

# Validity of Aortic Stiffness as Surrogate Marker of Cardiovascular Disease

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# Surrogate Markers of Cardiovascular Disease

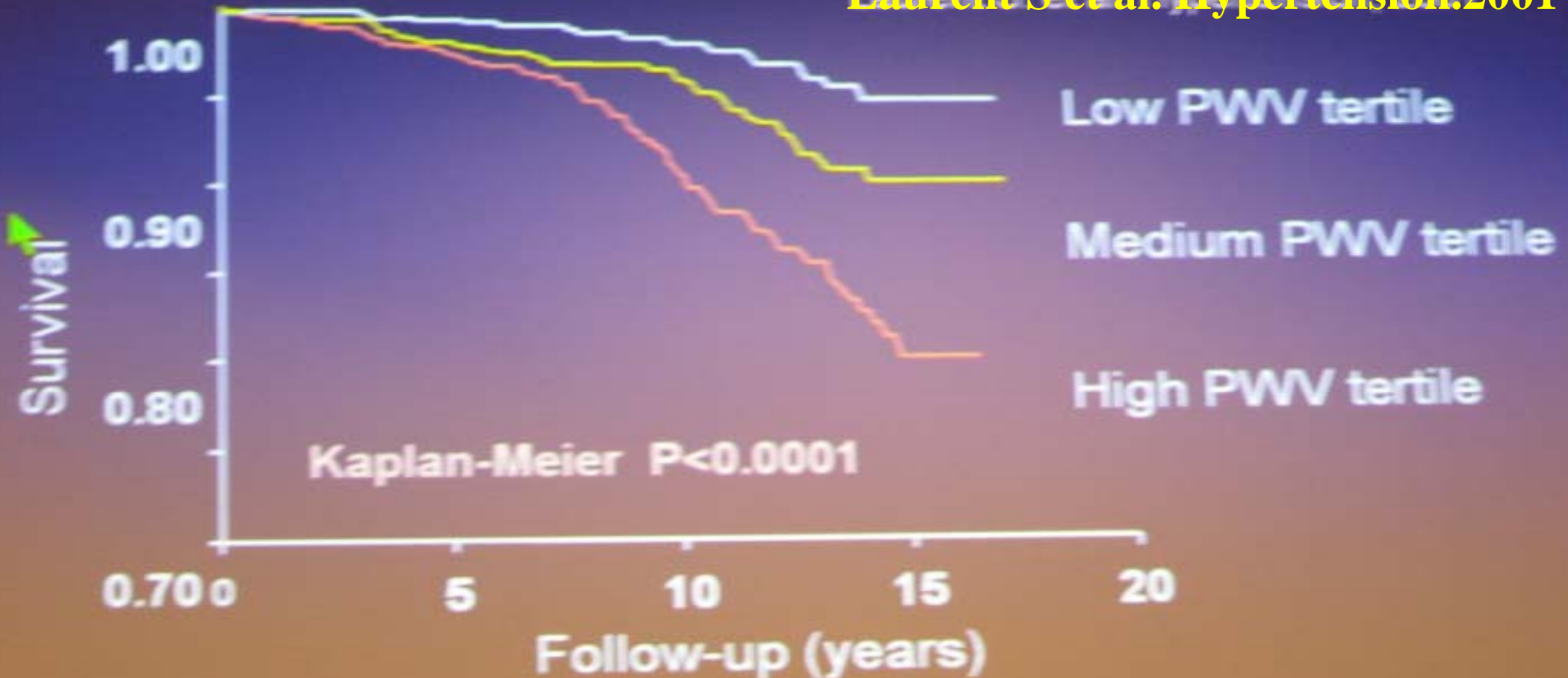
- Carotid IMT
- LVH
- Endothelial function(FMD)
- CRP
- Aortic stiffness

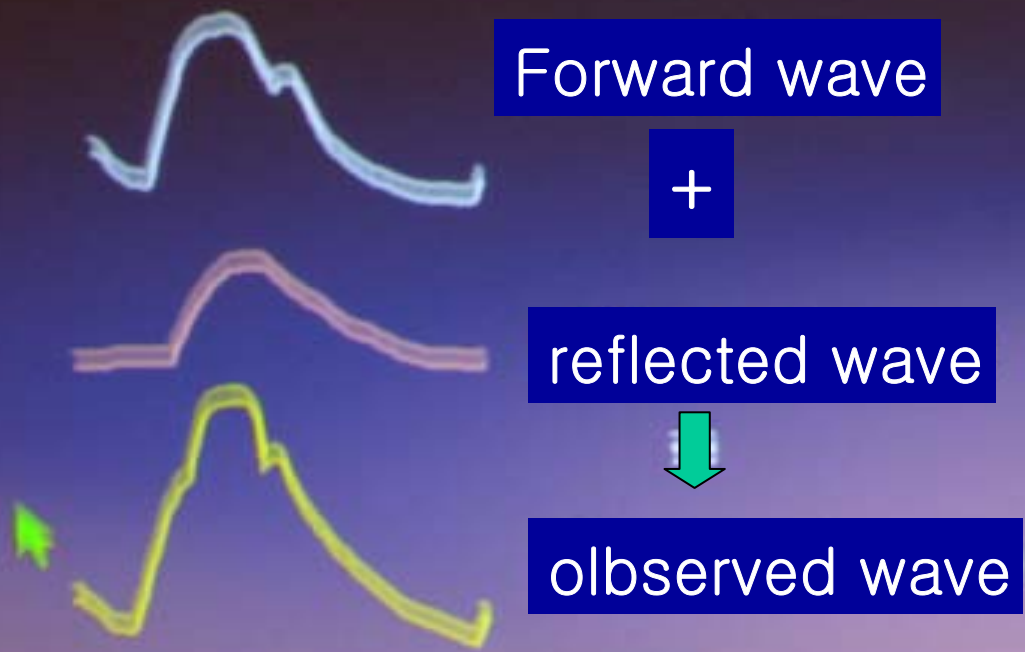
# Parameters of Aortic Stiffness

- Aortic strain
- Aortic distensibility
- Aortic stiffness index
- Augmentation index
- Pulse wave velocity

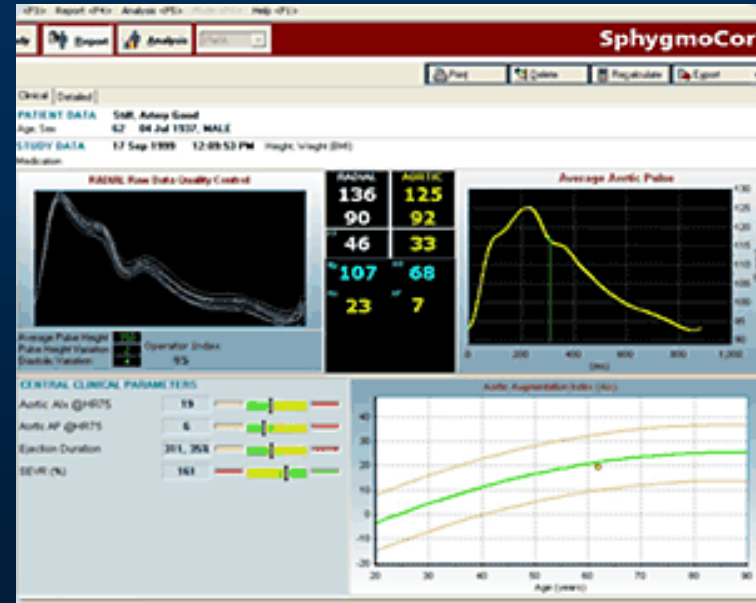
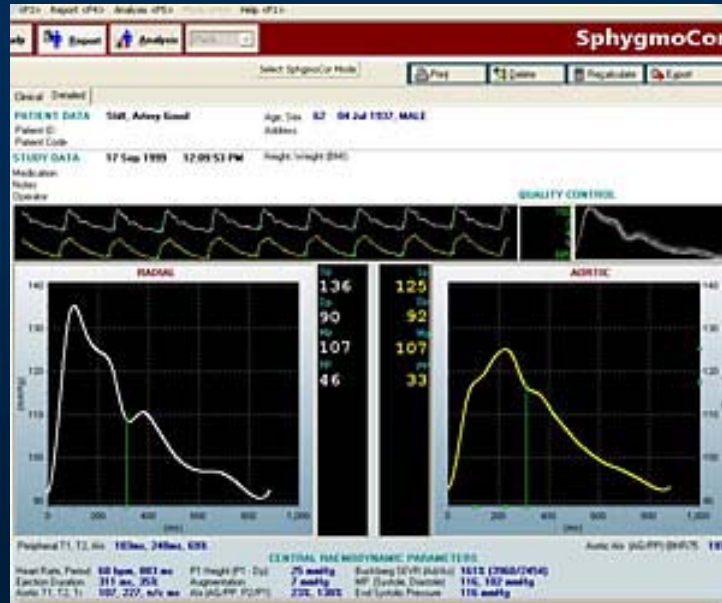
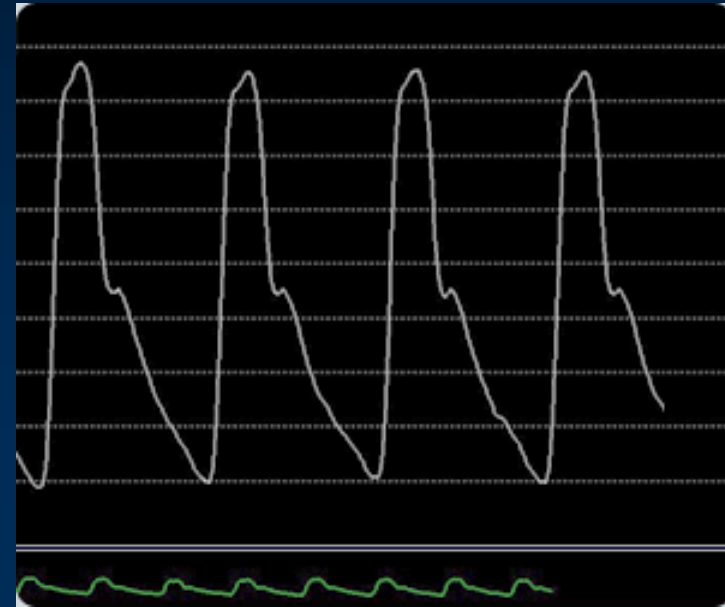
# Aortic stiffness and all-cause mortality in 1980 hypertensives

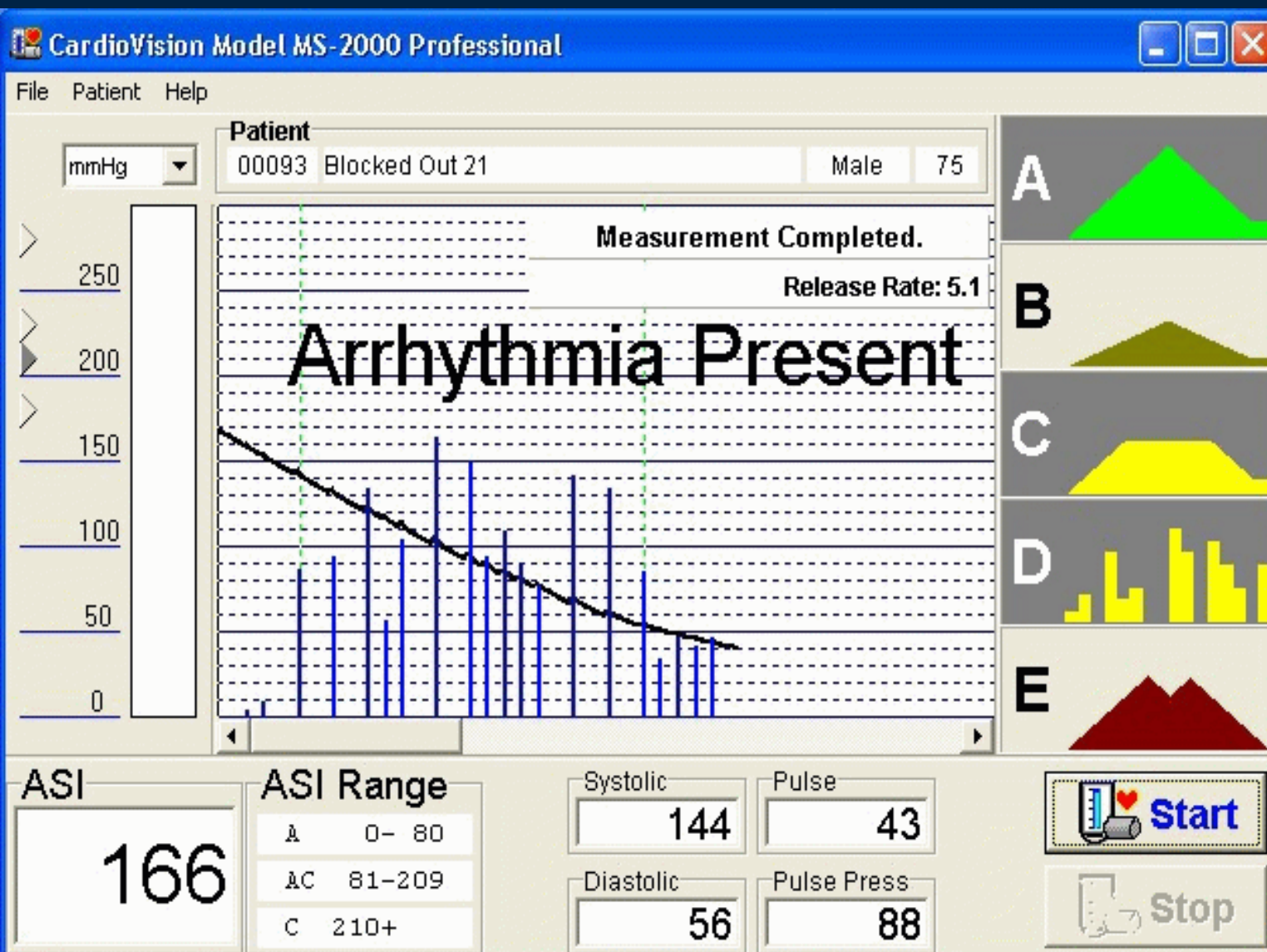
Laurent S et al. Hypertension. 2001



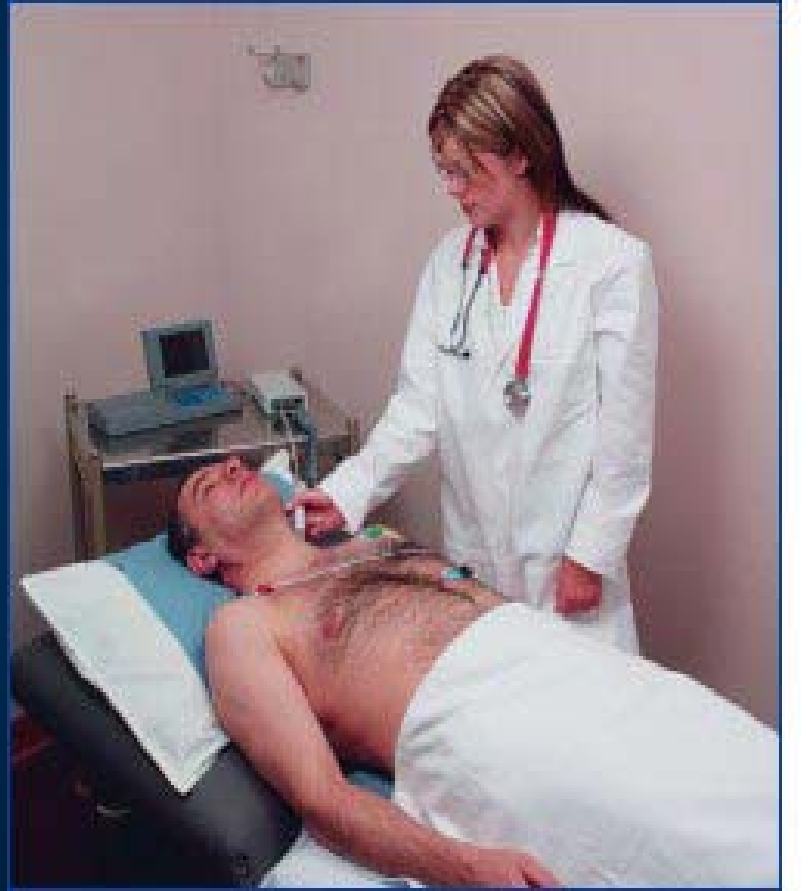


# Applanation tonometry (SphygmoCor)



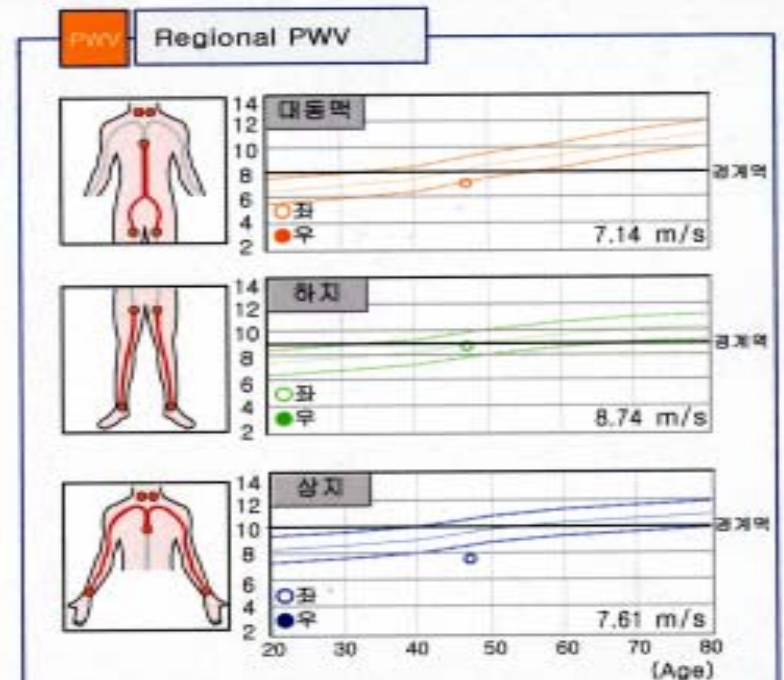
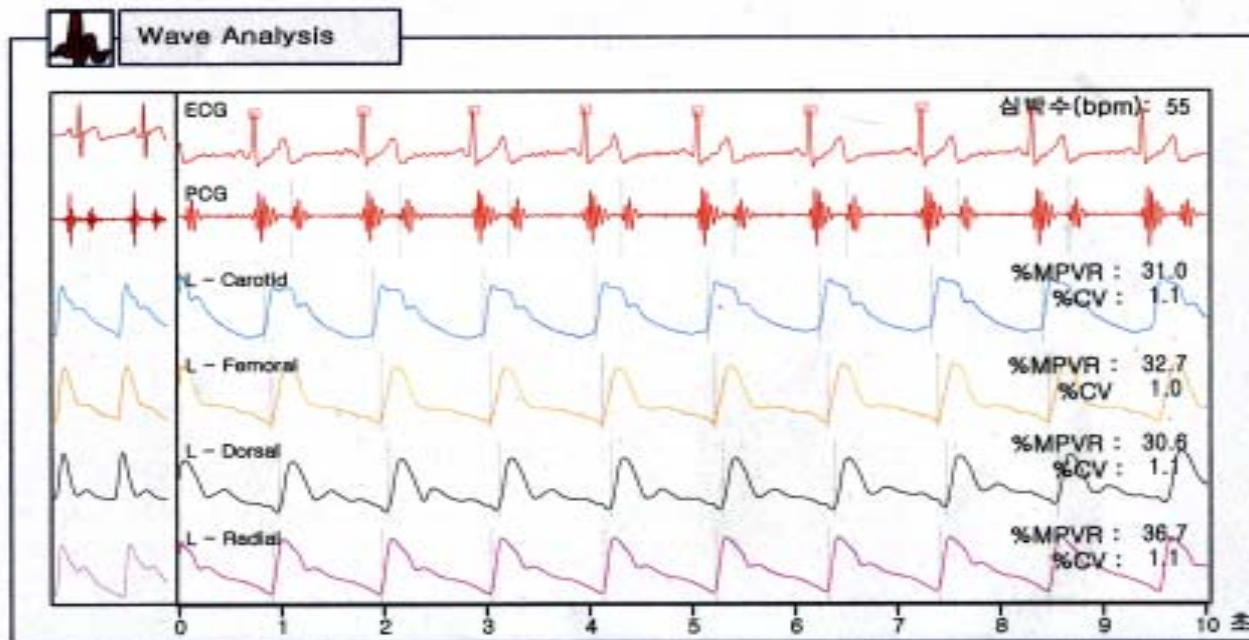


# Complior, Micro-system, Omron

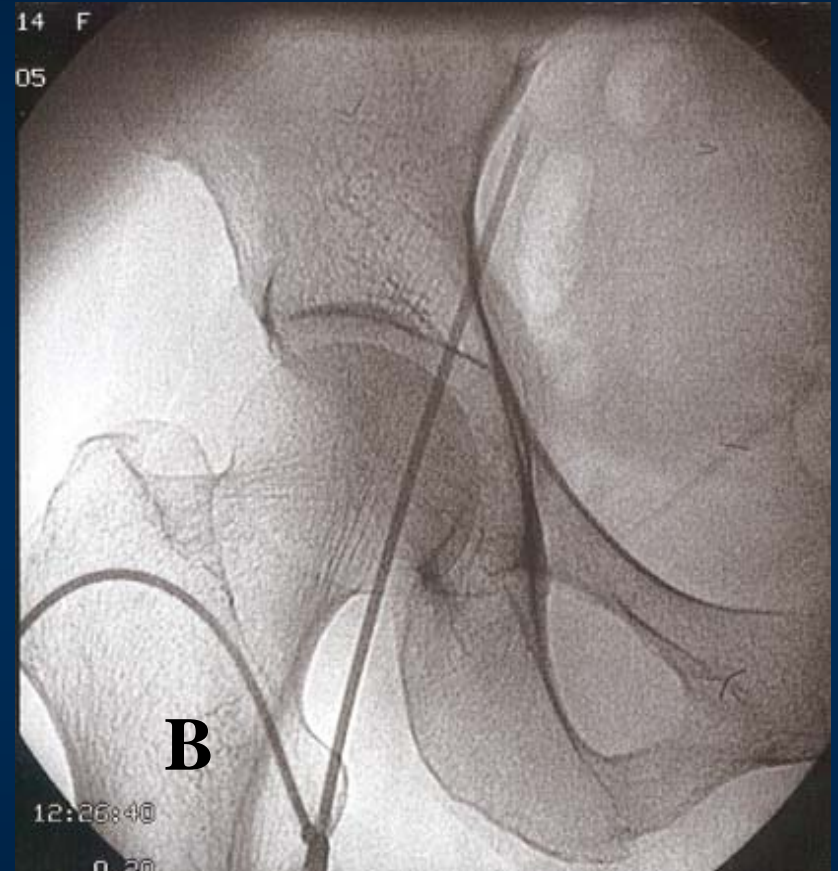
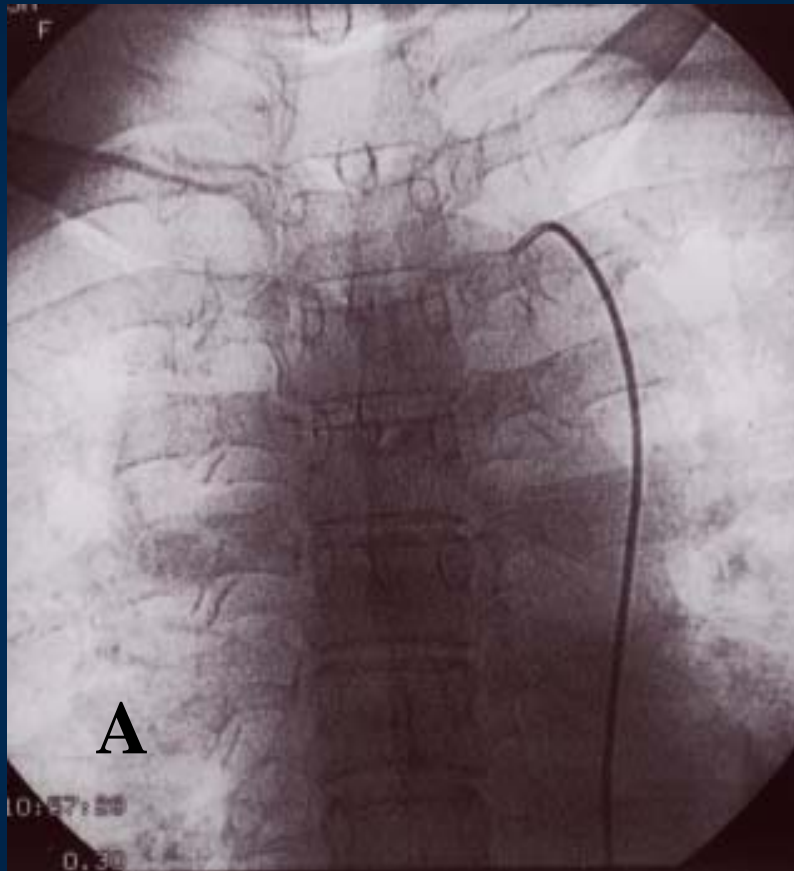




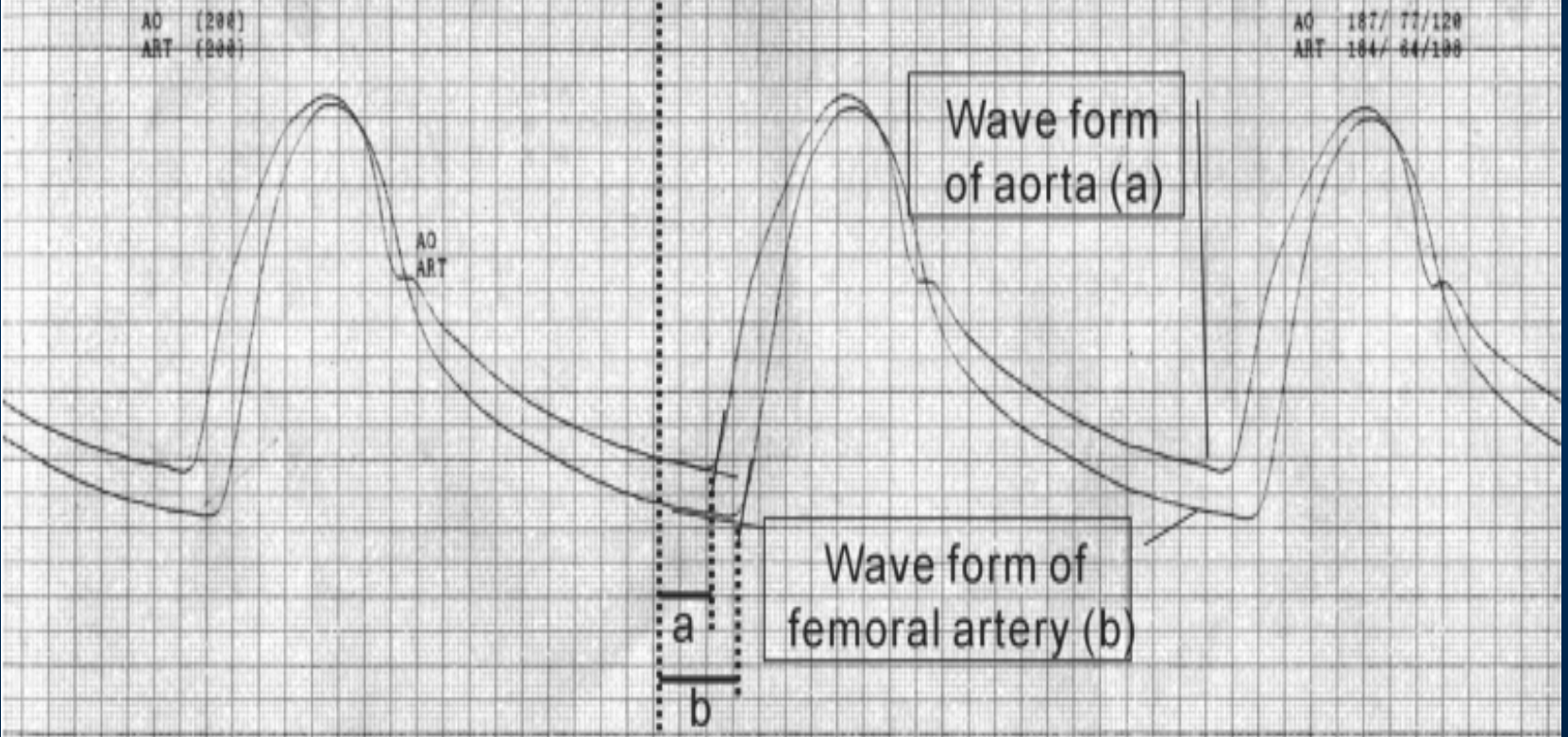
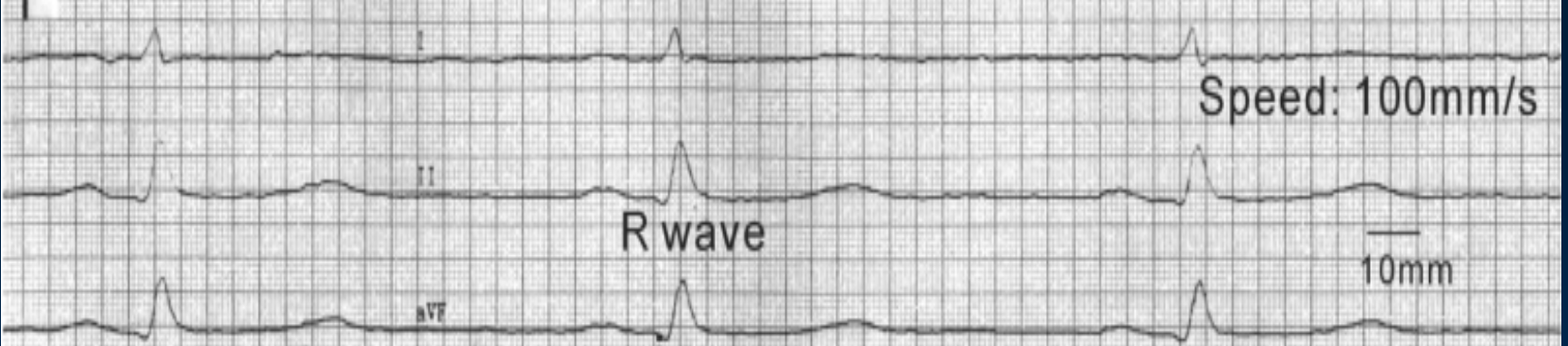
# BaPWV and ABI by Colin



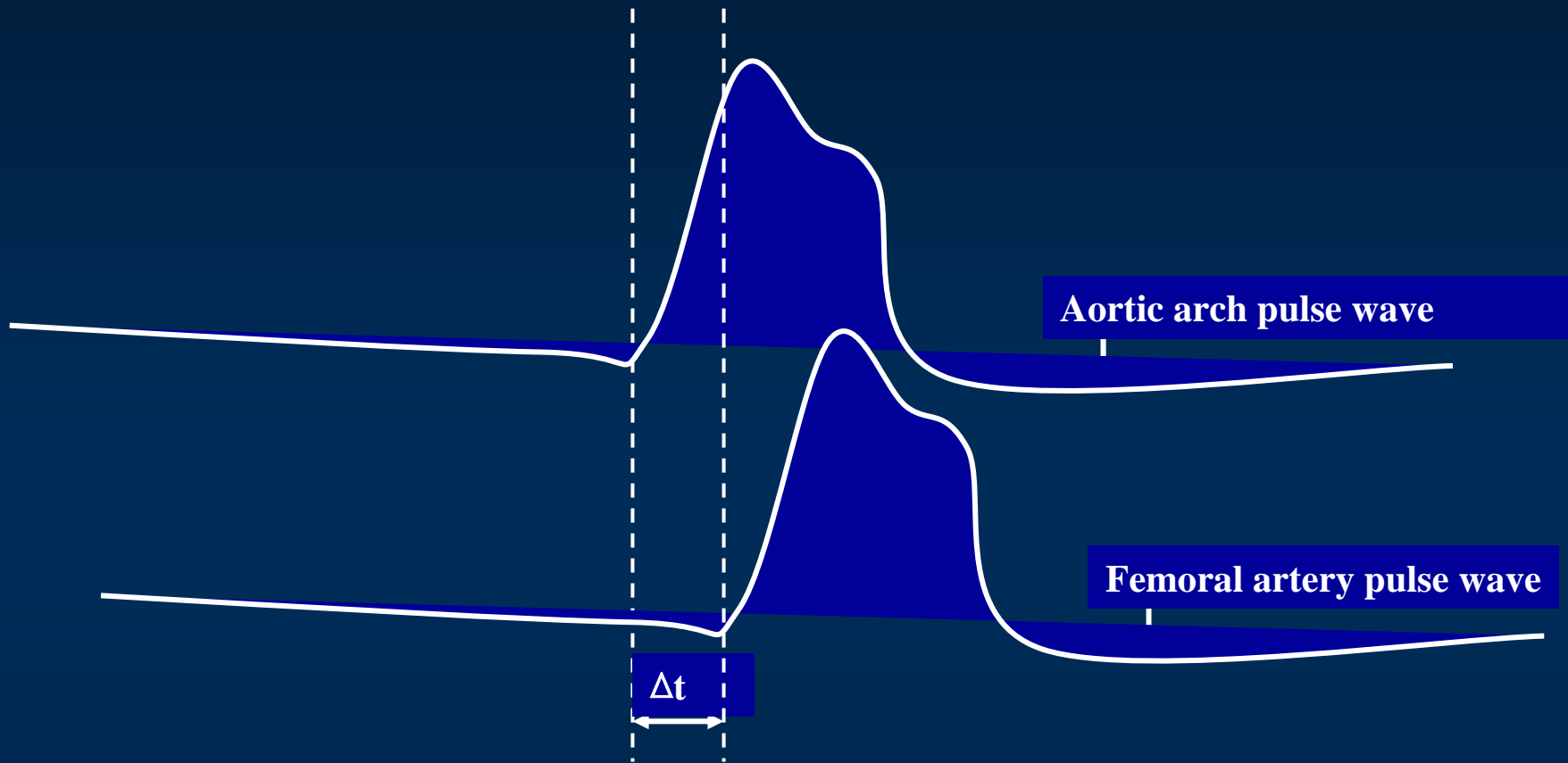
# PWV by Cardiac Catheterization



Aorto-iliac pulse wave velocity (PWV) was measured between aortic arch (A), just distal to the left subclavian artery ostium and right common iliac artery (B).

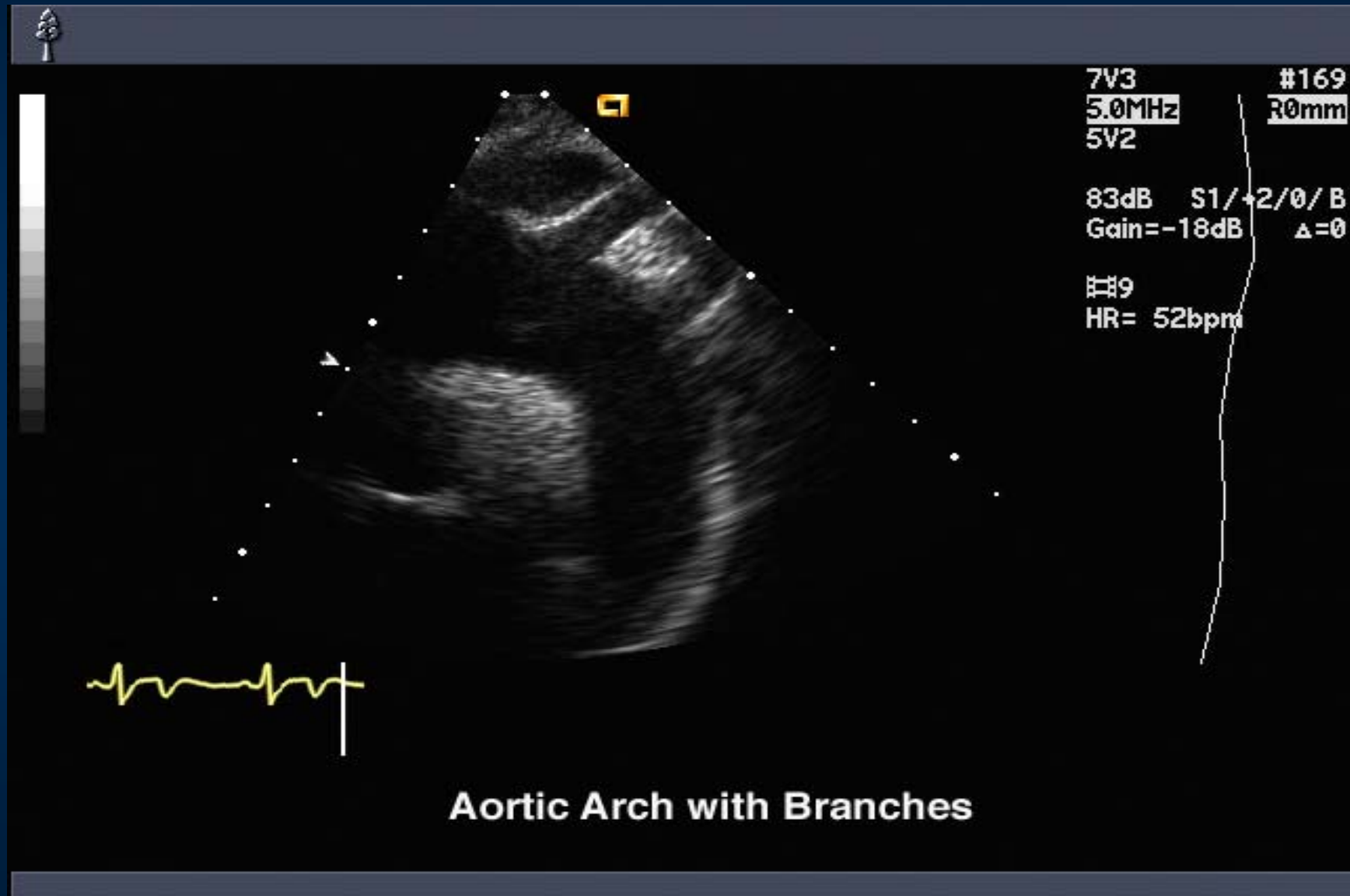


$$\text{Pulse wave velocity} = \frac{\text{Length of subclavian to femoral artery (m)}}{\text{Time duration } [(b-a) / 100] \text{ (second)}}$$



$\Delta t$  is the time interval between two pulse waves.

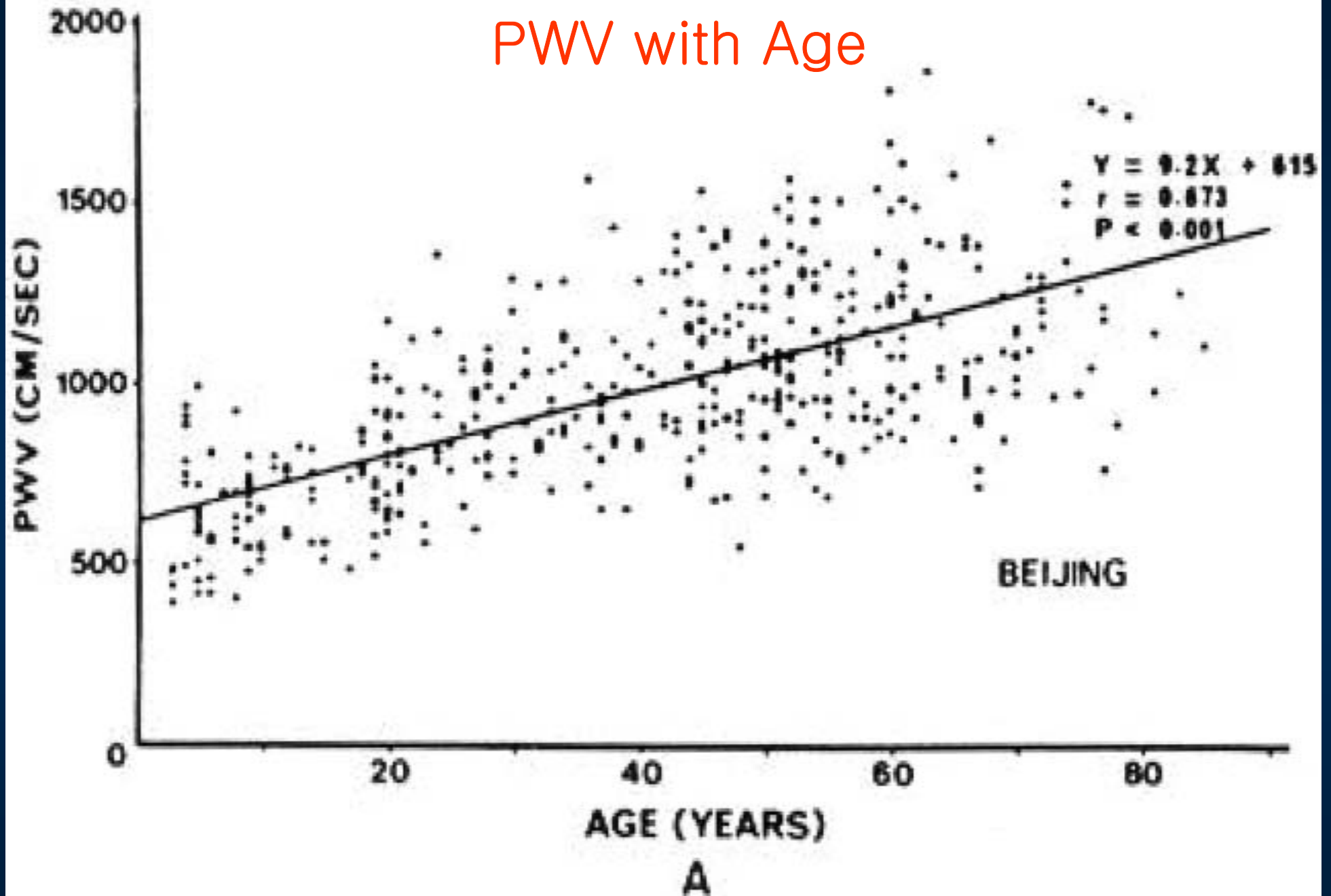
# PWV by Echocardiography



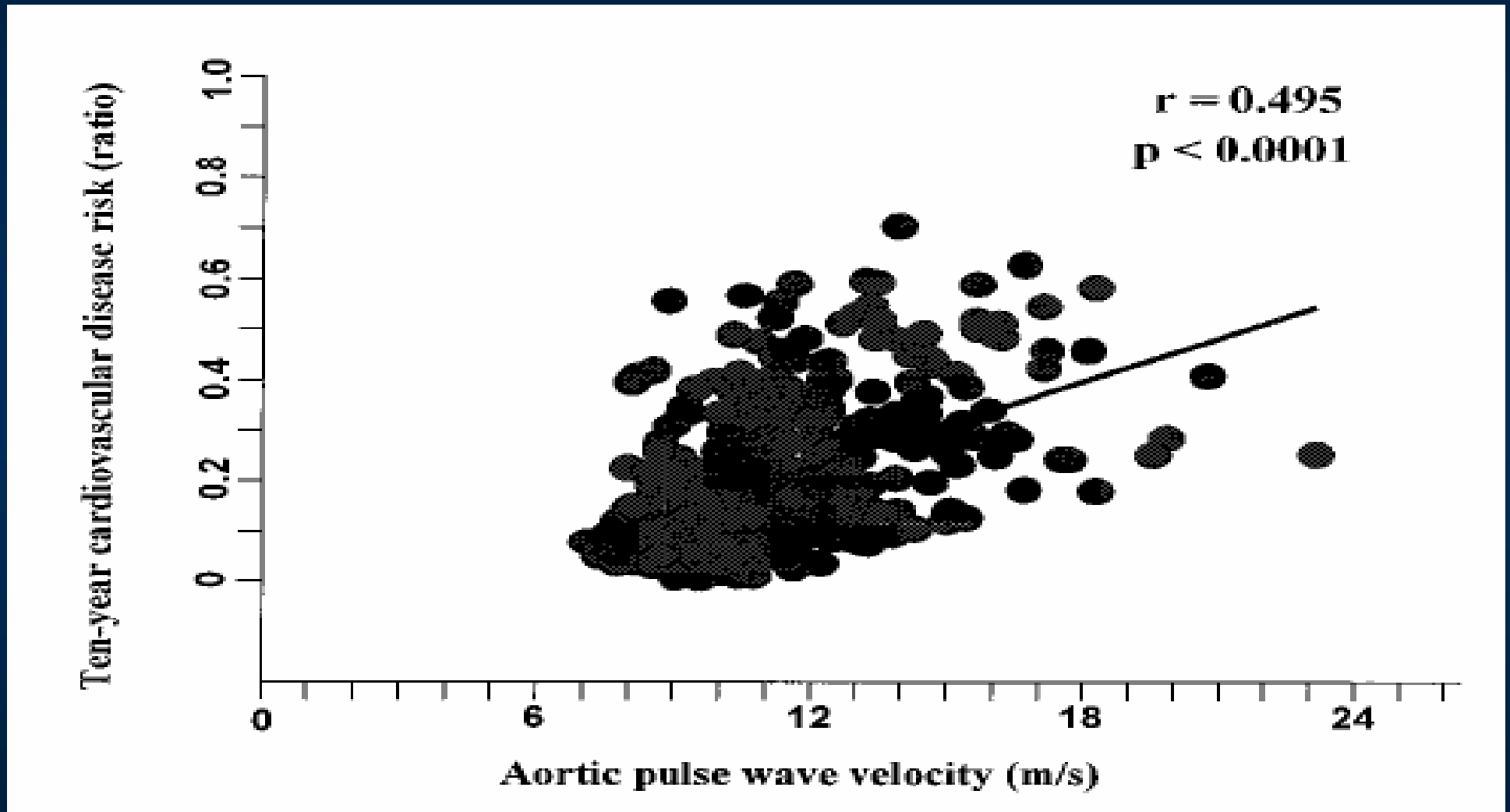


$$\text{PWV} = \frac{\text{length (Aortic arch-femoral a)}}{\text{T2} - \text{T1}} \text{ meter/ sec}$$

# PWV with Age



# Relation between 10-year CVD risk and Aortic PWV





# Odds Ratio of Being in High-Risk Group

Parameter	Odds Ratio of Being in High-Risk Group (95% CI)					Cardiovascular Mortality
	MI	CHD	CHD Mortality	Stroke	CVD	
Pulse wave velocity, >13.5 m/s	3.5 (2.3–5.5)	4.6 (2.9–7.2)	4.9 (3.1–7.8)	6.1 (3.8–9.6)	5.3 (3.4–8.4)	7.1 (4.5–11.3)
Gender, male	6.6 (4.4–9.9)	7.1 (4.5–11.2)	7.3 (4.3–12.7)	2.0 (1.3–3.1)	3.8 (2.6–5.7)	2.9 (1.9–4.3)
Age, >60 y	3.0 (2.0–4.4)	3.9 (2.6–5.9)	7.3 (4.5–11.9)	11.1 (6.7–18.2)	6.1 (4.0–9.2)	12.9 (8.1–20.5)
Plasma glucose, >7.0 mmol/L	8.1	5.9	5.5	7.1	8.4	4.7

	MI	CHD	CHD mortality	Stroke	CVD	CV mortality
PWV>13.5m/s	3.5	4.6	4.9	6.1	5.3	7.1

	(4.8–16.8)	(2.3–6.0)	(1.6–4.3)	(1.2–3.2)	(2.3–6.3)	(1.4–3.6)
Tobacco life-long dose, >20 pack-years	4.4 (2.6–7.2)	2.0 (1.3–3.2)	1.9 (1.2–3.2)	1.7 (1.1–2.8)	2.6 (1.6–4.1)	1.7 (1.1–2.8)
Total/HDL cholesterol, ratio >5	3.7 (2.5–5.5)	3.9 (2.6–5.9)	3.6 (2.3–5.7)	1.5 (1.0–2.3)	3.6 (2.3–5.4)	2.8 (1.9–4.3)
Left ventricular hypertrophy, yes-no	2.2 (1.2–4.1)	11.2 (5.3–23.8)	3.0 (1.6–5.6)	2.2 (1.2–4.0)	4.9 (2.5–9.5)	4.5 (2.4–8.4)
Plasma creatinine, >100 $\mu$ mol/L	1.8 (1.1–2.7)	2.5 (1.6–3.9)	2.7 (1.7–4.3)	1.7 (1.1–2.8)	1.8 (1.2–2.8)	1.8 (1.2–2.8)

Ten-year absolute MI risk >5%, 10-year absolute CHD risk >15%, 10-year absolute CHD mortality risk >5%, 10-year absolute stroke risk >5%, 10-year absolute CVD risk >20%, and 10-year cardiovascular mortality risk >5% were considered high.

## Odds Ratios of Developing Hypertension during 4 yrs F/U in Relation to Aortic Stiffness (Multivariate Logistic regression models)

Variable	Men		Women	
	Odds Ratio	P value	Odds Ratio	P value
Systolic BP	8.7619	< 0.001	8.6920	< 0.001
Diastolic BP	6.6947	< 0.001	6.7582	< 0.001
Age	8.8211	< 0.001	9.0145	< 0.001
<u>Aortic stiffness index</u>	1.2165	< 0.05	1.1981	< 0.05

## Aortic Pulse Wave Velocity as an Independent Marker of Coronary Artery Disease

H. EUY LIM, C. GYU PARK, S. HEE SHIN, J. CHEON AHN, H. SEOG SEO AND D. JOO OH

*From the Division of Cardiology, Cardiovascular Center, Korea University Hospital, Guro-Dong, Guro-Gu, Seoul KR-152-050, Korea*

**Euy Lim H, Gyu Park C, Hee Shin S, Cheon Ahn J, Seog Seo H, Joo Oh D.** *Aortic pulse wave velocity as an independent marker of coronary artery disease.* Blood Pressure 2004; 13: 000–000.

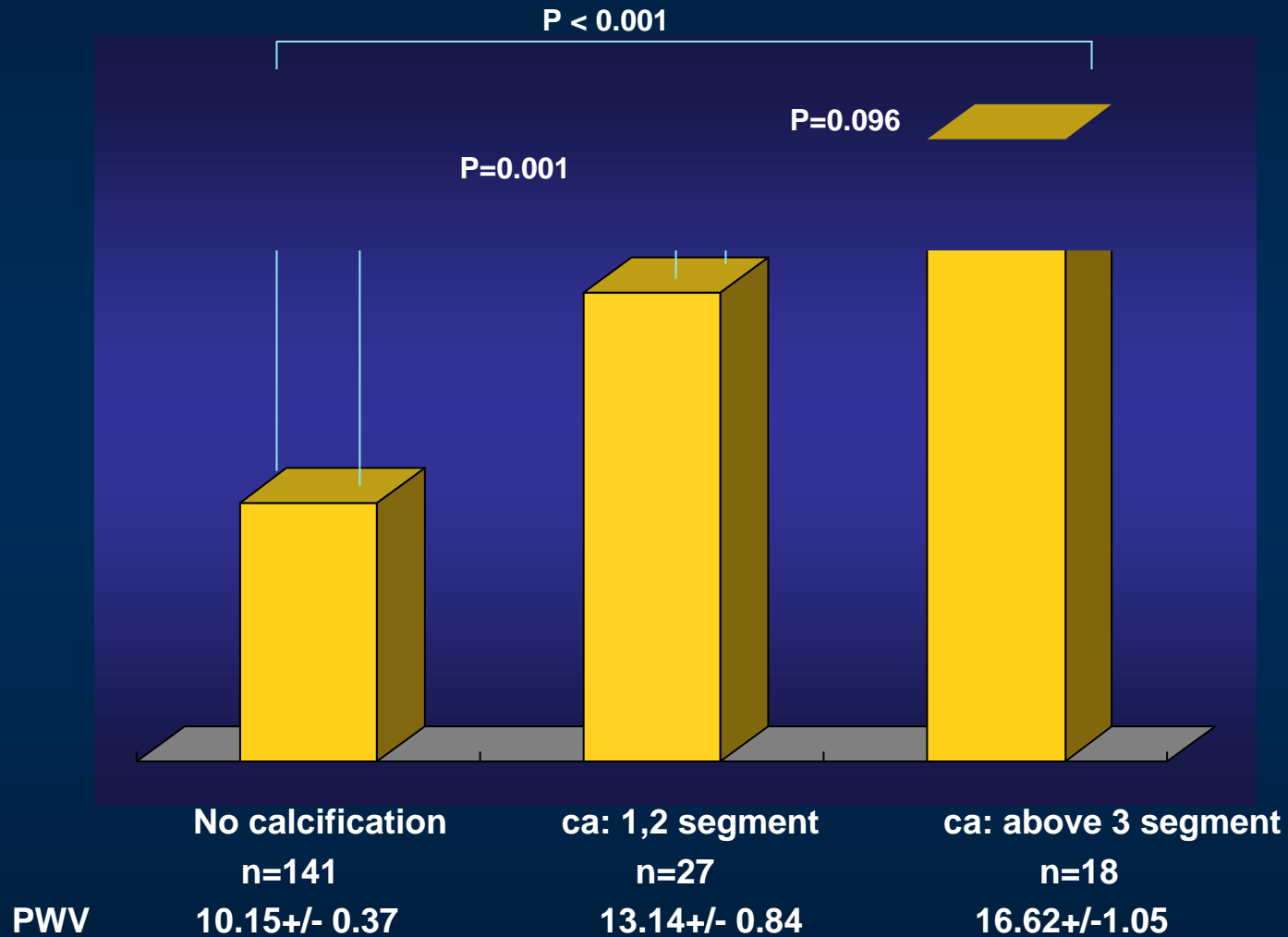
**Background:** Arterial stiffness has been known as a major contributory factor to cardiovascular (CV) morbidity and mortality in patients with hypertension. Pulse wave velocity (PWV), a surrogate measurement of large artery damage, has not been ascertained as an independent risk factor of coronary artery disease (CAD). The aim of this study was to assess whether PWV is associated with CV risk. **Methods and results:** We prospectively enrolled 326 consecutive patients undergoing coronary angiography for the assessment of suspected CAD. Arterial stiffness was assessed through aorto-femoral PWV using fluid-filled system. PWV was higher in patients with CAD than those without CAD ( $12.5 \pm 5.1$  vs  $10.2 \pm 3.1$  m/s,  $p < 0.001$ ). In multivariate logistic regression analysis, after entering for age, diabetes and other CV risk factors, PWV remained the significant independent variable for CAD ( $p = 0.050$ ). When the severity of CAD was expressed as one-, two- or three-vessel disease, PWV was a significantly associated with the severity of CAD ( $p < 0.001$ ). **Conclusions:** Our findings suggest that PWV is an independent risk marker for CAD, as well as strongly associated with the severity of CAD. **Key words:** *arterial stiffness, cardiovascular risk factor, coronary artery disease, pulse wave velocity.*

Table I. *Clinical characteristics of subjects with and without coronary artery disease*

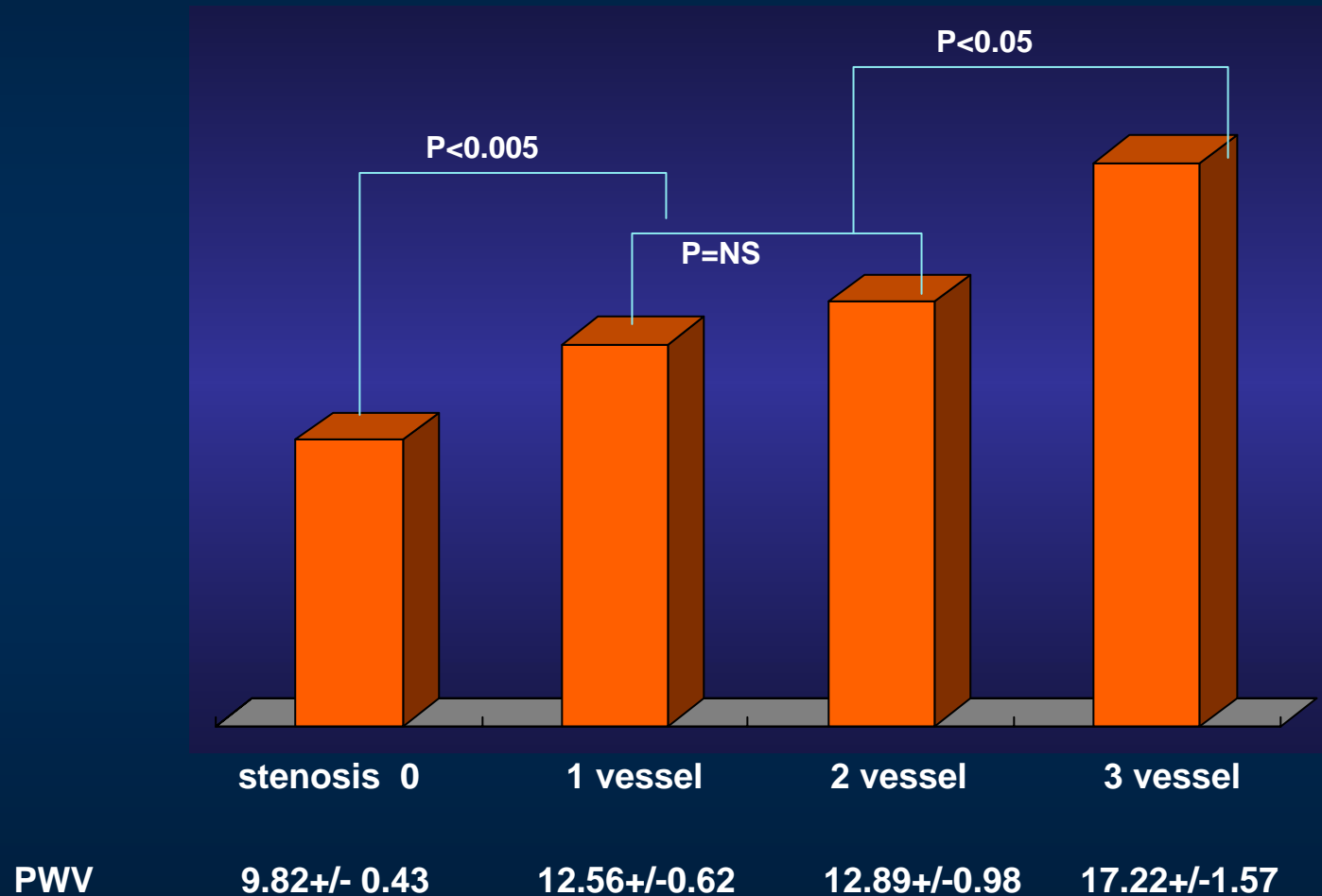
Parameters	CAD ( <i>n</i> = 146)	No CAD ( <i>n</i> = 180)	<i>p</i> -value
Age, years	61.9 ± 10.7	55.8 ± 10.3	<0.001
Gender, men/women	71/75	72/108	0.118
Diabetes, %	27.4% (40)	12.2% (22)	0.001
Cigarette smoking, %	41.1% (60)	28.3% (51)	0.016
Previous antihypertensive treatment, %	67.8% (99)	50.0% (90)	0.001
Body mass index, kg/m <sup>2</sup>	24.5 ± 3.3	24.7 ± 4.5	0.607
Weight-to-height ratio	0.02 ± 0.07	0.02 ± 0.08	0.004
<b>PWV, m/s</b>	<b>12.5</b>	<b>10.2</b>	<b>&lt;0.001</b>
HDL-C	43.5 ± 15.0	46.9 ± 15.5	0.022
TG	144.0 ± 69.3	129.0 ± 73.3	0.061
LDL-C	115.4 ± 36.9	105.9 ± 32.5	0.013
Office blood pressure, mmHg			
Systolic	145.2 ± 24.2	140.9 ± 25.4	0.114
Diastolic	87.1 ± 12.9	87.0 ± 13.5	0.984
Pulse	58.2 ± 18.7	53.8 ± 16.5	0.026
Pulse wave velocity, m/s	12.5 ± 5.1	10.2 ± 3.1	<0.001

CAD, coronary artery disease; HDL-C, high-density lipoprotein cholesterol; TG, Triglyceride; LDL-C, low-density lipoprotein cholesterol. Continuous variables are expressed as mean ± SD.

# Correlation between PWV and coronary artery calcification

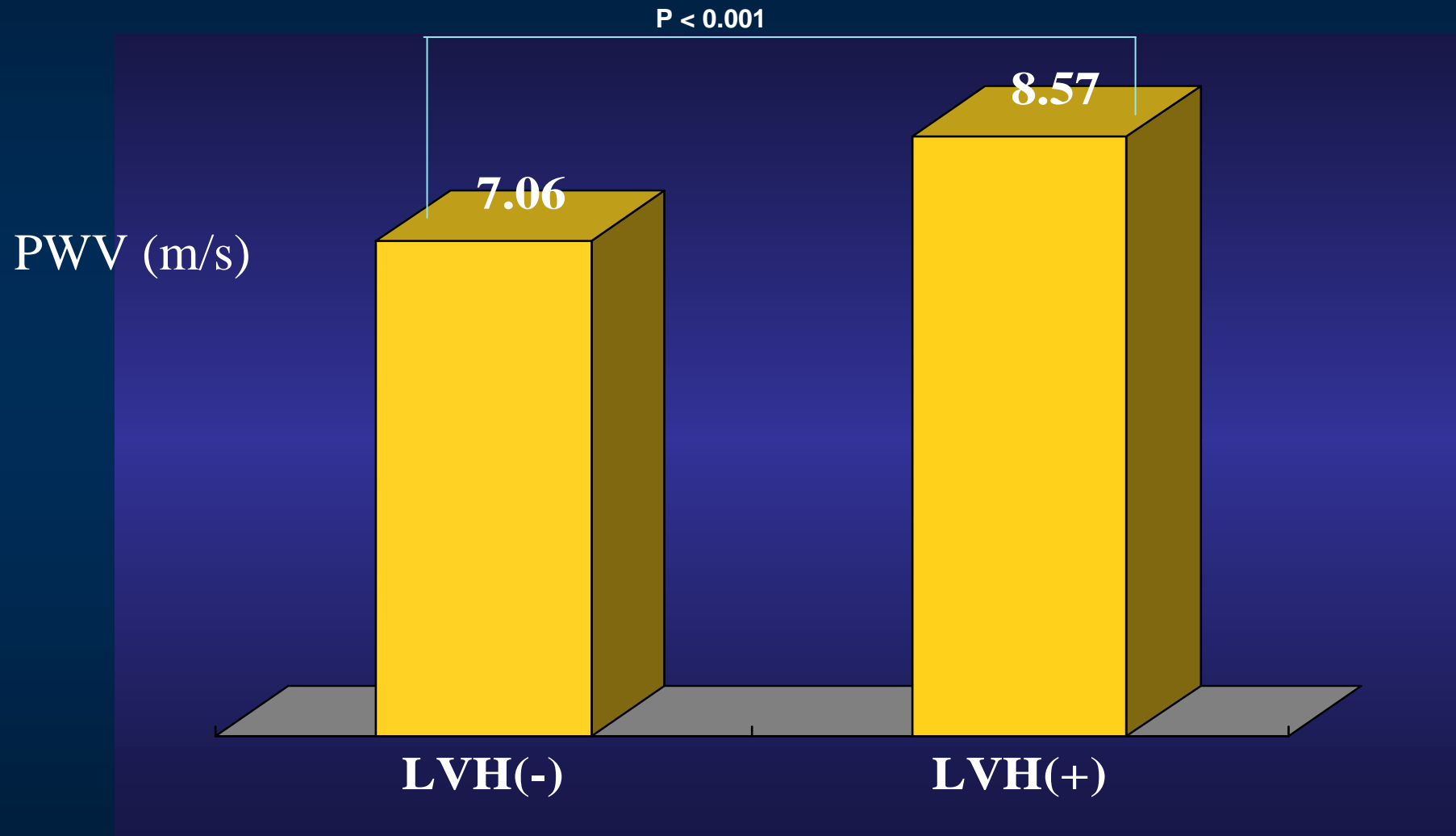


# Correlation between PWV and coronary artery stenosis



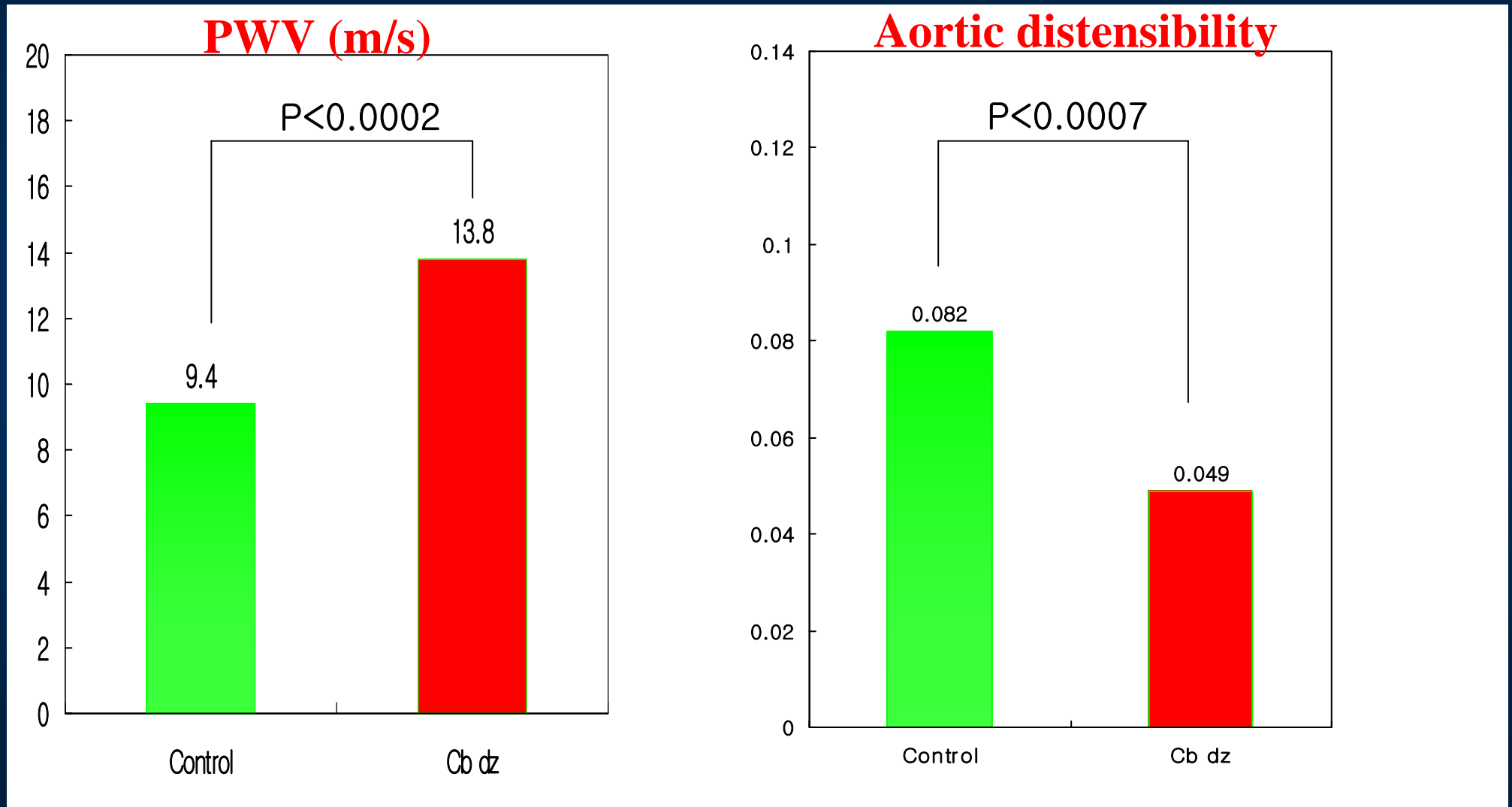
# Aortic Stiffness and LVH

# Correlation between PWV and LVMI



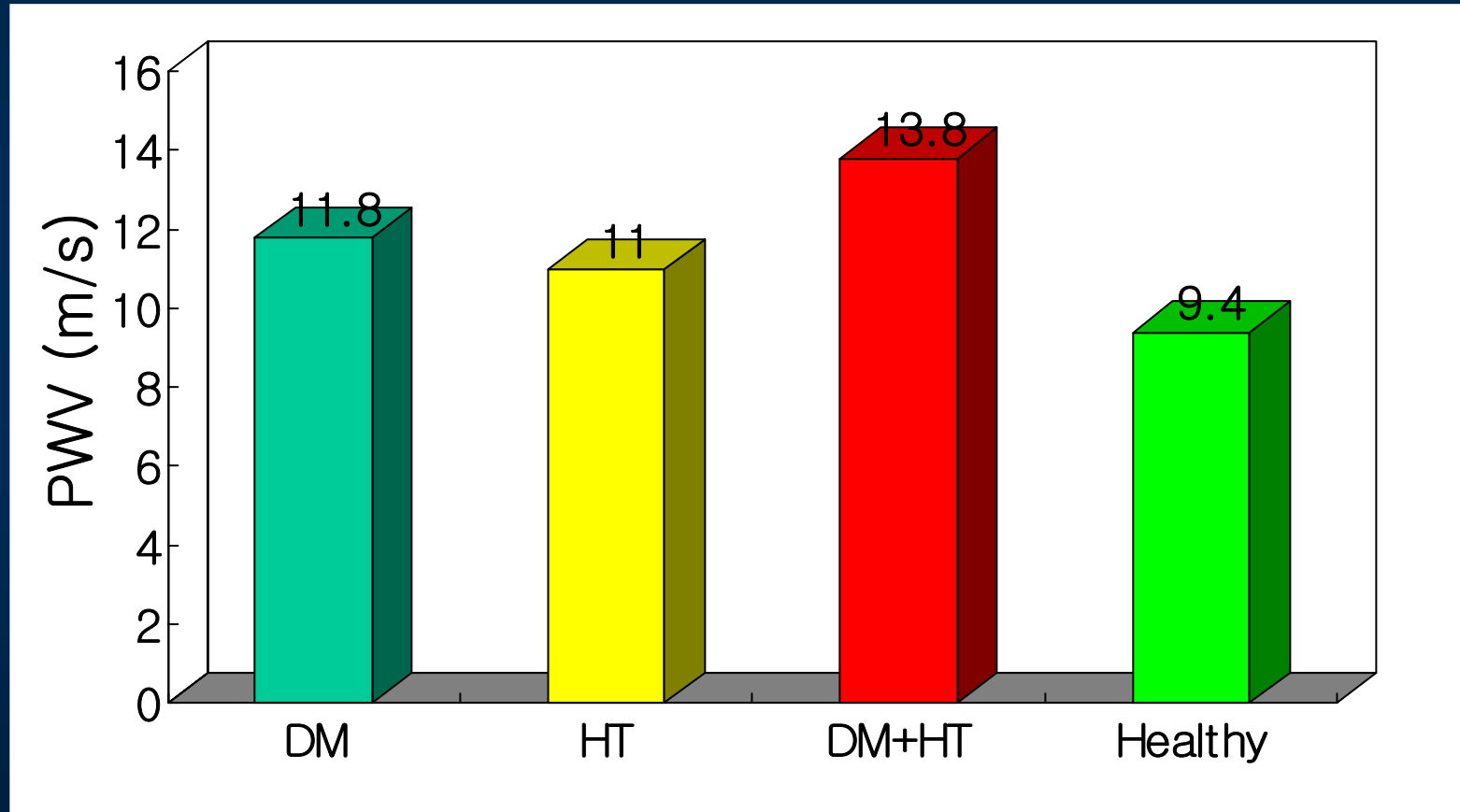


# Aortic Distensibility and Cerebrovascular Disease



# Aortic Stiffness and Diabetes

# Hypertension and type II DM on Aortic Stiffness



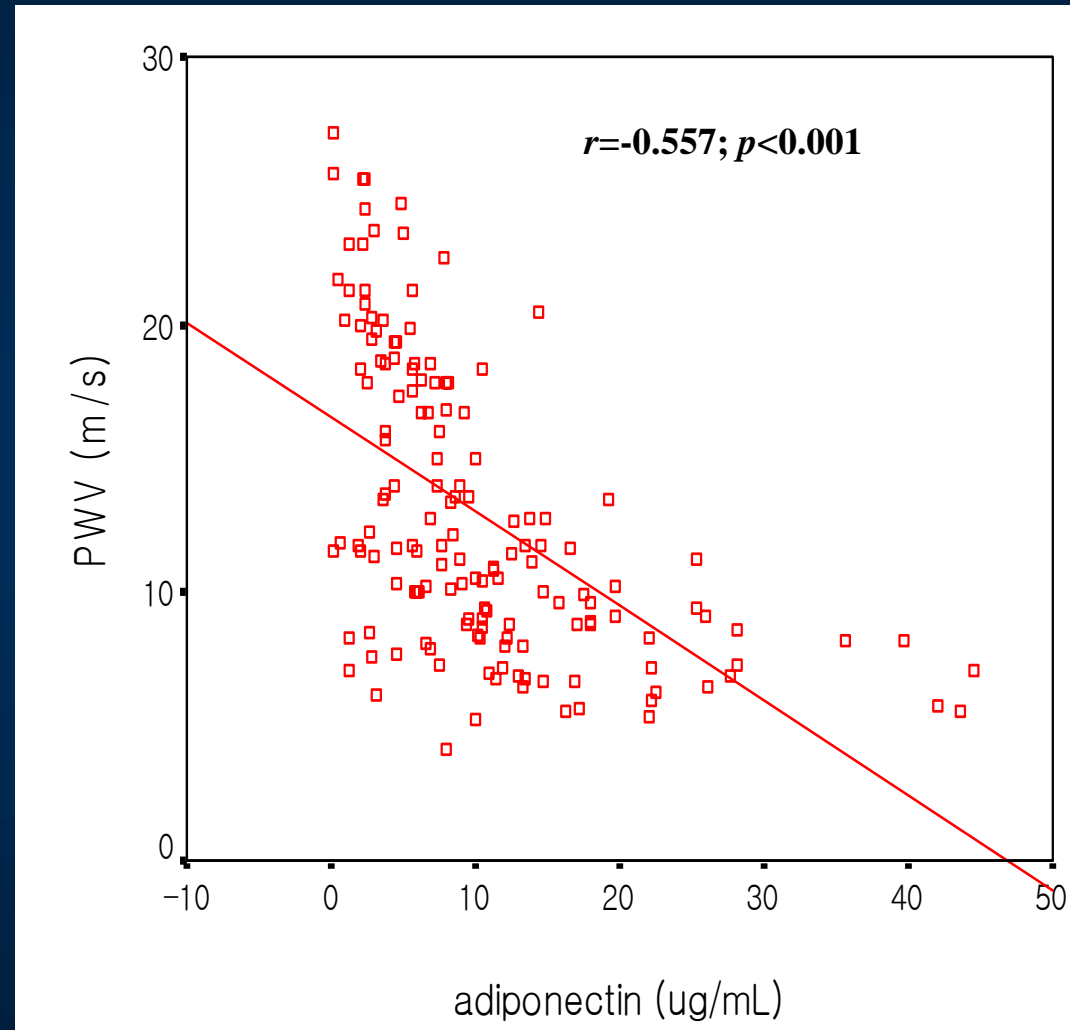
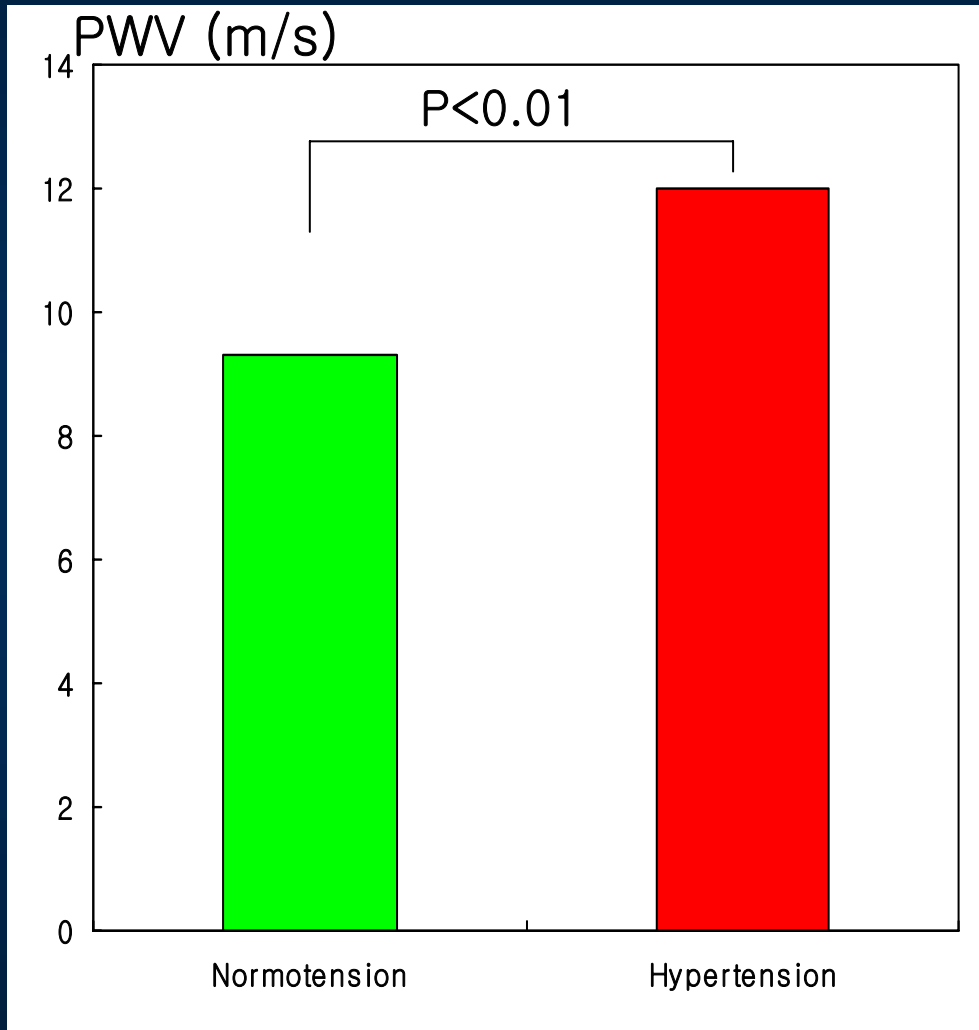
## Associations among Plasma Adiponectin, Hypertension, Left Ventricular Diastolic Function and Left Ventricular Mass Index

SOON JUN HONG, CHANG GYU PARK, HONG SEOG SEO, DONG JOO OH AND YOUNG MOO RO

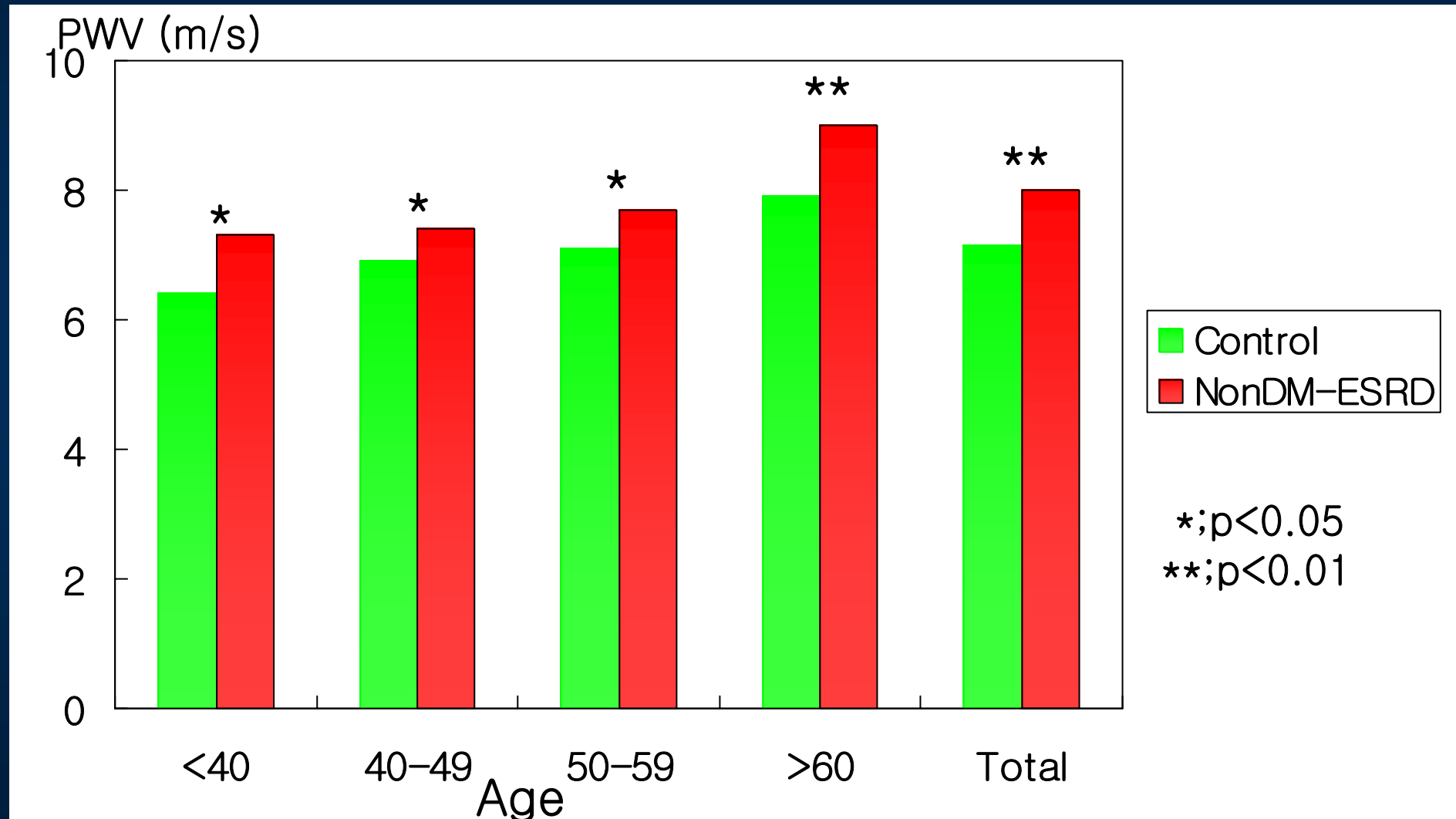
*From the Division of Cardiology, Cardiovascular Center, Korea University Hospital, Seoul, Korea*

**Hong SJ, Park CG, Seo HS, Oh DJ, Ro YM.** *Associations among plasma adiponectin, hypertension, left ventricular diastolic function and left ventricular mass index.* Blood Pressure 2004; 13: 236-242

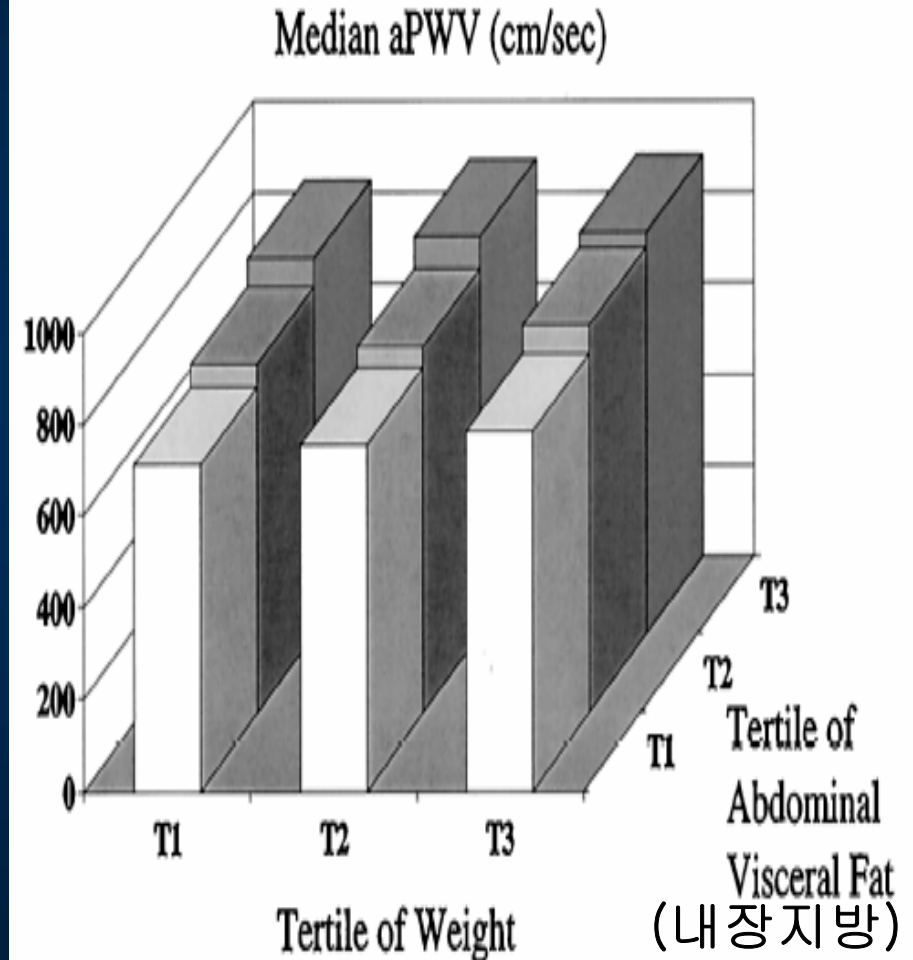
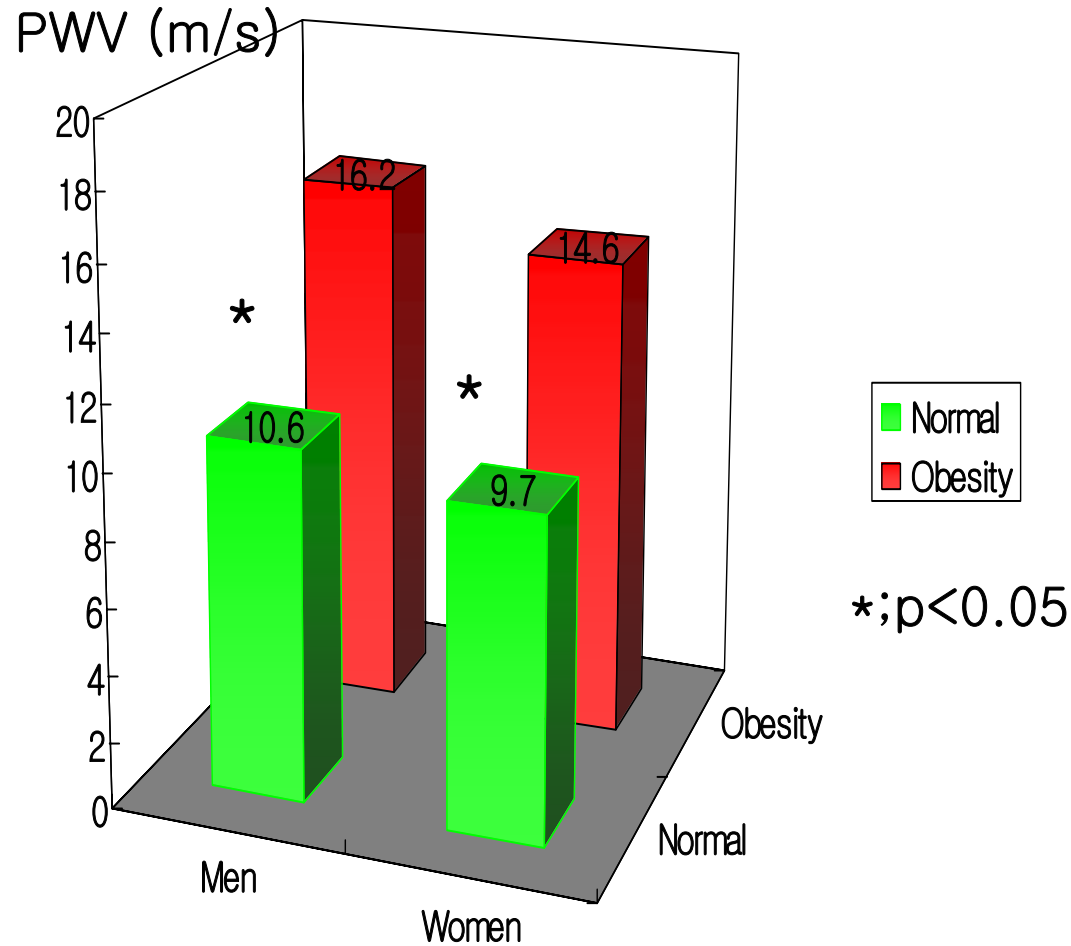
# Correlation between plasma adiponectin and PWV



# PWV in Non-Diabetic Hemodialysis Patients



# PWV and Obesity



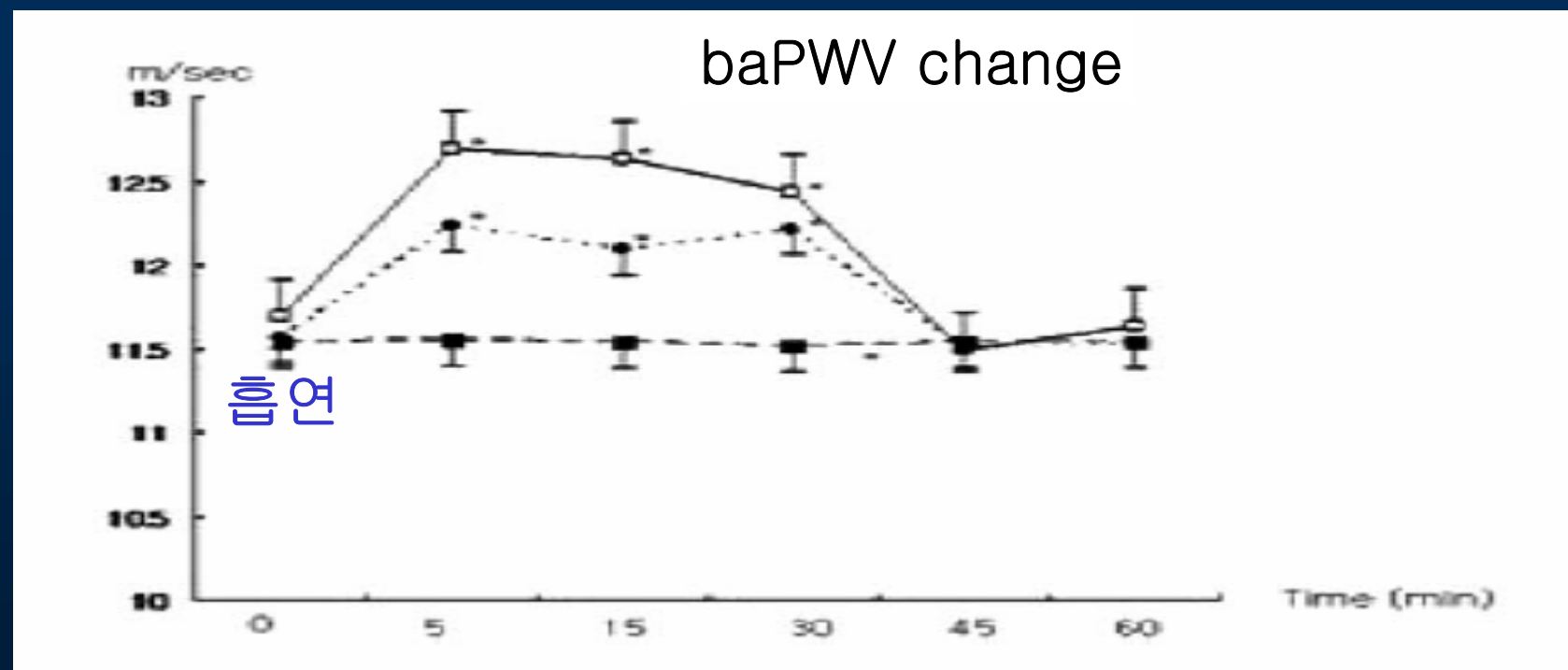
ORIGINAL ARTICLE

Acute and chronic effects of **cigarette smoking** on arterial stiffness

JIN WON KIM<sup>1</sup>, CHANG GYU PARK<sup>1</sup>, SEUNG JOON HONG<sup>1</sup>, SEONG MI PARK<sup>3</sup>,  
SEUNG WOON RHA<sup>1</sup>, HONG SEOG SEO<sup>1</sup>, DONG JOO OH<sup>1</sup> & YOUNG MOO RHO<sup>2</sup>

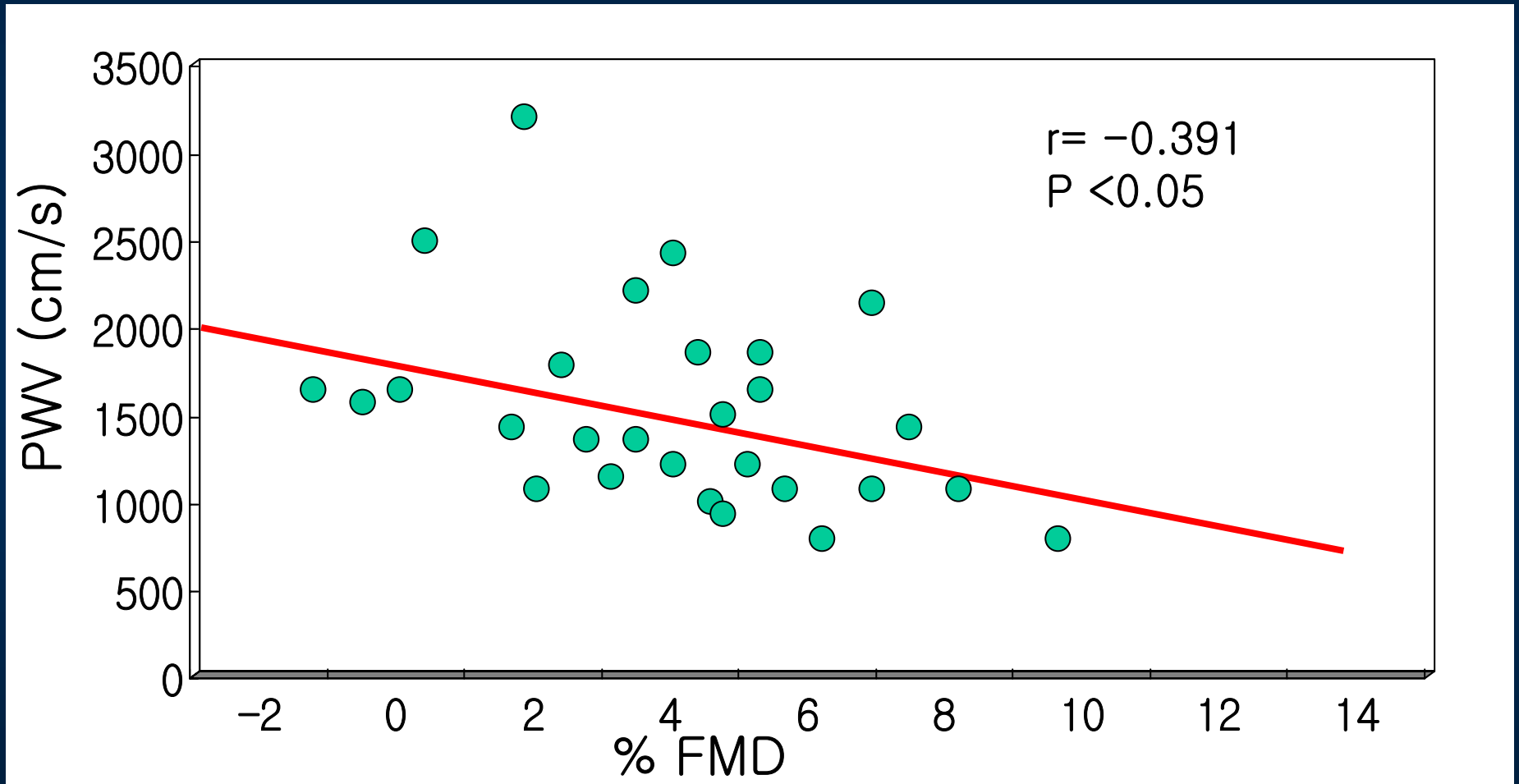
<sup>1</sup>Department of Cardiology, Korea University, Guro Hospital, Cardiovascular Center, Seoul, Korea,

<sup>2</sup>Department of Cardiology, Korea University, Anam Hospital, Cardiovascular Center, Seoul, Korea, and <sup>3</sup>Department of Cardiology, Inha University Hospital, Cardiovascular Center, Seoul, Korea





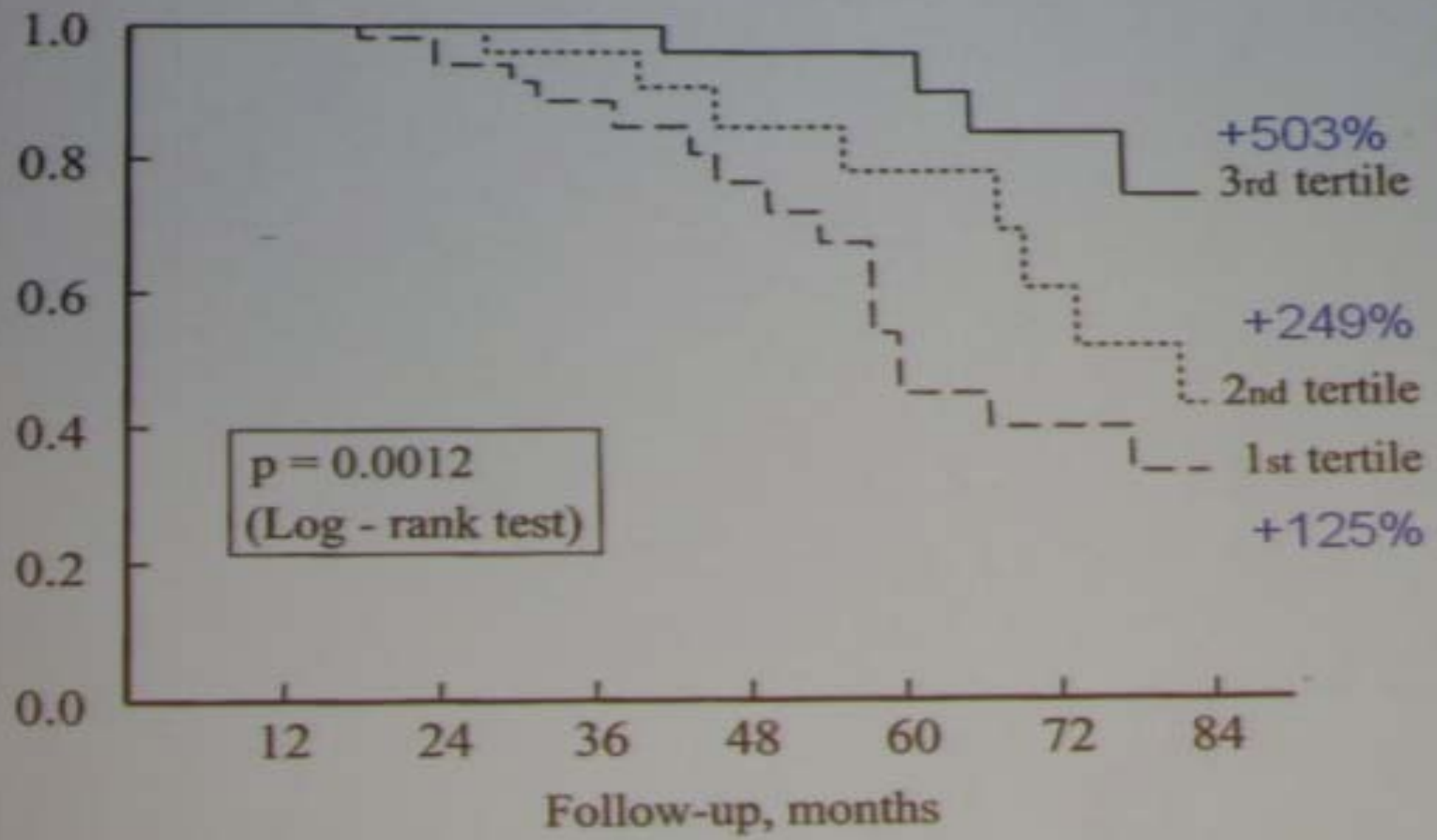
# PWV and Endothelial Function



# Prognostic Value of Forearm Vascular Endothelial Dysfunction (FMD) In hypertensive patients

Perticone F et al., Circulation 2001

Survival ratio



N= 225  
never-treated HT

Mean follow-up : 31.5 months

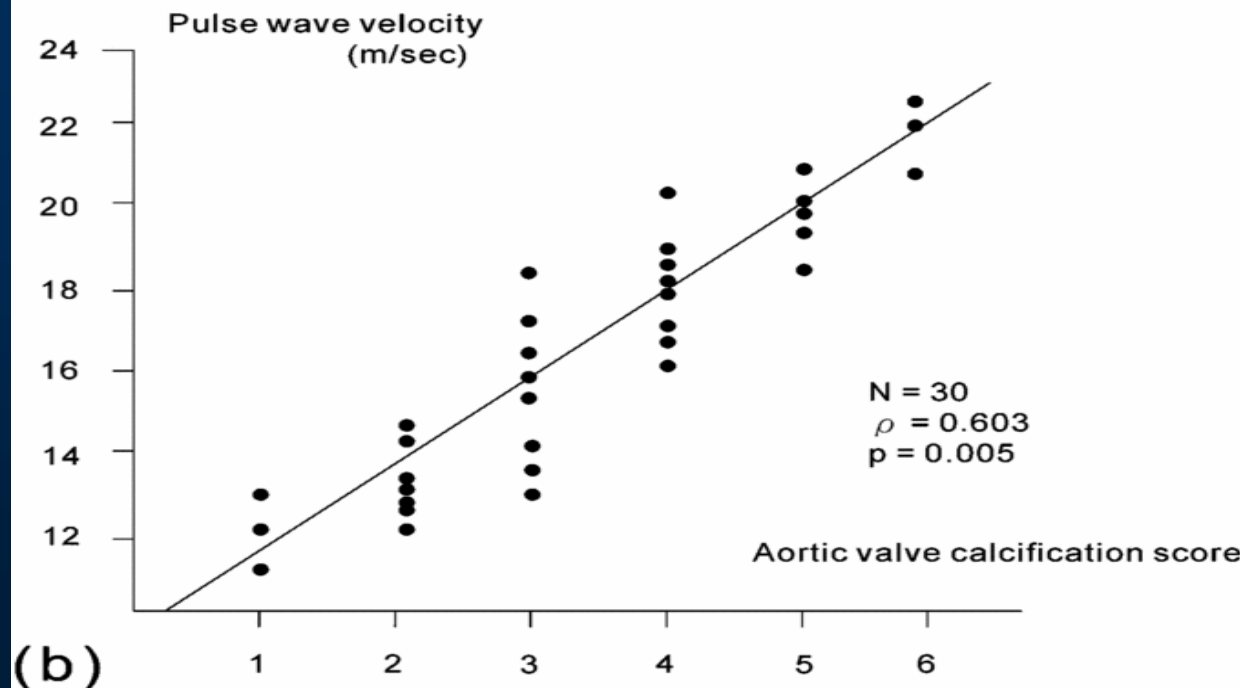
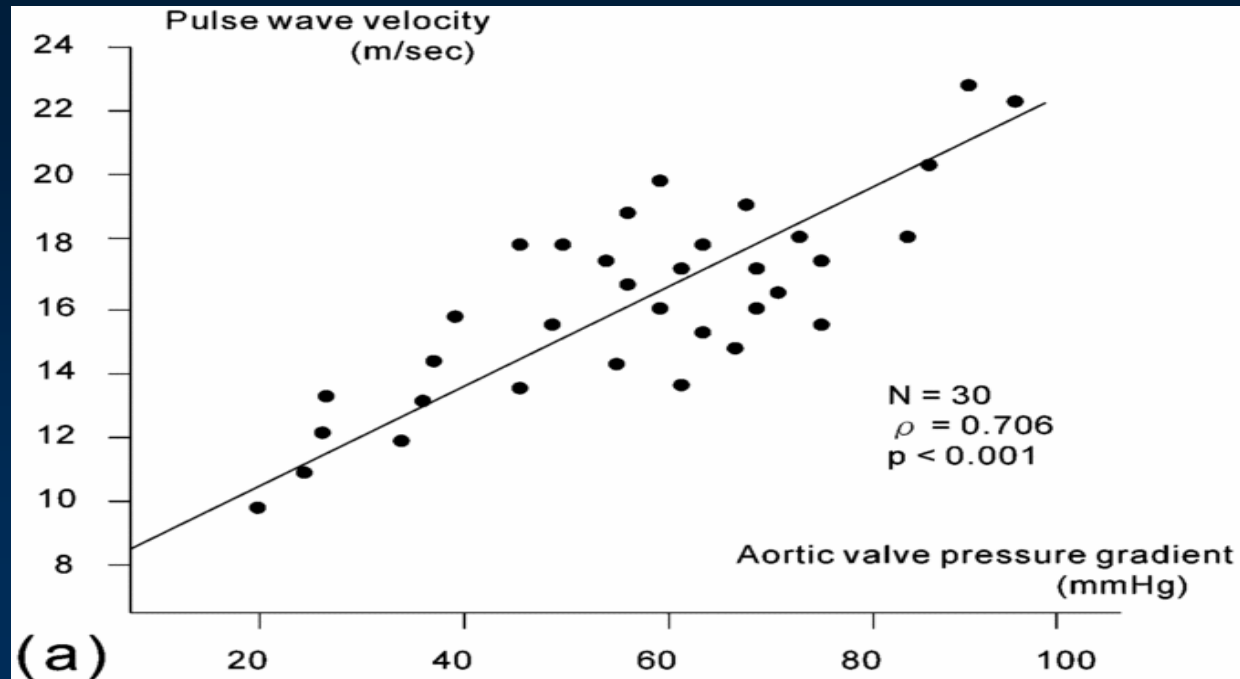
Ach infusion  
Forearm blood flow

Cox model :  
3rd vs 1st tertile  
RR = 2.08 (1.2-3.5)  
P<0.01

Number at risk

Follow-up (months)	225	164	132	73	52	41	27	10
0	225	164	132	73	52	41	27	10

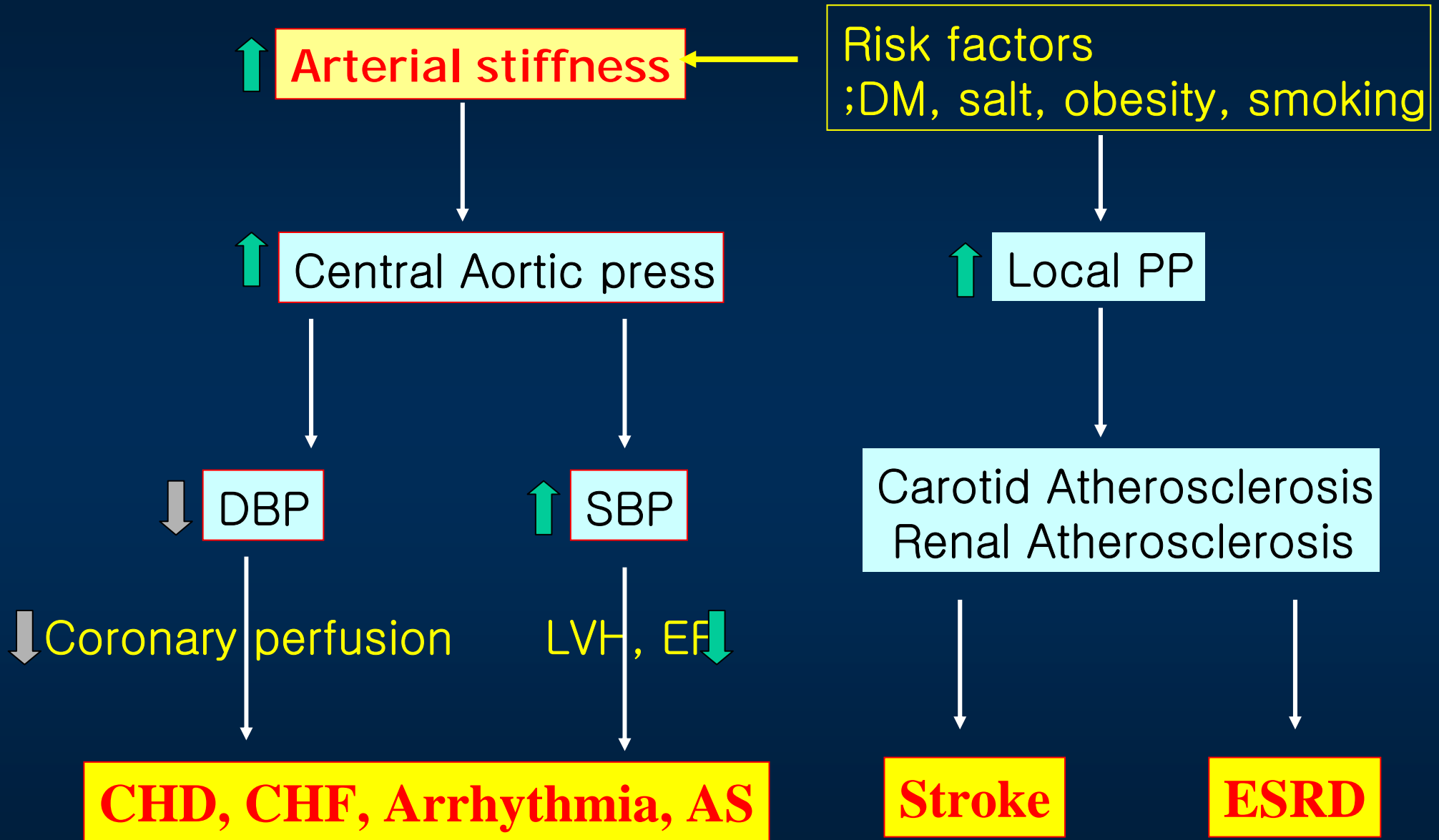
# Correlation between PWV and Aortic Valve Calcification, and Aortic Valve Stenosis



# Cox proportional analysis for significant AS

	Odds ratio	95% CI	P value
<b>Univariate analysis</b>			
Diabetes mellitus	3.4	1.5–6.1	0.01
<b>Increased PWV</b>	<b>3.1</b>	1.4–9.0	0.01
Systemic hypertension	2.6	1.3–6.6	0.01
Severe CAD ( two-vessel disease)	2.2	1.5–4.7	0.03
Increased AV calcification	2.2	1.3–2.9	0.01
Lower LV ejection fraction	2.0	0.8–3.8	0.1
Increased age	1.7	0.8–3.8	0.2
<b>Multivariate analysis</b>			
<b>Increased PWV</b>	<b>3.0</b>	1.3–9.9	0.01
Diabetes mellitus	2.3	1.6–10.5	0.01
Systemic hypertension	2.2	1.1–5.2	0.03

CI, confidence interval.

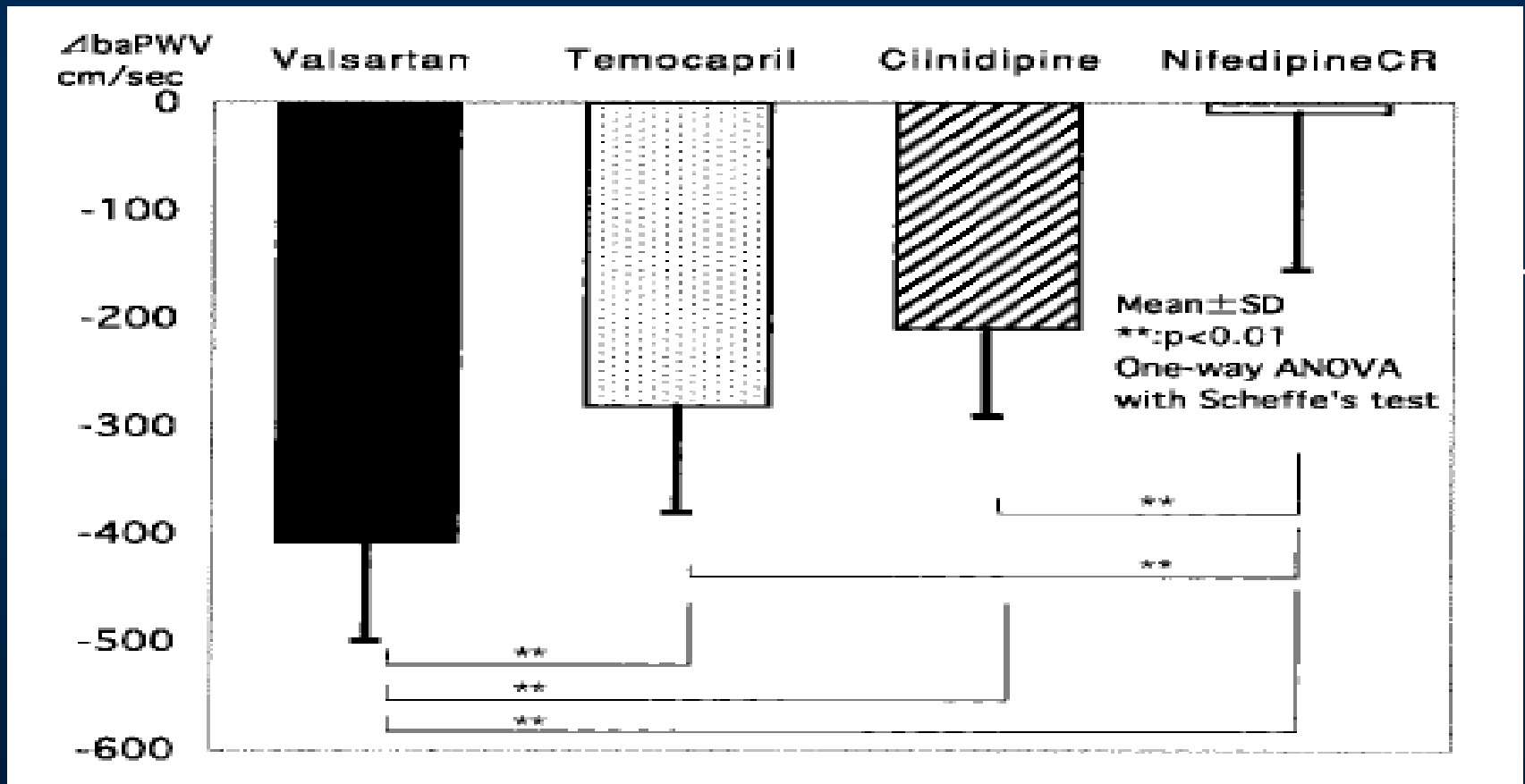


# Optimal treatment of Aortic Stiffness

# Various Antihypertensives ; Changes in Vascular Stiffness

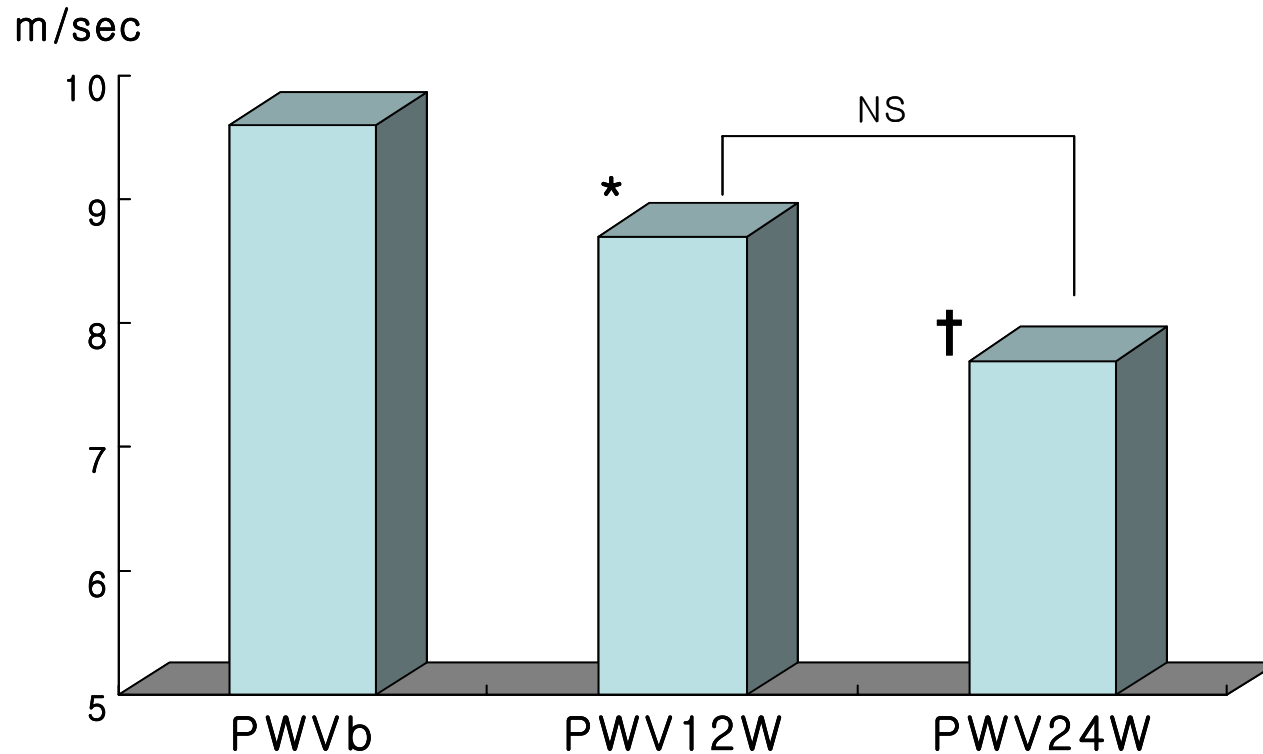
ARB	ACEI	Cilnidipine	L형 CCB
409 ± 90cm/s	281 ± 99cm/s	209 ± 82cm/s	9 ± 146cm/s

$\Delta$ baPWV  
(cm/sec)



# ARB on PWV

Figure 2. Change of PWV after Irbesartan treatment

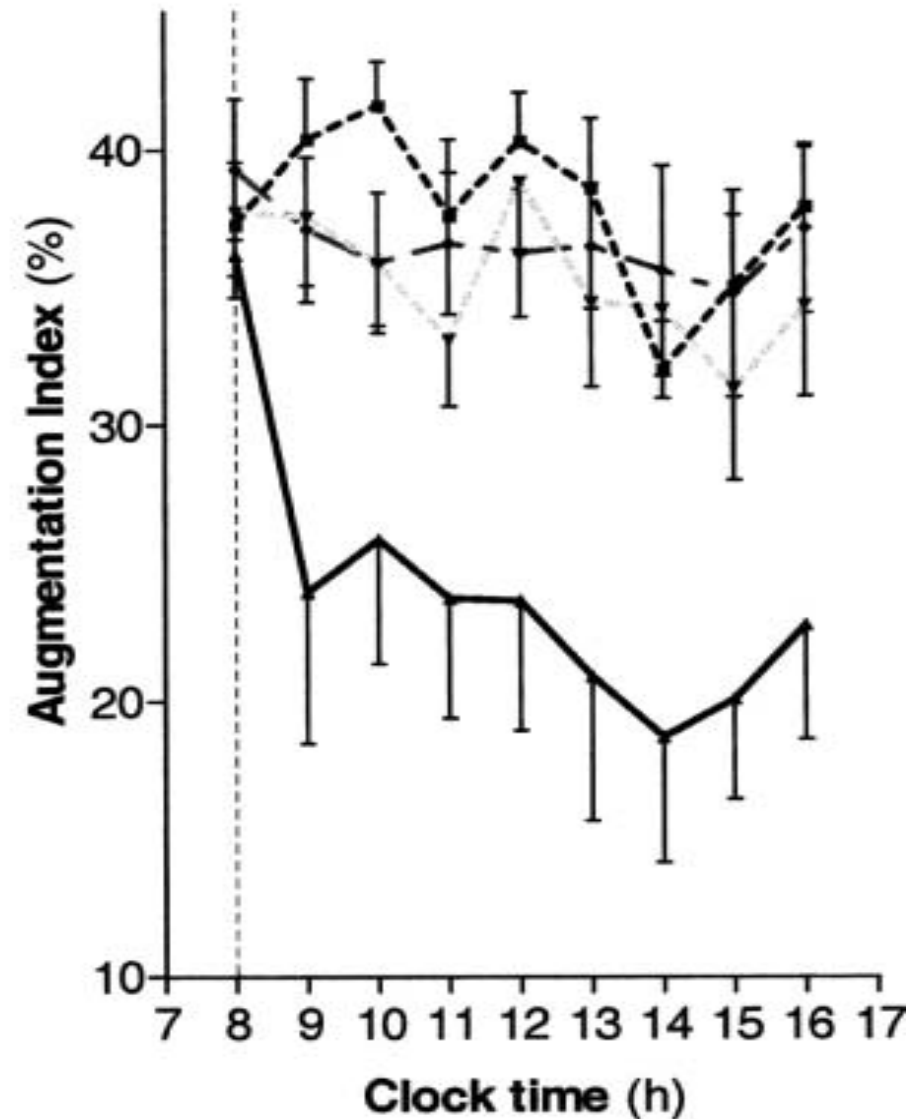
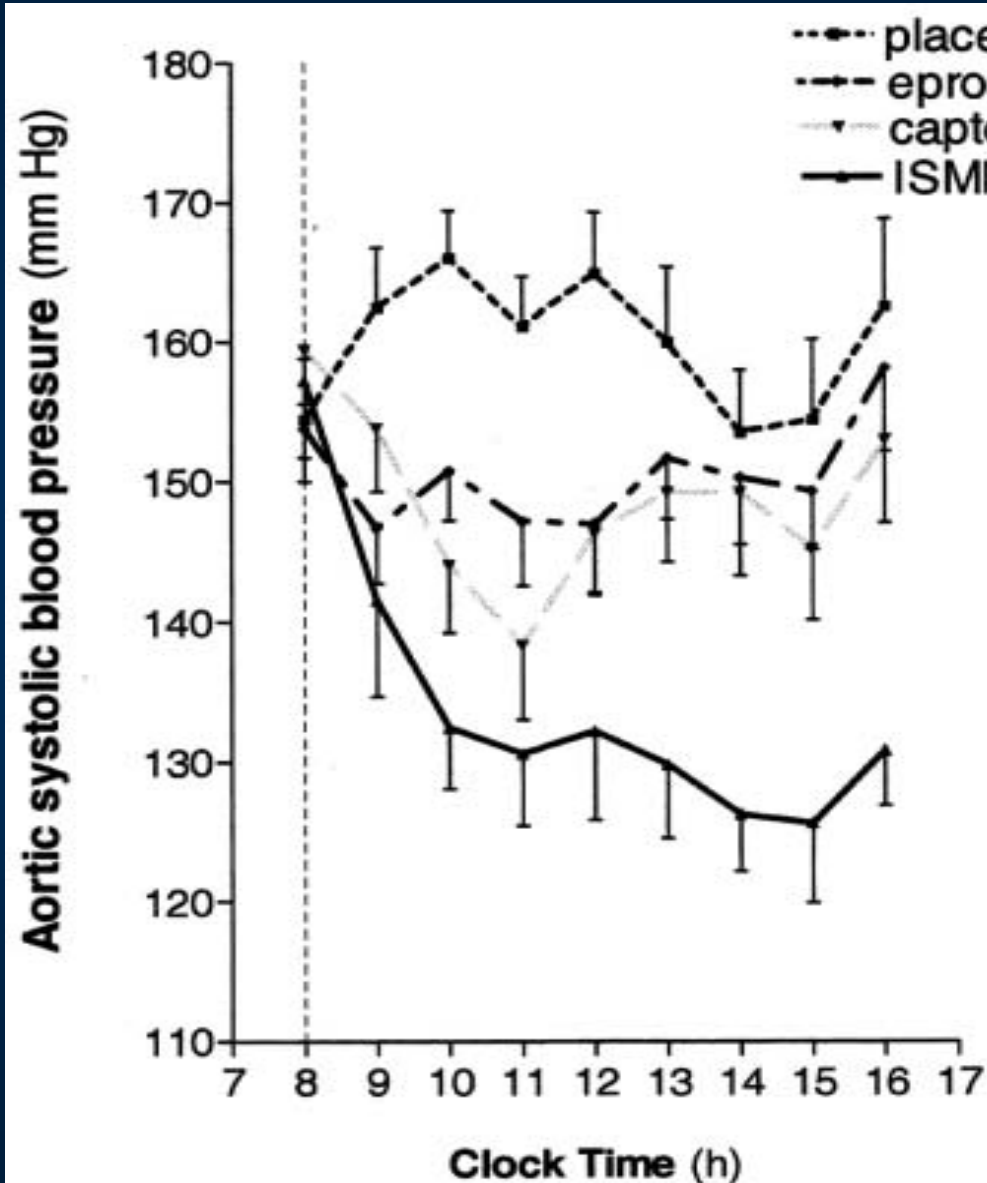


PWVb: before Irbesartan treatment, PWV12W: after 12 weeks of Irbesartan treatment, PWV24W: after 24 weeks of Irbesartan treatment

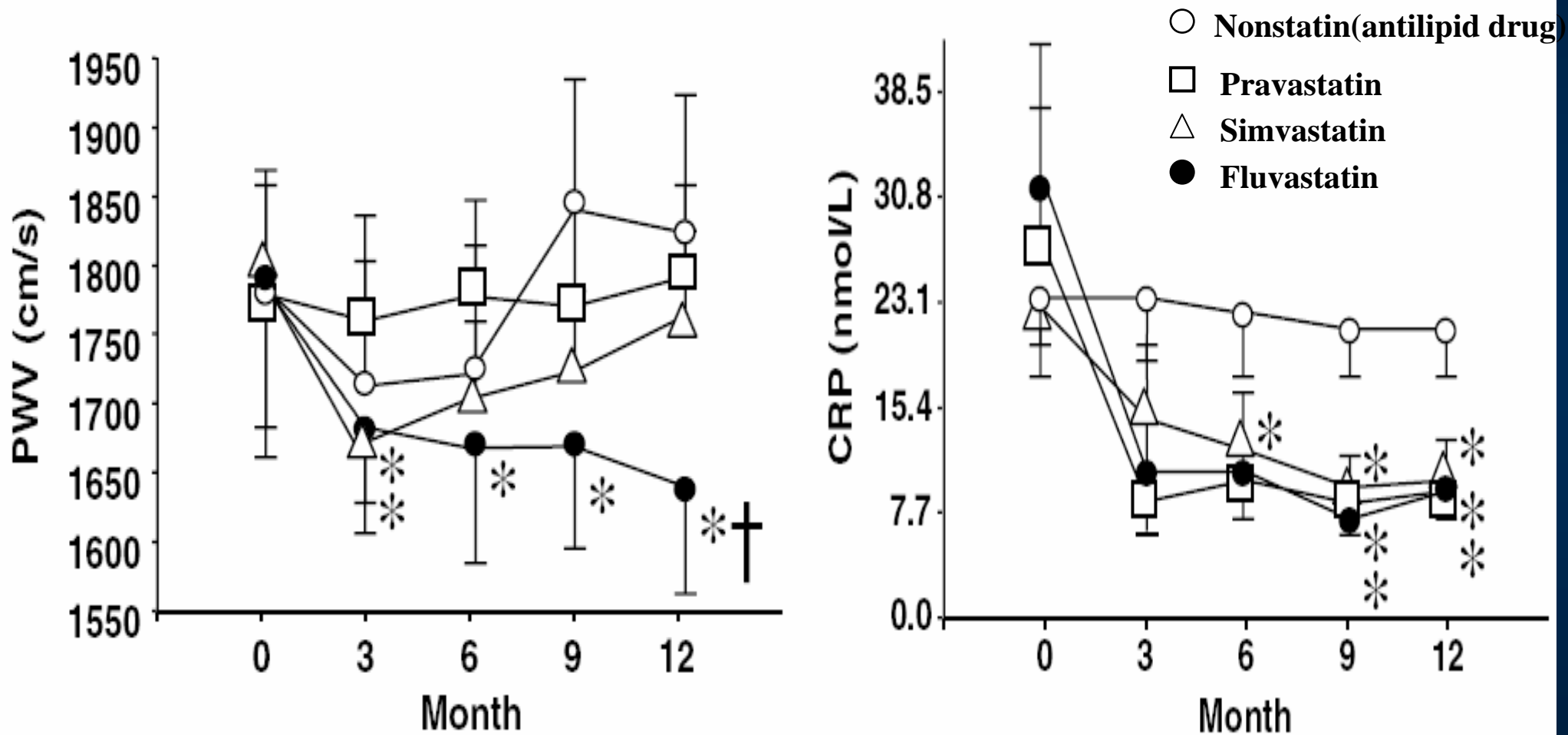
\* :  $p=0.06$  compared to PWVb, † :  $p=0.007$  compared to PWVb, NS:  $p>0.05$



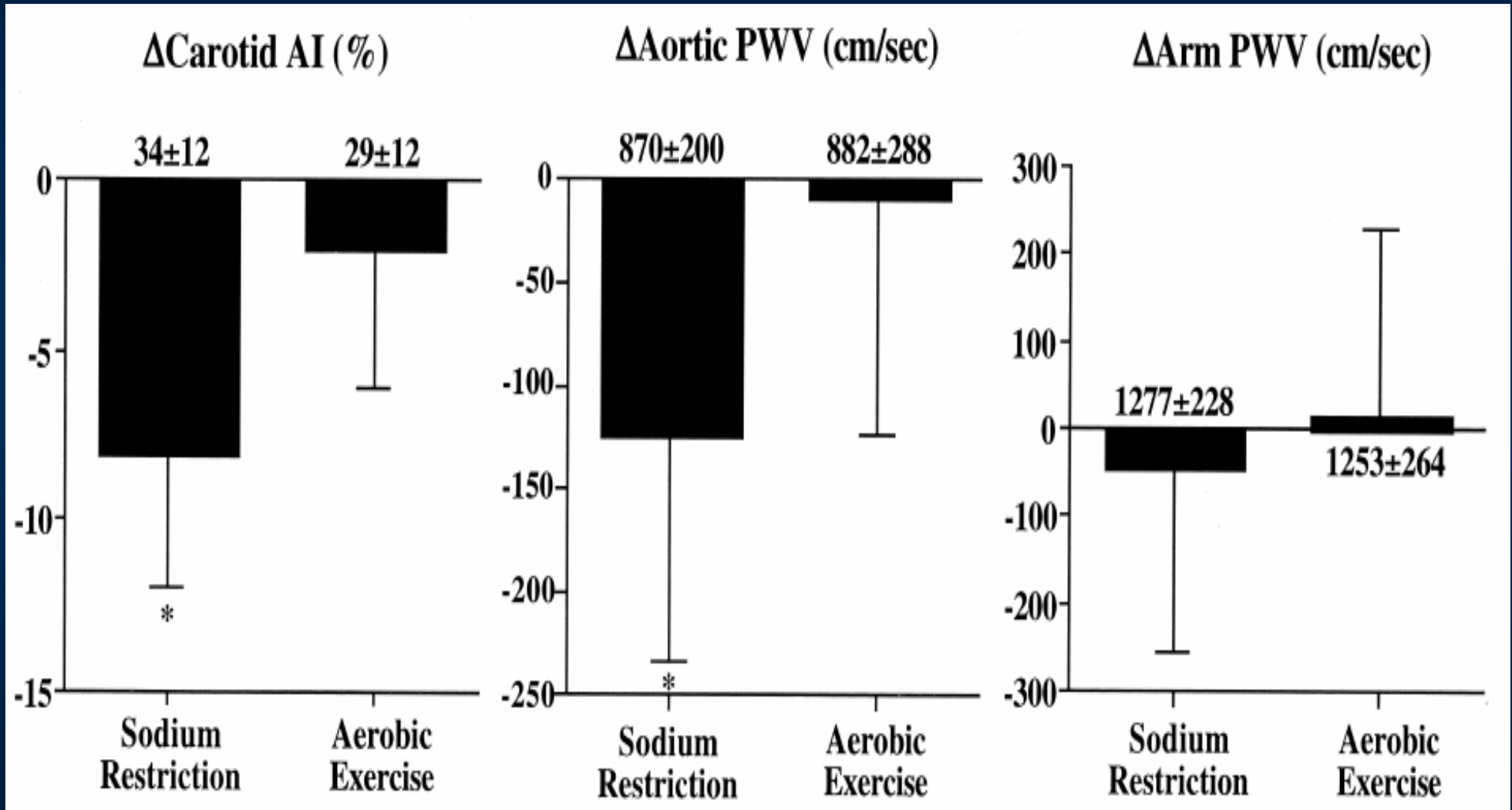
# Effects of Isosorbide Mononitrate and All Inhibition on Pulse Wave Reflection in Hypertension(SBP>150, DBP<100)



# Statin and PWV



# Aortic Stiffness: Na Reduction and Exercise

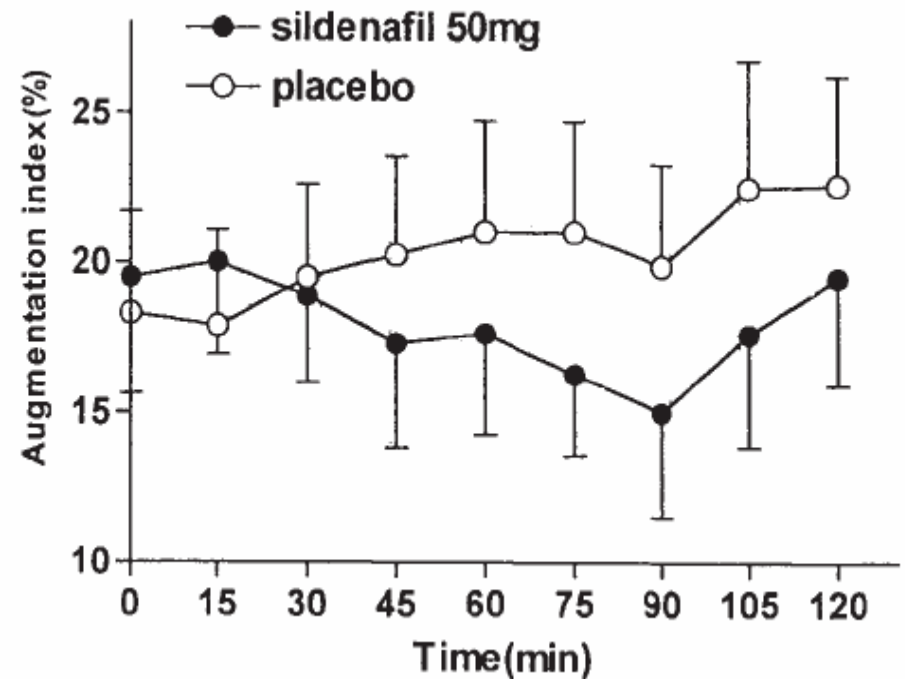
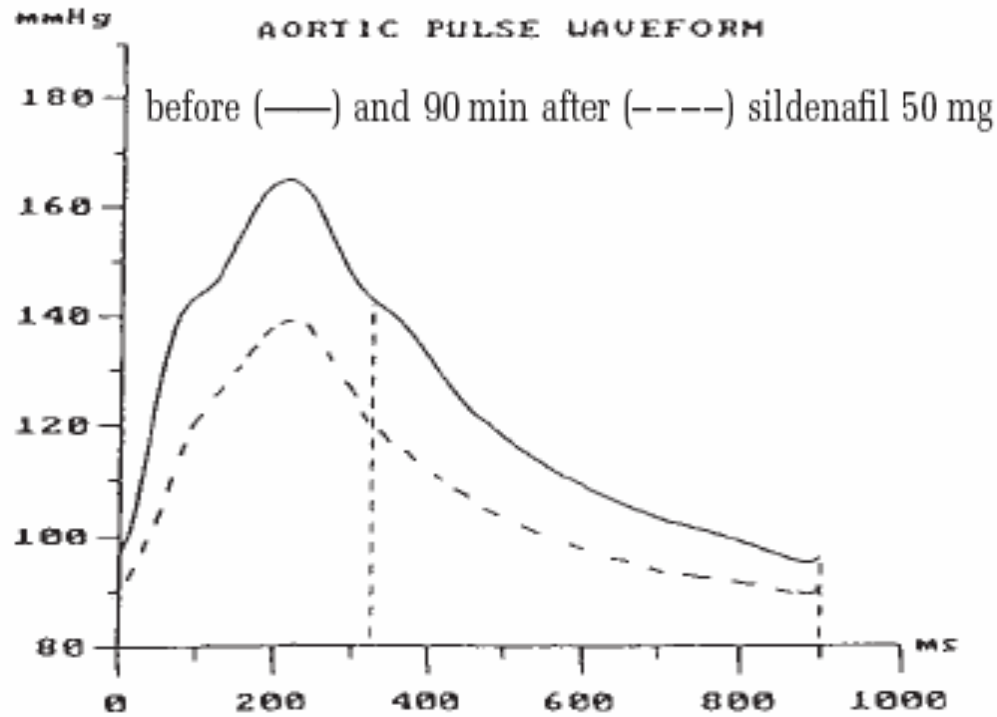


## ORIGINAL ARTICLE

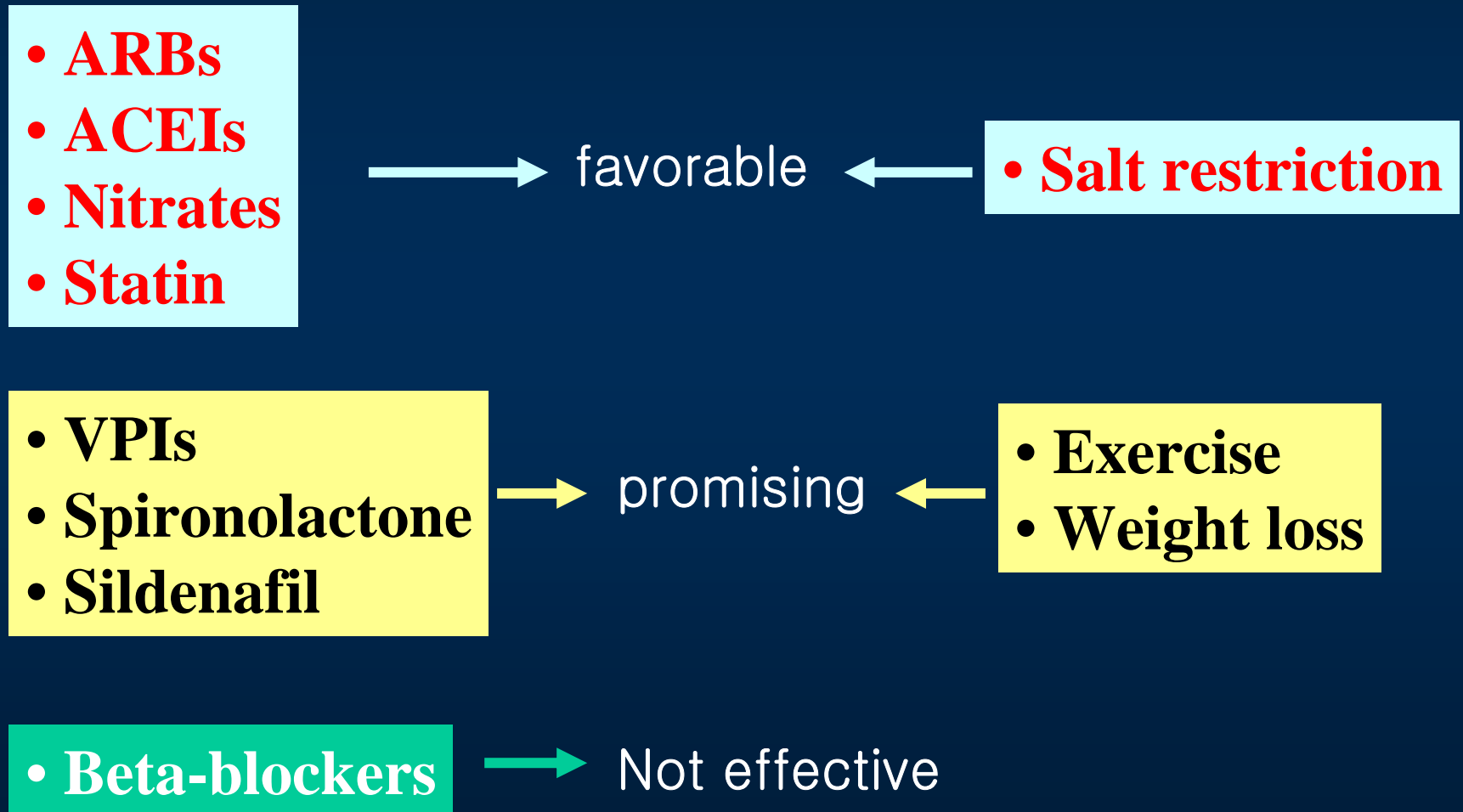
# Effect of sildenafil on blood pressure and arterial wave reflection in treated hypertensive men

A Mahmud, M Hennessy and J Feely

Department of Pharmacology and Therapeutics, Trinity Centre and Hypertension Clinic, St. James's Hospital, Dublin 8, Ireland



# Effect of various medications on aortic stiffness



Thank You for Your Attention !