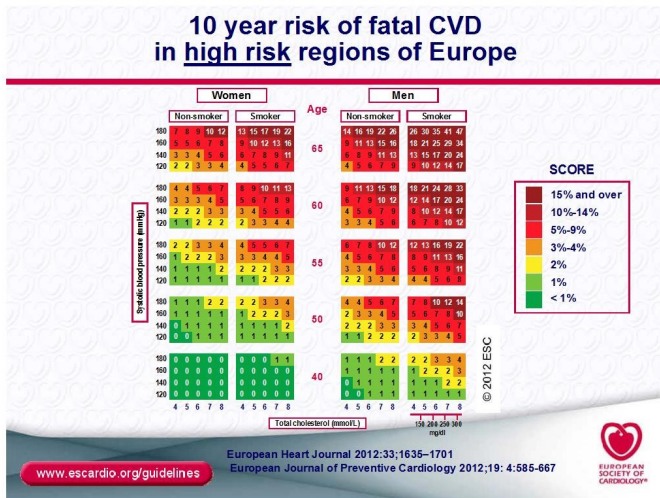




Нови инструментални методи за оценка на сърдечно-съдовия риск

*Наталия Спасова
УМБАЛ “Царица
Йоанна – ИСУЛ”*

Системи за оценка на сърдечно-съдов риск



PROCAM Score

LDL cholesterol (mg/dl)	HDL cholesterol (mg/dl)	Systolic blood pressure (mmHg)	Age (years)
≤100	≤35	<110	35-65
101-105	36-37	110-119	Age-35
106-110	38-39	120-129	Smoker
111-115	40-41	130-139	Never
116-120	42-43	140-149	Former
121-125	44-45	150-159	Current
126-130	46-47	≥160	Antihypertensive drug
131-135	48-49	<120	No
136-140	50-51	≥120	Yes
141-145	52-53	Diabetes	MI in family history
146-150	54-55		
151-155	>55		
156-160			
161-165			
166-170	Triglycerides (mg/dl)		
171-175	<100		
176-180	100-149		
181-185	150-199		
186-190	≥200		
191-195			
196-200			
>200			

Pooled Cohort Risk Assessment Equations

Predicts 10-year risk for a first atherosclerotic cardiovascular disease (ASCVD) event

Risk Factors for ASCVD

Gender: Male Female

Age: years

Race:

Total Cholesterol: mg/dL

HDL Cholesterol: mg/dL

Systolic BP: mmHg

Receiving treatment for high blood pressure (if SBP > 120 mmHg): No Yes

Diabetes: No Yes

Smoker: No Yes

US units

http://www.chd-taskforce.de/pdf/sk_procam_07e.pdf

- Много лесни и удобни за използване в клинична практика
- Оценяват взаимодействието на рисковите фактори
- Не оценяват промените в артериалните съдове

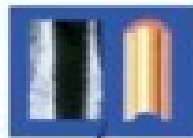
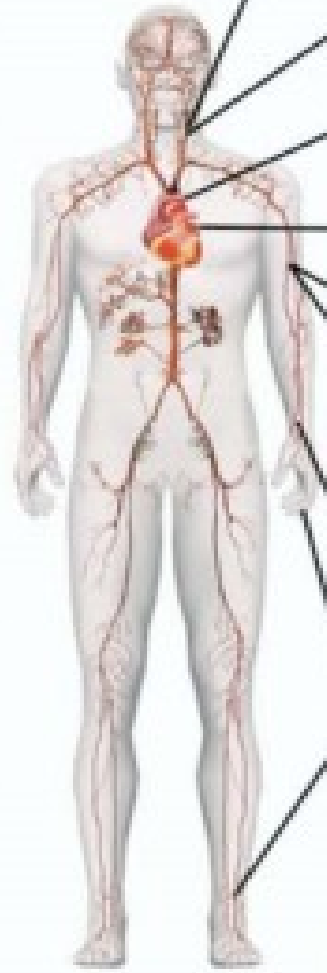
Screening for Atherosclerosis

Risk Factors vs Disease

Numerous Risk Factors

- High LDL →
- Low HDL →
- High BP →
- Diabetes →
- Smoking →
- CRP →
- Metabolic Syndrome →
- Lp(a) →
- Homocysteine →
- Dense LDL →
- Lp-PLA2 →
- ApoB/ApoA →
- Family History →
- Sedentary Life →
- Obesity →
- Stress →
- ... →
- ? →

Over 200 risk factors have been reported.



Carotid IMT and Plaque Measured by Ultrasound



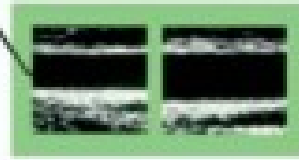
Aortic and Carotid Plaque Detected by MRI



Coronary Calcium Score Measured by CT



Ankle — Brachial Index



Brachial Vasoreactivity Measured by Ultrasound



Vascular Compliance Measured by Radial Tonometry



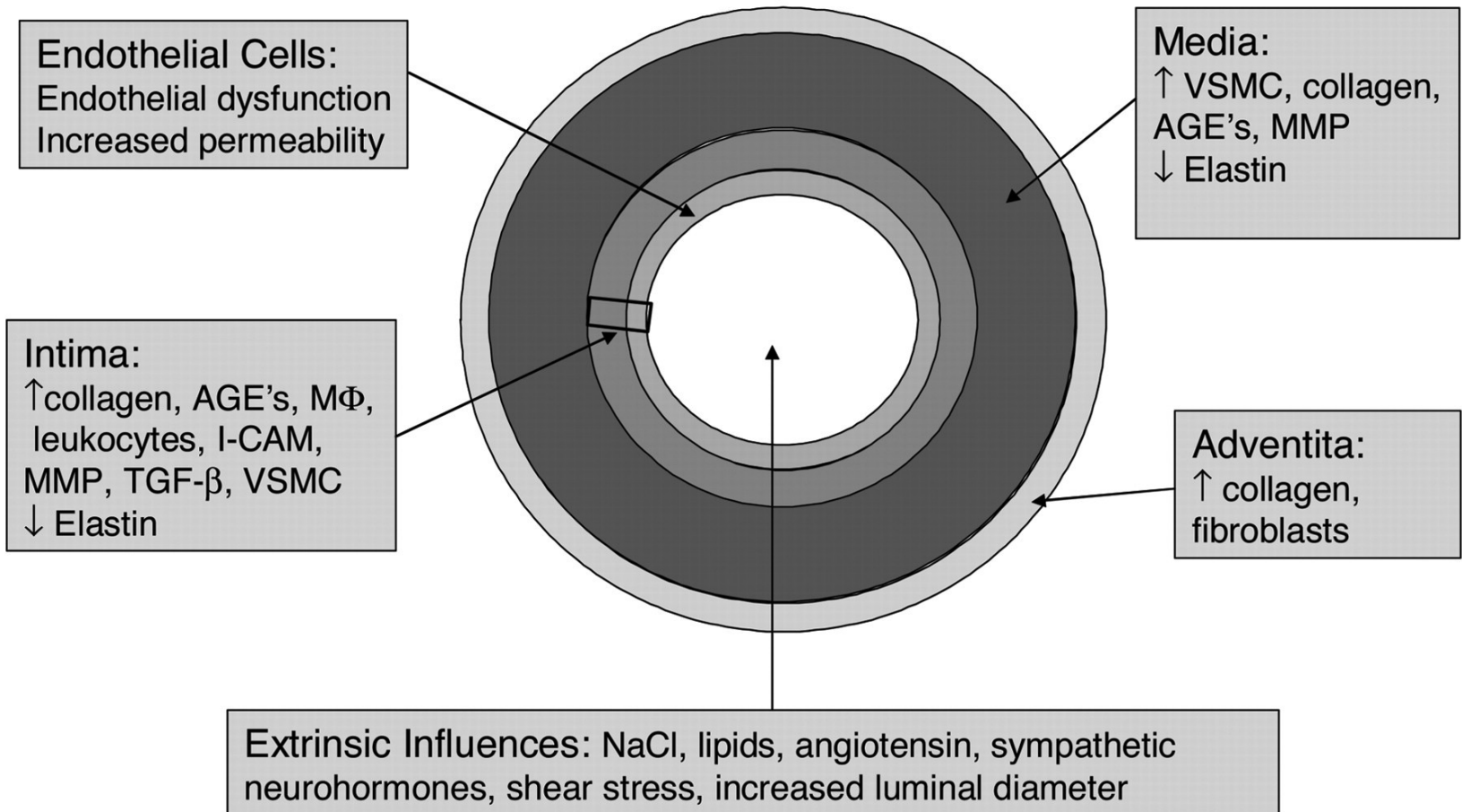
Microvascular Reactivity Measured by Fingertip Tonometry

Examples of Arterial Structure Tests

Examples of Arterial Function Tests

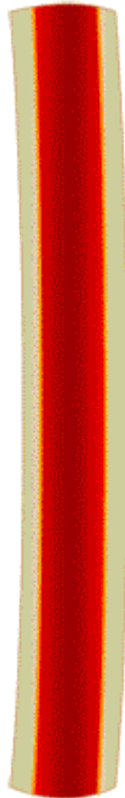
СЪДОВА РИГИДНОСТ

Figure 1. Summary of the multiple causes and locations of arterial stiffness.

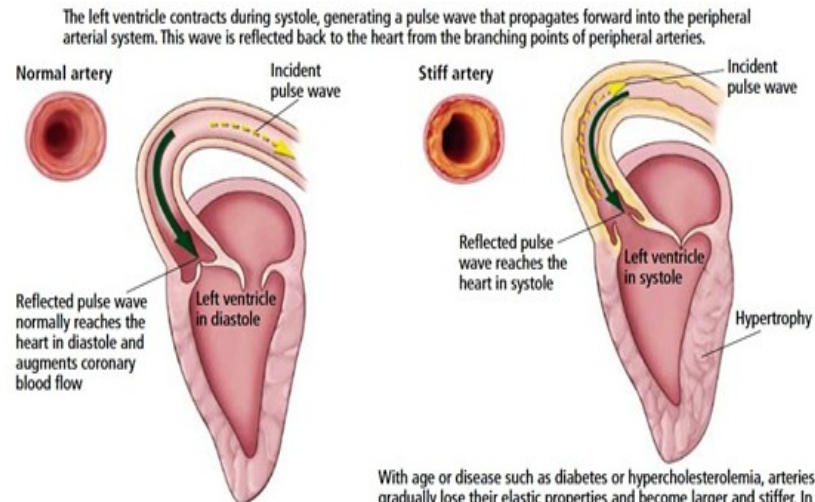


Zieman S J et al. *Arterioscler Thromb Vasc Biol.* 2005;25:932-943

Pulse Wave Velocity

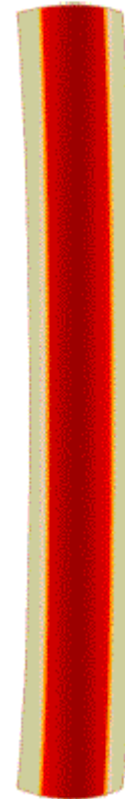


Elastic



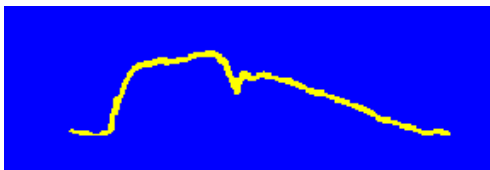
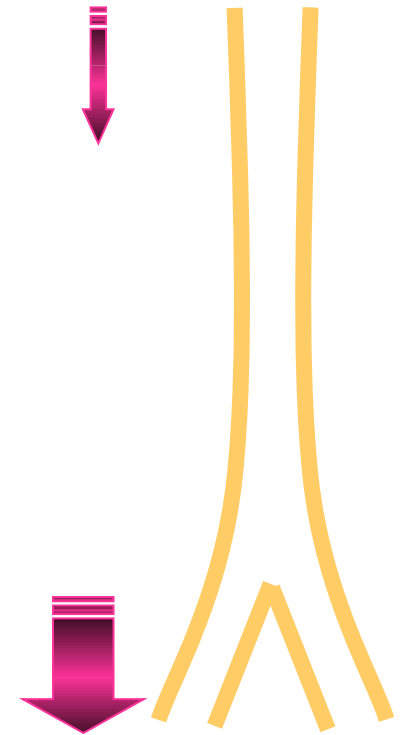
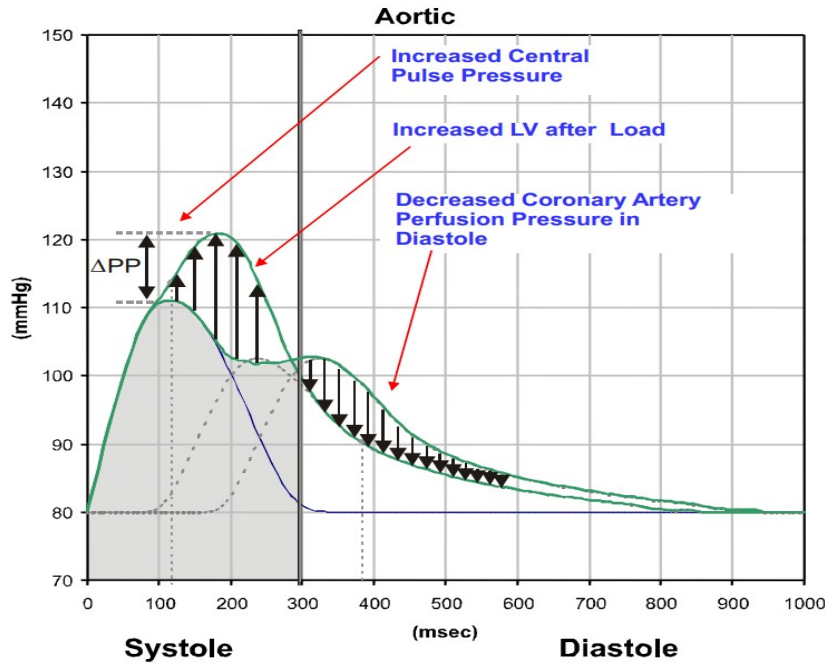
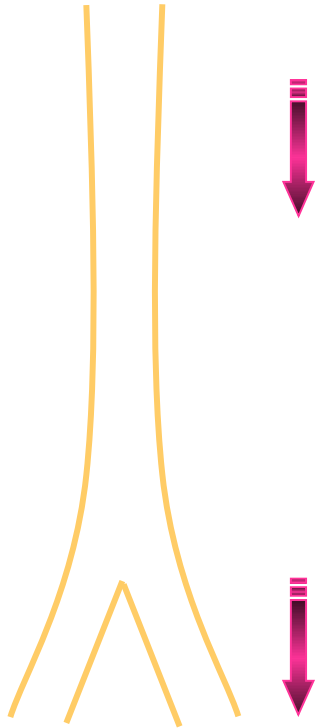
With age or disease such as diabetes or hypercholesterolemia, arteries gradually lose their elastic properties and become larger and stiffer. In these arteries, the reflected wave returns faster and merges with the incident wave in systole. This results in a higher left ventricular afterload and decreased perfusion of coronary arteries, leading to left ventricular hypertrophy and increased arterial and central blood pressure.

CCF

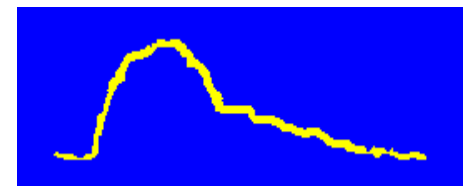


Stiff

Wave reflection



Elastic



Stiff

**Атериална
Хипертония**

**Захарен
Диабет**



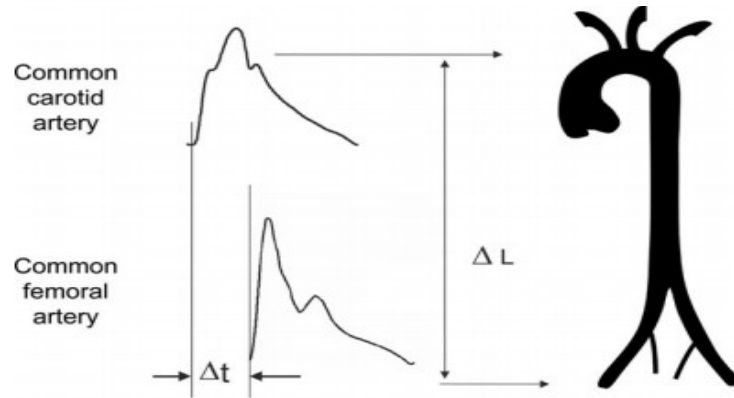
Методи за измерване на съдова ригидност

Expert consensus document on arterial stiffness: methodological issues and clinical applications

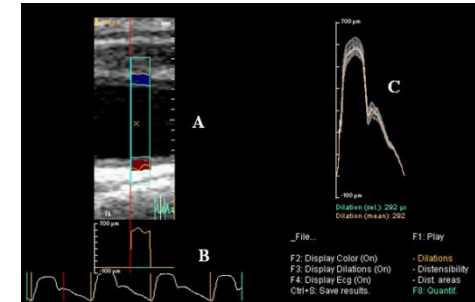


Измерване на каротидо-феморална скорост на пулсовата вълна, SphygmoCor, AtCor

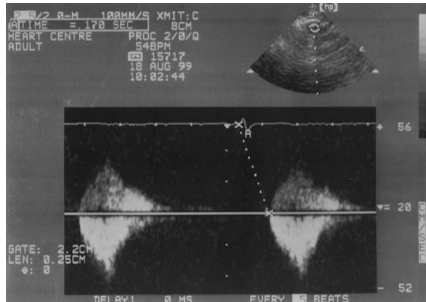
Position statement: PWV. Carotid-femoral PWV is considered as the 'gold-standard' measurement of arterial stiffness.



European Heart Journal (2006) 27, 2588-2605

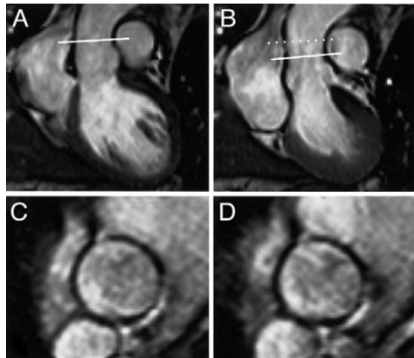


Измерване на параметрите на съдова ригидност с помощта на TDI Cardiovascular Ultrasound 2007



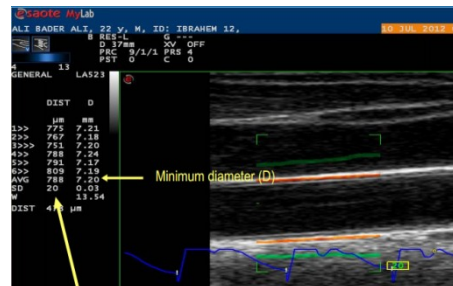
Pulsed Doppler на дясна обща феморална артерия. Изчисляване на интервала между началото на QRS комплекса и началото на систолната крива.

Локално измерване на съдова ригидност Journal of Human Hypertension 2003; 17:407-412



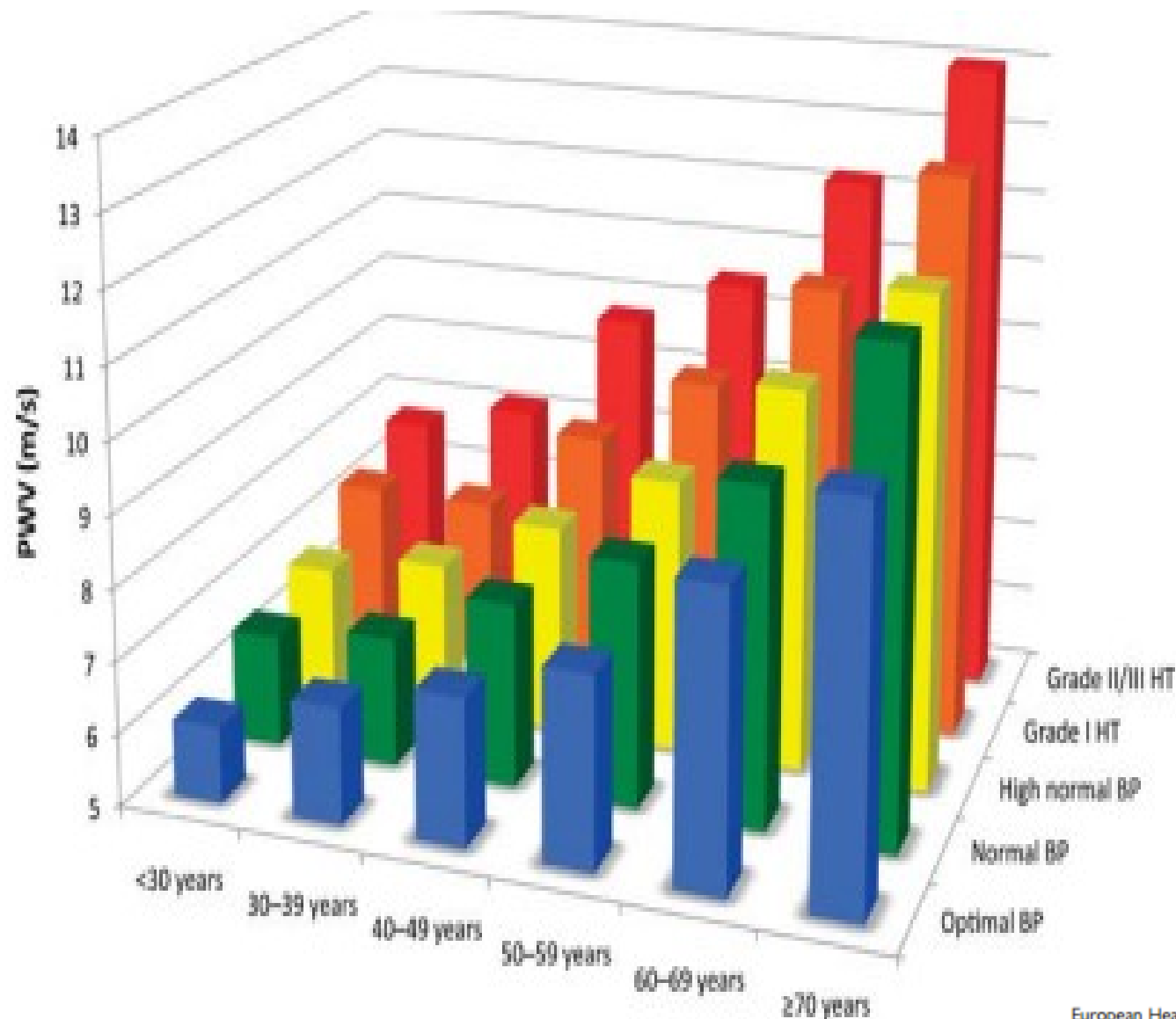
Distensibility measurement of the Aortic Root with Cardiac MRI

J Am Coll Cardiol 2011;57:1511-22



Измерване на локална съдова ригидност на каротидна артерия

Determinants of pulse wave velocity in healthy people and in the presence of cardiovascular risk factors: 'establishing normal and reference values'



EUROPEAN
SOCIETY OF
CARDIOLOGY®

Aortic Pulse Wave Velocity Improves Cardiovascular Event Prediction

An Individual Participant Meta-Analysis of Prospective Observational Data From 17,635 Subjects

Table 1

Pooled Adjusted Hazard Ratios (95% CIs) of a 1-SD Increase in Log_e-Transformed aPWV for All-Cause Mortality, CVD Mortality, CHD Events, Stroke Events, and CVD Events

	Model 1*	Model 2*	Model 3*
CHD events (n = 1,195)	1.35 (1.22-1.50)	1.32 (1.18-1.48)	1.23 (1.11-1.35)
CVD events (n = 1,785)	1.45 (1.30-1.61)	1.37 (1.23-1.52)	1.30 (1.18-1.43)
Stroke events (n = 641)	1.54 (1.34-1.78)	1.37 (1.21-1.54)	1.28 (1.16-1.42)
CVD mortality (n = 395)	1.41 (1.27-1.56)	1.35 (1.20-1.53)	1.28 (1.15-1.43)
All-cause mortality (n = 2,041)	1.22 (1.16-1.27)	1.20 (1.15-1.26)	1.17 (1.11-1.22)

*Model 1 adjusts for sex and age group; model 2 adjusts for sex, age group, and systolic blood pressure; and model 3 additionally adjusts for other risk factors (cholesterol, high-density lipoprotein cholesterol, smoking status, presence of diabetes, and antihypertensive medication), stratified by race in the Sutton-Tyrell study (27). Not all studies had data on every risk factor.

aPWV = aortic pulse wave velocity; CHD = coronary heart disease; CI = confidence interval; CVD = cardiovascular disease.

Net Reclassification Statistics Showing Percent Change in 5-Year Risk Prediction (and 5- and 10-Year Overall Reclassification) Associated With Including log_e aPWV as a Risk Factor in the Fully-Adjusted Model

Reclassification indices showed that the addition of aPWV improved risk prediction (13% for 10-year CVD risk for intermediate risk) for some subgroups.

Impact of Aortic Stiffness Attenuation on Survival of Patients in End-Stage Renal Failure

Alain P. Guerin, Jacques Blacher, Bruno Pannier, Sylvain J. Marchais, Michel E. Safar and
Gérard M. London

Circulation
JOURNAL OF THE AMERICAN HEART ASSOCIATION

150 пациенти
Проследяване 52 месеца

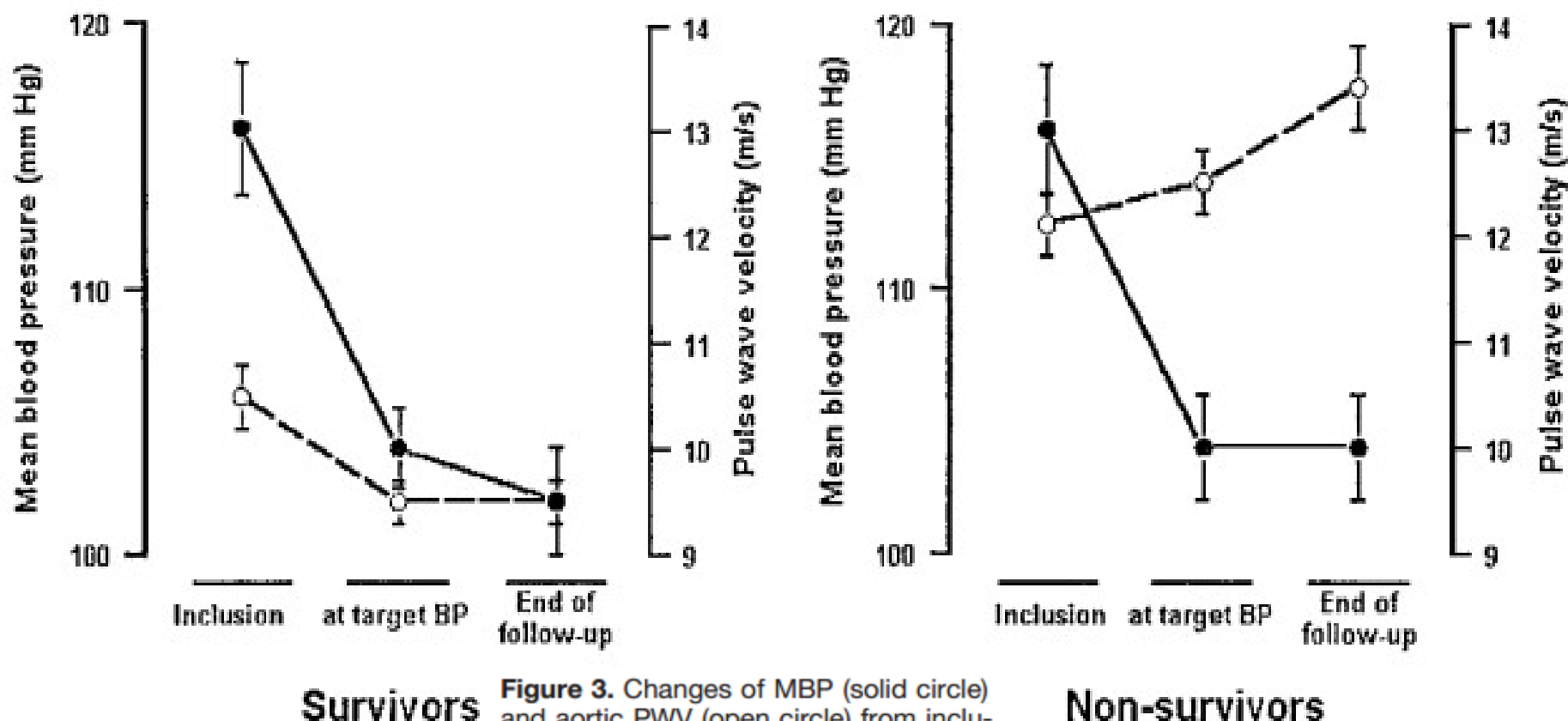


Figure 3. Changes of MBP (solid circle) and aortic PWV (open circle) from inclusion to end of follow-up for survivors and nonsurvivors. Values are mean \pm SEM.

Non-survivors

2013 ESH/ESC Guidelines for the management of arterial hypertension



Table 4 Factors—other than office BP—influencing prognosis; used for stratification of total CV risk in Figure 1

Asymptomatic organ damage

Pulse pressure (in the elderly) ≥ 60 mmHg

Electrocardiographic LVH (Sokolow–Lyon index >3.5 mV; RaVL >1.1 mV; Cornell voltage duration product >244 mV*ms), or

Echocardiographic LVH [LVM index: men >115 g/m²; women >95 g/m² (BSA)]^a

Carotid wall thickening (IMT >0.9 mm) or plaque

Carotid–femoral PWV >10 m/s

Ankle-brachial index <0.9

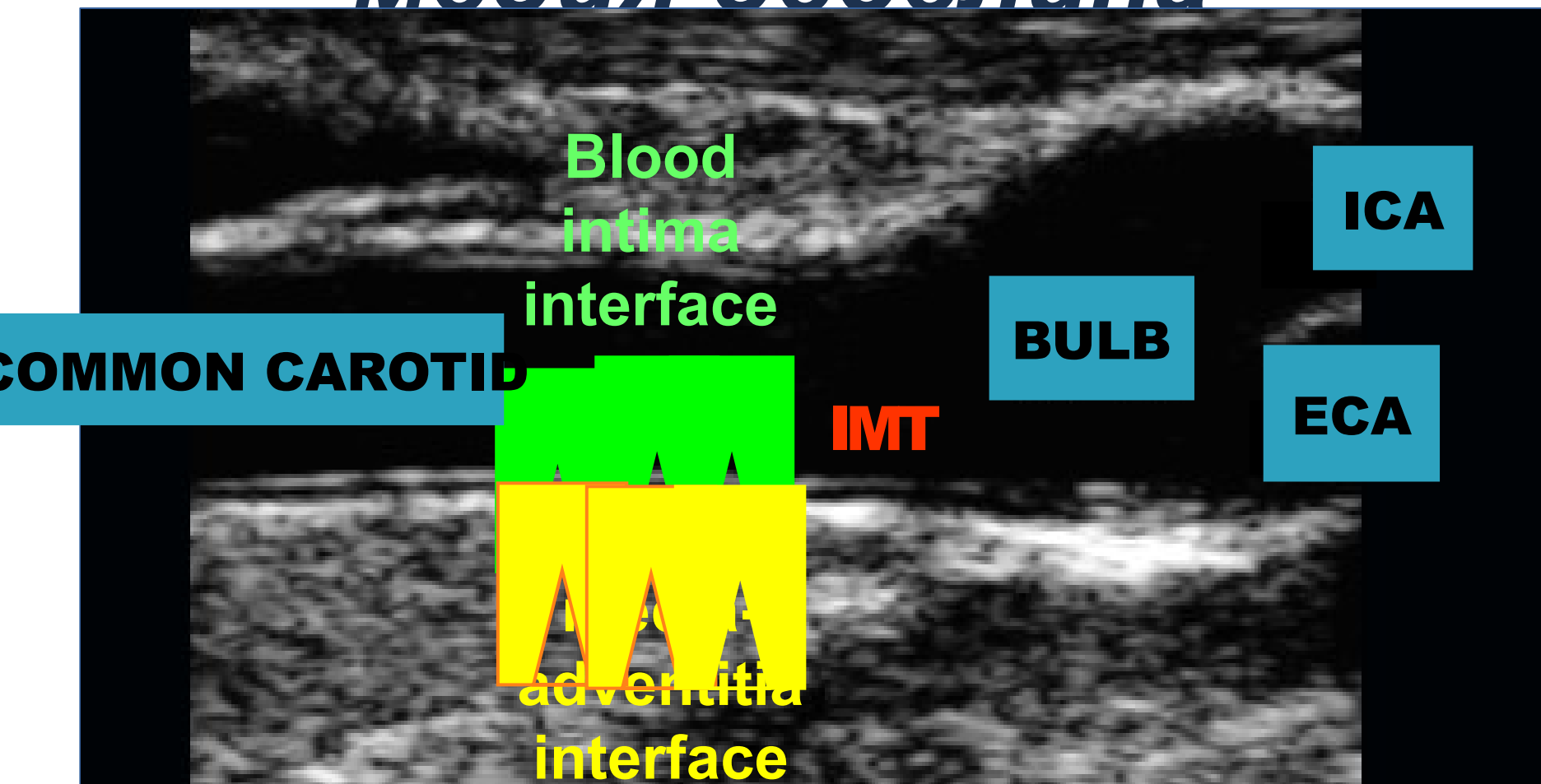
CKD with eGFR 30–60 mL/min/1.73 m² (BSA)

Microalbuminuria (30–300 mg/24 h), or albumin–creatinine ratio (30–300 mg/g; 3.4–34 mg/mmol) (preferentially on morning spot urine)

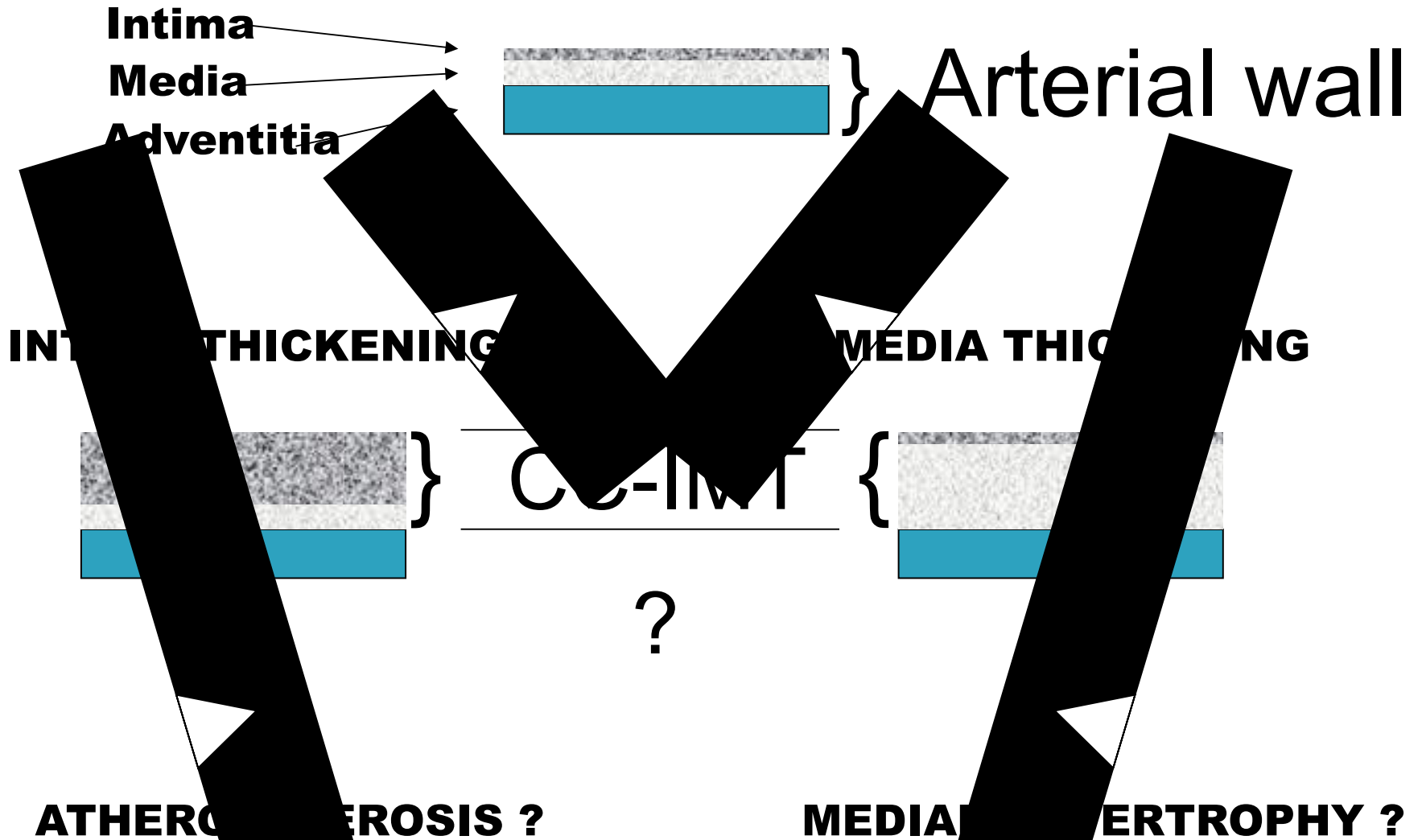
Total cardiovascular risk assessment

Recommendations	Class ^a	Level ^b	Ref. ^c
In asymptomatic subjects with hypertension but free of CVD, CKD, and diabetes, total CV risk stratification using the SCORE model is recommended as a minimal requirement.	I	B	43
As there is evidence that OD predicts CV death independently of SCORE, a search for OD should be considered, particularly in individuals at moderate risk.	IIa	B	51, 53

Измерване на интима-медия дебелина



Ultrasonography cannot distinguish



Carotid Intima-Media Thickness and Presence or Absence of Plaque Improves Prediction of Coronary Heart Disease Risk

The ARIC (Atherosclerosis Risk In Communities) Study

13145 пациенти
15 години просл.

Journal of the American College of Cardiology

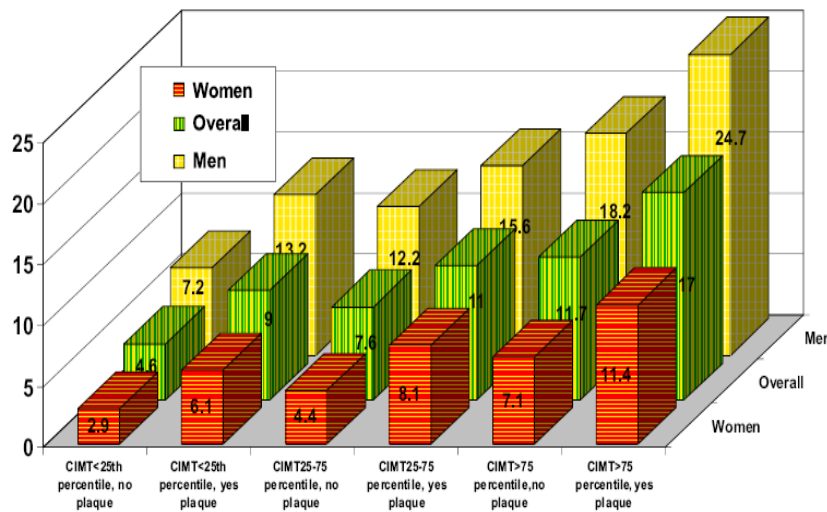


Figure 1 Adjusted Coronary Heart Disease Incidence Rate per 1,000 Person-Years Adjusted by CIMT Categories With and Without Plaque

For every carotid intima-media thickness (CIMT) category (i.e., <25th percentile, 25th to 75th percentile, and >75th percentile), for the overall group (green bars), men (yellow bars), or women (orange bars), having carotid artery plaque is associated with a higher incidence of coronary heart disease.

Table 2 Adjusted AUC for Different Models, With 95% Confidence Interval for Difference in Adjusted AUC Comparing Various Models With TRF-Only Model

Model	Overall	Men	Women
TRF only	0.742	0.674	0.759
TRF + CIMT	0.750 (0.005 to 0.012)	0.690 (0.009 to 0.022)	0.762 (-0.002 to 0.006)
TRF + plaque	0.751 (0.006 to 0.013)	0.686 (0.005 to 0.017)	0.770 (0.005 to 0.016)
TRF + CIMT + plaque	0.755 (0.008 to 0.017)	0.694 (0.011 to 0.027)	0.770 (0.005 to 0.017)
TRF + CIMT + plaque vs. TRF + IMT	(0.001 to 0.006)	(-0.001 to 0.006)	(0.003 to 0.012)
TRF + IMT + plaque vs. TRF + plaque	(0.001 to 0.005)	(0.002 to 0.011)	(-0.002 to 0.002)

AUC = area under the curve; CI = confidence interval; CIMT = carotid intima-media thickness; TRF = traditional risk factors.

Table 6 NRI Using Various Comparison Models in Overall Sample, Men, and Women

Model	Overall		Men		Women	
	NRI	Clinical NRI	NRI	Clinical NRI	NRI	Clinical NRI
TRF vs. TRF + CIMT	7.1 (2.2 to 10.6)	16.7 (9.3 to 22.4)	8.9 (3.4 to 15.1)	15.8 (8.6 to 24.6)	6.1 (-2.3 to 9.4)	15.9 (1 to 23.3)
TRF vs. TRF + plaque	7.7 (2.3 to 11.4)	17.7 (10.9 to 24.7)	4.2 (0.2 to 12.2)	10.5 (4.5 to 20.5)	10.2 (0.7 to 15.4)	25.6 (7.8 to 37.6)
TRF vs. TRF + CIMT + plaque	9.9 (3.8 to 13.5)	21.7 (13.4 to 28.2)	8.9 (4.1 to 17.1)	16.4 (9.5 to 27)	9.8 (1.1 to 15.4)	25.4 (9 to 37)
TRF + CIMT vs. TRF + CIMT + plaque	2.8 (-1.2 to 6.4)	10.6 (3.8 to 16.5)	0.03 (-2.6 to 6.3)	5.1 (0.3 to 13.2)	3.6 (-1.7 to 11.6)	12.8 (2.5 to 28.6)
TRF + plaque vs. TRF + CIMT + plaque	2.1 (-1.1 to 5.3)	7.9 (2.6 to 13.3)	4.8 (-0 to 10)	10.7 (4.3 to 19)	-0.3 (-3.7 to 3.6)	2.5 (-3.5 to 10.3)

European Guidelines on cardiovascular disease prevention in clinical practice (version 2012)

European Heart Journal (2012) **33**, 1635–1701

Recommendations	Class ^a	Level ^b	GRADE	Ref ^c
Measurement of carotid intima-media thickness and/or screening for atherosclerotic plaques by carotid artery scanning should be considered for cardiovascular risk assessment in asymptomatic adults at moderate risk.	IIa	B	Strong	130–132

2013 ACC/AHA Guideline on the Assessment of Cardiovascular Risk

A Report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines

<p>5. CIMT is not recommended for routine measurement in clinical practice for risk assessment for a first ASCVD event.</p>	<p>N (No Recommendation For or Against)</p>	<p>Appendix 1</p>	<p>III: No Benefit[†]</p>	<p>B (12,16,18)</p>
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*Journal of the American College of
Cardiology (2014), doi:
10.1016/j.jacc.2013.11.005*

Common Carotid Intima-Media Thickness Measurements in Cardiovascular Risk Prediction

A Meta-analysis

45828 пациенти

Table 3. Summary of the Indices of Added Value in the Total USE-IMT Cohort and in the Intermediate-Risk Categories, by Sex

	All	USE-IMT	Men	Women
NRI, % (95% CI)	0.8 (0.1 to 1.6)		0.9 (-0.2 to 1.9)	0.8 (-0.2 to 1.6)
IDI (95% CI)	0.0024 (0.0012 to 0.0036)		0.0024 (0.0004 to 0.0041)	0.0025 (0.0009 to 0.0040)
Relative IDI, %	3.6		3.6	3.7
USE-IMT, Intermediate-Risk Group (5% to <20%)				
NRI, % (95% CI)	3.6 (2.7 to 4.6)		3.2 (2.3 to 4.4)	3.9 (2.7 to 4.9)
IDI (95% CI)	0.0024 (0.0012 to 0.0036)		0.0019 (0.0003 to 0.0034)	0.0031 (0.0013 to 0.0048)
Relative IDI, %	3.6		2.7	4.6

Abbreviations: IDI, integrated discrimination improvement; NRI, net reclassification improvement; USE-IMT, USE Intima-Media Thickness collaboration.

Conclusion The addition of common CIMT measurements to the Framingham Risk Score was associated with small improvement in 10-year risk prediction of first-time myocardial infarction or stroke, but this improvement is unlikely to be of clinical importance.

JAMA. 2012;308(8):796-803

www.jama.com

Измерване на тиббио-брахиален индекс



How to calculate the ankle-brachial index

Right arm:

Systolic pressure 120 mm Hg

Left arm:

Systolic pressure 100 mm Hg

Right ankle: Systolic pressure

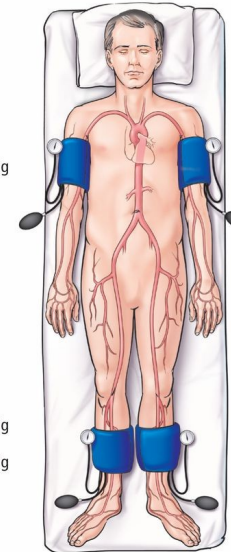
Posterior tibial (PT) 68 mm Hg

Dorsalis pedis (DP) 64 mm Hg

Left ankle: Systolic pressure

Posterior tibial (PT) 136 mm Hg

Dorsalis pedis (DP) 132 mm Hg



Right ABI equals ratio of:

Higher of the right ankle pressure (PT or DP)
Higher arm pressure (right or left arm)

$$\frac{68 \text{ mm Hg}}{120 \text{ mm Hg}} = 0.57^*$$

Left ABI equals ratio of:

Higher of the left ankle pressure (PT or DP)
Higher arm pressure (right or left arm)

$$\frac{136 \text{ mm Hg}}{120 \text{ mm Hg}} = 1.13^*$$

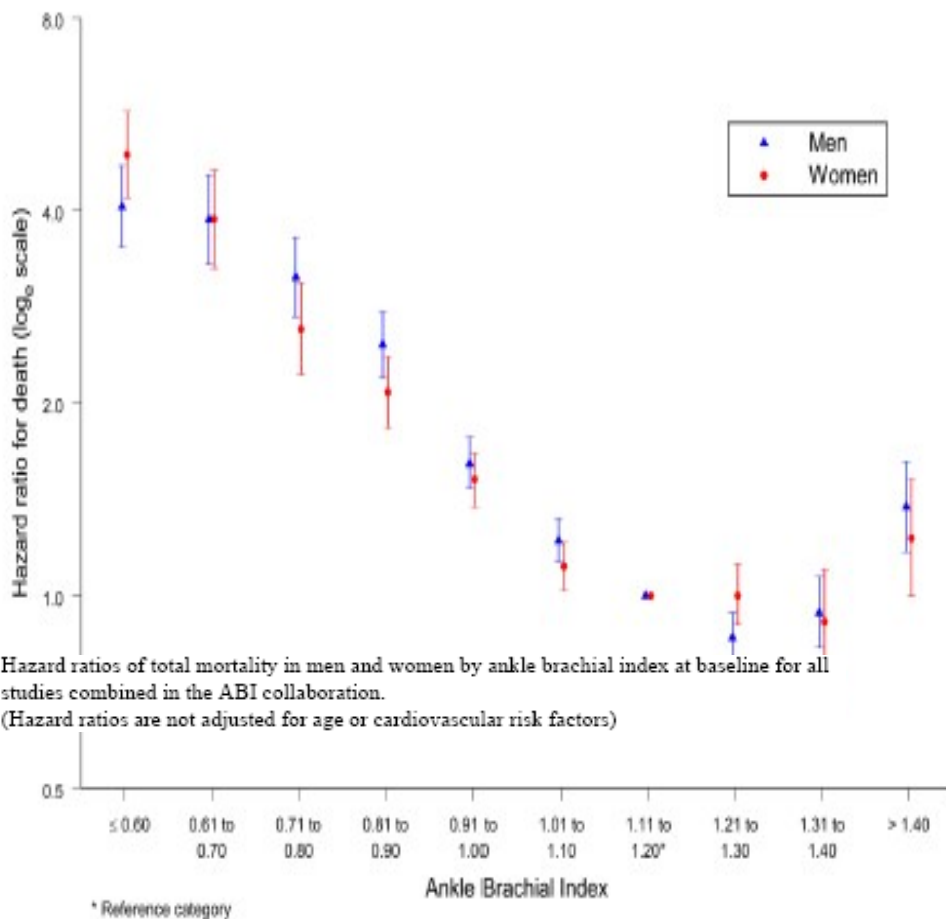
*The lower of these numbers is the patient's overall ankle-brachial index. Overall ankle-brachial index = 0.57

Medical Illustrator: Joseph Pangrace
©2012 CCF

Ankle Brachial Index Combined with Framingham Risk Score to Predict Cardiovascular Events and Mortality: A Meta-analysis

Ankle Brachial Index Collaboration

>48 294 пациенти без установено ССЗ



Ankle Brachial Index:	≤ 0.90	0.91 to 1.00	1.11 to 1.40	>1.40	Overall
Total Coronary Heart Disease ³ % (n)					
<u>Men</u>					
Framingham Risk Category ²					
Low <10%	8 (76)	5 (1076)	4 (4255)	5 (216)	5 (5643)
Intermediate 10-19%	16 (245)	12 (2069)	12 (4815)	8 (203)	13 (7392)
High ≥20%	40 (1149)	21 (3406)	18 (3888)	14 (1173)	23 (8398)
<u>Women</u>					
Framingham Risk Category ²					
Low <10%	21 (1083)	10 (8191)	9 (7009)	11 (321)	11 (15905)
Intermediate 10-19%	25 (338)	12 (2429)	11 (2433)	13 (143)	13 (5563)
High ≥20%	44 (200)	21 (598)	22 (577)	34 (43)	27 (1418)

Изследването на ABI в допълнение към FRS рекласифицира 19 % от мъжете и 36 % от жените

European Guidelines on cardiovascular disease prevention in clinical practice (version 2012)

Measurement of ankle-brachial index should be considered for cardiovascular risk assessment in asymptomatic adults at moderate risk.



European Heart Journal (2012) **33**, 1635–1701

2013 ACC/AHA Guideline on the Assessment of Cardiovascular Risk

A Report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines

3. If, after quantitative risk assessment, a risk-based treatment decision is uncertain, assessment of 1 or more of the following—family history, hs-CRP, CAC score, or ABI—may be considered to inform treatment decision making.

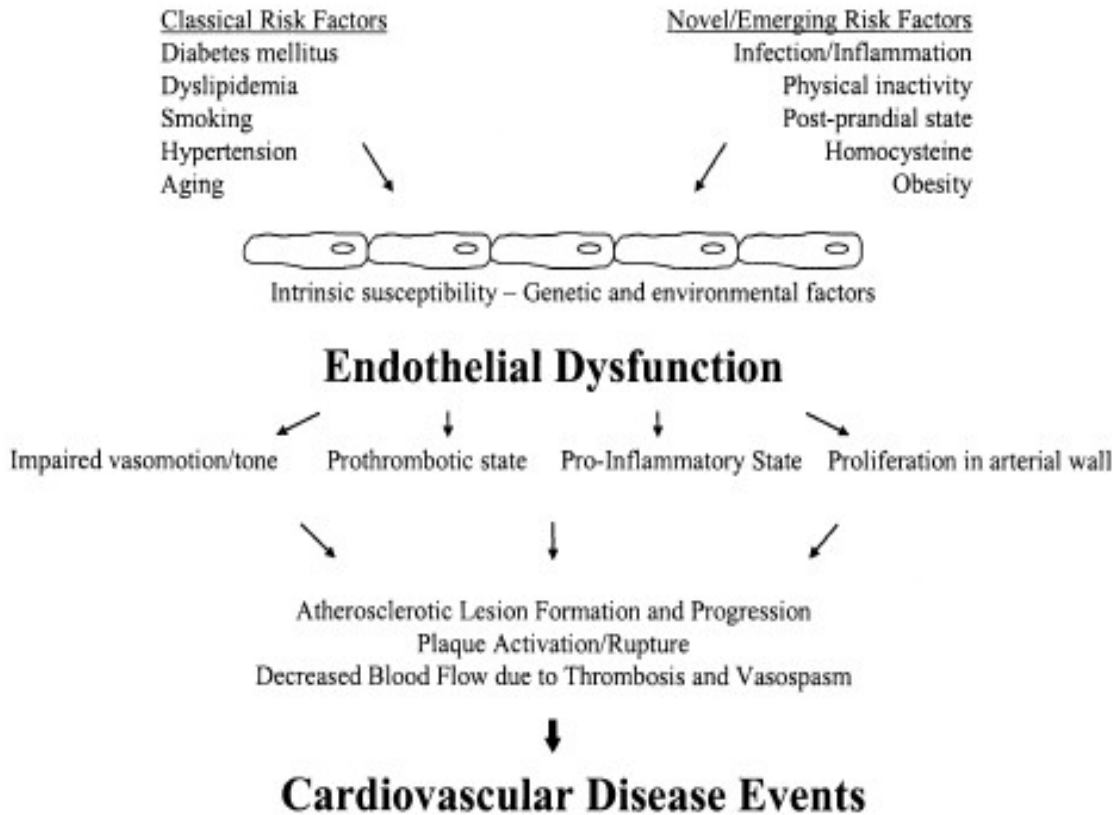
E (Expert Opinion)

Appendix 1

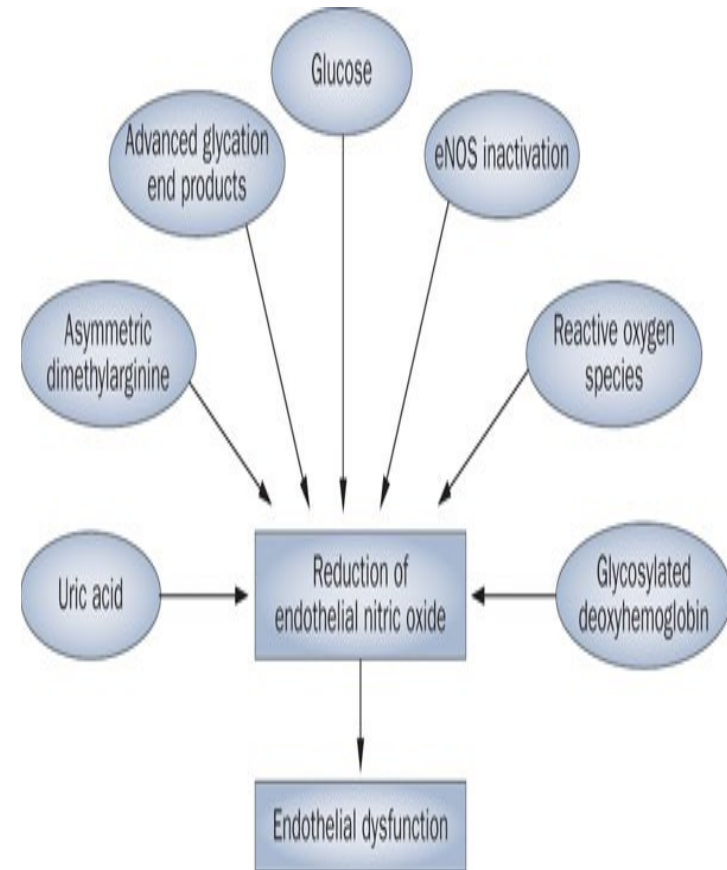
IIb†

Journal of the American College of Cardiology (2014), doi: 10.1016/j.jacc.2013.11.005

Ендотелна дисфункция

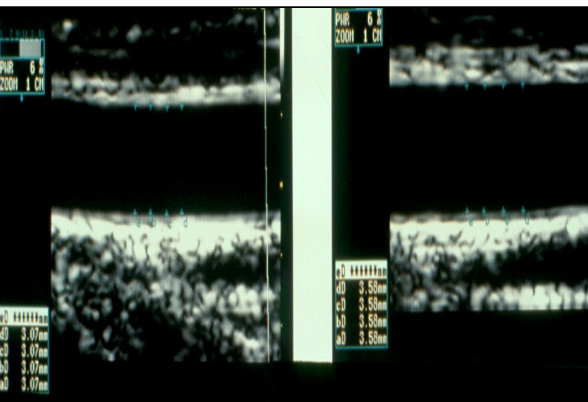


J Am Coll Cardiol 2003;42:1149–60

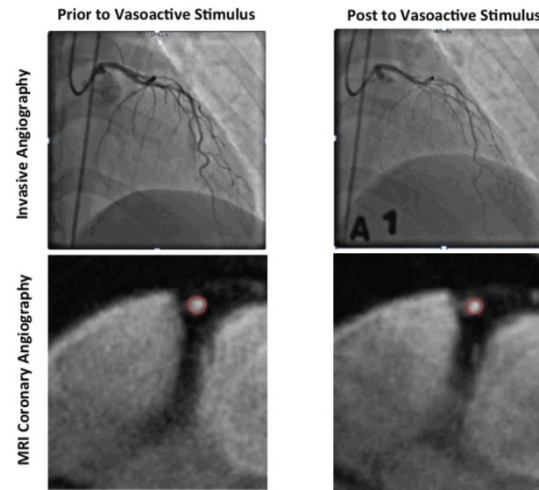


Nature Reviews Nephrology 7, 36-44 (January 2011)

Измерване на ендотелната функция



Поток-медирана вазодилатация на брахиална артерия

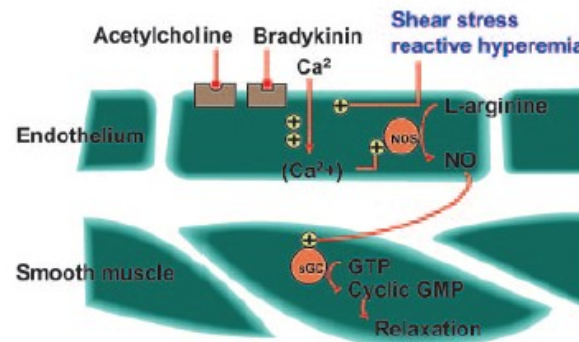


Корелация между инвазивна оценка на ендотелна функция и MRI
JACC April 1, 2014



Endo-PAT2000

Периферна артериална тонометрия



Circulation. 2005;111:363-368

Заклучение

- Неинвазивните методи за изследване на периферните атериалните съдове помагат да се оценят биологичните промени в съдовете и са маркери за субклинична органична увреда
- Приложението им е особено полезно при пациенти с интермедиерен риск за сърдечно-съдови усложнения
- Повишена съдова ригидност, централно аортно и пулсово налягане, ИМД, АВІ, САС, и ендотелна дисфункция са предиктори за неблагоприятна прогноза
- Дали намаляването на съдовата ригидност, ИМД, подобряване на ендотелната функция ще подобри прогнозата на пациентите все още предстои да бъде изследвано