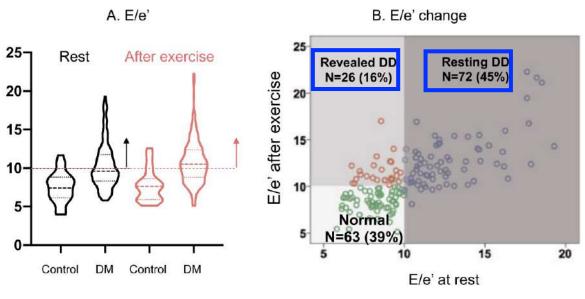
Diastolic dysfunction: at rest **45%**, after exercise \rightarrow **57%** ... associated with **impaired exercise capacity** (lower exercise peak METs)

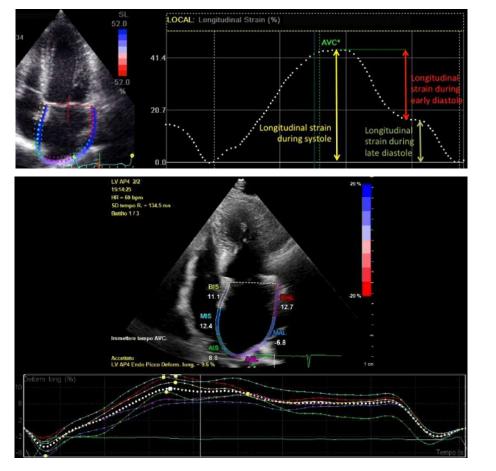
	Normal (N = 63)	Resting DD (N = 72)	Revealed DD ($N = 26$)	P-value
Peak METs	9.0 ± 2.6	7.3 ± 2.3*	7.3 ± 2.1*	<0.001
Percent-predicted METs (%)	105 ± 28	98 ± 29	91 ± 30	0.13
<85% ppMETs, n (%)	15 (26)	25 (37)	12 (52)	0.07





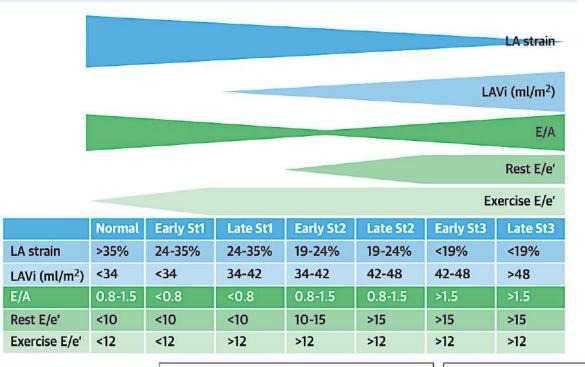


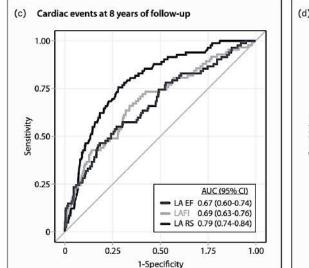


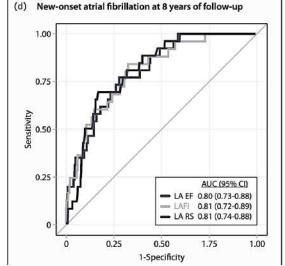


Left atrial deformation

CENTRAL ILLUSTRATION Evolution of LV Diastolic Parameters With Increasing Disease Severity







Left atrial deformation vs. traditional assessment of diastolic dysfunction

- More sensitive marker of early phases of DD, improved stratification of stage
- Independent predictor of <u>all-cause mortality and hospitalization</u>
- Independent predictor of <u>CV events (HR 2.10)</u> and <u>Atrial Fibrillation (HR 6.45)</u>

Α

CVD-Free Survival

8

06.0

0.80

0.70

09.0

0.50

1.00

06.0

0.80

0.70

0.60

0.50

HR 0.95 per %

Follow-up (years)

В

CVD-Free Survival



536 asymptomatic DM patients, assessed with cardiac magnetic resonance

Left atrial size and function impairment associated with incident CV events (Heart Failure, Myocardial infarction, Atrial Fibrillation, Stroke)

Impact of LA EDVi stronger than ESVi

(MESA) Timothy M. Markman¹, Mohammadali Habibi², Bharath Ambale Venkatesh³ Minimum LA Volume **Passive LA Emptying Fraction** С 8 <10 mm3/m2 0.00 Survival >10 mm3/m2 0.80 CVD-Free ≤14% 0.70 0.60 HR 1.12 per mm³/m² HR 0.97 per % 0.50 10 Follow-up (years) Follow-up (years) **Total LA Emptying Fraction** Active LA Emptying Fraction D 0.90 >25% CVD-Free Survival 0.80 ≤25% 0.70 09.0

0.50

ESC

European Society doi:10.1093/ehic/jew332

European Heart Journal - Cardiovascular Imaging (2017) 18, 1138-1144

Association of left atrial structure and function and incident cardiovascular disease in patients with diabetes mellitus: results from multi-ethnic study of atherosclerosis

HR 0.98 per %

Follow-up (years)

10

open Access Full Text Article

ORIGINAL RESEARCH

Increased Left Atrial Stiffness is Significantly Associated with Paroxysmal Atrial Fibrillation in Diabetic Patients

Diana-Aurora Arnautu (1)^{1,2}, Sergiu-Florin Arnautu (1)^{1,3}, Mirela-Cleopatra Tomescu (1)^{1,3},

60 DM patients (50% with paroxysmal atrial fibrillation), assessed with <u>speckle-tracking echocardiography</u>

LA stiffness (E/A ratio / LA reservoir) **had strongest association with paroxysmal atrial fibrillation** (OR=5.2)

Parameter	Univariate Analysis OR (95% CI)	P value	Multivariate Analysis OR (95% CI)	P value
LAS	0.73 (0.56–0.96)	0.02	0.58 (0.37–0.92)	0.02
LA-pool strain	0.78 (0.64–0.94)	<0.01	0.74 (0.58–1.95)	0.018
LA- pump strain	0.78 (0.64–0.94)	<0.01	0.68 (0.50–0.94)	0.019
LAsf	0.73 (0.56–0.96)	0.01	5.2 (4.05–7.17)	0.007

Abbreviations: OR, Odds ratio; CI, confidence interval; LAS, left atrial global strain; LASf, left atrial stiffness.

	Diabetes with PAF n = 30	Diabetes without PAF n = 30	P value	
LVEDV (mL)	97.8±21.0	102.5±17.9	0.34	
LVESD (mL)	46.1 ± 10.2	49.3±12.6	0.28	
LVMI (g/m2)	143 ± 41	139 ± 35	0.68	
LVEF (%)	57 ± 2	60 ± 14	0.25	
Mitral E/A-ratio	1.3 ± 0.4	1.2 ± 0.2	0.22	
Septal E/E' average ratio	13.9 ± 4.2	13.1 ± 3.6	0.43	
TAPSE (cm)	2.5 ± 0.43	2.4 ± 0.49	0.40	
GLS (%)	-14.4 ± 3.1	-15.2 ± 3.2	0.32	
LAVI (mL/m2)	42.1 ± 9.3	43.5 ± 10.3	0.58	
LAEF (%)	47.7 ± 4.7	49.7 ± 5.9	0.15	
LA global strain (%)	14.9 ± 1.9	16.8 ± 2.5	<0.01	
LA-pool strain (%)	20.0 ± 4.3	23.8 ± 3.6	<0.001	
LA- pump strain (%)	9.6 ± 3.0	12.3 ± 3.4	<0.01	
LAsf (%)	0.42±0.09	0.35±0.11	0.02	

Table 2 Echocardiography Parameters in Diabetic Patients

Abbreviations: LVEDV, left ventricular end diastolic volume; LVESV, left ventricular end systolic volumes; LVEF, left ventricular ejection fraction; LVMI, left ventricular mass index; E, peak transmitral early diastolic inflow; A, peak transmitral late diastolic inflow; TAPSE, tricuspid annular plane systolic excursion, GLS, global longitudinal strain, LA, left atrium; LAVI, indexed left atrial volume; LAEF, left atrial total emptying fraction; LASF, left atrial stiffness.

25-

3D-Echocardiography

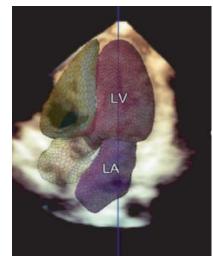
42838384	Contents lists available at ScienceDirect
	Journal of Diabetes and Its Complications
ELSEVIER	journal homepage: www.elsevier.com/locate/jdiacomp

Diabetic microvascular complications are associated with left atrial structural alterations in asymptomatic type 2 diabetes patients: A cross-sectional study *

Mingxia Gong^a, Min Xu^{a,*}, Jun Meng^a, Shu Jiang^a, Xiaohong Jiang^b

319 asymptomatic DM patients, assessed with <u>3D</u> <u>echocardiography</u> with semi-automatic quantification of LA volume over time

LA size (LAVi min and LAVIpre) associated with no. of microvascular complications

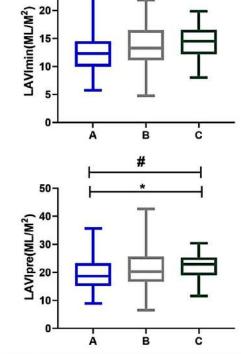


Gong, Journal of Diabetes, 2023; Spitzer, CFR 2017; Linden, JASE 2019

Characteristics of echocardiographic parameters grouped by microvascular complications in 279patients.

Table 2

Group by number of microvascular complications	Group A (n = 144)	Group B (n = 96)	Group C (n = 39)	P- value	
LVDd (mm)	$\textbf{47.2} \pm \textbf{4.0}$	47.2 ± 3.6	$\textbf{46.2} \pm \textbf{3.8}$	0.364	
LVSd (mm)	$\textbf{30.8} \pm \textbf{3.1}$	$\textbf{30.8} \pm \textbf{2.8}$	$\textbf{30.4} \pm \textbf{2.8}$	0.770	
IVSD (mm)	$\textbf{10.4} \pm \textbf{1.2}$	$\textbf{10.8} \pm \textbf{1.5}$	11.4 ± 2.0	0.445	
PWD (mm)	$\textbf{10.3} \pm \textbf{1.0}$	$\textbf{10.6} \pm \textbf{1.2}$	11.1 ± 1.7	0.307	
LVEF (%)	63.8 ± 2.5	63.5 ± 3.3	63.5 ± 2.5	0.615	
GLS	$-17.25 \pm$	$-16.74 \pm$	$-16.54 \pm$	0.293	
	2.57	2.46	2.49		
E (cm/s)	$\textbf{69.9} \pm \textbf{15.2}$	66.9 ±	72.9 ±	0.089	
		14.7	16.8		
A (cm/s)	$\textbf{91.1} \pm \textbf{16.3}$	89.8 \pm	90.2 \pm	0.797	
		13.9	14.7		
E/A	0.8 ± 0.2	0.7 ± 0.1	0.8 ± 0.2	0.085	
e' septal (cm/s)	6.3 ± 1.9	5.9 ± 1.8	5.7 ± 1.2	0.094	
e' lateral (cm/s)	7.9 ± 2.1	7.8 ± 2.4	$\textbf{7.7} \pm \textbf{1.8}$	0.796	
E/e'	$\textbf{9.9} \pm \textbf{3.4}$	$\textbf{10.1} \pm \textbf{3.4}$	$\textbf{10.8} \pm \textbf{2.5}$	0.310	
LAVImin (ml/m²)	$\textbf{12.5} \pm \textbf{3.6}$	$\textbf{13.7} \pm \textbf{3.9}$	14.5 ± 3.1	0.004	
LAVImax (ml/m ²)	26.4 ± 7.0	$\textbf{28.8} \pm \textbf{8.4}$	29.2 ± 5.9	0.047	
LAVIpre (ml/m ²)	19.6 ± 5.9	21.2 ± 6.5	22.2 ± 4.5	0.022	



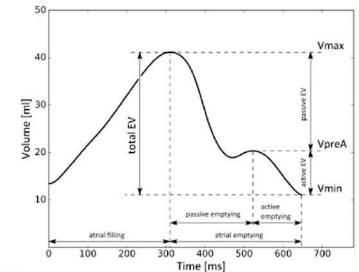


Fig. 2. Distribution of echocardiographic parameters in diabetes geometries by microvascular complications. A: No microvascular complications; B: 1 microvascular complication; C: 2–3 microvascular complications.

BACKGROUND

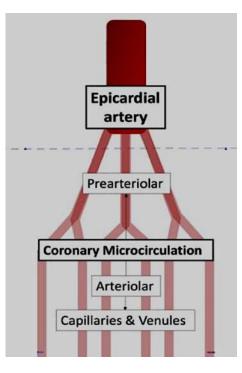
DIABETES



<u>High prevalence</u>: 1/10 worldwide (undiagnosed in > 1/3) <u>Increased risk</u> (2-5 fold) of **Major CV events** (HF, AF/stroke, CAD)



Myocardial ischemia (CAD / INOCA)



IDF Diabetes Atlas 2022 Reports; Dandamudi, J Card Fail 2014; Muhlestein JAMA 2014; From Am J Card 2009; Rajagopalan JACC 2005, Beagley J, et al. Diabetes Res Clin Pract 2014; Sarwar N, et al. Lancet 2010

(2) Early identification of MYOCARDIAL ISCHEMIA

Stress-Echocardiography

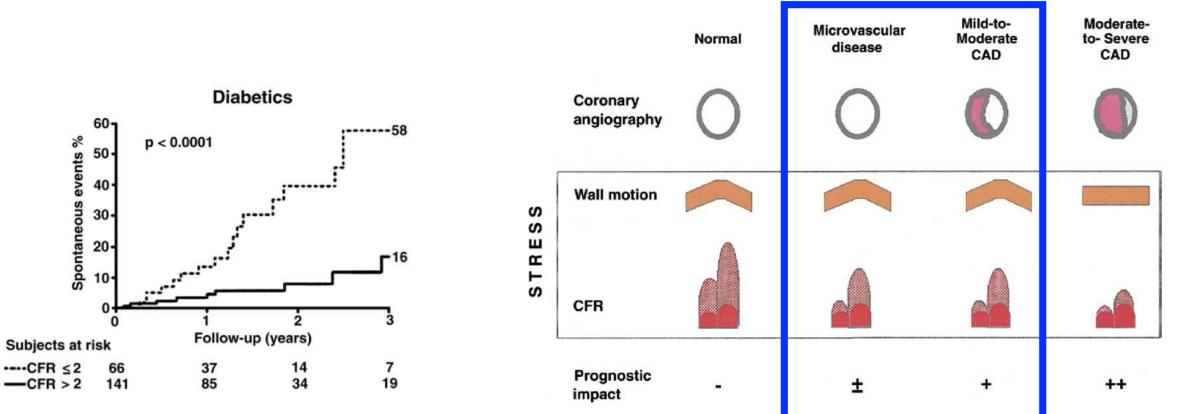
Abnormal coronary flow reserve (≤2) present in 27% DM patients (and negative stressechocardiography)

..associated with **increased risk of MACE (HR 1.50**, 95% CI 1-2.25)



Additional Prognostic Value of Coronary Flow Reserve in Diabetic and Nondiabetic Patients With Negative Dipyridamole Stress Echocardiography by Wall Motion Criteria

Lauro Cortigiani, MD,* Fausto Rigo, MD, FESC,† Sonia Gherardi, MD,‡



Computed Tomography

Original Article

I Atheroscler Thromb, 2021; 28: 1052-1062. http://doi.org/10.5551/jat.59386

Coronary Artery Calcium Score Predicts Long-Term Cardiovascular **Outcomes in Asymptomatic Patients with Type 2 Diabetes**

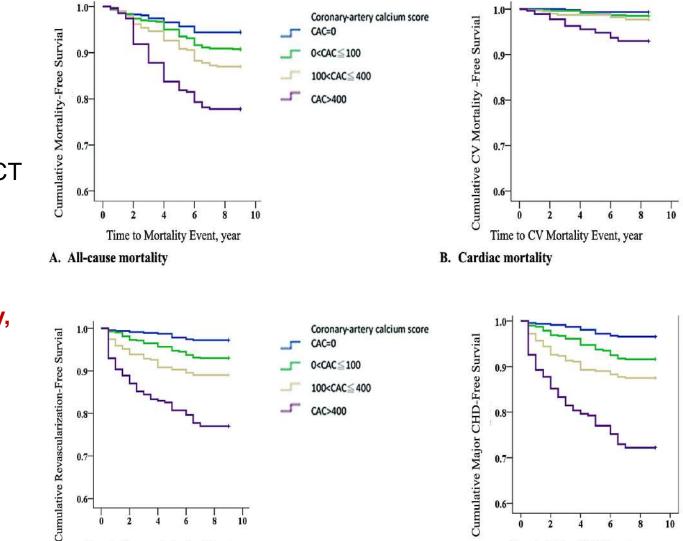
Meng-Huan Lei¹, Yu-Lin Wu², Sheng-Liang Chung¹, Chao-Chin Chen¹, Wei-Cheng Chen¹ and Yu-Chen Hsu¹

2162 asymptomatic DMT2 patients assessed with CT coronary calcium score scan

(low radiation 1-2 mSV, no contrast media).

Patients stratified by Agatston Score High risk: score > 400, HR 8.67 for cardiac mortality, HR = 10.83 for coronary revascularization

	I	All cause	mortality	t	1	Cardiac 1	mortality		Ne	on-cardia	c mortali	ity				
CAC score category	Hazard Ratio	(95%	6 CI)	p value	Hazard Ratio	CARACTER ACCOUNTS IN		Hazard Ratio	(95%	6 CI)	p value					
CAC=0		1 (refe	rence)			1 (refe	rence)		1 (reference)							
0 <cac≤100< th=""><th>1.07</th><th>0.63</th><th>1.81</th><th>.797</th><th>2.16</th><th>0.46</th><th>10.11</th><th>.329</th><th>0.95</th><th>0.54</th><th>1.67</th><th>.862</th><th></th><th></th><th></th><th></th></cac≤100<>	1.07	0.63	1.81	.797	2.16	0.46	10.11	.329	0.95	0.54	1.67	.862				
$100 < CAC \le 400$	1.45	0.84	2.52	.184	3.35	0.69	16.30	.134	1.22	0.68	2.22	.505				
CAC>400	2.08	1.18	3.66	.011	8.67	1.87	40.27	.006	1.38	0.73	2.59	.324				
	C	ox mode	l: p< .00)1	Cox model: $p < .001$		C	ox mode	l: <i>p</i> < .00	1						
		Major	CHD		Cord	Coronary revascularization		AMI				Ischemic stroke				
CAC score category	Hazard Ratio	(95%	6 CI)	<i>p</i> value	Hazard Ratio	(95%	6 CI)	<i>p</i> value	Hazard Ratio	100000 (NOSCO-0508) A DAM		p value	Hazard Ratio	(95%	OCI)	p value
CAC=0		1 (refe	rence)			1 (refe	rence)			1 (refe	rence)			1 (reference)		
0 <cac≤100< td=""><td>3.14</td><td>1.54</td><td>6.41</td><td>.002</td><td>3.39</td><td>1.52</td><td>7.58</td><td>.003</td><td>5.71</td><td>0.73</td><td>44.44</td><td>.096</td><td>1.19</td><td>0.63</td><td>2.25</td><td>.590</td></cac≤100<>	3.14	1.54	6.41	.002	3.39	1.52	7.58	.003	5.71	0.73	44.44	.096	1.19	0.63	2.25	.590
100 < CAC ≤ 400	4.18	1.99	8.80	< .001	4.57	1.98	10.52	< .001	6.31	0.76	52.23	.087	1.18	0.57	2.43	,652
CAC>400	10.52	5.07	21.83	< .001	10.83	4.76	24.62	< .001	4.19	0.44	39.51	.211	2.11	2.11	4.32	.410
	C	ox mode	1: p< .00)1	C	ox mode	l: p< .00)1	C	ox mode	l: p< .04	7	Cox model: $p < .001$			



Assessment of coronary artery calcium score with computed tomography may be considered as a risk modifier^c in the cardiovascular risk assessment of asymptomatic subjects. 449,457

Time to Revascularization Event, year

C. Coronary revascularization

All models were adjusted for age, diabetes duration and the number of risk factors.

IIb

Time to Major CHD Event, year

0.6

D. Major coronary heart disease

Control

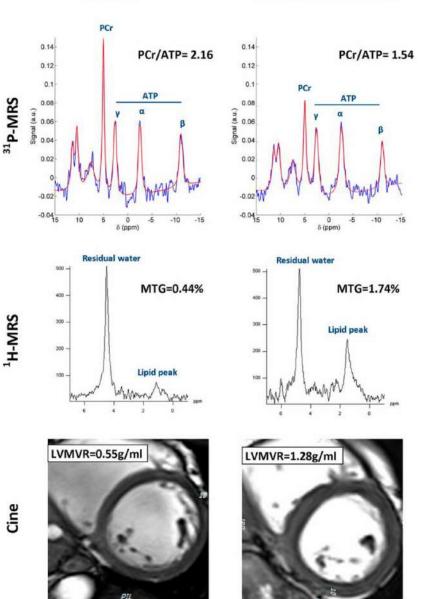
Patient with T2DM

³¹P Cardiac magnetic resonance spectroscopy

46 T2DM patients (non hypertensive) vs. controls:

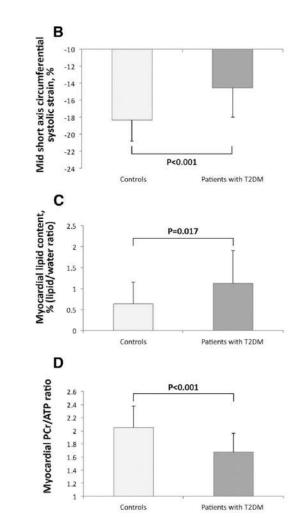
Myocardial steatosis (DM had 2fold increase in myocardial trygliceride content) was independent predictor of LV remodeling and reduced systolic strain.

.. Associated with impairement of myocardial energetics (-18% reduction in myocardial phosphocreatine to ATP ratio)



Eylem Levelt,1,2 Masliza Mahmod,1 Stefan K. Piechnik,1 Rina Ariga,1 Jane M. Francis,¹ Christopher T. Rodgers,¹ William T. Clarke,¹ Nikant Sabharwal,³ Jurgen E. Schneider,¹ Theodoros D. Karamitsos,^{1,4} Kieran Clarke,² Oliver J. Rider,¹ and Stefan Neubauer¹

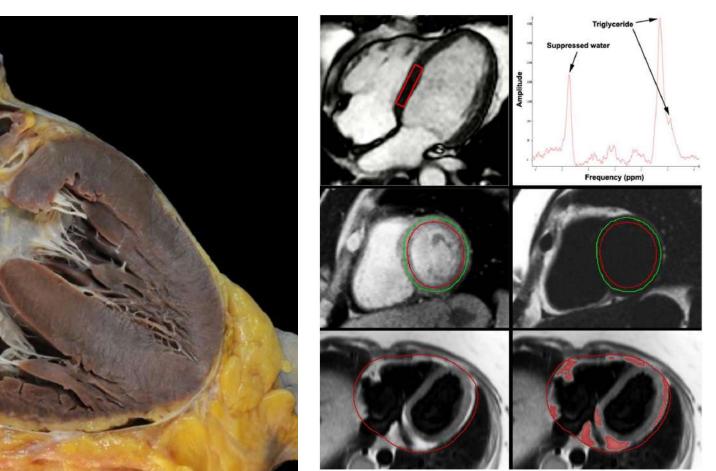
Relationship Between Left Ventricular Structural and Metabolic Remodeling in Type 2 Diabetes



Cine

³¹P Cardiac magnetic resonance spectroscopy

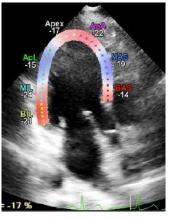
Epicardial adipose tissue volume and insulin-resistance both independently associated with -increased myocardial fat content -higher burden of interstitial fibrosis -impaired LV longitudinal strain

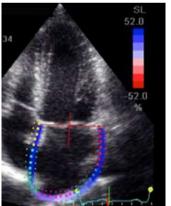


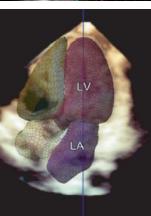
Circulation: Cardiovascular Imaging

ORIGINAL ARTICLE

Impact of Epicardial Adipose Tissue, Left Ventricular Myocardial Fat Content, and Interstitial Fibrosis on Myocardial Contractile Function







What's the role of CARDIAC IMAGING in <u>asymptomatic</u> diabetic patients ?

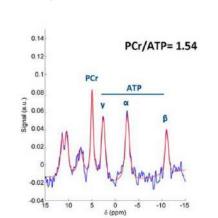
Detection of preclinical:

- ✓ systolic and/or diastolic dysfunction
- ✓ impaired exercise capacity
- \checkmark cardiac remodeling
- $\checkmark\,$ focal and/or diffuse fibrosis
- ✓ risk of microvascular complications
- $\checkmark\,$ impaired left atrial deformation predisposing to AF
- ✓ cardiac steatosis, epicardial adipose tissue volume

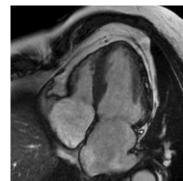
Risk stratification (adverse remodeling, CV events, mortality, hospitalizations)











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Thank you for your attention!