



XXIX CONGRESSO NAZIONALE ANCE

10 - 13 Ottobre 2019
Centro Congressi
Hilton Sorrento Palace
Sorrento (NA)





UNIVERSITÀ
DEGLI STUDI
DI PALERMO



A.O.U.P.
«Paolo Giaccone»

«Scuola di Medicina

LETTURA

La Valutazione del Danno Subclinico Cardiaco e Vascolare nel Paziente Iperteso e i Target Pressori raccomandati dalle Linee Guida AHA-ACC 2017 ed ESC-ESH 2018

Salvatore Novo (Palermo)

Presenta: M. Di Franco (Palermo)

Centro Congressi Hilton Sorrento Palace, Sabato 12 Ottobre 2019, ore 18.45-19.15

2018 ESC/ESH GUIDELINES ON ARTERIAL HYPERTENSION

Hypertension is defined as office SBP ≥ 140 mmHg and/or DBP ≥ 90 mmHg. This statement is based on evidence from multiple RCTs demonstrating that treatment of patients with these BP values is beneficial.

Category	Systolic (mmHg)		Diastolic (mmHg)
Optimal	< 120	and	< 80
Normal	120-129	and/or	80-84
High normal	130-139	and/or	85-89
Grade 1	140-159	and/or	90-99
Grade 2	160-179	and/or	100-109
Grade 3	≥ 180	and/or	≥ 110
Isolated systolic hypertension	≥ 140	and	< 90

2017 ACC/AHA GUIDELINES ON ARTERIAL HYPERTENSION

Table 6. Categories of BP in Adults*

BP Category	SBP		DBP
Normal	<120 mm Hg	and	<80 mm Hg
Elevated	120–129 mm Hg	and	<80 mm Hg
Hypertension			
Stage 1	130–139 mm Hg	or	80–89 mm Hg
Stage 2	≥140 mm Hg	or	≥90 mm Hg

2017 ACC/AHA GUIDELINES ON ARTERIAL HYPERTENSION

Table 7. Prevalence of Hypertension Based on 2 SBP/DBP Thresholds*†

	SBP/DBP \geq 130/80 mm Hg or Self-Reported Antihypertensive Medication†		SBP/DBP \geq 140/90 mm Hg or Self-Reported Antihypertensive Medication‡	
	Men (n=4717)	Women (n=4906)	Men (n=4717)	Women (n=4906)
Overall, crude	46%		32%	
Overall, age-sex adjusted	48%	43%	31%	32%
Age group, y				
20–44	30%	19%	11%	10%
45–54	50%	44%	33%	27%
55–64	70%	63%	53%	52%
65–74	77%	75%	64%	63%
75+	79%	85%	71%	78%
Race-ethnicity§				
Non-Hispanic white	47%	41%	31%	30%
Non-Hispanic black	59%	56%	42%	46%
Non-Hispanic Asian	45%	36%	29%	27%
Hispanic	44%	42%	27%	32%

**VI SONO EVIDENZE CLINICHE CHE
GIUSTIFICANO UN ABBASSAMENTO DEI
LIVELLI DI PRESSIONE ARTERIOSA PER
INIZIARE UN TRATTAMENTO?**

SUB-CLINICAL CARDIOVASCULAR DAMAGE IN ARTERIAL HYPERTENSION

Really, the evolution of BP from normal level to hypertension has been found to be associated with increased CV events and other risk factors.

Both office and home BP have an independent and continuous relationship with the incidence of CV events (haemorrhagic or ischaemic stroke, myocardial infarction, sudden death, heart failure, and PAD), as well as end-stage renal disease.

Moreover, recent studies show that also newly diagnosed arterial hypertension can be associated with subclinical CV organ damage.

SUB-CLINICAL VASCULAR DAMAGE IN ARTERIAL HYPERTENSION

Morning BP **is associated with carotid IMT** in pre-hypertensive patients.

Alpaydin S et al. *Blood Press Monit.* 2017; 22: 131-6

Pre-Hypertension **is associated with increased carotid ATS plaque** in the community population of Southern China.

Hong H et al. *BMC Cardiovasc Disord.* 2013; 13-20.

Body Mass Index impact on Arterial Stiffness in Young Pre-hypertensives:
A Cross Sectional Study

Deepika V et al. *J Res Health Sci.* 2017; 18: e00402.

Pre-hypertension and high **serum uric acid increase arterial stiffness.**

Thitiwuthikiat Pet al. *Scand J Clin Lab Invest.* 2017; 77: 673-8.

SUB-CLINICAL VASCULAR DAMAGE IN ARTERIAL HYPERTENSION

High Central Aortic rather than Brachial BP is Associated with Carotid Wall Remodeling and **Increased Arterial Stiffness** in Childhood.

Peluso G et al. High Blood Press Cardiovasc Prev. 2017; 24: 49-60

Ascending Aorta Elasticity is related to Pre-hypertension.

Jia CF et al. Am J Hypertens 2017; 30: 61-6.

Pulse pressure increase reduce LV global longitudinal strain in normotensive and newly diagnosed, untreated, hypertensive patients.

Lembo M et al. J Hypertens. 2016; 34: 1201-7

ASYMPTOMATIC CAROTID LESIONS IN A SAMPLE OF 755 SUBJECTS IN RELATION WITH A SINGLE RISK FACTOR

	NORMALS (%)	IMT (%)	A.C.P. (%)	IMT+ACP (%)
R.F. -	83,87	10,75	5,37	16,2
R.F. +	52,9	32,5 *	14,7 *	47,2 *
HYPERTENS.	34,48	44,82 *	20,68 *	65,5 *
NIDDM	58,30	25,00 *	16,7 *	41,7 *
HYPER-C	73,33	13,33	13,33 *	26,6 *
FAMILY H. CVD	80,64	12,90	6,46	19,4
HYPER-TG	75,00	25,00	0	25,00
SMOKING	66,66	33,33 *	0	33,3 *
OBESITY	95,23	4,76	0	4,7

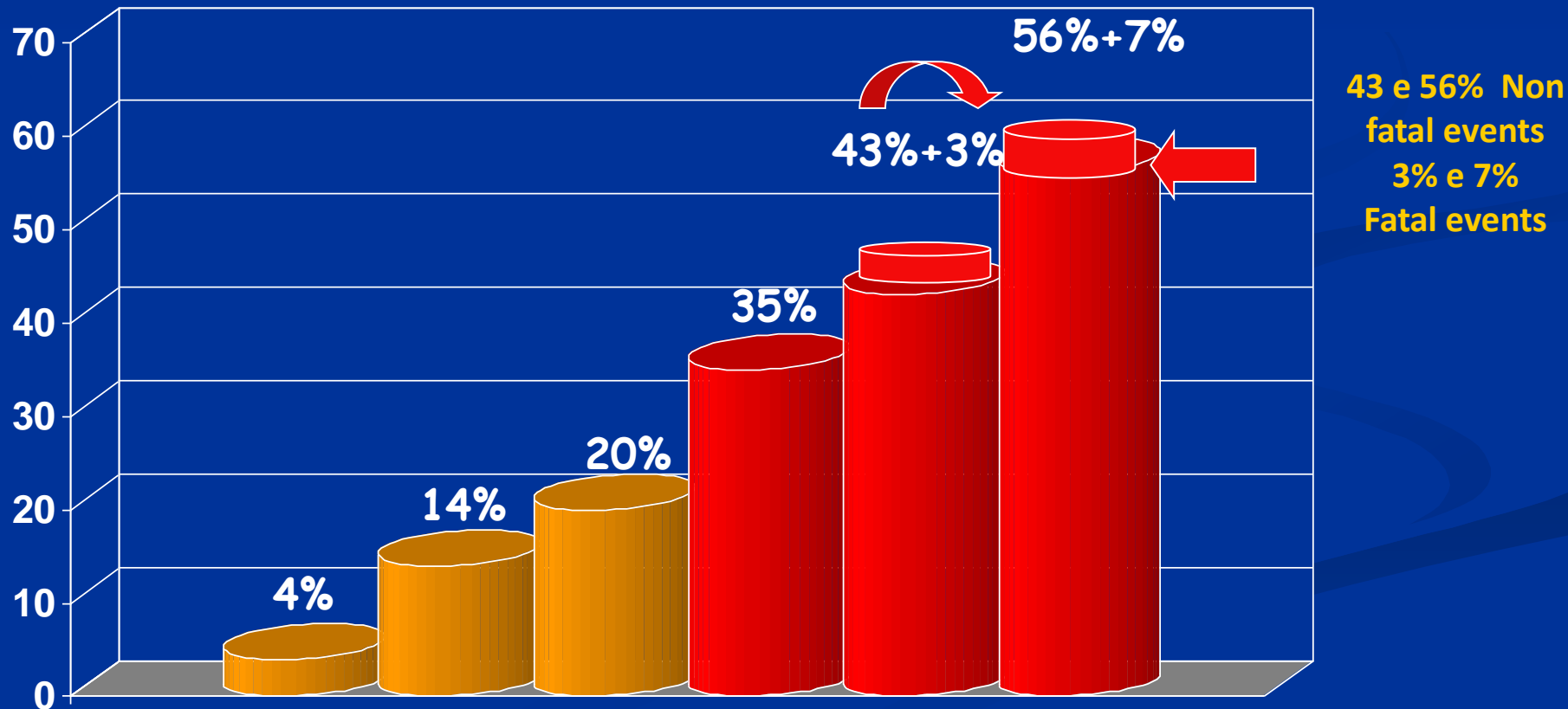
**A LIVELLO VASCOLARE L'IPERTENSIONE
ARTERIOSA, ANCHE LA PRE-IPERTENSIONE E
L'IPERTENSIONE ARTERIOSA DI RECENTE
DIAGNOSI, SI ASSICIA AD UN AUMENTO
DELL'ARTERIAL STIFFNESS E DELL'INCIDENZA
DI IMT/PLACCA CAROTIDEA**

IMT INDICATOR OF CARDIOVASCULAR EVENTS

- **Increased IMT is an independent RF for MI and stroke**
 - **Finnish study (1257 men)**
 - ↑IMT for 0.1mm → ↑11% risk for AMI
 - **ARIC study (15800 m,w)**
 - ↑IMT is associated with 2-4 fold increase of coronary events
 - **Rotterdam study (7983 m, w)**
 - ↑ IMT – 1SD (0.163 mm) → MI – OR - 1.57
Stroke – OR -1.51
 - **CHS study (4776 m,w)**
 - ↑IMT → 4 fold increase of risk for AMI and stroke in highest quintile

SUB-CLINICAL ATS ADD TO PREDICTION OF CV RISK: A TEN YEARS FOLLOW-UP STUDY IN 558 PATIENTS

Novo S, Visconti C, Amoroso GR, Corrado E, Muratori I, Fazio G, Novo G
Eur J Cardiovasc Prev & Rehabilitation 2010; 17: 514-8



SUB-CLINICAL ATS AND METS INCREASE CARDIO- AND CEREBRO-VASCULAR EVENTS RATE: A 20-YEAR FOLLOW UP.

Novo S et al. Cardiovasc Diabetol. 2013; 23: 155-62

RESULTS

251 patients suffering from MetS

278 healthy patients

144 total CV adverse events

98 total CV adverse events

$P < 0,0001$

108 in pts with Carotid ATS

36 in pts without Carotid ATS

71 in pts with Carotid ATS

27 in pts without Carotid ATS

$P < 0,0099$ – RR 3.0

$P < 0,0003$ – RR 2.6

Risk categories

Very high-risk	<p>Subjects with any of the following:</p> <ul style="list-style-type: none">• Documented CVD, clinical or unequivocal on imaging. Documented clinical CVD includes previous AMI, ACS, coronary revascularization and other arterial revascularization procedures, stroke and TIA, aortic aneurysm and PAD. Unequivocally documented CVD on imaging includes significant plaque on coronary angiography or carotid ultrasound. It does NOT include some increase in continuous imaging parameters such as intima-media thickness of the carotid artery.• DM with target organ damage such as proteinuria or with a major risk factor such as smoking or marked hypercholesterolaemia or marked hypertension.• Severe CKD (GFR <30 mL/min/1.73 m²).• A calculated SCORE =/>10%.
High-risk	<p>Subjects with:</p> <ul style="list-style-type: none">• Markedly elevated single risk factors, in particular cholesterol >8 mmol/L (>310 mg/dL) (e.g. in familial hypercholesterolaemia) or BP ≥180/110 mmHg.• Most other people with DM (with the exception of young people with type 1 DM and without major risk factors that may be at low or moderate risk).• Moderate CKD (GFR 30–59 mL/min/1.73 m²).• A calculated SCORE ≥5% and <10%.
Moderate-risk	SCORE is ≥1% and <5% at 10 years. Many middleaged subjects belong to this category.
Low-risk	SCORE <1%.

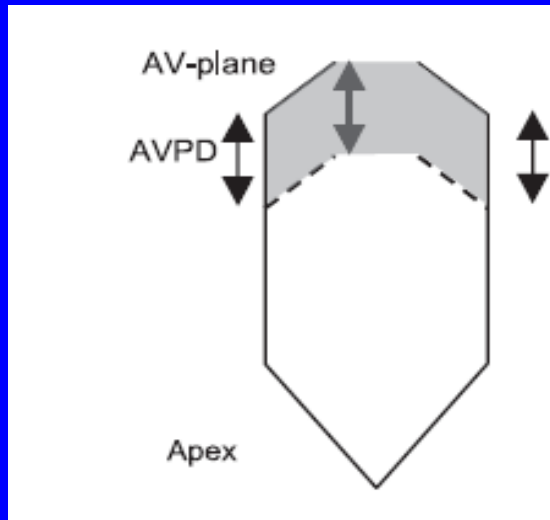
SUB-CLINICAL CARDIAC DAMAGE

Abnormal LV geometry of hypertensive patients is frequently associated with **diastolic dysfunction** which can be further evaluated by a combination of **transmitral flow and tissue Doppler studies**.

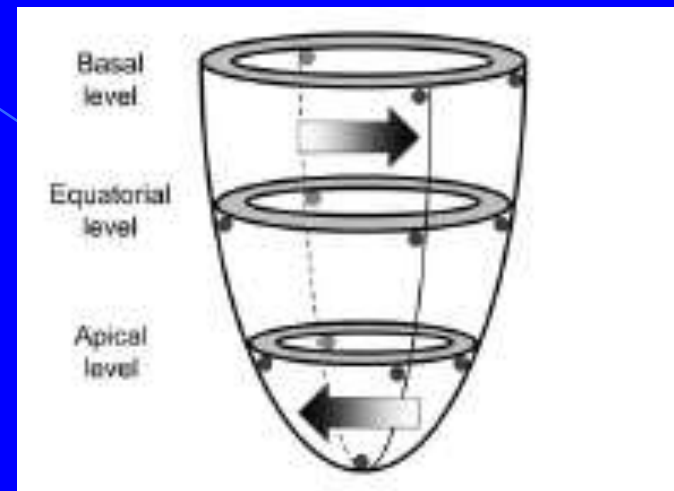
LA size is also frequently increased in hypertensive patients; it is related to **diastolic dysfunction** and associated with the **incidence of Atrial Fibrillation and CV events**.

CARDIAC MECHANIC

Longitudinal function



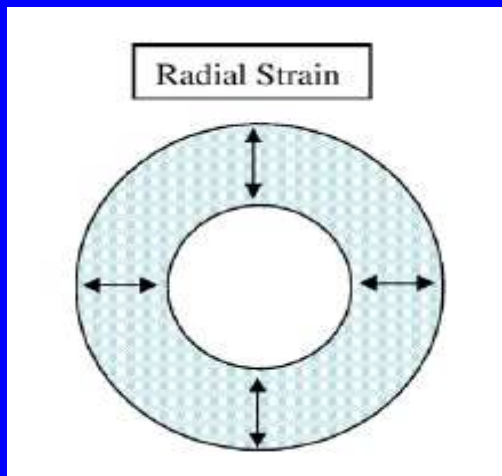
TWIST



Movement of base versus apex

Clockwise rotation of the base and anticlockwise rotation of the apex

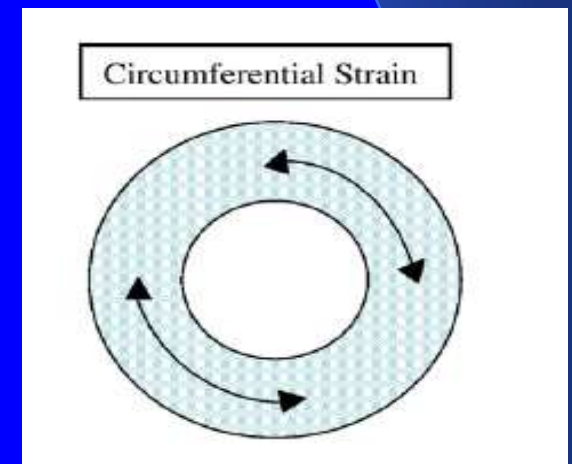
Radial function

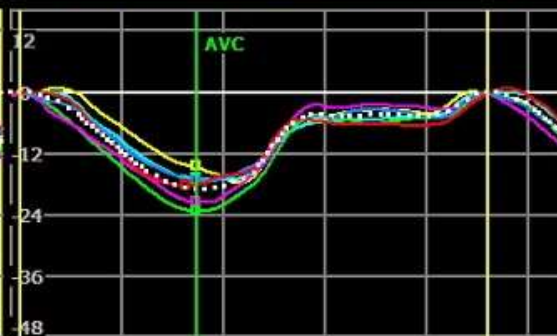
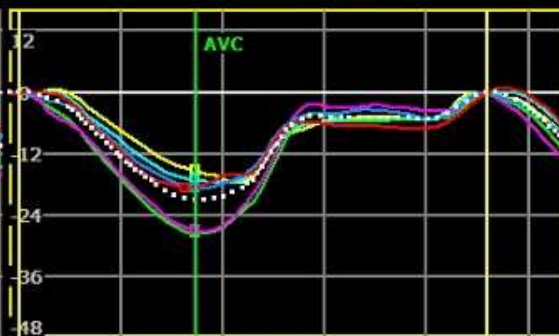
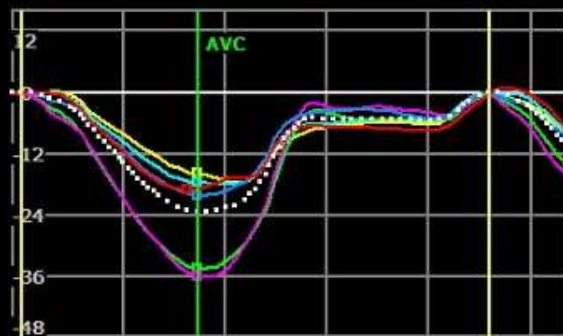
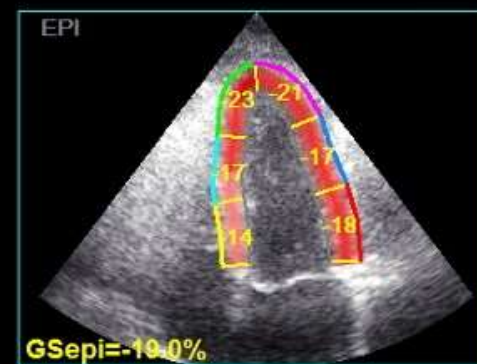
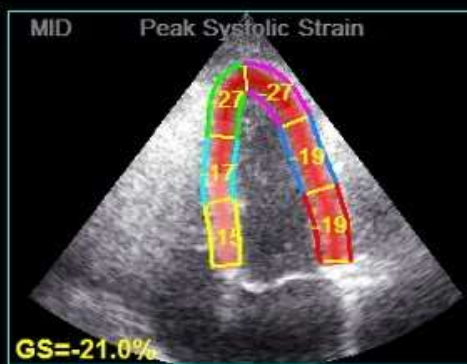
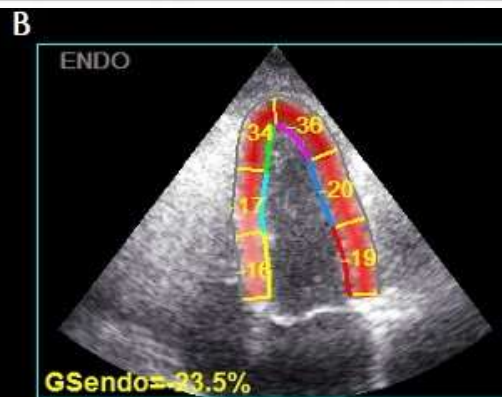
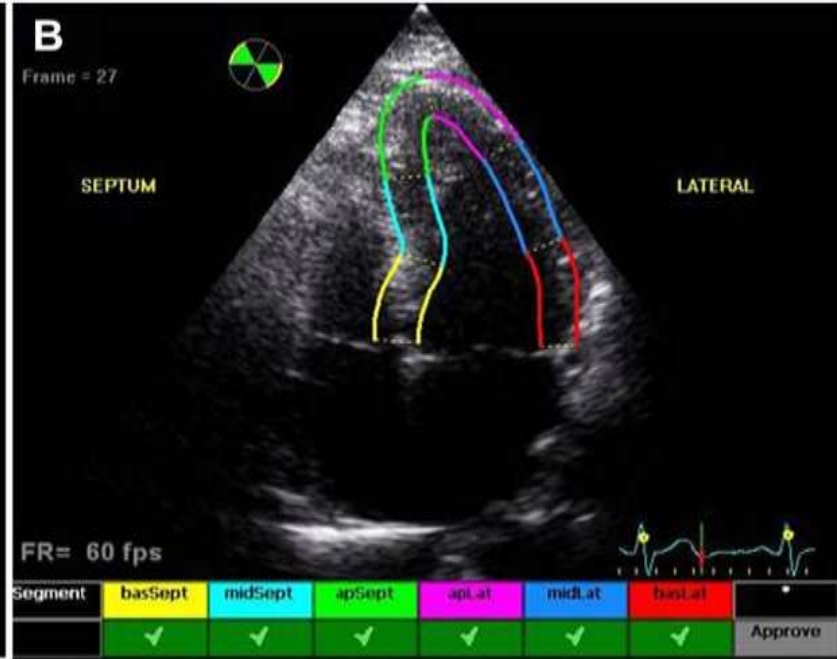
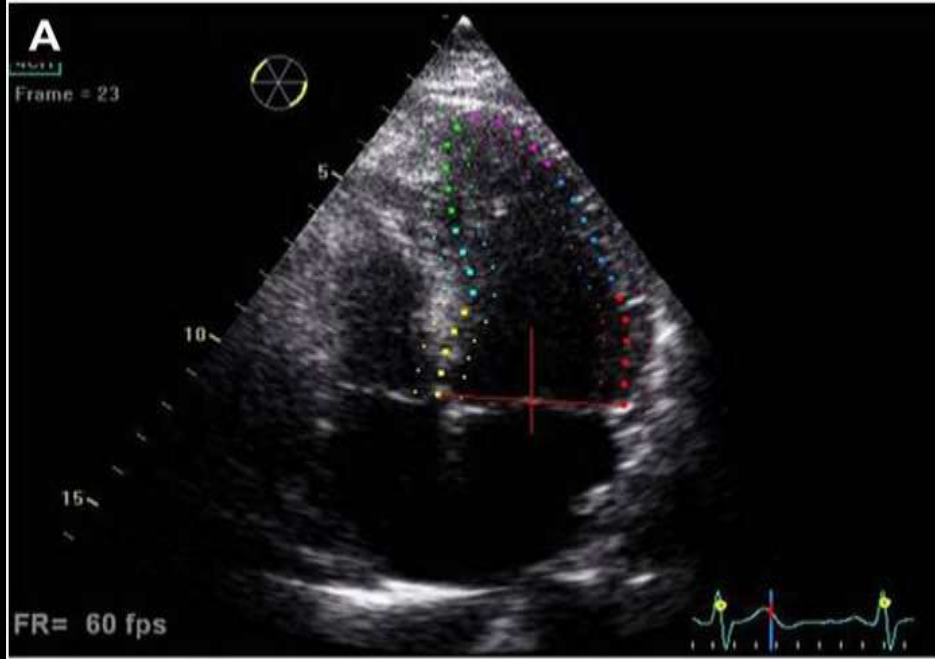


Thickening of myocardium

Length variation along the circular perimeter

Circumferential function





The increase of Clinic and 24-h systolic BP is associated **with LV reduced GLS** evaluated with 3DE strain analysis, independently of LV structure and diastolic function.

Tadic et al. Int J Cardiovasc Imaging 2014; 30: 699-711

Early LA strain alterations could be linked to exertional dyspnea, often present in newly diagnosed arterial hypertension.

Braunauer et al. Int J Cardiovasc Imaging 2018; 34: 701-11

Finally, **RV structure, function, and deformation** are significantly changed in subjects with subclinical hypertension.

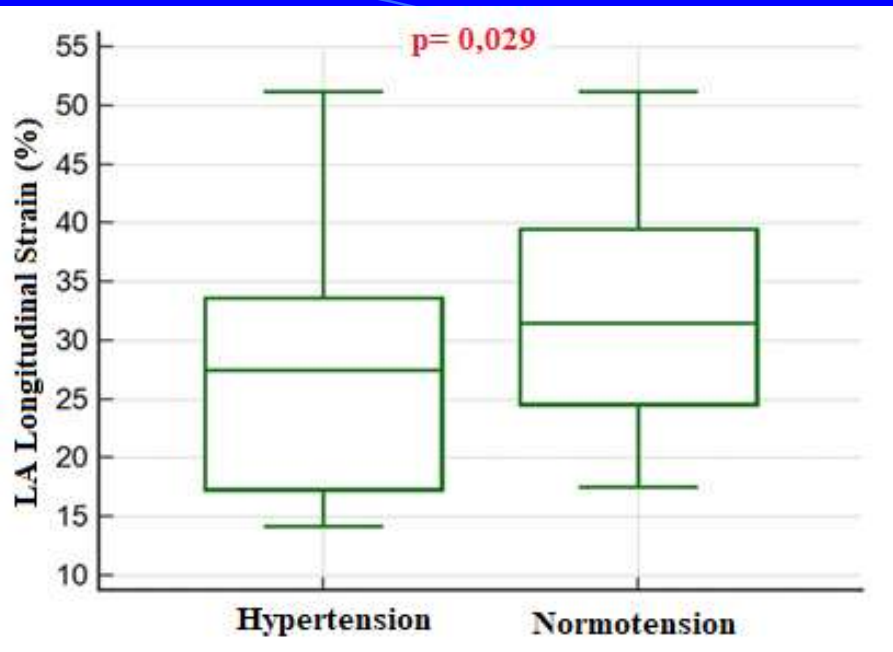
Tadic et al. J Clin Hypertens 2018; 20: 400-7

OUR RECENT EXPERIENCE

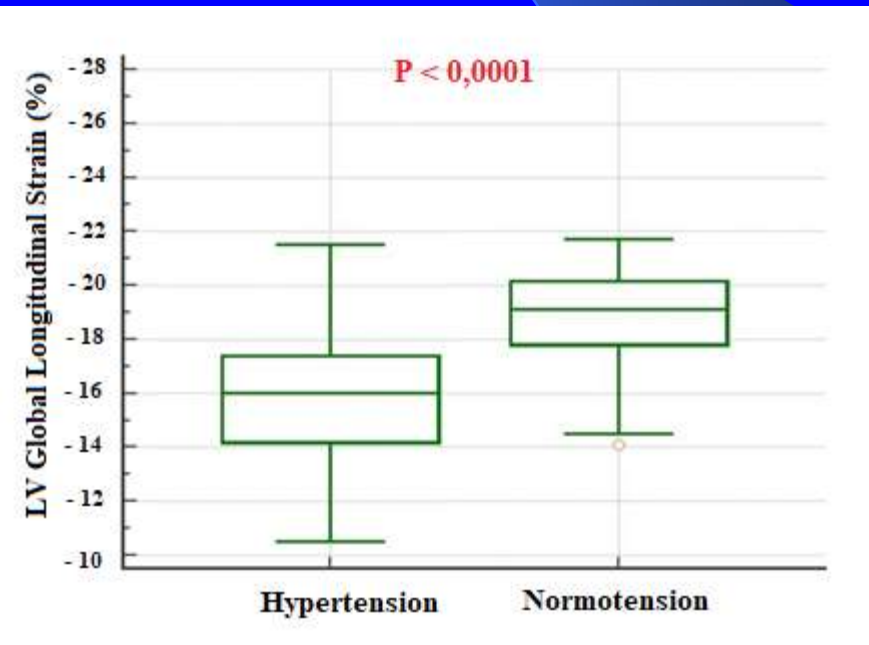
49 Patients with newly diagnosed Arterial Hypertension

- * Asymptomatic
- * Recent diagnosis (< 1 month)
- * ABPM confirmed diagnosis of hypertension
- * **No** LA dilatation – LVH – IHD – Valvulopathies – Cardiomyopathies – Nephropaty – COLD – Other diseases

43 Normotensive Control Subjects



Anche i soggetti ipertesi di recente diagnosi mostrano una riduzione dello strain longitudinale dell'Asin. e dello strain longitudinale globale del VS!!



NEWSLY DIAGNOSED ARTERIAL HYPERTENSION AND BEHAVIOR OF PAP

In early stage of uncomplicated arterial hypertension, **increased pulmonary vascular resistance (PVR)** has been reported at rest and more rarely during exercise.

Vriz (**Can J Cardiol. 2015; 31: 537-43**) described that resting and exercise Pulmonary Vascular Resistance are increased in uncomplicated hypertension, without this being related to increased Pulmonary Venous Pressure or resistive vessel stiffness, **suggesting an early increase in pulmonary vascular tone.**

OUR STUDY ON THE BEHAVIOR OF PAP IN UNCOMPLICATED HYPERTENSIVES

- * 69 patients with hypertension, with (n=24) or without (n=45) LV diastolic dysfunction, and BP > 140/90 mm Hg.
- * **All patients underwent physical ecostress**, using an ergometer bed, increasing the work load of 25 watt every 2 min.
- * Echocardiographic parameters, BP, and ECG were registered at rest and during each step of the exercise until the maximal HR or muscle exhaustion or symptoms.

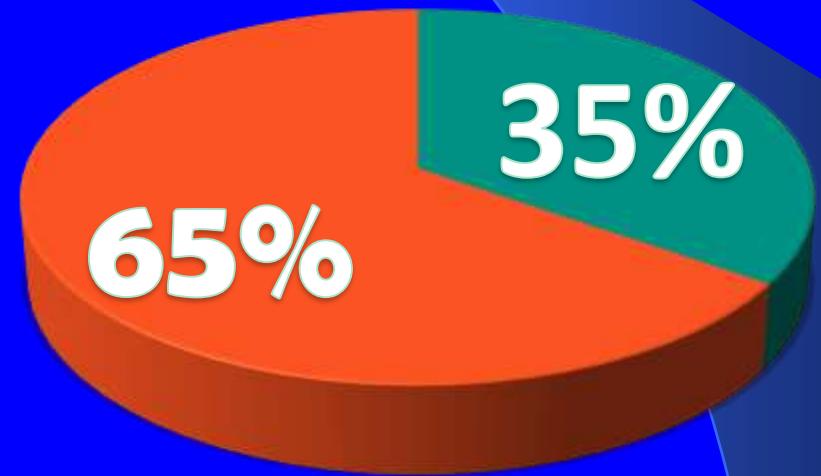
CHARACTERISTICS OF PATIENTS WITH OR WITHOUT DIASTOLIC DYSFUNCTION

	Diastolic Dysfunction	No Diastolic Dysfunction	P <
Patients	24 (34,8%)	45 (65,2%)	0,16
Age	63,04±10	60,4±8,5	0,16
Female Gender	9 (13%)	14 (20,3%)	0,6
Pressure/Rate Product	25024	22729	0,03
ACE-I /ARBs	18 (26%)	30 (43,4%)	0,38
B-blockers	14 (20,3%)	25 (36,2%)	0,75
Ca++ channel blockers	7 (10,1%)	11 (15,9%)	0,63
Diuretics	6 (8,6%)	13 (18,8%)	0,84
Hypercolesterolemia	14 (20,3%)	29 (42%)	0,62
Smoking	5 (7,25%)	20 (29%)	0,05
Diabetes	4 (5,8%)	8 (11,5%)	0,9

Patients divided in two groups:

- 1) With increase of the E/E' ratio > 14 during exercise (index of diastolic dysfunction (EACVI/ASE);
- 2) With the ratio E/E' < 14 during exercise as index of normal LV function.

Patients at peak of exercise



■ Diastolic dysfunction

■ No diastolic dysfunction

RESULTS

Mean values registered at rest in the two Groups, with or without LV dysfunction

	DIASTOLIC DYSFUNCTION	NO DIASTOLIC DYSFUNCTION	P
E/E'	11,86 [11,18;14,57]	7,33 [5,85;8,9]	<0,0001
E/A	0,87 [0,76;1,25]	0,81 [0,75;1,01]	0,20
HR	69,7±14	70,5±11	0,16
SBP	128,7±14,4	127,9±15,8	0,8
DBP	80,8±7,4	70,5±11,1	0,043
E'	0,06 [0,04;0,07]	0,08 [0,07;0,1]	<0,0001
PAPS	29,5 [23,5;33]	24 [20;29]	0,01
VMAXIT	2,4±0,4	2,1±0,3	0,4
TAPSE	23,2±3,7	24,1±3,6	0,9
WALL THICKENING	21 (30,4%)	16 (23,1%)	<0,0001

Mean values registered in the two Groups, with or without LV dysfunction, at peak of exercise

	DIASTOLIC DYSFUNCTION	NO DIASTOLIC DYSFUNCTION	P
E/E'	16,1 [15;20,5]	11,2 [8,4;12,8]	<0,0001
E/A	0,93 [0,79;1,19]	0,85 [0,75;1,06]	0,3
HR	122±19	123±14	0,08
SBP	204,3±17,3	183,1±18,1	<0,0001
DBP	90,2±9,8	83,3±11,5	0,01
E'	0,07 [0,05;0,08]	0,09 [0,08;0,12]	<0,0001
PAP _s	46,12±8,84	36,13±10,09	0,0001
V _{MAXIT}	3,24±0,38	2,73±0,49	<0,0001
TAPSE	26,2±4,6	26,3±4,4	0,9

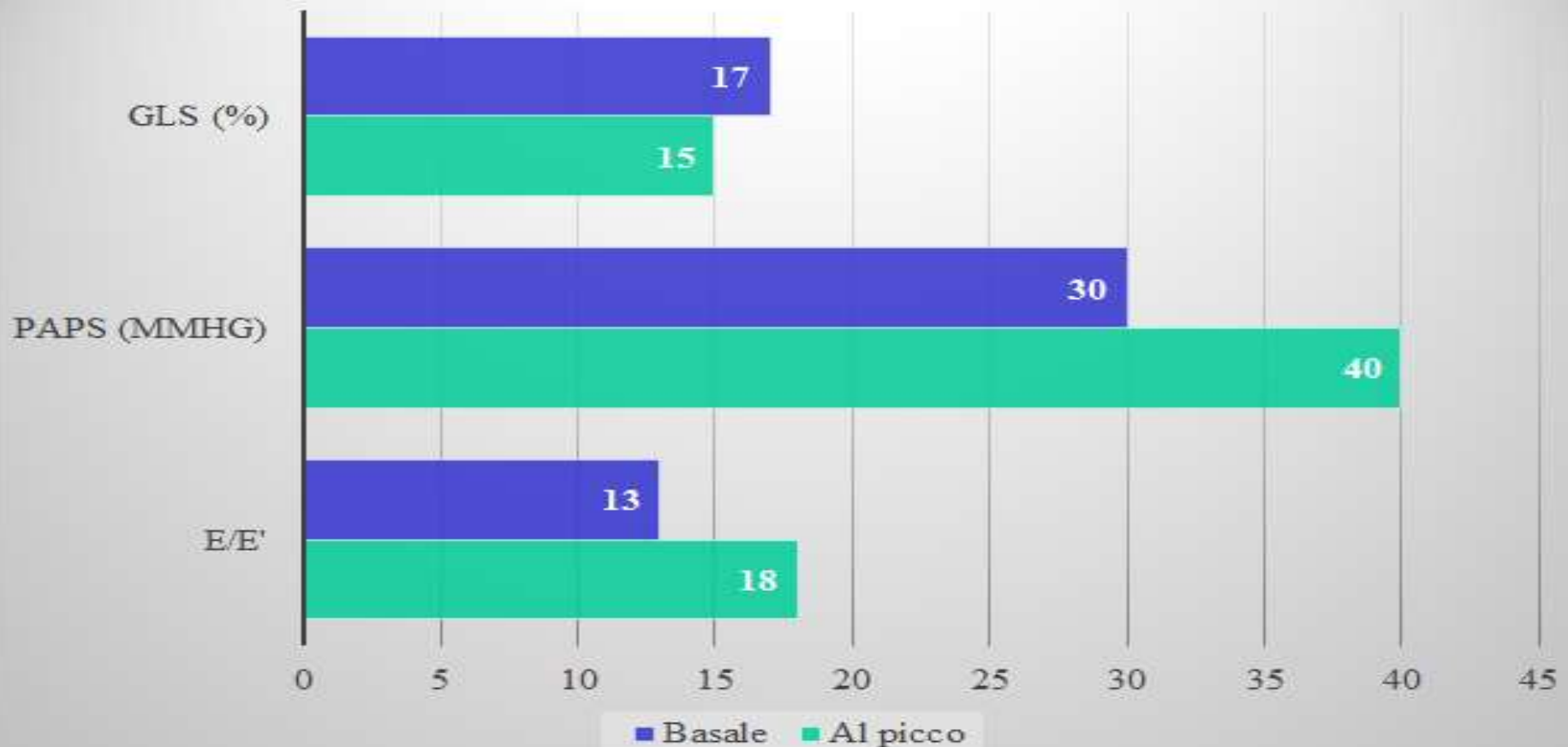


Why Italy works

STRESSECHO 2020

**Feasibility and Utility of the Ecostress,
beyond the Coronaropathy**

In a preliminary sample of patients with newly diagnosed hypertension and diastolic dysfunction we observed a reduction in GLS under stress and a simultaneous increase in PAPs and LV filling pressures



CONSIDERAZIONI

Nell'Ipertensione Arteriosa modificazioni della Stiffness arteriosa e della morfologia di parete (IMT/Placca) possono verificarsi anche precocemente.

Analogamente, cambiamenti della deformazione miocardica, sia ventricolare che atriale, possono svilupparsi in una fase precoce.

L'utilità clinica della valutazione degli indici di deformazione miocardica è sicuramente meritevole di ulteriore attenzione e ricerca.

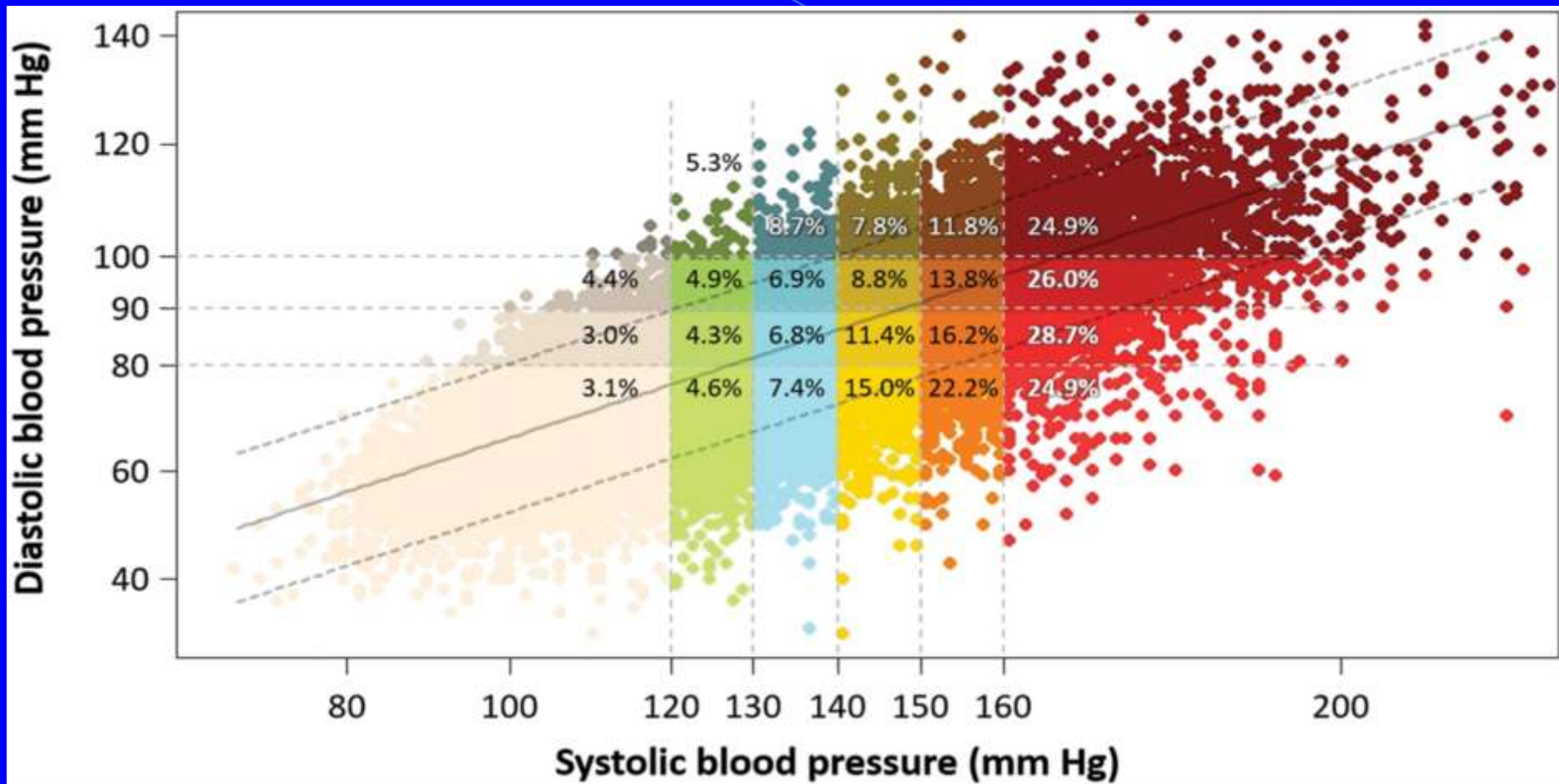
Lo scopo è quello di ottenere **indicatori** più precoci, sensibili e affidabili attraverso i quali delineare razionalmente il profilo prognostico e terapeutico dei pazienti ipertesi.

TAKE HOME MESSAGE

Le Linee Guida ESC/ESH 2018 sull'Ipertensione Arteriosa e, soprattutto, le Linee Guida ACC/AHA 2017, quando analizzano gli aspetti terapeutici, **suggeriscono più attenzione agli stadi iniziali dell'Ipertensione Arteriosa e una strategia terapeutica più precoce al fine di prevenire o ridurre il danno a carico degli organi bersaglio.**

L'uso delle tecniche di imaging vascolare e cardiaco, note (eco vascolare) e meno note (ecocardio-GLS) possono migliorare la stratificazione del rischio dei pazienti e guidare **nella scelta dei pazienti che devono essere trattati più precocemente e nei target di pressione arteriosa da raggiungere con il trattamento.**

DISTRIBUTION OF SYSTOLIC AND DIASTOLIC BLOOD PRESSURE AND 10-YEAR RATES OF MAJOR CV EVENTS



**Every coin has its reverse!!!
(Worsening of Mood)**

ESC/ESH 2018 GUIDELINES ON MANAGEMENT OF ARTERIAL HYPERTENSION

High normal BP
BP 130-139/85-89 mmHg

Lifestyle advice

Consider drug treatment in very high risk patients with CVD, especially CAD

Grade 1 Hypertension
BP 140-159/90-99 mmHg

Lifestyle advice

Immediate drug treatment in high or very high risk patients with CVD, renal disease or HMOD

Drug treatment in low moderate risk patients without CVD, renal disease or HMOD after 3-6 months of lifestyle intervention if BP not controlled

Grade 2 Hypertension
BP 160-179/100-109 mmHg

Lifestyle advice

Immediate drug treatment in all patients

Aim for BP control within 3 months

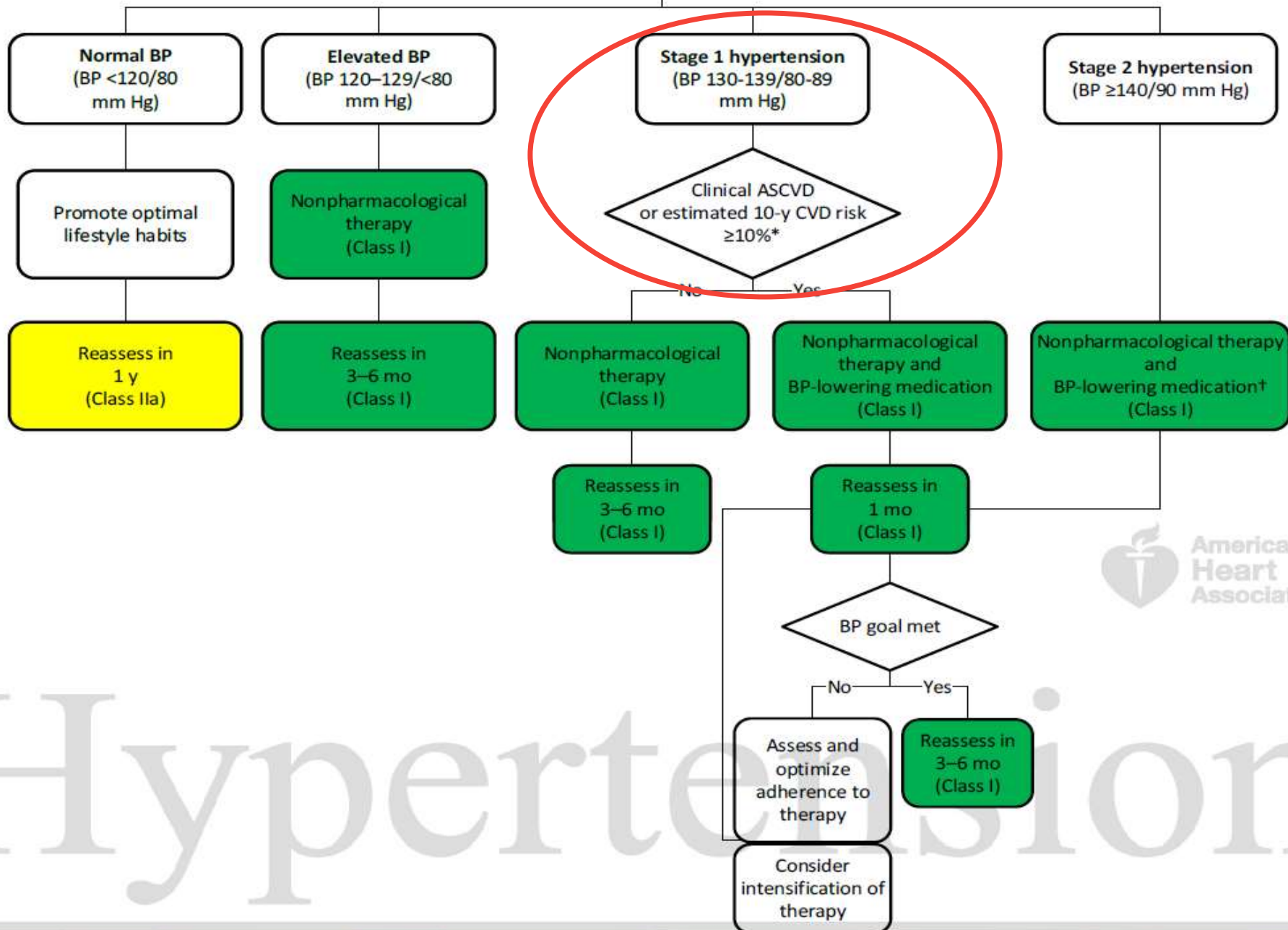
Grade 3 Hypertension
BP \geq 180/110 mmHg

Lifestyle advice

Immediate drug treatment in all patients

Aim for BP control within 3 months

BP thresholds and recommendations for treatment and follow-up



Hypertension

2017 ACC/AHA GUIDELINES ON ARTERIAL HYPERTENSION

8.1.5. BP Goal for Patients With Hypertension

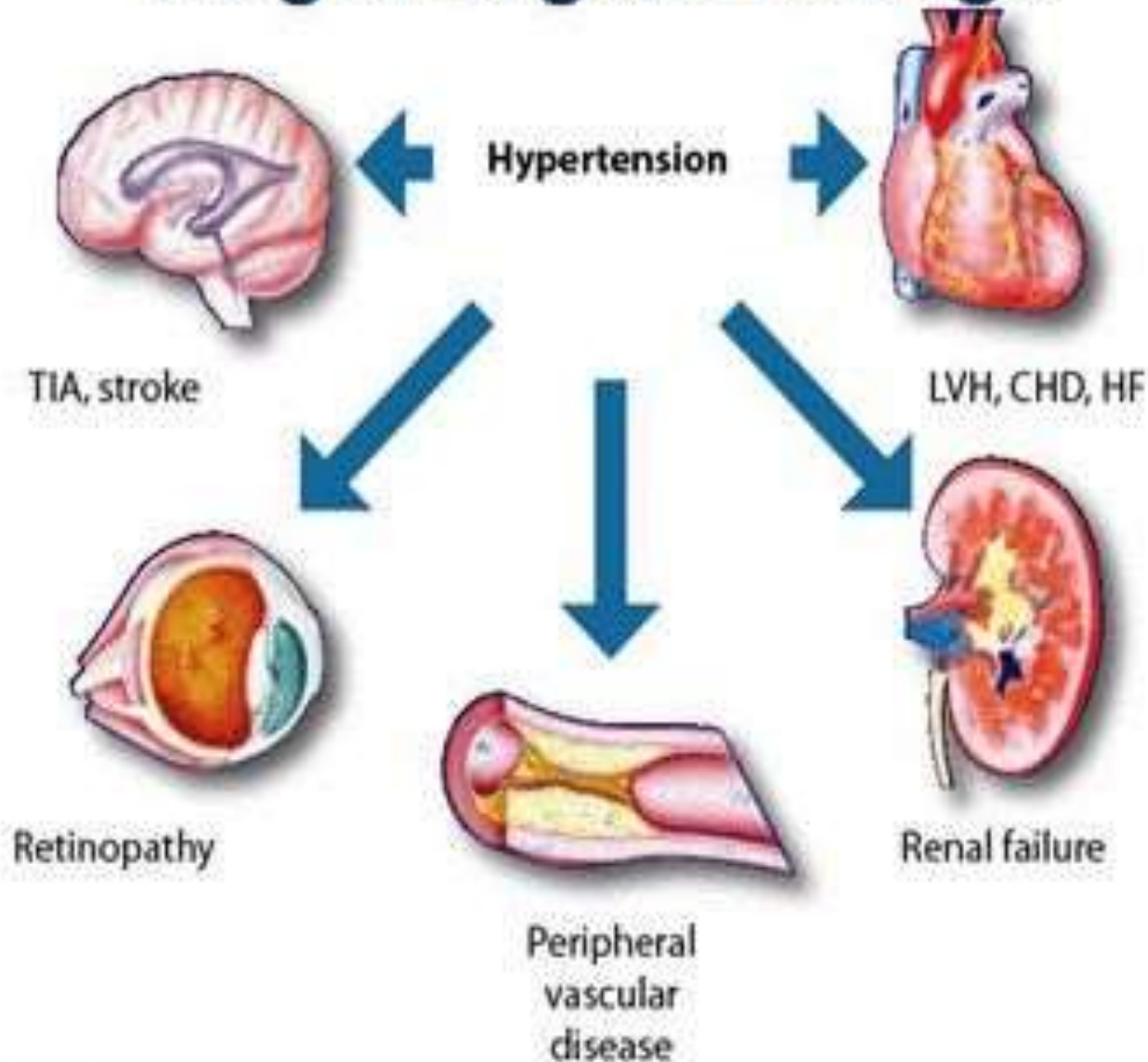
Recommendations for BP Goal for Patients With Hypertension

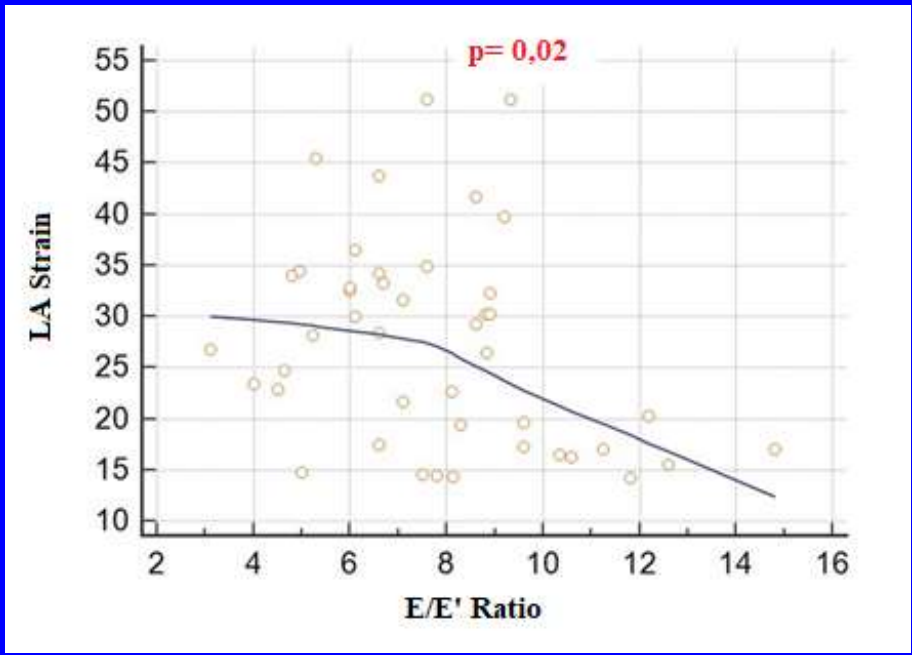
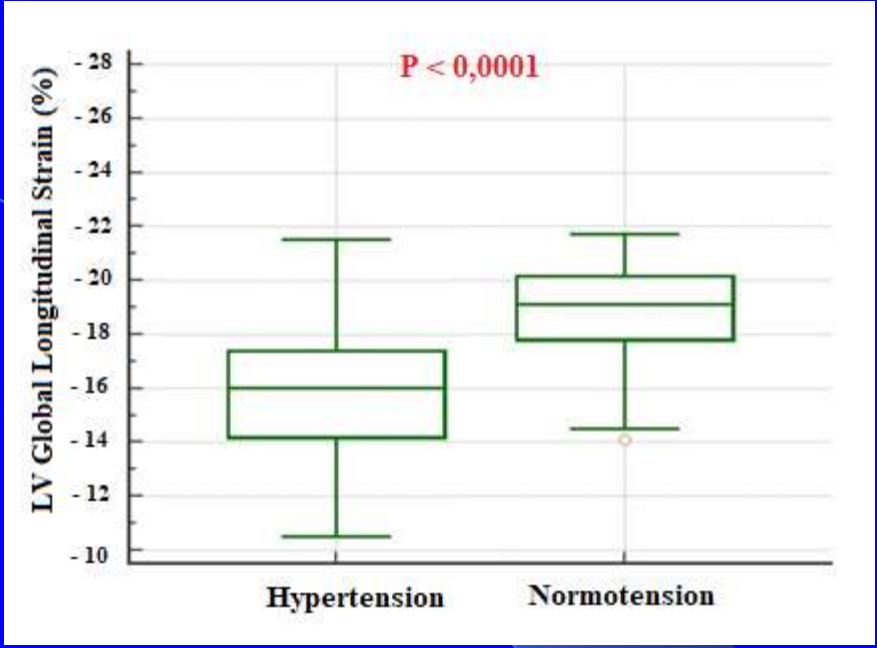
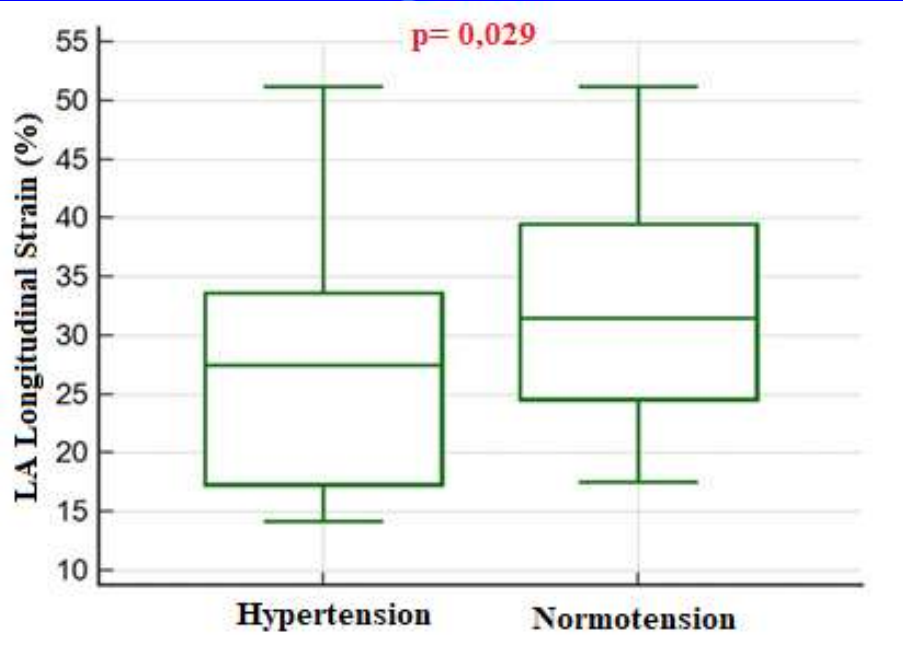
References that support recommendations are summarized in Online Data Supplement 26 and Systematic Review Report.

COR	LOE	Recommendations
I	SBP: B-R ^{SR}	1. For adults with confirmed hypertension and known CVD or 10-year ASCVD event risk of 10% or higher (see Section 8.1.2), a BP target of less than 130/80 mm Hg is recommended (1-5).
	DBP: C-EO	
IIb	SBP: B-NR	2. For adults with confirmed hypertension, without additional markers of increased CVD risk, a BP target of less than 130/80 mm Hg may be reasonable (6-9).
	DBP: C-EO	

SR indicates systematic review.

Complications of Hypertension: Target-Organ Damage



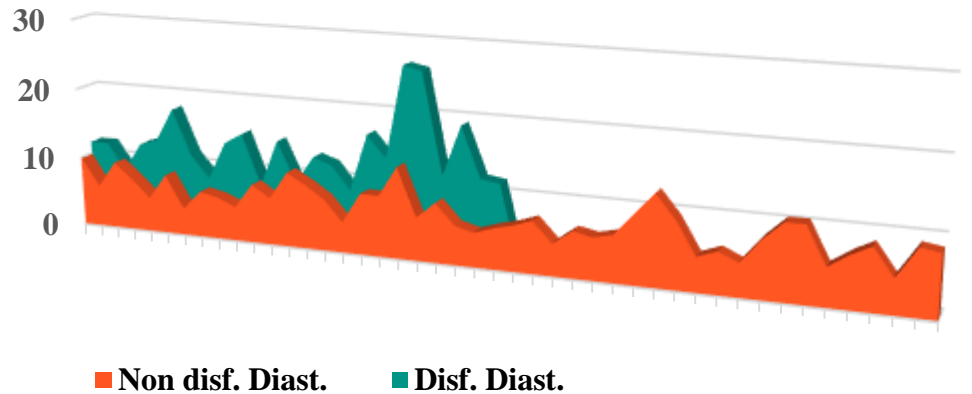


RESULTS

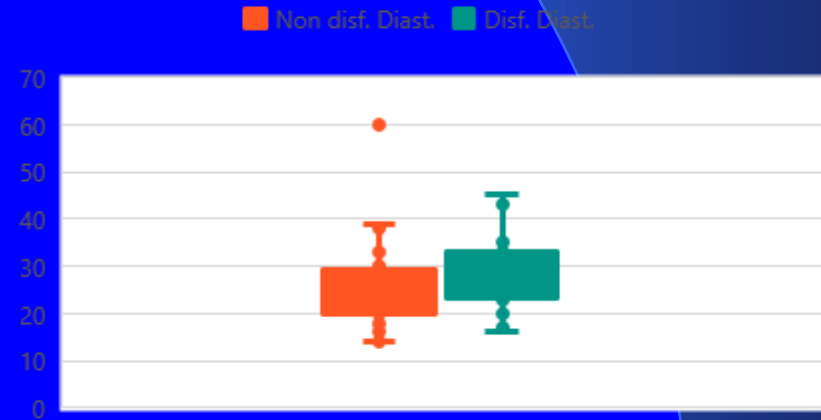
Groups, with or without LV dysfunction

	DIASTOLIC DYSFUNCTION	NO DIASTOLIC DYSFUNCTION	P
E/E'	11,86[11,18;14,57]	7,33[5,85;8,9]	<0,0001
E/A	0,87[0,76;1,25]	0,81[0,75;1,01]	0,20
HR	69,7±14	70,5±11	0,16
SBP	128,7±14,4	127,9±15,8	0,8
DBP	80,8±7,4	70,5±11,1	0,043
E'	0,06[0,04;0,07]	0,08[0,07;0,1]	<0,0001
PAPs	29,5[23,5;33]	24[20;29]	0,01
V _{MAXIT}	2,4±0,4	2,1±0,3	0,4
TAPSE	23,2±3,7	24,1±3,6	0,9
WALL THICKENING	21 (30,4%)	16(23,1%)	<0,0001

BASELINE E/E' RATIO



PAPs basale



Mean values registered in the two Groups, with or without LV dysfunction, at peak of exercise

	DIASTOLIC DYSFUNCTION	NO DIASTOLIC DYSFUNCTION	P
E/E'	16,1[15;20,5]	11,2[8,4;12,8]	<0,0001
E/A	0,93[0,79;1,19]	0,85[0,75;1,06]	0,3
HR	122±19	123±14	0,08
SBP	204,3±17,3	183,1±18,1	<0,0001
DBP	90,2±9,8	83,3±11,5	0,01
E'	0,07[0,05;0,08]	0,09[0,08;0,12]	<0,0001
PAPs	46,12±8,84	36,13±10,09	0,0001
V _{MAXIT}	3,24±0,38	2,73±0,49	<0,0001
TAPSE	26,2±4,6	26,3±4,4	0,9

E/E' at peak of exercise

