



Università  
degli Studi  
di Ferrara

Anno Accademico 2019-2020  
C.L. Infermieristica – C.L. Ostetricia

Corso di  
Semeiotica Medica e Medicina Interna

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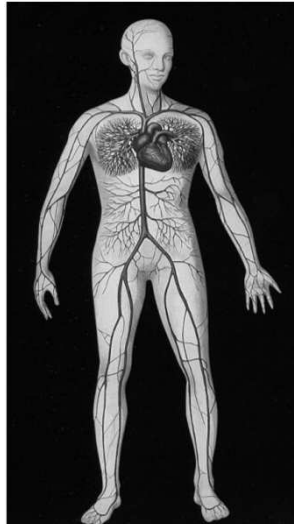
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## Malattie dell'apparato cardiovascolare I



- Ipertensione arteriosa
- Patologia cardiovascolare aterosclerotica (cardiopatia ischemica, angina e infarto)
- Differenze di genere
- Sindrome Takotsubo
- Scompenso cardiaco

## Ipertensione arteriosa



## Pressione arteriosa

Forma di energia potenziale che deriva dalla contrazione del cuore e serve per consentire al sangue di scorrere contro la resistenza offerta dalle arteriole periferiche.

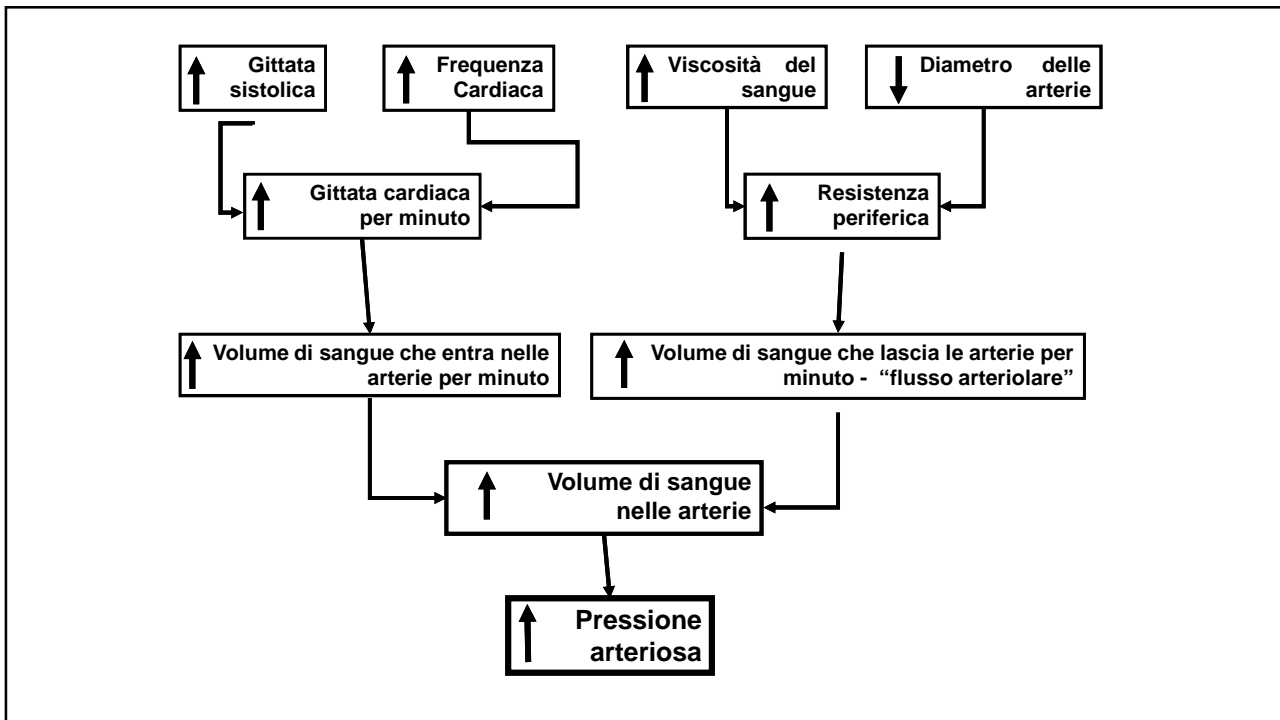
Viene espressa in mmHg: altezza raggiunta da colonna di mercurio, se si applica a questa la forza del sistema arterioso.

Da cosa è determinata?

$$P = V \times R$$

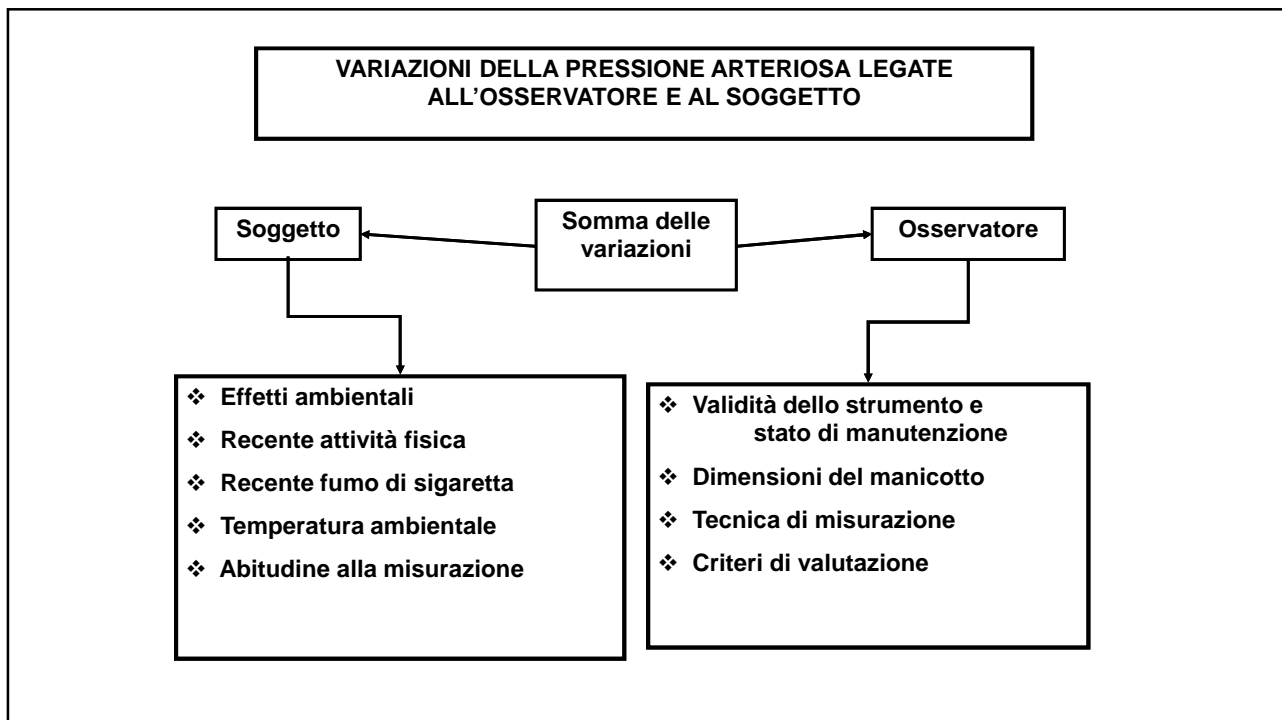
**V**= volume minuto cardiaco= vol sistolico x freq. cardiaca

**R**= resistenza periferica totale= lunghezza sistema arterioso x viscosità plasmatica/raggio del vaso<sup>4</sup>



## Pressione arteriosa

- Pressione arteriosa sistolica (PAS): picco della pressione intravascolare raggiunto durante la sistole ventricolare per l'eiezione del sangue ventricolare
- Pressione arteriosa diastolica (PAD): pressione intravascolare raggiunta alla fine della diastole ventricolare = resistenza vascolare periferica
- Pressione differenziale: differenza tra le due



## Regolazione della pressione arteriosa

- (1) Sistema nervoso autonomo
- (2) Sistema renina angiotensina aldosterone (SAA)
- (3) Endotelio vasale
- (4) Ormone antidiuretico (ADH)

## (1) Sistema Nervoso Autonomo

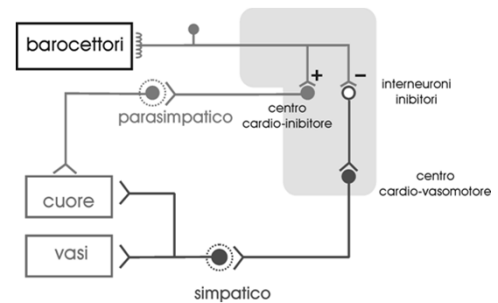
Regola l'omeostasi pressoria attraverso la modulazione:

(1) simpatica (recettori  $\beta$  renali), della diuresi e del volume circolante, sia direttamente che mediante il SRAA;

(2) adrenergica (recettori  $\alpha$ ) del tono vasocostrittore periferico

(3) parasimpatica e simpatica (recettori muscarinici sensibili all'acetilcolina (ACh) e recettori  $\beta$  sensibili alla noradrenalina (NA) della frequenza, della gittata cardiaca, dell'inotropismo cardiaco.

Il tono simpatico dipende da influenze nervose centrali, dalla regolazione riflessa barocettiva e chemocettiva e dalla secrezione periferica di NA

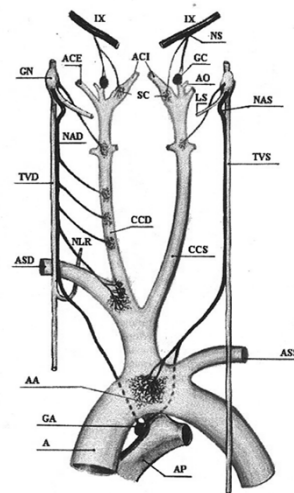


## (1) Sistema Nervoso Autonomo

I barocettori sono terminazioni sensitive localizzate nei seni carotidei e nell'arco aortico.

Captano variazioni pressorie e inviano stimoli ai centri nervosi cerebrali.

Se PA  $\downarrow$   $\rightarrow$   $\uparrow$  scarica vagale sul nodo sinusale  $\rightarrow$   $\uparrow$  FC e  $\uparrow$  attività nervi simpatici  $\rightarrow$  vasocostrizione periferica e rilascio renina  $\rightarrow$   $\uparrow$  pressione arteriosa



## (1) Sistema Nervoso Autonomo

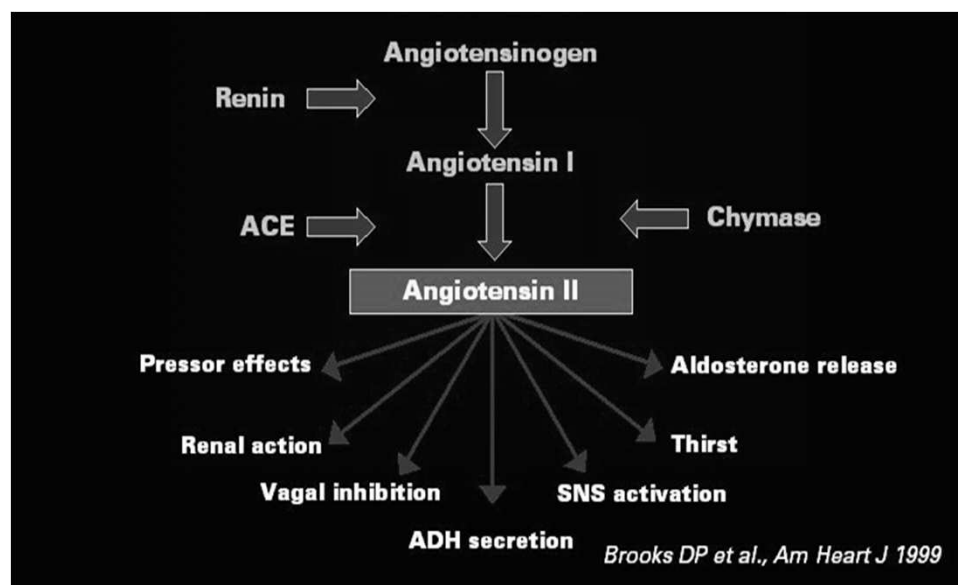
I chemocettori sono localizzati nell'arco aortico e nei corpi carotidei. Sono attivati da: **ipossia, acidosi e ipocapnia**

nelle situazioni di ipotensione grave.

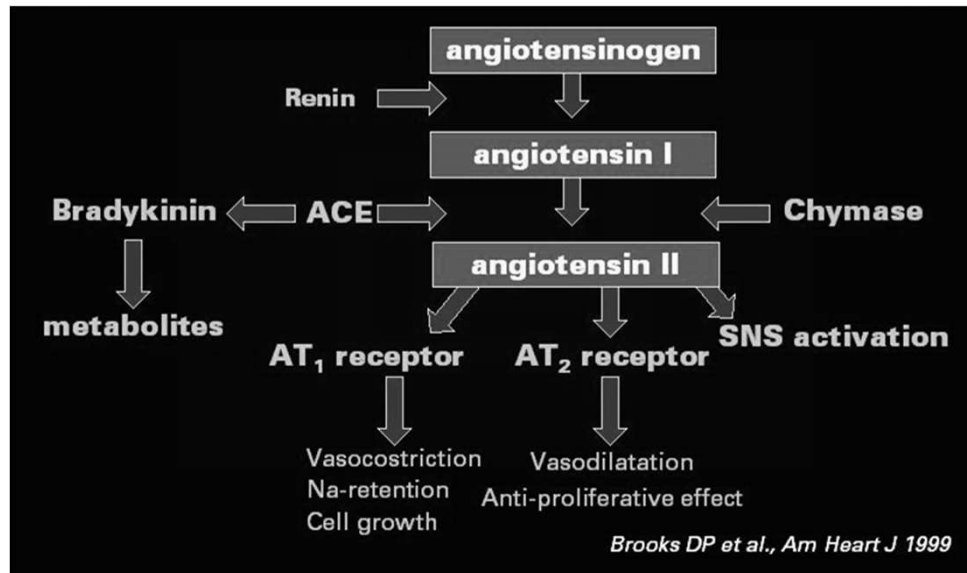
Inducono aumento della ventilazione se si riduce  $PO_2$  nel sangue arterioso

Attivazione → ↑ scarica simpatica → vasocostrizione arteriolare → ↑ FC → ↑ distensione polmonare

## (2) Sistema Renina-Angiotensina-Aldosterone (SRAA)



## (2) Sistema Renina-Angiotensina-Aldosterone (SRAA)



## (3) Endotelio vasale

Le cellule endoteliali secernono mediatori chimici ad azione paracrina che possono favorire o inibire la contrazione delle cellule muscolari lisce

**Vasodilatatori:** ossido nitrico (NO), prostaglandina I<sub>2</sub> (PGI<sub>2</sub>), fattore iperpolarizzante endotelio attivato

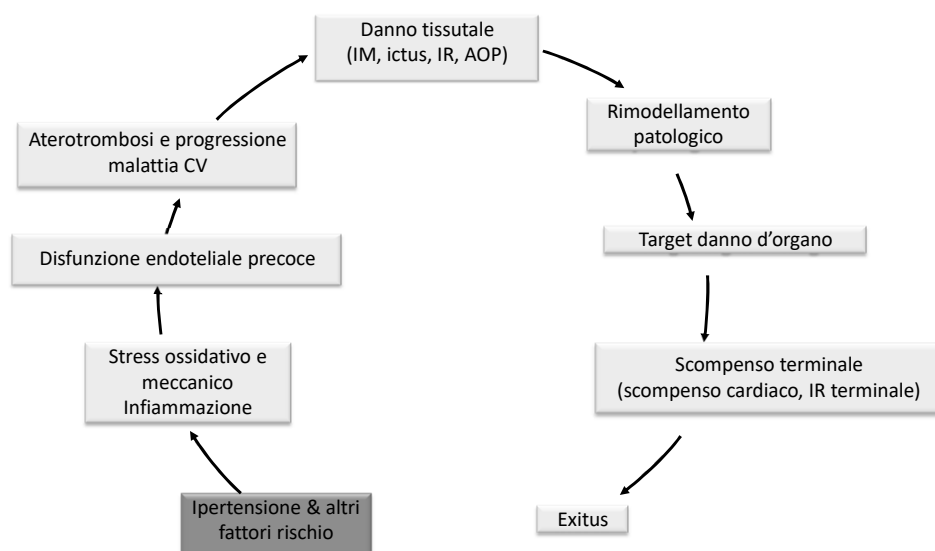
**Vasocostrittori:** endoteline (endotelina 1) trombossano A<sub>2</sub> (TXA<sub>2</sub>), prostaglandina H<sub>2</sub> (PGH<sub>2</sub>)

## (4) Ormone antidiuretico

Prodotto dall'ipotalamo, agisce modificando la permeabilità dell'epitelio del nefrone distale all'acqua e dunque la concentrazione urinaria. La sua liberazione dipende da modificazioni dell'osmolalità e dell'ipovolemia.

↑ Permeabilità all'acqua → passaggio di acqua nell'interstizio  
 → riduzione del volume urinario → ↑ concentrazione intratubulare di NaCl → ↑ riassorbimento di NaCl.

## Dall'ipertensione al continuum delle patologie cardiovascolari





Circulation



**Heart Disease and Stroke Statistics—2017 Update: A Report From the American Heart Association**  
 Emelia J. Benjamin, Michael J. Blaha, Stephanie E. Chirve, Mary Cushman, Sandeep R. Das, Rajat Deo, Sarah D. de Ferranti, James Floyd, Myrriam Fornage, Cathleen Gillespie, Carmen R. Isasi, Monik C. Jiménez, Lori Chaffin Jordan, Suzanne E. Judd, Daniel Lackland, Judith H. Lichtman, Lynda Lissabeth, Simin Liu, Chris T. Longenecker, Rachel H. Mackey, Kamihito Matsushita, Dariusz Mozaffarian, Michael E. Mussolino, Kharram Nasir, Robert W. Neumar, Latha Palaniappan, Dilip K. Pandey, Ravi K. Thiagarajan, Matthew J. Reeves, Matthew Ritchey, Carlos J. Rodriguez, Gregory A. Roth, Wayne D. Rosamond, Cornelia Sasson, Amyris Towfigh, Connie W. Tsao, Melanie B. Turner, Salim S. Virani, Jennifer H. Voeks, Joshua Z. Willey, John T. Wilkins, Jason HY. Wu, Heather M. Alger, Sally S. Wong, Paul Muntner and On behalf of the American Heart Association Statistics Committee and Stroke Statistics Subcommittee

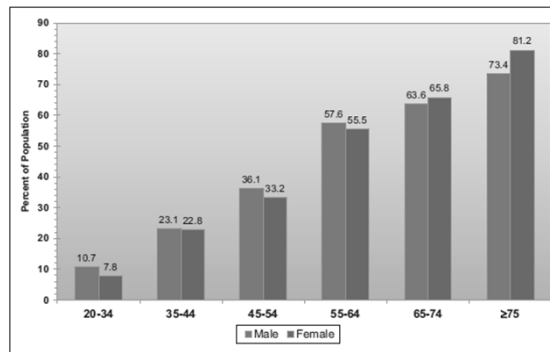
*Circulation*, published online January 25, 2017;  
 Circulation is published by the American Heart Association, 7272 Greenville Avenue, Dallas, TX 75231  
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 Print ISSN: 0009-7322, Online ISSN: 1524-4539

**AHA STATISTICAL UPDATE**

**Heart Disease and Stroke Statistics—2017 Update**  
**A Report From the American Heart Association**

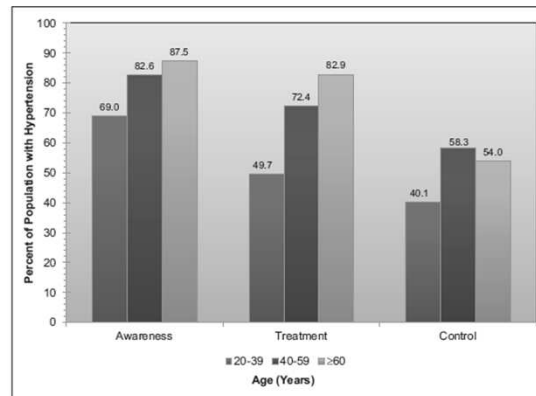
*AHA Statistical 2016 Update, Circulation 2017*

**Prevalenza dell’ipertensione negli adulti (NHANES 2007-2012)**



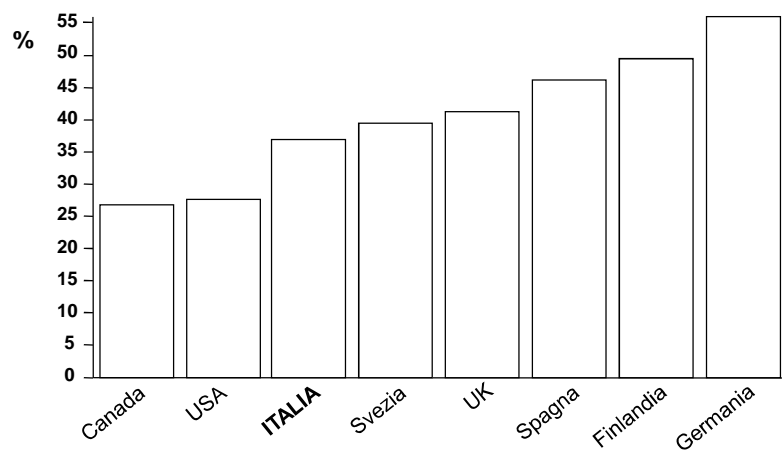
*AHA Statistical 2017 Update, Circulation 2017*

## Prevalenza dell'ipertensione negli adulti (NHANES 2007-2012)



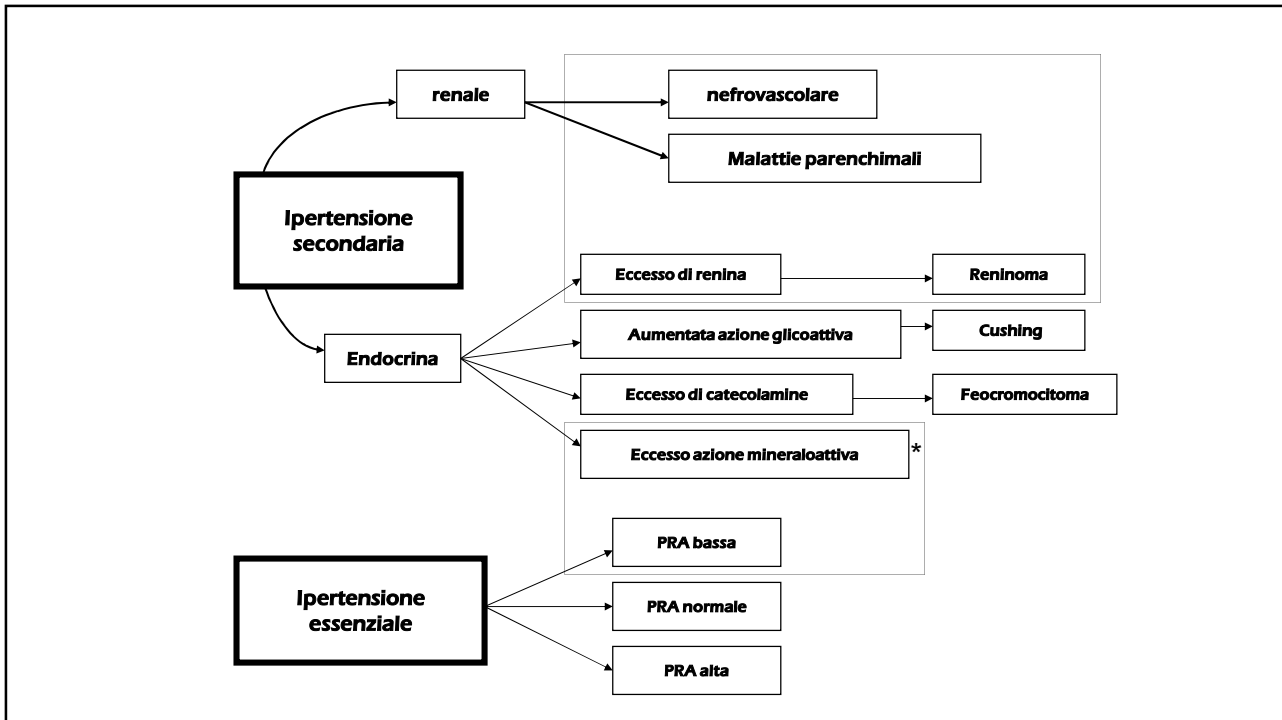
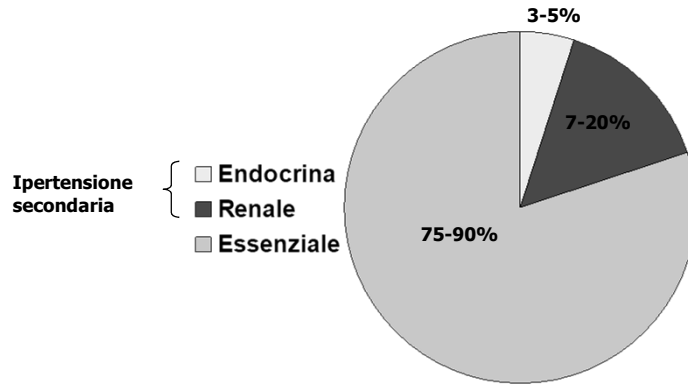
*AHA Statistical 2017 Update, Circulation 2017*

## Prevalenza dell'ipertensione nella popolazione italiana



*Wolf-Maier K et al, JAMA 2003*

## Cause di ipertensione



## 2018 ESC/ESH Guidelines for the management of arterial hypertension

The Task Force for the management of arterial hypertension of the European Society of Cardiology (ESC) and the European Society of Hypertension (ESH)

Table 1 ESC Classes of recommendations

| Classes of recommendations | Definition   | Suggested wording to use    |
|----------------------------|--|-----------------------------|
| Class I                    | Evidence and/or general agreement that a given treatment or procedure is beneficial, useful, effective.                        | Is recommended/is indicated |
| Class II                   | Conflicting evidence and/or a divergence of opinion about the usefulness/efficacy of the given treatment or procedure.         |                             |
| Class IIa                  | <i>Weight of evidence/opinion is in favour of usefulness/efficacy.</i>   | Should be considered        |
| Class IIb                  | <i>Usefulness/efficacy is less well established by evidence/opinion.</i>   | May be considered           |
| Class III                  | Evidence or general agreement that the given treatment or procedure is not useful/effective, and in some cases may be harmful. | Is not recommended          |

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Table 2 ESC Levels of evidence

|                     |  |
|---------------------|--|
| Level of evidence A | Data derived from multiple randomized clinical trials or meta-analyses.                      |
| Level of evidence B | Data derived from a single randomized clinical trial or large non-randomized studies.        |
| Level of evidence C | Consensus of opinion of the experts and/or small studies, retrospective studies, registries. |

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### New concepts

#### BP measurement

- **Wider use of out-of-office BP measurement with ABPM and/or HBPM, especially HBPM**, as an option to confirm the diagnosis of hypertension, detect white-coat and masked hypertension, and monitor BP control.

#### Less conservative treatment of BP in older and very old patients

- **Lower BP thresholds and treatment targets for older patients**, with emphasis on considerations of biological rather than chronological age (i.e. the importance of frailty, independence, and the tolerability of treatment).
- Recommendation that **treatment should never be denied or withdrawn on the basis of age**, provided that treatment is tolerated.

#### A SPC treatment strategy to improve BP control

- **Preferred use of two-drug combination therapy** for the initial treatment of most people with hypertension.
- **A single-pill treatment strategy for hypertension** with the preferred use of SPC therapy for most patients.
- **Simplified drug treatment algorithms** with the preferred use of an ACE inhibitor or ARB, combined with a CCB and/or a thiazide/thiazide-like diuretic, as the core treatment strategy for most patients, with beta-blockers used for specific indications.

#### New target ranges for BP in treated patients

- **Target BP ranges for treated patients** to better identify the recommended BP target and **lower safety boundaries for treated BP**, according to a patient's age and specific comorbidities.

#### Detecting poor adherence to drug therapy

- A strong emphasis on the **importance of evaluating treatment adherence** as a major cause of poor BP control.

#### A key role for nurses and pharmacists in the longer-term management of hypertension

- **The important role of nurses and pharmacists** in the education, support, and follow-up of treated hypertensive patients is emphasized as part of the overall strategy to improve BP control.

ABPM = ambulatory blood pressure monitoring; ACE = angiotensin-converting enzyme; AF = atrial fibrillation; ARB = angiotensin receptor blocker; BP = blood pressure; CCB = calcium channel blocker; CV = cardiovascular; CVD = cardiovascular disease; HBPM = home blood pressure monitoring; HMOD = hypertension-mediated organ damage; SCORE = Systematic COronary Risk Evaluation; SPC = single-pill combination.

### 2018 ESC/ESH Guidelines for the management of arterial hypertension

The Task Force for the management of arterial hypertension of the European Society of Cardiology (ESC) and the European Society of Hypertension (ESH)

**Table 3** Classification of office blood pressure<sup>a</sup> and definitions of hypertension grade<sup>b</sup>

| 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 hypertension                        | 140–159         | and/or | 90–99            |
| Grade 2 hypertension                        | 160–179         | and/or | 100–109          |
| Grade 3 hypertension                        | ≥180            | and/or | ≥110             |
| Isolated systolic hypertension <sup>b</sup> | ≥140            | and    | <90              |

BP = blood pressure; SBP = systolic blood pressure.

<sup>a</sup>BP category is defined according to seated clinic BP and by the highest level of BP, whether systolic or diastolic.

<sup>b</sup>Isolated systolic hypertension is graded 1, 2, or 3 according to SBP values in the ranges indicated.

The same classification is used for all ages from 16 years.

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| Hypertension disease staging      | Other risk factors, HMOD, or disease                                  | BP (mmHg) grading                       |                                     |                                       |                                    |
|-----------------------------------|---|---|-------------------------------------|---------------------------------------|------------------------------------|
|                                   |   | High normal<br>SBP 130–139<br>DBP 85–89 | Grade 1<br>SBP 140–159<br>DBP 90–99 | Grade 2<br>SBP 160–179<br>DBP 100–109 | Grade 3<br>SBP ≥180<br>or DBP ≥110 |
| Stage 1<br>(uncomplicated)        | No other risk factors   | Low risk                                | Low risk                            | Moderate risk                         | High risk                          |
|                                   | 1 or 2 risk factors   | Low risk                                | Moderate risk                       | Moderate to high risk                 | High risk                          |
|                                   | ≥3 risk factors   | Low to Moderate risk                    | Moderate to high risk               | High Risk                             | High risk                          |
| Stage 2<br>(asymptomatic disease) | HMOD, CKD grade 3, or diabetes mellitus without organ damage          | Moderate to high risk                   | High risk                           | High risk                             | High to very high risk             |
| Stage 3<br>(established disease)  | Established CVD, CKD grade ≥4, or diabetes mellitus with organ damage | Very high risk                          | Very high risk                      | Very high risk                        | Very high risk                     |

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**Figure 1** Classification of hypertension stages according to blood pressure levels, presence of cardiovascular risk factors, hypertension-mediated organ damage, or comorbidities. CV risk is illustrated for a middle-aged male. The CV risk does not necessarily correspond to the actual risk at different ages. The use of the SCORE system is recommended for formal estimation of CV risk for treatment decisions. BP = blood pressure; CKD = chronic kidney disease; CV = cardiovascular; DBP = diastolic blood pressure; HMOD = hypertension-mediated organ damage; SBP = systolic blood pressure; SCORE = Systematic COronary Risk Evaluation.

## Fattori di rischio

### Stili di Vita

Dieta Ipercalorica o ricca in Acidi grassi Saturi e Colesterolo

Fumo

Consumo eccessivo di Alcol

Sedentarietà

### FRC Modificabili

Iperensione Arteriosa

Elevato LDL-Colesterolo

Ridotto HDL-Colesterolo

Elevati Trigliceridi

Iperglicemia/Diabete

Obesità

### FRC Non Modificabili

Età

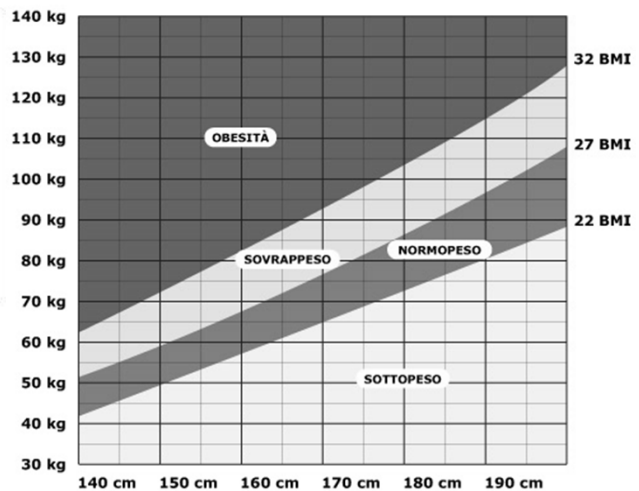
Sesso

Storia familiare di CHD o altra malattia aterosclerotica precoce (maschi <55 a, femmine <65 a)

Anamnesi positiva per CHD o altra malattia aterosclerotica

## Fattori di rischio

| Classificazione    | BMI (kg/m <sup>2</sup> ) |
|--------------------|--------------------------|
|                    | Cut-off principali       |
| Sottopeso          | < 18,50                  |
| Magrezza severa    | < 16,00                  |
| Magrezza moderata  | 16,00 - 16,99            |
| Magrezza leggera   | 17,00 - 18,49            |
| Normopeso          | 18,50 - 24,99            |
| Sovrappeso         | ≥ 25,00                  |
| Pre-obesità        | 25,00 - 29,99            |
| Obesità            | ≥ 30,00                  |
| Obesità I classe   | 30,00 - 34,99            |
| Obesità II classe  | 35,00 - 39,99            |
| Obesità III classe | ≥ 40,00                  |

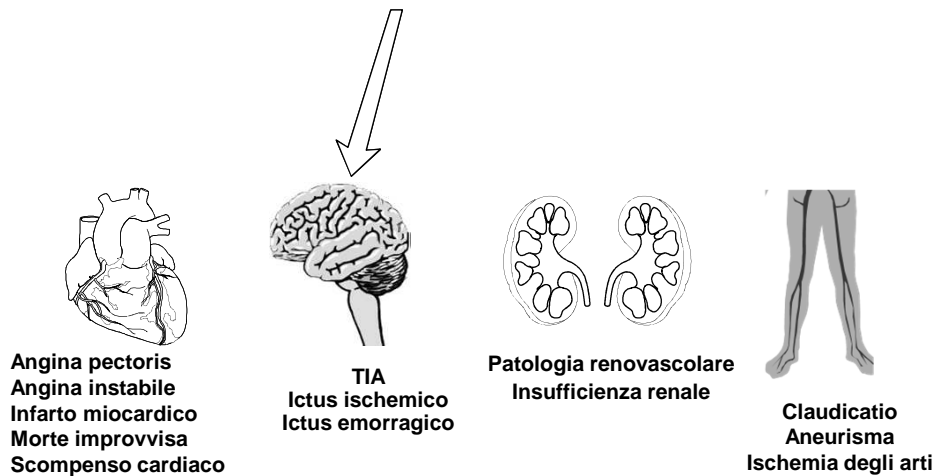


## Stratificazione del rischio

| Risk factors   | Asymptomatic organ damage   | Diabetes mellitus   |
|--|---|---|
| Male sex   | Pulse pressure (in the elderly) $\geq 60$ mmHg  | Fasting plasma glucose $\geq 7.0$ mmol/L (126 mg/dL) on two repeated measurements, and/or   |
| Age (men $\geq 55$ years; women $\geq 65$ years)   | Electrocardiographic LVH (Sokolow-Lyon index $> 3.5$ mV; RaVL $> 1.1$ mV; Cornell voltage duration product $> 244$ mV*ms), or       | HbA <sub>1c</sub> $> 7\%$ (53 mmol/mol), and/or   |
| Smoking  | Echocardiographic LVH [LVM index: men $> 115$ g/m <sup>2</sup> ; women $> 95$ g/m <sup>2</sup> (BSA)] <sup>a</sup>                  | Post-load plasma glucose $> 11.0$ mmol/L (198 mg/dL)  |
| Dyslipidaemia  | Carotid wall thickening (IMT $> 0.9$ mm) or plaque  | <b>Established CV or renal disease</b>  |
| Total cholesterol $> 4.9$ mmol/L (190 mg/dL), and/or   | Carotid-femoral PWV $> 10$ m/s  | Cerebrovascular disease: ischaemic stroke; cerebral haemorrhage; transient ischaemic attack |
| Low-density lipoprotein cholesterol $> 3.0$ mmol/L (115 mg/dL), and/or                                       | Ankle-brachial Index $< 0.9$  | CHD: myocardial infarction; angina; myocardial revascularization with PCI or CABG           |
| High-density lipoprotein cholesterol: men $< 1.0$ mmol/L (40 mg/dL), women $< 1.2$ mmol/L (46 mg/dL), and/or | CKD with eGFR 30–60 mL/min/1.73 m <sup>2</sup> (BSA)  | Heart failure, including heart failure with preserved EF                                    |
| Triglycerides $> 1.7$ mmol/L (150 mg/dL)   | Microalbuminuria (30–300 mg/24 h), or albumin-creatinine ratio (30–300 mg/g; 3.4–34 mg/mmol) (preferentially on morning spot urine) | Symptomatic lower extremities peripheral artery disease                                     |
| Fasting plasma glucose 5.6–6.9 mmol/L (102–125 mg/dL)  |   | CKD with eGFR $< 30$ mL/min/1.73m <sup>2</sup> (BSA); proteinuria ( $> 300$ mg/24 h).       |
| Abnormal glucose tolerance test  |   | Advanced retinopathy; haemorrhages or exudates, papilloedema                                |
| Obesity [BMI $\geq 30$ kg/m <sup>2</sup> (height <sup>2</sup> )]   |   |   |
| Abdominal obesity (waist circumference: men $\geq 102$ cm; women $\geq 88$ cm) (in Caucasians)               |   |   |
| Family history of premature CVD (men aged $< 55$ years; women aged $< 65$ years)                             |   |   |

## Danno d'organo

### Ipertensione



## Danno d'organo



## Valutazione diagnostica

### SCOPI

- (a) Stabilire i livelli pressori
- (b) Identificare le cause di ipertensione secondaria
- (c) Stratificazione prognostica del rischio

### PROCEDURE

- (d) Misurazione ripetuta della pressione arteriosa
- (e) Valutazione della storia clinica
- (f) Esame fisico
- (g) Valutazione strumentale e di laboratorio



## Definizione dei valori pressori

**Table 9** Definitions of hypertension according to office, ambulatory, and home blood pressure levels

| Category                    | SBP (mmHg) |        | DBP (mmHg) |
|-----------------------------|------------|--------|------------|
| Office BP <sup>a</sup>      | ≥140       | and/or | ≥90        |
| Ambulatory BP               |            |        |            |
| Daytime (or awake) mean     | ≥135       | and/or | ≥85        |
| Night-time (or asleep) mean | ≥120       | and/or | ≥70        |
| 24 h mean                   | ≥130       | and/or | ≥80        |
| Home BP mean                | ≥135       | and/or | ≥85        |

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BP = blood pressure; DBP = diastolic blood pressure; SBP = systolic blood pressure.

<sup>a</sup>Refers to conventional office BP rather than unattended office BP.

## Monitoraggio non invasivo della PA 24-ore

### INDICAZIONI

- Considerevole variabilità tra i valori pressori ambulatoriali
- Elevati valori pressori in pz. con rischio cardiovascolare basso
- Marcata discrepanza tra valori pressori ambulatoriali e domiciliari
- Sospetta ipertensione secondaria
- Monitoraggio terapia (efficacia antipertensiva – copertura 24 ore)
- Dipping notturno
- Finalità di ricerca

## Routine diagnostica e laboratoristica

**Table 14 Routine workup for evaluation of hypertensive patients**

| Routine laboratory tests  |
|---|
| Haemoglobin and/or haematocrit  |
| Fasting blood glucose and glycated HbA <sub>1c</sub>  |
| Blood lipids: total cholesterol, LDL cholesterol, HDL cholesterol   |
| Blood triglycerides   |
| Blood potassium and sodium  |
| Blood uric acid   |
| Blood creatinine and eGFR   |
| Blood liver function tests  |
| Urine analysis: microscopic examination; urinary protein by dipstick test or, ideally, albumin:creatinine ratio |
| 12-lead ECG   |

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eGFR = estimated glomerular filtration rate; ECG = electrocardiogram; HbA<sub>1c</sub> = haemoglobin A1c.

## Valutazione del danno d'organo

**Table 15 Assessment of hypertension-mediated organ damage**

| Basic screening tests for HMOD           | Indication and interpretation  |
|--|--|
| 12-lead ECG                              | Screen for LVH and other possible cardiac abnormalities, and to document heart rate and cardiac rhythm   |
| Urine albumin:creatinine ratio           | To detect elevations in albumin excretion indicative of possible renal disease   |
| Blood creatinine and eGFR                | To detect possible renal disease   |
| Fundoscopy                               | To detect hypertensive retinopathy, especially in patients with grade 2 or 3 hypertension  |
| More detailed screening for HMOD         |  |
| Echocardiography                         | To evaluate cardiac structure and function, when this information will influence treatment decisions   |
| Carotid ultrasound                       | To determine the presence of carotid plaque or stenosis, particularly in patients with cerebrovascular disease or vascular disease elsewhere   |
| Abdominal ultrasound and Doppler studies | <ul style="list-style-type: none"> <li>To evaluate renal size and structure (e.g. scarring) and exclude renal tract obstruction as possible underlying causes of CKD and hypertension</li> <li>Evaluate abdominal aorta for evidence of aneurysmal dilatation and vascular disease</li> <li>Examine adrenal glands for evidence of adenoma or pheochromocytoma (CT or MRI preferred for detailed examination); see section 8.2 regarding screening for secondary hypertension</li> <li>Renal artery Doppler studies to screen for the presence of renovascular disease, especially in the presence of asymmetric renal size</li> </ul> |
| PWV                                      | An index of aortic stiffness and underlying arteriosclerosis   |
| ABI                                      | Screen for evidence of LEAD  |
| Cognitive function testing               | To evaluate cognition in patients with symptoms suggestive of cognitive impairment   |
| Brain imaging                            | To evaluate the presence of ischaemic or haemorrhagic brain injury, especially in patients with a history of cerebrovascular disease or cognitive decline  |

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ABI = ankle-brachial index; CKD = chronic kidney disease; CT = computed tomography; ECG = electrocardiogram; eGFR = estimated glomerular filtration rate; HMOD = hypertension-mediated organ damage; LEAD = lower extremity artery disease; LVH = left ventricular hypertrophy; MRI = magnetic resonance imaging; PWV = pulse wave velocity.

**Table 18 Sensitivity to detect treatment-induced changes, reproducibility and operator independence, time to changes, and prognostic value of changes provided by markers of hypertension-mediated organ damage**

| Marker of HMOD            | Sensitivity to changes | Reproducibility and operator independence | Time to changes        | Prognostic value of the change |
|---------------------------|------------------------|---|------------------------|--------------------------------|
| LVH by ECG                | Low                    | High                                      | Moderate (>6 months)   | Yes                            |
| LVH by echocardiogram     | Moderate               | Moderate                                  | Moderate (>6 months)   | Yes                            |
| LVH by CMR                | High                   | High                                      | Moderate (>6 months)   | No data                        |
| eGFR                      | Moderate               | High                                      | Very slow (years)      | Yes                            |
| Urinary protein excretion | High                   | Moderate                                  | Fast (weeks to months) | Moderate                       |
| Carotid IMT               | Very low               | Low                                       | Slow (>12 months)      | No                             |
| PWV                       | High                   | Low                                       | Fast (weeks to months) | Limited data                   |
| Ankle-brachial index      | Low                    | Moderate                                  | Slow (>12 months)      | Moderate                       |

CMR = cardiac magnetic resonance; ECG = electrocardiogram; eGFR = estimated glomerular filtration rate; HMOD = hypertension-mediated organ damage; IMT = intima-media thickness; LVH = left ventricular hypertrophy; PWV = pulse wave velocity.

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## Obiettivi del trattamento

L'obiettivo primario del trattamento del pz. iperteso è quello di ottenere la **massima riduzione del rischio cardiovascolare globale a lungo termine**.

E' perciò richiesto il trattamento di tutti i fattori di rischio reversibili identificati, una gestione appropriata delle condizioni cliniche associate, oltre al trattamento dell'ipertensione stessa

## Modifiche degli stili di vita (lifestyle changes)

Abolizione del fumo

Calo ponderale

Riduzione del consumo di alcolici, se eccessivo

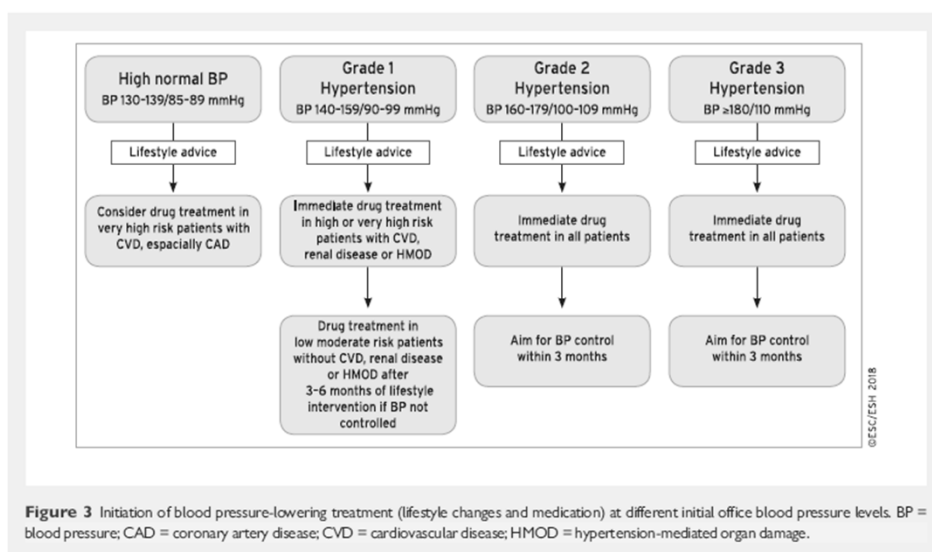
Esercizio fisico

Dieta iposodica

Incremento di apporto alimentare di frutta e verdura

Riduzione dei grassi totali alimentari

## Trattamento iniziale



**Figure 3** Initiation of blood pressure-lowering treatment (lifestyle changes and medication) at different initial office blood pressure levels. BP = blood pressure; CAD = coronary artery disease; CVD = cardiovascular disease; HMOD = hypertension-mediated organ damage.

## Strategie terapeutiche

Nella maggior parte dei casi il trattamento dovrebbe essere iniziato gradualmente e valutato nell'arco di alcune settimane

A seconda dei valori pressori di base e della presenza o meno di altri fattori, è ragionevole iniziare la terapia con uno o due farmaci, entrambi a basso dosaggio

E' probabile che per raggiungere il target pressorio sia necessario l'impiego di una terapia di associazione tra due o più farmaci.

Entrambe le strategie terapeutiche presentano vantaggi e svantaggi.

| Other risk factors,<br>asymptomatic organ damage<br>or disease | Blood Pressure (mmHg)                       |   |  |   |
|--|---|---|--|---|
|  | High normal<br>SBP 130–139<br>or DBP 85–89  | Grade 1 HT<br>SBP 140–159<br>or DBP 90–99                                       | Grade 2 HT<br>SBP 160–179<br>or DBP 100–109                                    | Grade 3 HT<br>SBP ≥180<br>or DBP ≥110                         |
| No other RF  | • No BP Intervention                        | • Lifestyle changes for several months<br>• Then add BP drugs targeting <140/90 | • Lifestyle changes for several weeks<br>• Then add BP drugs targeting <140/90 | • Lifestyle changes<br>• Immediate BP drugs targeting <140/90 |
| 1–2 RF   | • Lifestyle changes<br>• No BP Intervention | • Lifestyle changes for several weeks<br>• Then add BP drugs targeting <140/90  | • Lifestyle changes for several weeks<br>• Then add BP drugs targeting <140/90 | • Lifestyle changes<br>• Immediate BP drugs targeting <140/90 |
| ≥3 RF  | • Lifestyle changes<br>• No BP intervention | • Lifestyle changes for several weeks<br>• Then add BP drugs targeting <140/90  | • Lifestyle changes<br>• BP drugs targeting <140/90                            | • Lifestyle changes<br>• Immediate BP drugs targeting <140/90 |
| OD, CKD stage 3 or diabetes                                    | • Lifestyle changes<br>• No BP intervention | • Lifestyle changes<br>• BP drugs targeting <140/90                             | • Lifestyle changes<br>• BP drugs targeting <140/90                            | • Lifestyle changes<br>• Immediate BP drugs targeting <140/90 |
| Symptomatic CVD,<br>CKD stage ≥4 or<br>diabetes with OD/RFs    | • Lifestyle changes<br>• No BP intervention | • Lifestyle changes<br>• BP drugs targeting <140/90                             | • Lifestyle changes<br>• BP drugs targeting <140/90                            | • Lifestyle changes<br>• Immediate BP drugs targeting <140/90 |

BP = blood pressure; CKD = chronic kidney disease; CV = cardiovascular; CVD = cardiovascular disease; DBP = diastolic blood pressure; HT = hypertension;  
OD = organ damage; RF = risk factor; SBP = systolic blood pressure.

## Soglie di trattamento

Table 19 Summary of office blood pressure thresholds for treatment

| Age group                                    | Office SBP treatment threshold (mmHg) |            |       |                   |                   | Office DBP treatment threshold (mmHg) |
|--|---------------------------------------|------------|-------|-------------------|-------------------|---------------------------------------|
|  | Hypertension                          | + Diabetes | + CKD | + CAD             | + Stroke/TIA      |                                       |
| 18 - 65 years                                | ≥140                                  | ≥140       | ≥140  | ≥140 <sup>a</sup> | ≥140 <sup>a</sup> | ≥90                                   |
| 65 - 79 years                                | ≥140                                  | ≥140       | ≥140  | ≥140 <sup>a</sup> | ≥140 <sup>a</sup> | ≥90                                   |
| ≥80 years                                    | ≥160                                  | ≥160       | ≥160  | ≥160              | ≥160              | ≥90                                   |
| <b>Office DBP treatment threshold (mmHg)</b> | ≥90                                   | ≥90        | ≥90   | ≥90               | ≥90               |                                       |

BP = blood pressure; CAD = coronary artery disease; CKD = chronic kidney disease; DBP = diastolic blood pressure; SBP = systolic blood pressure; TIA = transient ischaemic attack.

<sup>a</sup>Treatment may be considered in these very high-risk patients with high-normal SBP (i.e. SBP 130–140 mmHg).

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## Trattamento farmacologico e scelte terapeutiche

- I principali benefici dipendono non solo dalla riduzione dei valori di PA ma anche dal tipo di farmaco impiegato
- Le classi principali di farmaci antipertensivi: **diuretici, β-bloccanti, calcioantagonisti diidropiridinici, ACE-inibitori, bloccanti recettoriali dell'AT II** sono tutte indicate come scelte terapeutiche con cui iniziare e proseguire il trattamento
- Identificare una classe come prima scelta è di relativa importanza vista la frequente necessità di associare due o più farmaci
- I farmaci si differenziano però per alcuni specifici effetti farmacologici con conseguenti indicazioni differenti
- Esiste una diversa tollerabilità da paziente a paziente

## Trattamento farmacologico e scelte terapeutiche

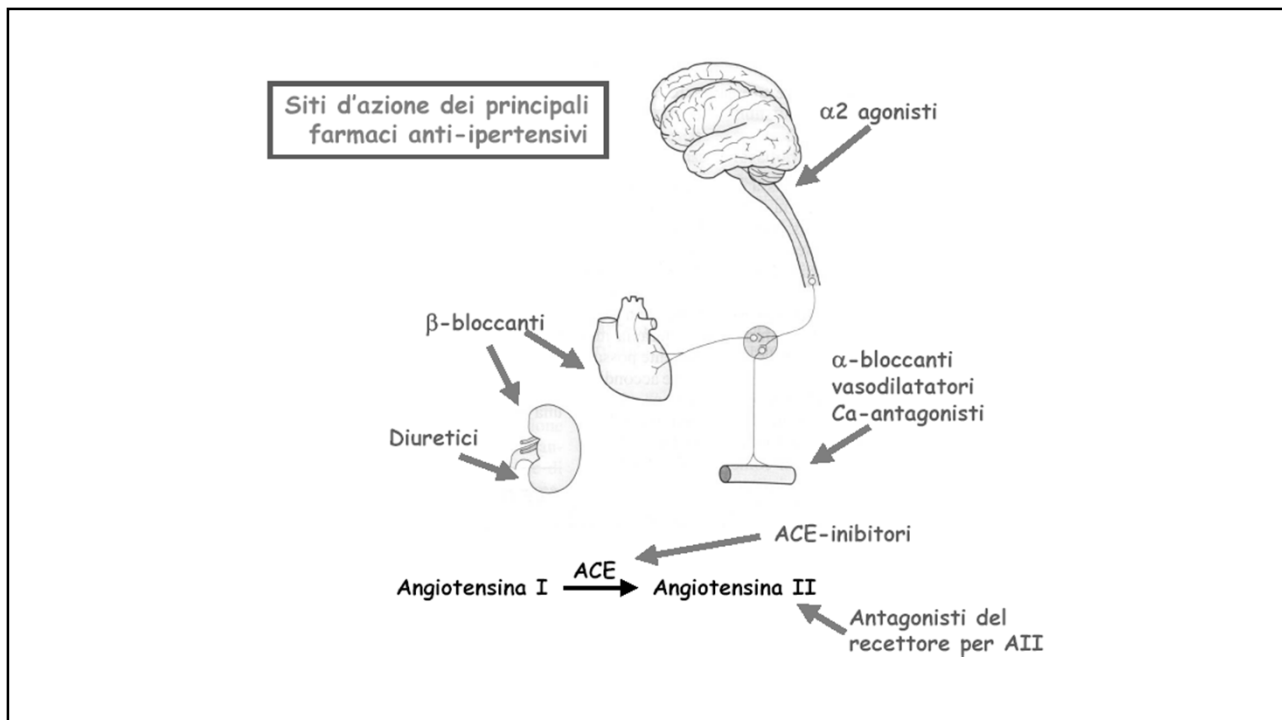
### Altri farmaci

- $\alpha$ -bloccanti
- farmaci che agiscono a livello del SNC (clonidina, metildopa)
- farmaci agenti sui recettori imidazolinici  $I_1$
- agonisti dei recettori  $\alpha_2$  adrenergici

Non sono consigliati come prima scelta

Consigliabile invece il loro utilizzo in terapia di associazione

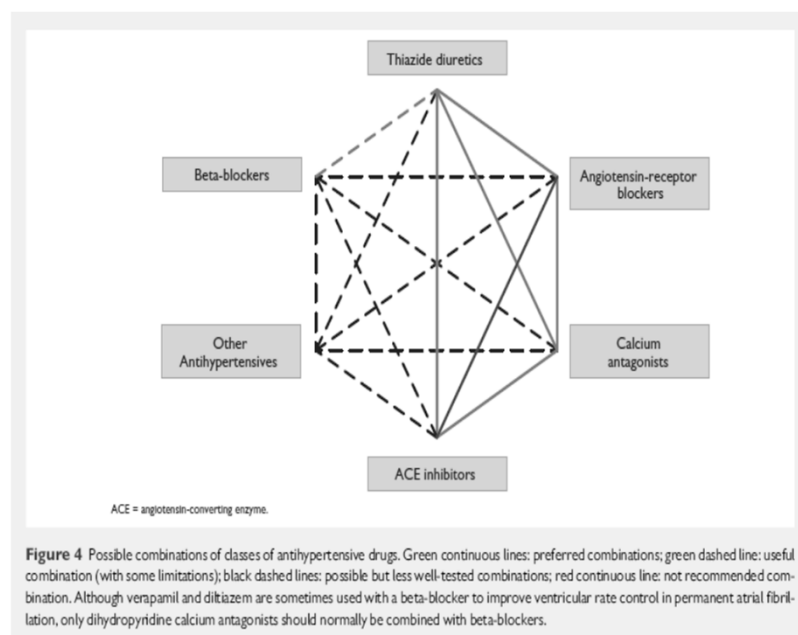
**Alcuni di questi hanno indicazioni ed usi specifici**



## Trattamento farmacologico e scelte terapeutiche

La scelta terapeutica può influenzata da numerosi fattori:

- I farmaci impiegati in passato;
- Il profilo di rischio del paziente;
- La presenza o meno di danno d'organo (TOD), diabete e complicanze (CCA);
- Il costo dei vari farmaci;
- L'eventuale preferenza espressa dal paziente;
- Effetti positivi o negativi del trattamento combinato con classi diverse





## Controindicazioni principali

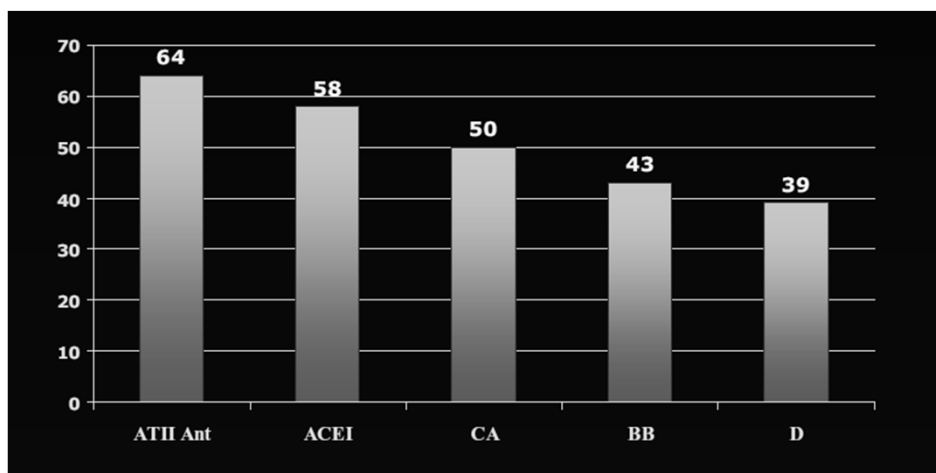
Table 20 Compelling and possible contraindications to the use of specific antihypertensive drugs

| Drug  | Contraindications  |  |
|---|--|--|
|   | Compelling   | Possible   |
| Diuretics (thiazides/thiazide-like, e.g. chlorthalidone and indapamide) | <ul style="list-style-type: none"> <li>Gout</li> </ul>   | <ul style="list-style-type: none"> <li>Metabolic syndrome</li> <li>Glucose intolerance</li> <li>Pregnancy</li> <li>Hypercalcaemia</li> <li>Hypokalaemia</li> </ul> |
| Beta-blockers   | <ul style="list-style-type: none"> <li>Asthma</li> <li>Any high-grade sinoatrial or atrioventricular block</li> <li>Bradycardia (heart rate &lt;60 beats per min)</li> </ul>   | <ul style="list-style-type: none"> <li>Metabolic syndrome</li> <li>Glucose intolerance</li> <li>Athletes and physically active patients</li> </ul>                 |
| Calcium antagonists (dihydropyridines)                                  |  | <ul style="list-style-type: none"> <li>Tachyarrhythmia</li> <li>Heart failure (HFrEF, class III or IV)</li> <li>Pre-existing severe leg oedema</li> </ul>          |
| Calcium antagonists (verapamil, diltiazem)                              | <ul style="list-style-type: none"> <li>Any high-grade sinoatrial or atrioventricular block</li> <li>Severe LV dysfunction (LV ejection fraction &lt;40%)</li> <li>Bradycardia (heart rate &lt;60 beats per min)</li> </ul> | <ul style="list-style-type: none"> <li>Constipation</li> </ul>   |
| ACE inhibitors  | <ul style="list-style-type: none"> <li>Pregnancy</li> <li>Previous angioneurotic oedema</li> <li>Hyperkalaemia (potassium &gt;5.5 mmol/L)</li> <li>Bilateral renal artery stenosis</li> </ul>                              | <ul style="list-style-type: none"> <li>Women of child-bearing potential without reliable contraception</li> </ul>  |
| ARBs  | <ul style="list-style-type: none"> <li>Pregnancy</li> <li>Hyperkalaemia (potassium &gt;5.5 mmol/L)</li> <li>Bilateral renal artery stenosis</li> </ul>   | <ul style="list-style-type: none"> <li>Women of child-bearing potential without reliable contraception</li> </ul>  |

ACE = angiotensin-converting enzyme; ARB = angiotensin receptor blocker; HFrEF = heart failure with reduced ejection fraction; LV = left ventricular.

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## Persistenza in terapia (a 1 anno)



## Ipertensione resistente al trattamento

Elevati valori basali di PA

Età avanzata

Obesità

Eccessivo introito di sale

Malattia renale cronica

Diabete

### AHA Scientific Statement

#### Resistant Hypertension: Diagnosis, Evaluation, and Treatment

##### A Scientific Statement From the American Heart Association Professional Education Committee of the Council for High Blood Pressure Research

David A. Calhoun, MD, FAHA, Chair; Daniel Jones, MD, FAHA; Stephen Textor, MD, FAHA; David C. Goff, MD, FAHA; Timothy P. Murphy, MD, FAHA; Robert D. Toto, MD, FAHA; Anthony White, PhD; William C.ushman, MD, FAHA; William White, MD; Domenic Sica, MD, FAHA; Keith Ferdinand, MD; Thomas D. Giles, MD; Bonita Falkner, MD, FAHA; Robert M. Carey, MD, MACP, FAHA

Chaloun DA et al, Hypertension 2008

## Ipertensione resistente al trattamento

Table 28 Medications and other substances that may increase blood pressure<sup>397</sup>

| Medication/substance                         |   |
|--|---|
| Oral contraceptive pill                      | Especially oestrogen containing; cause hypertension in ~5% of women, usually mild but can be severe   |
| Diet pills                                   | For example, phenylpropanolamine and sibutramine  |
| Nasal decongestants                          | For example, phenylephrine hydrochloride and naphazoline hydrochloride  |
| Stimulant drugs                              | Amphetamine, cocaine, and ecstasy; these substances usually cause acute rather than chronic hypertension  |
| Liquorice                                    | Chronic excessive liquorice use mimics hyperaldosteronism by stimulating the mineralocorticoid receptor and inhibiting cortisol metabolism                    |
| Immunosuppressive medications                | For example, cyclosporin A (tacrolimus has less effect on BP and rapamycin has almost no effect on BP) and steroids (e.g. corticosteroids and hydrocortisone) |
| Antiangiogenic cancer therapies              | Antiangiogenic drugs such as VEGF inhibitors (e.g. bevacizumab), tyrosine kinase inhibitors (e.g. sunitinib), and sorafenib have been reported to increase BP |
| Other drugs and substances that may raise BP | Anabolic steroids, erythropoietin, non-steroidal anti-inflammatory drugs, and herbal remedies (e.g. ephedra and ma huang)                                     |

BP = blood pressure; VEGF = vascular endothelial growth factor.

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## Condizioni particolari

8.9 Women, pregnancy, oral contraception, and hormone-replacement therapy

### Management of hypertension in pregnancy

| Recommendations   | Class <sup>a</sup> | Level <sup>b</sup>    |
|---|--------------------|-----------------------|
| In women with gestational hypertension, pre-existing hypertension superimposed by gestational hypertension, or with hypertension and subclinical organ damage or symptoms, initiation of drug treatment is recommended when SBP is $\geq 140$ mmHg or DBP $\geq 90$ mmHg. | I                  | C                     |
| In all other cases, initiation of drug treatment is recommended when SBP is $\geq 150$ mmHg or DBP is $\geq 95$ mmHg.   | I                  | C                     |
| Methyldopa, labetalol, and CCBs are recommended as the drugs of choice for the treatment of hypertension in pregnancy. <sup>407,418</sup>   | I                  | B (methyldopa)        |
|   | I                  | C (labetalol or CCBs) |
| ACE inhibitors, ARBs, or direct renin inhibitors are not recommended during pregnancy.  | III                | C                     |
| SBP $\geq 170$ mmHg or DBP $\geq 110$ mmHg in a pregnant woman is an emergency, and admission to hospital is recommended.   | I                  | C                     |
| In severe hypertension, drug treatment with i.v. labetalol, oral methyldopa, or nifedipine is recommended.  | I                  | C                     |
| The recommended treatment for hypertensive crisis is i.v. labetalol or nicardipine and magnesium.   | I                  | C                     |
| In pre-eclampsia associated with pulmonary oedema, nitroglycerin given as an i.v. infusion is recommended.  | I                  | C                     |
| In women with gestational hypertension or mild pre-eclampsia, delivery is recommended at 37 weeks. <sup>453</sup>   | I                  | B                     |
| It is recommended to expedite delivery in pre-eclampsia with adverse conditions, such as visual disturbances or haemostatic disorders.  | I                  | C                     |

ACE = angiotensin-converting enzyme; ARB = angiotensin receptor blocker; CCB = calcium channel blocker; DBP = diastolic blood pressure; i.v. = intravenous; SBP = systolic blood pressure.  
<sup>a</sup>Class of recommendation.  
<sup>b</sup>Level of evidence.

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## Percentuale di controllo della PA (pz in trattamento)

