

# Hypertension & Dyslipidimia in the Spotlight: Bad fellows in CV Health

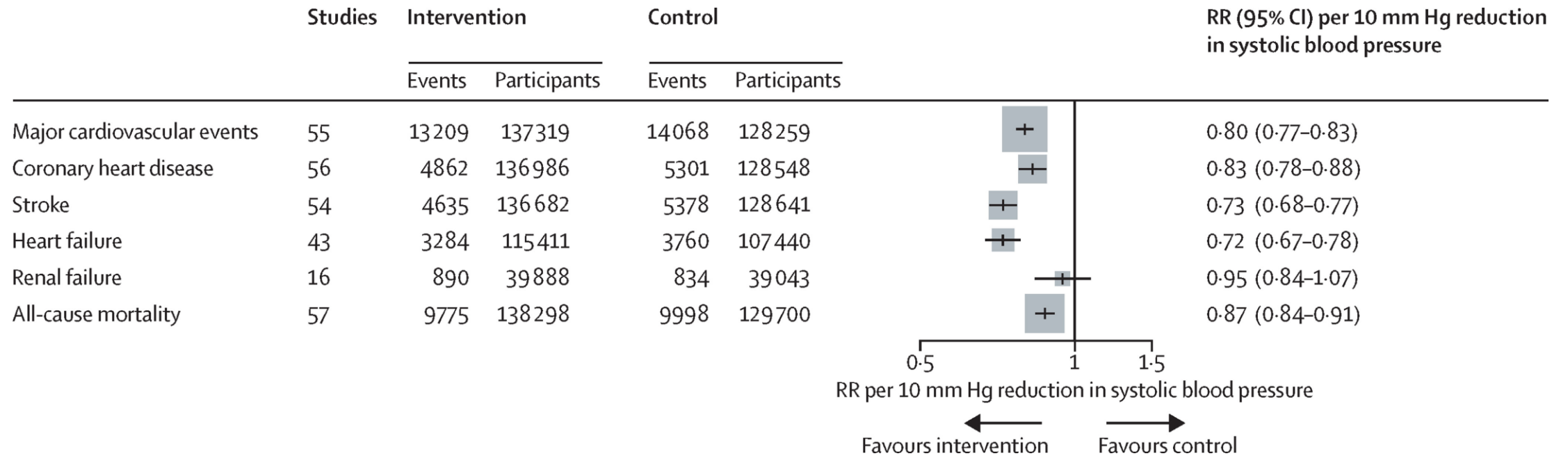
- Introduction
- Update in Guideline recommendations
- Clinical cases

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# Introduction

- Cardiovascular disease (CVD) accounts for >4 million deaths / year in Europe
- Uncontrolled hypertension is responsible for 9.4 million documented deaths worldwide
- Importance of ASCVD prevention remains undisputed
- More patients are surviving their first CVD event and are at high-risk of recurrences
- Hypertension & Dyslipidemia are important CV risk factors that plays a key role in ASCVD

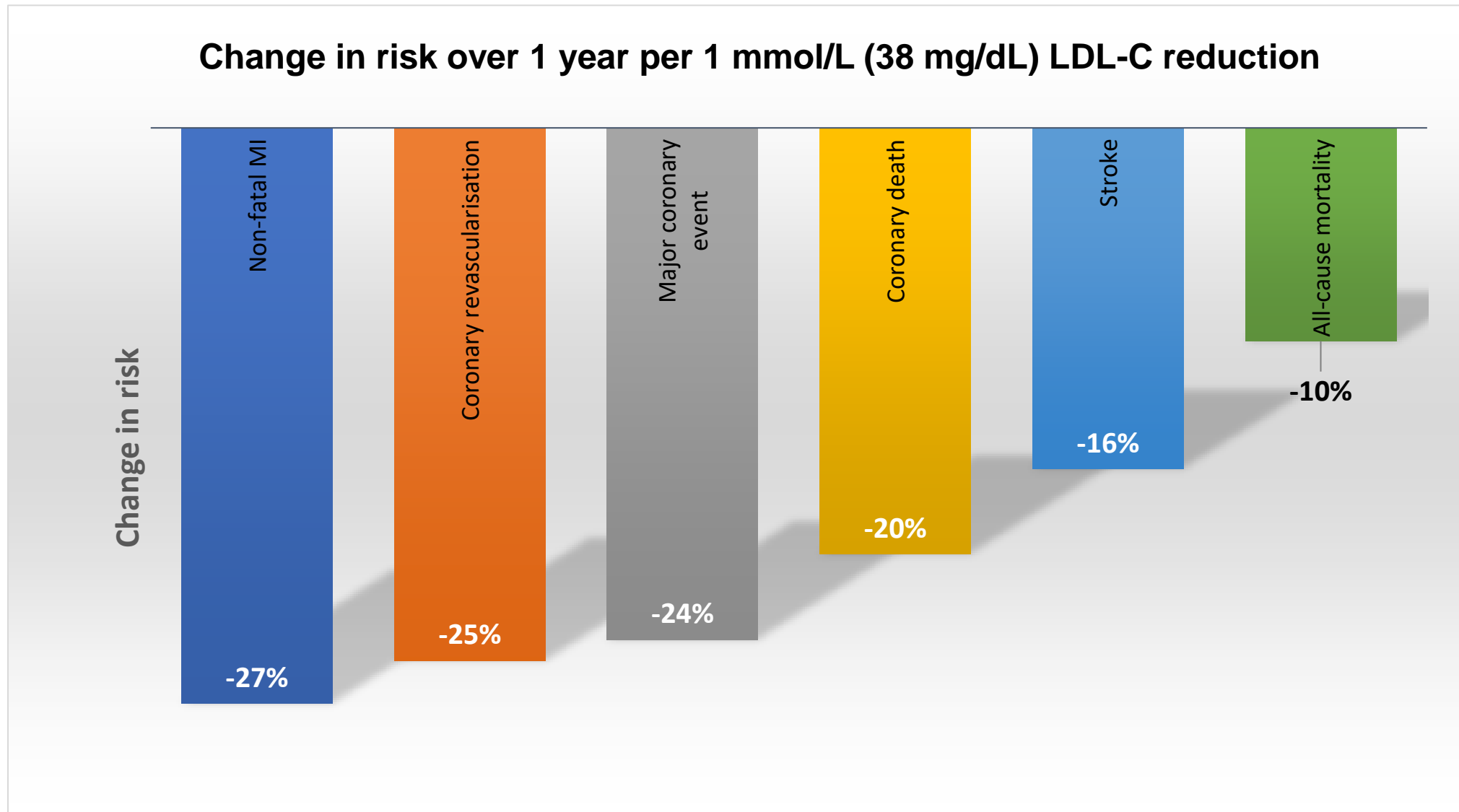
# Benefits of Blood Pressure Lowering are well documented



Reference: Ettehad, D., et al., Blood pressure lowering for prevention of cardiovascular disease and death: a systematic review and meta-analysis.

*The Lancet*, 387(10022), 957-967. doi:10.1016/S0140-6736(15)01225-8

# Benefits of Intensive Lipid Lowering Therapy are well documented



CV, cardiovascular; LDL-C, low-density lipoprotein cholesterol; MI, myocardial infarction.

Adapted from Cholesterol Treatment Trialists' (CTT) Collaboration. *Lancet* 2010;376:1670-81.

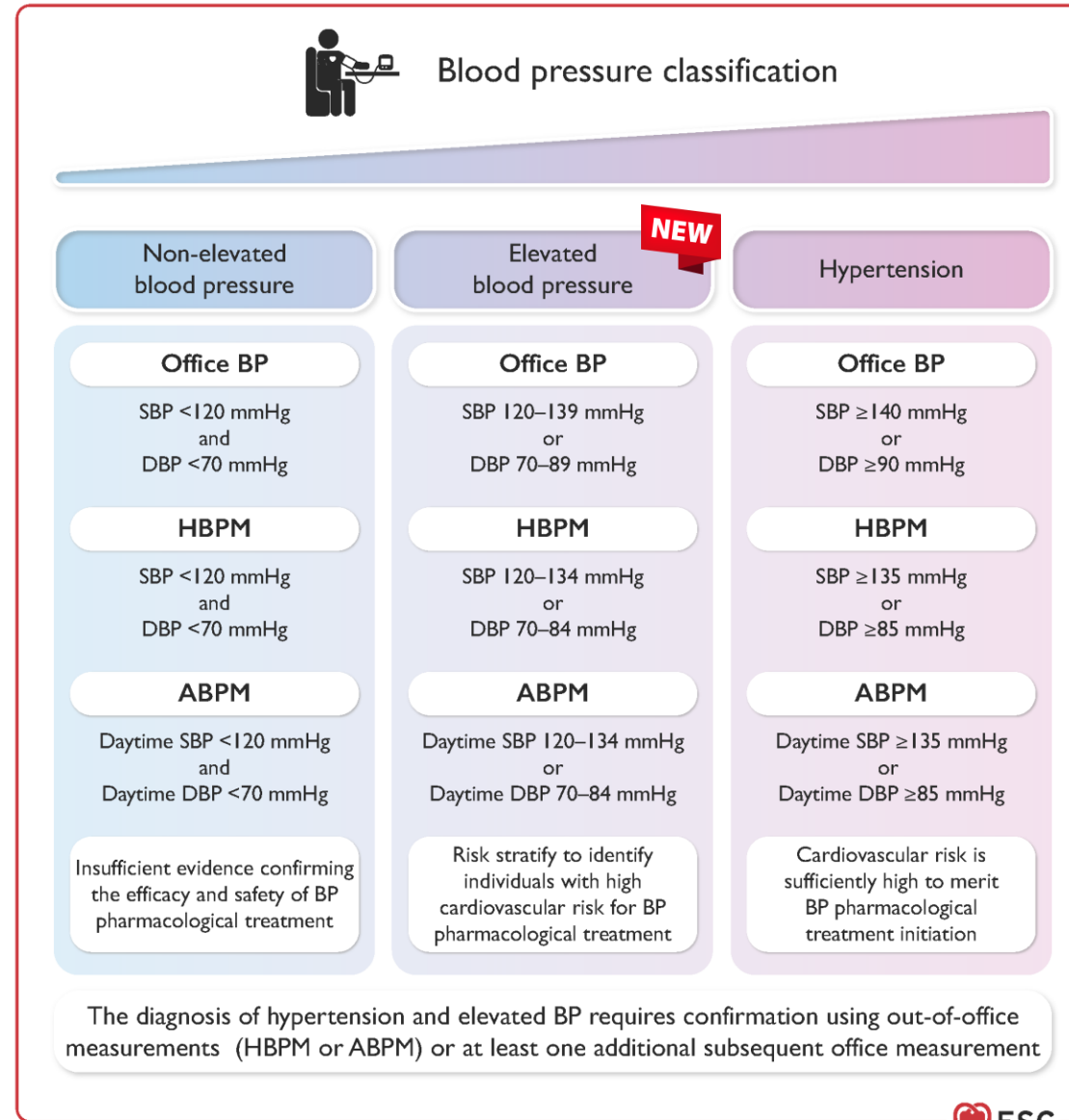
# Revised recommendations (1)

2018 Guidelines	Class	Level	2024 Guidelines	Class	Level
<b><i>Definition and classification of elevated blood pressure and hypertension</i></b>					
It is recommended that BP be classified as optimal, normal, high-normal, or grades 1–3 hypertension, according to office BP.	<b>I</b>	<b>C</b>	It is recommended that BP be categorized as non-elevated BP, elevated BP, and hypertension to aid treatment decisions.	<b>I</b>	<b>B</b>
CV risk assessment with the SCORE system is recommended for hypertensive patients who are not already at high or very high risk due to established CVD, renal disease, or diabetes, a markedly elevated single risk factor (e.g. cholesterol), or hypertensive LVH.	<b>I</b>	<b>B</b>	SCORE2 is recommended for assessing 10-year risk of fatal and non-fatal CVD among individuals aged 40–69 years with elevated BP who are not already considered at increased risk due to moderate or severe CKD, established CVD, HMOD, diabetes mellitus, or familial hypercholesterolaemia.	<b>I</b>	<b>B</b>

# Blood pressure categories

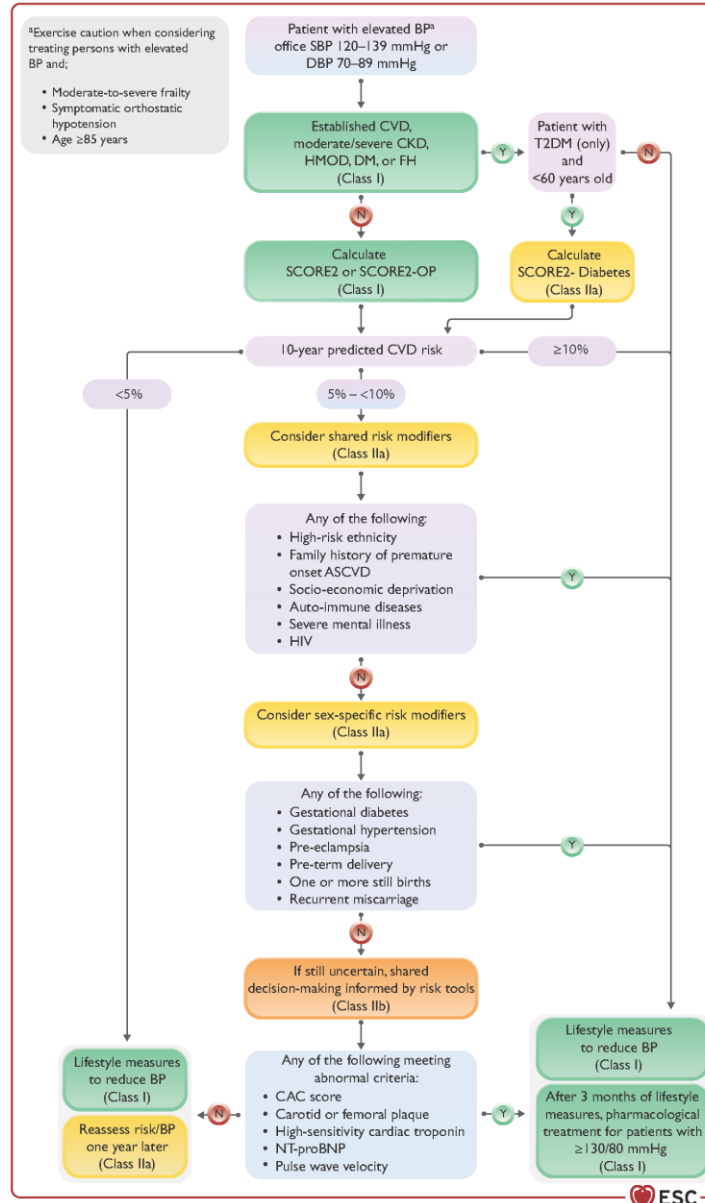
**NEW**

- Simplified Blood pressure categories
- New group of patients with elevated bp
- No hypertension grades defined
- For “Elevated bp”: risk stratify to identify high CV risk for BP pharmacological treatment



**NEW**

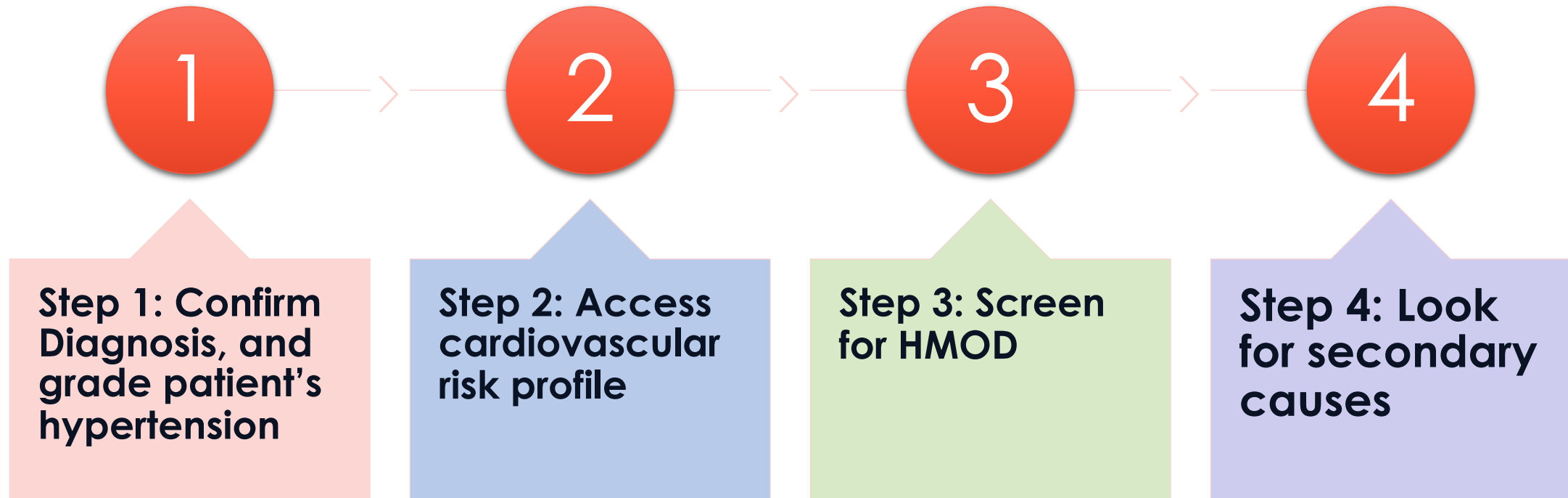
# Summary of cardiovascular disease risk-stratification approach for blood pressure treatment in adults with elevated blood pressure



ESC 2024 guidelines introduced **Cardiovascular disease risk-stratification approach in patients with elevated BP**

**ASCVD:** atherosclerotic cardiovascular disease. **BP:** blood pressure. **CAC:** coronary artery calcium. **CV:** cardiovascular. **CVD:** cardiovascular disease. **CKD:** chronic kidney disease. **DM:** diabetes mellitus. **HMOD:** hypertension mediated organ damage. **HIV:** human immunodeficiency virus. **NT-proBNP:** N-terminal pro-brain natriuretic peptide. **SCORE-2/2-OP:** Systematic COronary Risk Evaluation 2/Older Persons

# Key steps in management of new hypertension patients





## Step 1: Confirm Diagnosis

- 📌 Repeat BP in clinic (over a few days)
- 📌 Home BP monitoring (patient keeps a diary)
- 📌 Ambulatory 24-hour BP measurement

## Step 2: Assess Cardiovascular risk profile

📌 Lipid profile (cholesterol levels)

📌 HbA1c (diabetes screening)

📌 BMI measurement

📌 Smoking history & Alcohol intake

📌 Unhealthy diet assessment

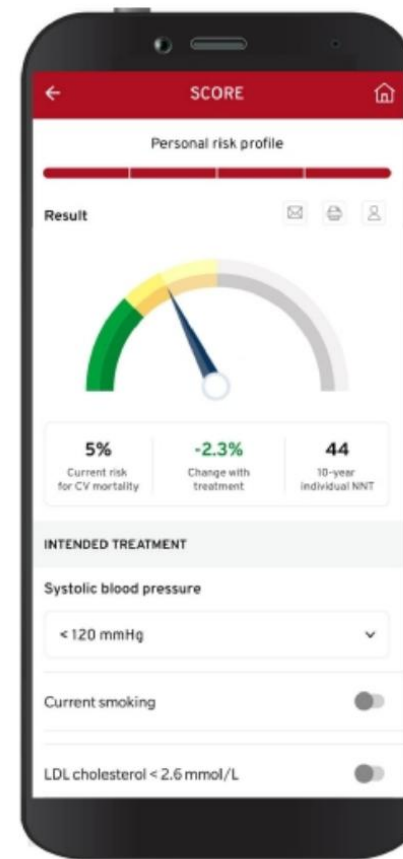
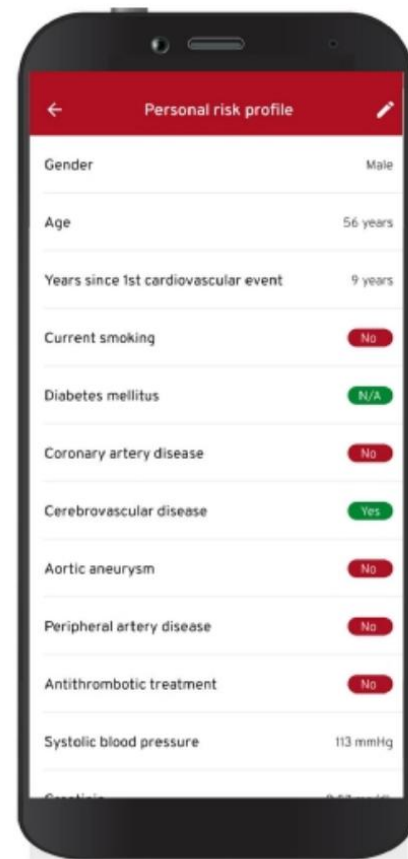
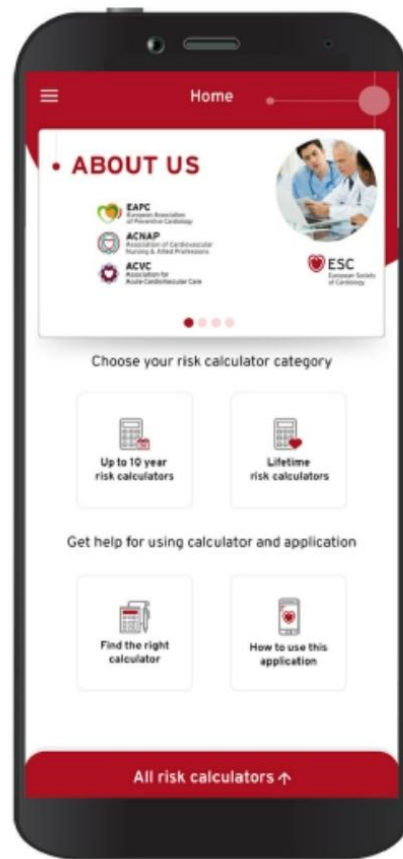
📌 Salt intake

📌 Sedentary lifestyle

📌 Stress levels

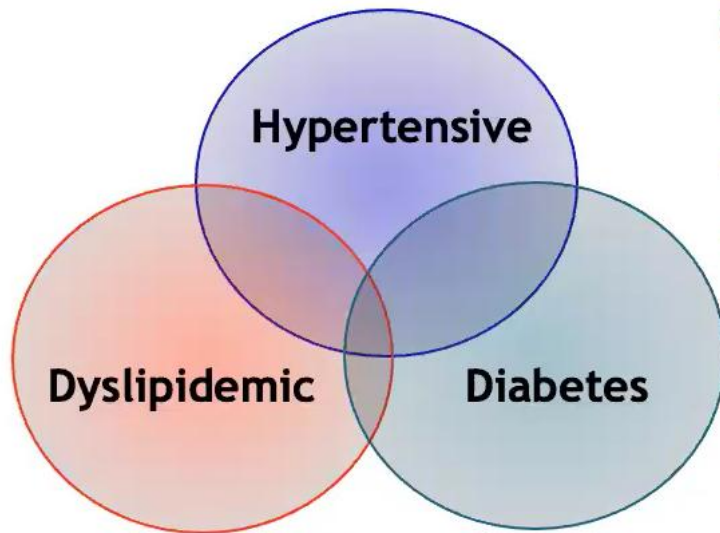
# CV Risk Calculators

- SCORE2
- SCORE2-OP
- SCORE2-Diabetes
- ASCVD
- ADVANCE
- SMART
- SMART-REACH\*
- DIAL\*
- LIFE-CVD\*



# Most patients have overlapping CV risk factors

Multiple comorbidities increase risk 400-700%



Of all people with hypertension:

- 65% have dyslipidemia
- 16% have type 2 diabetes
- 45% are overweight/obese



Of all people with dyslipidemia:

- 48% have hypertension
- 14% have type 2 diabetes
- 35% are overweight/obese

Of all people with type 2 diabetes:

- 60% have hypertension
- 60% have dyslipidemia
- 90% are overweight/obese

## Step 3: Check for Hypertension-Mediated Organ Damage (HMOD)

 heart assessment - ECG +/- Echo







 kidney function - Urinalysis, Cr, ACR

 eye damage check - Fundoscopy

## Step 4: Look for Secondary Causes of Hypertension



### Consider further investigations in patients with:

-  **Resistant hypertension** (uncontrolled on 3+ medications)
-  **Endocrine features** (e.g., Cushing's, pheochromocytoma signs)
-  **Young onset hypertension** (<40 years)
-  **Hypokalemia or metabolic alkalosis** (suggesting hyperaldosteronism)
-  **Renal bruit** (possible renovascular hypertension)
-  **Obstructive sleep apnea symptoms** (snoring, daytime fatigue)

# 2019 ESC/EAS Dyslipidemia Classification

<b>Very high risk</b>	<p>People with any of the following:</p> <ul style="list-style-type: none"><li>• Documented ASCVD, either clinical or unequivocal on imaging</li><li>• DM with target organ damage, or at least three major risk factors, or early onset of T1DM of long duration (&gt; 20 years)</li><li>• Severe CKD (eGFR &lt; 30 mL/min/1.73 m<sup>2</sup>)</li><li>• A calculated SCORE ≥ 10% for 10-year risk of fatal CVD</li><li>• FH with ASCVD or with another major risk factor</li></ul>
<b>High risk</b>	<p>People with:</p> <ul style="list-style-type: none"><li>• Markedly elevated single risk factors, in particular TC &gt; 8 mmol/L (310 mg/dL), LDL-C &gt; 4.9 mmol/L (190 mg/dL), or BP ≥ 180/110 mmHg</li><li>• Patients with FH without other major risk factors</li><li>• Patients with DM without target organ damage, with DM duration ≥ 10 years or another additional risk factor</li><li>• Moderate CKD (eGFR 30-59 mL/min/1.73 m<sup>2</sup>)</li><li>• A calculated SCORE ≥ 5% and &lt; 10% for 10-year risk of fatal CVD</li></ul>
<b>Moderate risk</b>	<ul style="list-style-type: none"><li>• Young patients (T1DM &lt; 35 years; T2DM &lt; 50 years) with DM duration &lt; 10 years, without other risk factors</li><li>• Calculated SCORE ≥ 1% and &lt; 5% for 10-year risk of fatal CVD</li></ul>
<b>Low-risk</b>	<ul style="list-style-type: none"><li>• Calculated SCORE &lt; 1% for 10-year risk of fatal CVD</li></ul>

# Clinical Cases



# Clinical Case – pt AT












- 52 year old
- Episodes of atypical chest pain which precipitated hospital review
- Represented with atypical chest pain
- Strong FH of MI
- Father 56 year old died of MI
- PMH Nil
- DH Nil
- Ex-smoker
- Non drinker
- Office bp: 148/87mmHg, HR 79/bpm
- LDL-c 5.13mmol/L
- Negative EST

# Clinical Case – pt AT

- Exercise stress test

EST carried out according to BRUCE protocol 90% of target heart rate reached during stage 3. No symptoms reported. No significant ST changes. No arrhythmias. Normal BP response.

# Tests and criteria for defining hypertension-mediated target organ damage and considerations for their use in clinical practice

Why measure?	Which organ?	What to measure?	How to diagnose HMOD?
 <p>Support decision to start or intensify BP-lowering treatment for:</p> <ul style="list-style-type: none"> <li>Individuals with elevated BP with SCORE2/SCORE2-OP risk of 5–&lt;10%</li> <li>Uncertain situations (i.e. BP or risk close to thresholds, masked or white-coat hypertension, non-traditional CVD risk factors)</li> <li>Individuals &lt;40 years old with elevated blood pressure</li> <li>Assistance overcoming patient and physician inertia</li> </ul>	<b>Kidney</b> 	 <p>eGFR ACR</p>	<b>Moderate-to-severe kidney disease</b> <ul style="list-style-type: none"> <li>eGFR &lt;60 mL/min/1.73 m<sup>2</sup> irrespective of albuminuria</li> <li>Albuminuria ≥30 mg/g irrespective of eGFR</li> </ul>
	<b>Heart</b> 	 <p>ECG</p>	<b>LVH</b> <ul style="list-style-type: none"> <li>Sokolow–Lyon: SV1+RV5 &gt;35 mm</li> <li>RaVL ≥11 mm</li> <li>Cornell voltage: SV3+RaVL&gt;28 mm (men) SV3+RaVL&gt;20 mm (women)</li> </ul>
		 <p>Echocardiography</p>	<b>LVH</b> <ul style="list-style-type: none"> <li>LV mass/height<sup>2.7</sup>(g/m<sup>2.7</sup>): &gt;50 (men) &gt;47 (women)</li> <li>LV mass/BSA(g/m<sup>2</sup>): &gt;115 (men) &gt;95 (women)</li> <li>LV concentric geometry: RWT ≥0.43</li> </ul>
		 <p>Cardiac biomarkers</p>	<b>Diastolic dysfunction</b> <ul style="list-style-type: none"> <li>LA volume/height<sup>2</sup> (mL/m<sup>2</sup>): &gt;18.5 (men) &gt;16.5 (women)</li> <li>LA volume index (mL/m<sup>2</sup>): 34</li> <li>e' &lt;7cm; E/e' &gt;14</li> </ul>
	<b>Arteries</b> 	 <p>Carotid or femoral ultrasound</p>	<b>Plaque (focal wall thickening &gt;1.5 mm)</b>
		 <p>Pulse wave velocity</p>	<ul style="list-style-type: none"> <li>Carotid-femoral PWV &gt;10 m/s</li> <li>Brachial-ankle PWV &gt;14 m/s</li> </ul>
 <p>Cardiac CT</p>		<b>Coronary artery calcium score &gt;100 Agatston units</b>	

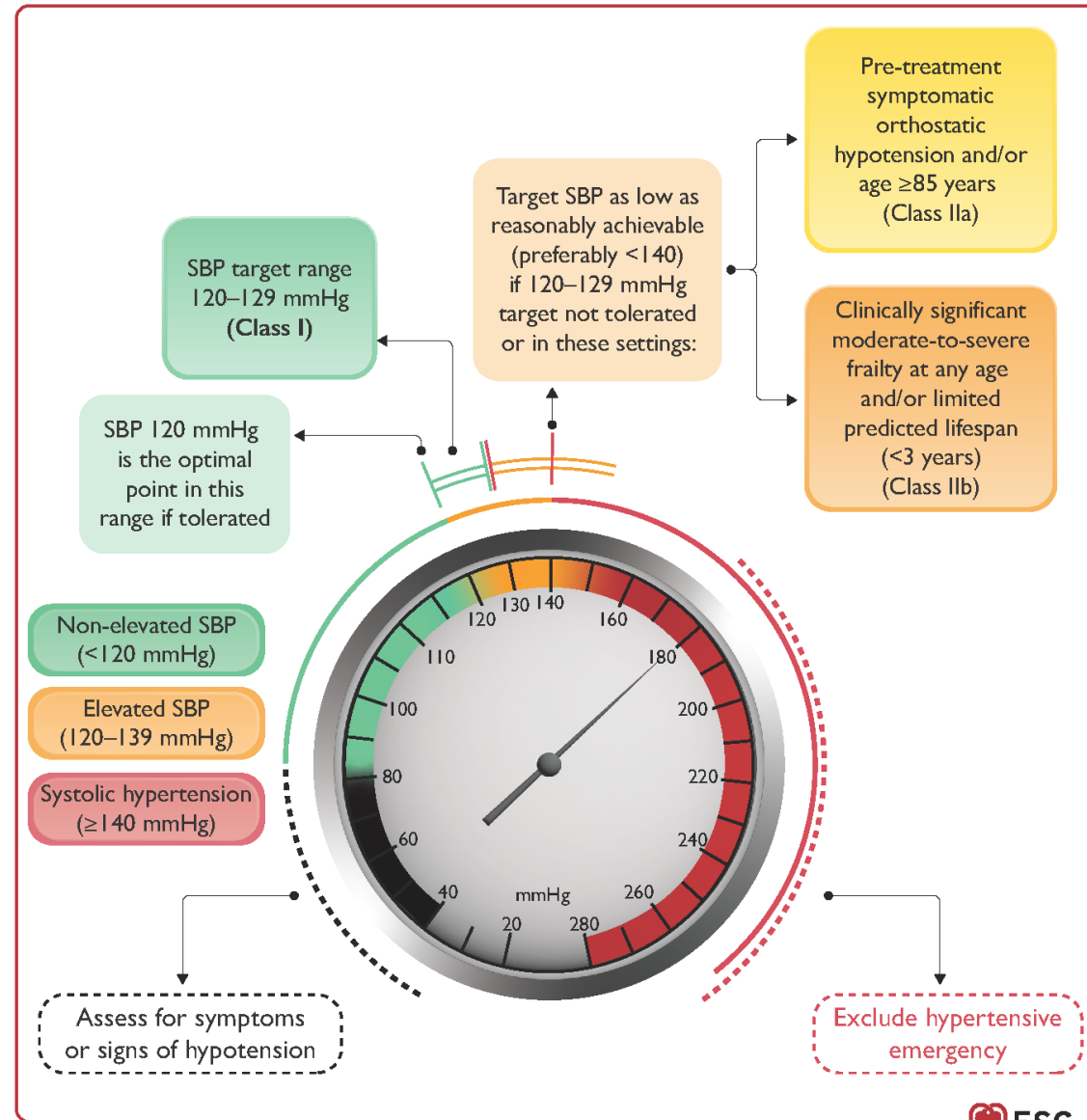
# Revised recommendations (11)

2018 Guidelines	Class	Level	2024 Guidelines	Class	Level
<b><i>Preventing and treating elevated blood pressure (blood pressure targets)</i></b>					
It is recommended that the first objective of treatment should be to lower BP to <140/90 mmHg in all patients and, provided that the treatment is well tolerated, treated BP values should be targeted to 130/80 mmHg or lower in most patients.	<b>I</b>	<b>A</b>	To reduce CVD risk, it is recommended that treated systolic BP values in most adults be targeted to <b>120–129 mmHg</b> , provided the treatment is well tolerated.	<b>I</b>	<b>A</b>
A diastolic BP target of <80 mmHg should be considered for all hypertensive patients, independent of the level of risk and comorbidities.	<b>IIa</b>	<b>B</b>	In cases where on-treatment systolic BP is at or below target (120–129 mmHg) but diastolic BP is not at target ( $\geq 80$ mmHg), intensifying BP-lowering treatment to achieve an on-treatment diastolic BP of <b>70–79 mmHg</b> may be considered to reduce CVD risk.	<b>IIb</b>	<b>C</b>

# Revised recommendations (12)

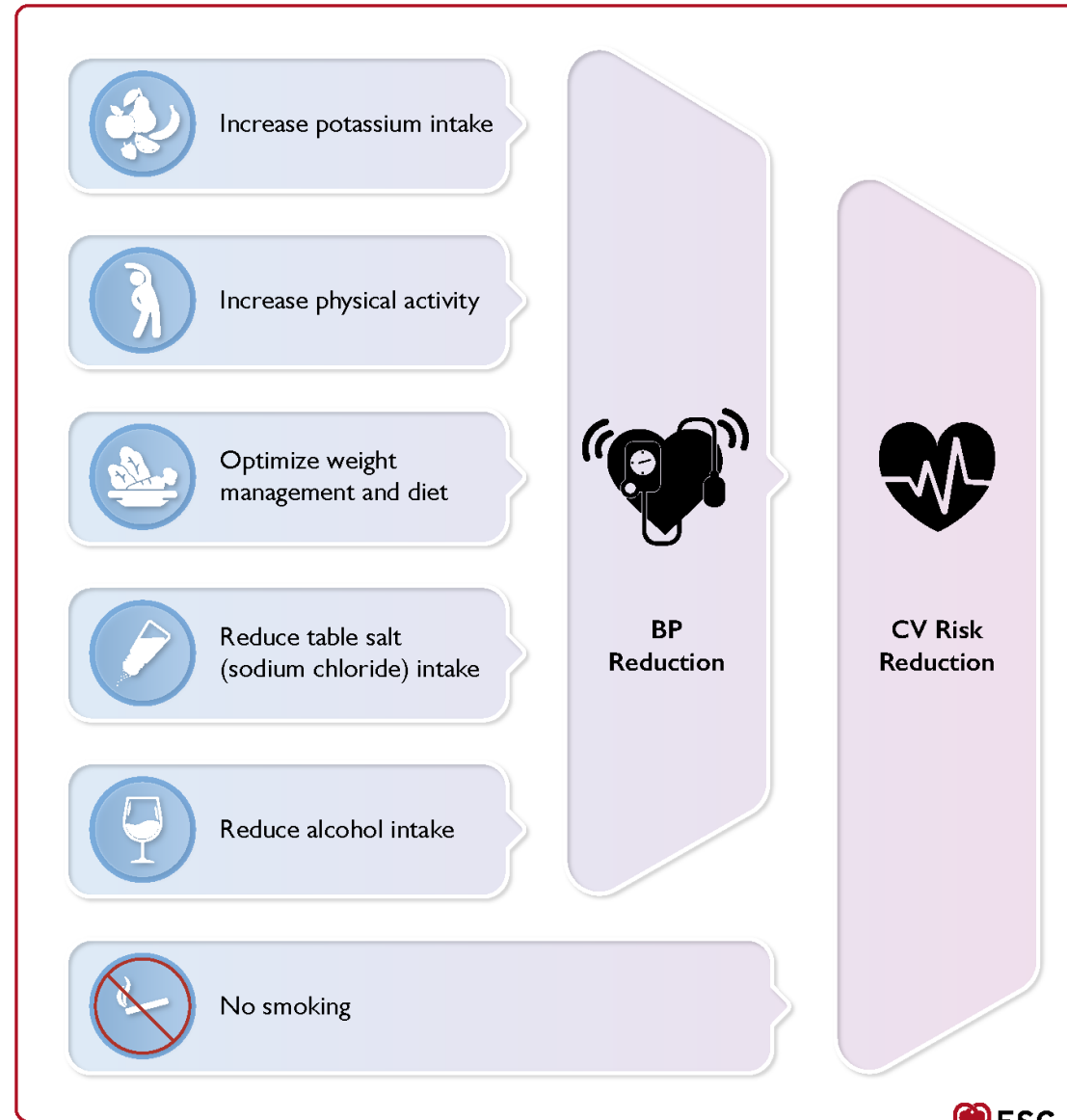
2018 Guidelines	Class	Level	2024 Guidelines	Class	Level
<b><i>Preventing and treating elevated blood pressure (blood pressure targets) cont.</i></b>					
<p>In older patients (aged <math>\geq 65</math> years) receiving BP-lowering drugs:</p> <ul style="list-style-type: none"> <li>It is recommended that systolic BP should be targeted to a BP range of 130–139 mmHg.</li> </ul>	<b>I</b>	<b>A</b>	<p>Because the CVD benefit of an on-treatment systolic BP target of 120–129 mmHg may not generalize to the following specific settings, personalized and more lenient systolic BP targets (e.g. <math>&lt; 140</math> mmHg): should be considered among patients meeting the following criteria:</p> <ul style="list-style-type: none"> <li>pre-treatment, symptomatic, orthostatic hypotension;</li> <li>and/or age <math>\geq 85</math> years.</li> </ul>	<b>IIa</b>	<b>C</b>

# Systolic blood pressure categories and treatment target range



## Effects of main lifestyle factors on blood pressure and cardiovascular risk reduction.

- Increased K<sup>+</sup> intake, higher physical activity, optimized weight management: reduce BP and are associated with lower overall CV risk (short arrows).
- Salt reduction reduces BP & (for persons with high baseline intake) reduces cardiovascular risk.
- Smoking cessation reduces overall cardiovascular risk but not BP (long arrow).



# Recommendations for pharmacological treatment of hypertension (1)

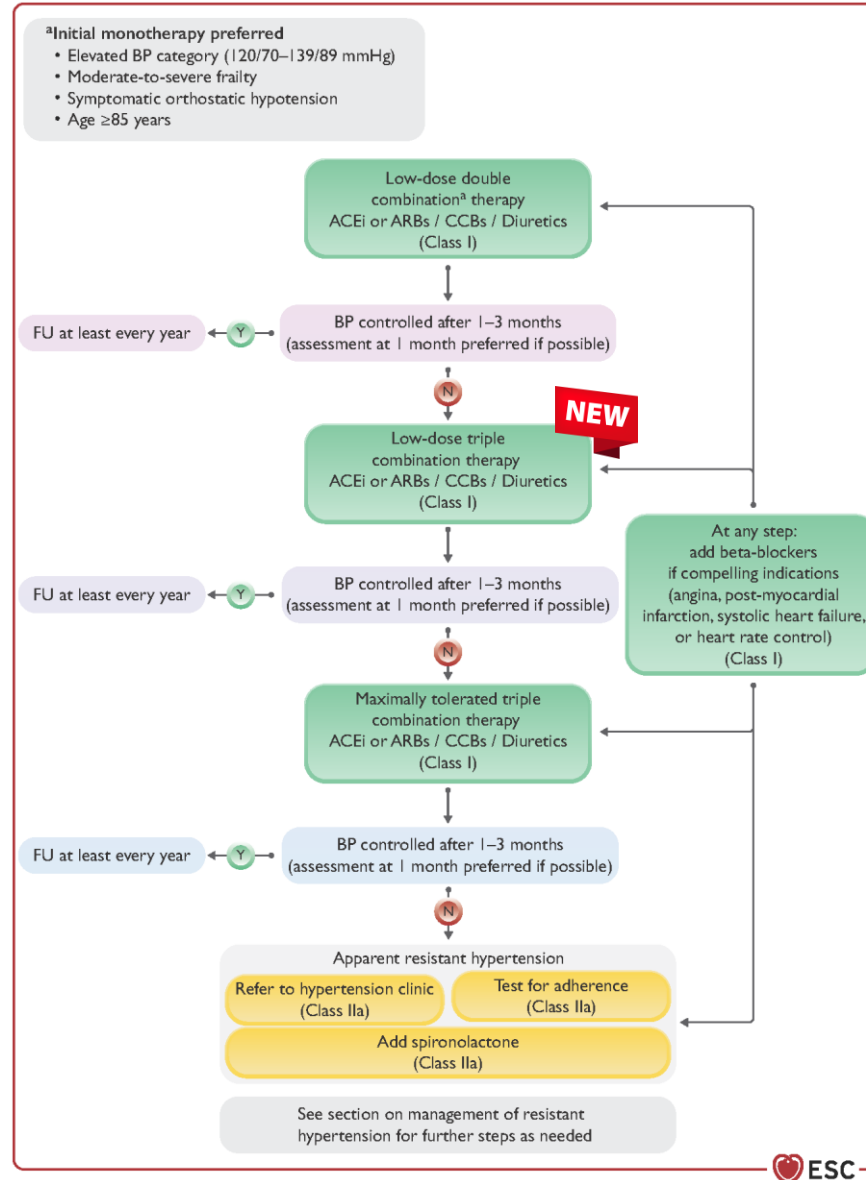
Recommendations	Class	Level
Among all BP-lowering drugs, ACE inhibitors, ARBs, dihydropyridine CCBs, and diuretics (thiazides and thiazide-like drugs such as chlorthalidone and indapamide) have demonstrated the most effective reduction of BP and CVD events, and are therefore recommended as first-line treatments to lower BP.	I	A
It is recommended that beta-blockers are combined with any of the other major BP-lowering drug classes when there are other compelling indications for their use, e.g. angina, post-myocardial infarction, heart failure with reduced ejection fraction, or for heart rate control.	I	A
It is recommended to take medications at the <u>most convenient time of day</u> for the patient to establish a habitual pattern of medication taking to improve adherence.	I	B
In patients receiving combination BP-lowering treatment, <u>fixed-dose single-pill</u> combination treatment is recommended.	I	B



# Recommendations for pharmacological treatment of hypertension (2)

Recommendations cont.	Class	Level
Given trial evidence for more effective BP control vs. monotherapy, <u>combination</u> BP-lowering treatment is recommended for most patients with confirmed hypertension (BP $\geq 140/90$ mmHg) as initial therapy. Preferred combinations are a RAS blocker (either an ACE inhibitor or an ARB) with a dihydropyridine CCB or diuretic. Exceptions to consider include patients aged $\geq 85$ years, those with symptomatic orthostatic hypotension, moderate-to-severe frailty, or elevated BP (systolic BP 120–139 mmHg or diastolic BP 70–89 mmHg) with a concomitant indication for treatment.	I	B
If BP is not controlled with a two-drug combination, increasing to a three-drug combination is recommended, usually a RAS blocker with a dihydropyridine CCB and a thiazide/thiazide-like diuretic, and preferably in a single-pill combination.	I	B
If BP is not controlled with a three-drug combination, adding spironolactone should be considered.	IIa	B

# Practical algorithm for pharmacological blood pressure lowering



← Treatment initiation with Dual combination

← Earlier use of Triple combination

# 2019 ESC/EAS Guidelines

LDL < 1.0 mmol/ L VERY VERY High Risk

LDL < 1.4 mmol/ L

**Very high risk**

People with any of the following:

- Documented ASCVD, either clinical or unequivocal on imaging
- DM with target organ damage, or at least three major risk factors, or early onset of T1DM of long duration (> 20 years)
- Severe CKD (eGFR < 30 mL/min/1.73 m<sup>2</sup>)
- A calculated SCORE ≥ 10% for 10-year risk of fatal CVD
- FH with ASCVD or with another major risk factor

LDL < 1.8 mmol/ L

**High risk**

People with:

- Markedly elevated single risk factors, in particular TC > 8 mmol/L (310 mg/dL), LDL-C > 4.9 mmol/L (190 mg/dL), or BP ≥ 180/110 mmHg
- Patients with FH without other major risk factors
- Patients with DM without target organ damage, with DM duration ≥ 10 years or another additional risk factor
- Moderate CKD (eGFR 30-59 mL/min/1.73 m<sup>2</sup>)
- A calculated SCORE ≥ 5% and < 10% for 10-year risk of fatal CVD

- Father 56 year old died of MI  
- LDL-c 5.13mmol/L

LDL < 2.6 mmol/ L

**Moderate risk**

- Young patients (T1DM < 35 years; T2DM < 50 years) with DM duration < 10 years, without other risk factors
- Calculated SCORE ≥ 1% and < 5% for 10-year risk of fatal CVD

LDL < 3 mmol/ L

**Low-risk**

- Calculated SCORE < 1% for 10-year risk of fatal CVD

ASCVD, atherosclerotic cardiovascular disease; BP, blood pressure; CKD, chronic kidney disease; CVD, cardiovascular disease; DM, diabetes mellitus; eGFR, estimated glomerular filtration rate; FH, familial hypercholesterolaemia; LDL-C, low-density lipoprotein cholesterol; SCORE, Systematic Coronary Risk Estimation; T1DM, type 1 diabetes mellitus; T2DM, type 2 diabetes mellitus; TC, total cholesterol. Adapted from Mach F, et al. *Eur Heart J* 2020;41(1):111-88.

### Intensity of lipid-lowering treatment

Treatment	Average LDL-C reduction
Moderate-intensity statin	≈ 30%
High-intensity statin	≈ 50%
High-intensity statin plus ezetimibe	≈ 65%
PCSK9 inhibitor	≈ 60%
PCSK9 inhibitor plus high-intensity statin	≈ 75%
PCSK9 inhibitor plus high-intensity statin plus ezetimibe	≈ 85%



European Heart Journal (2021) **42**, 3227 – 3337  
doi:10.1093/eurheartj/ehab484

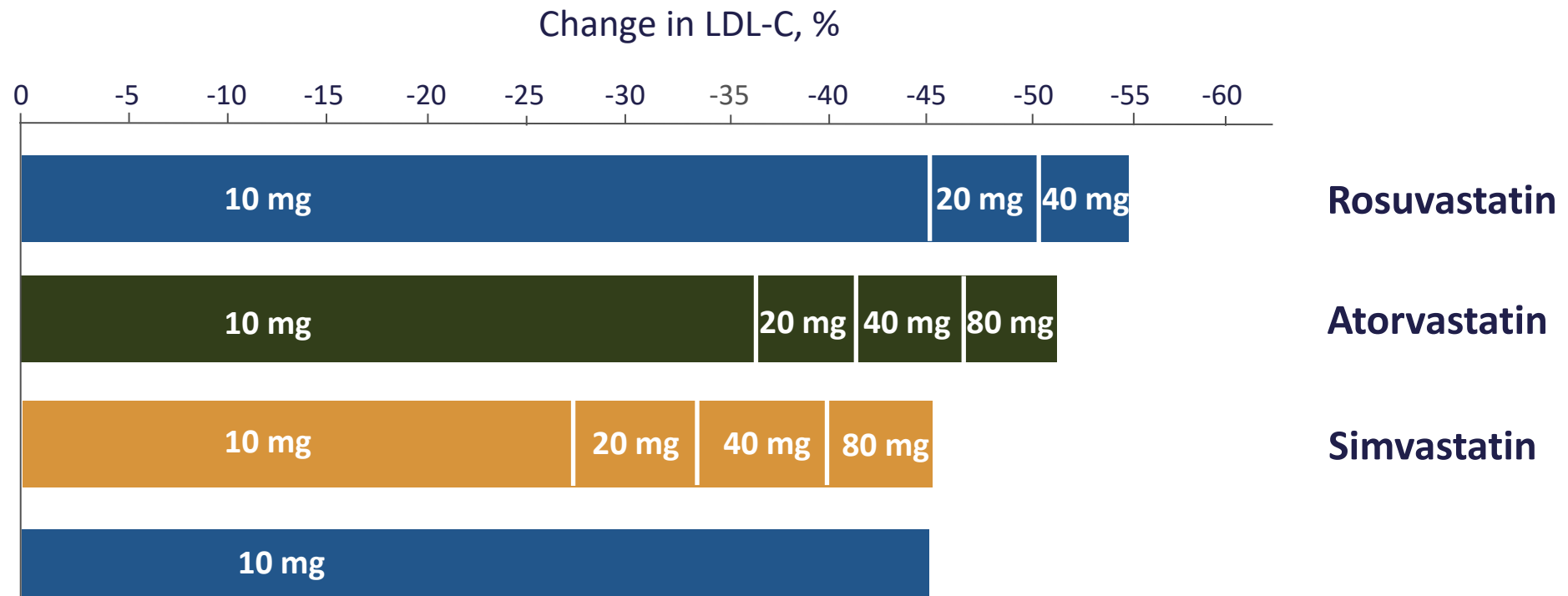
ESC GUIDELINES

## 2021 ESC Guidelines on cardiovascular disease prevention in clinical practice

# Statin intensity

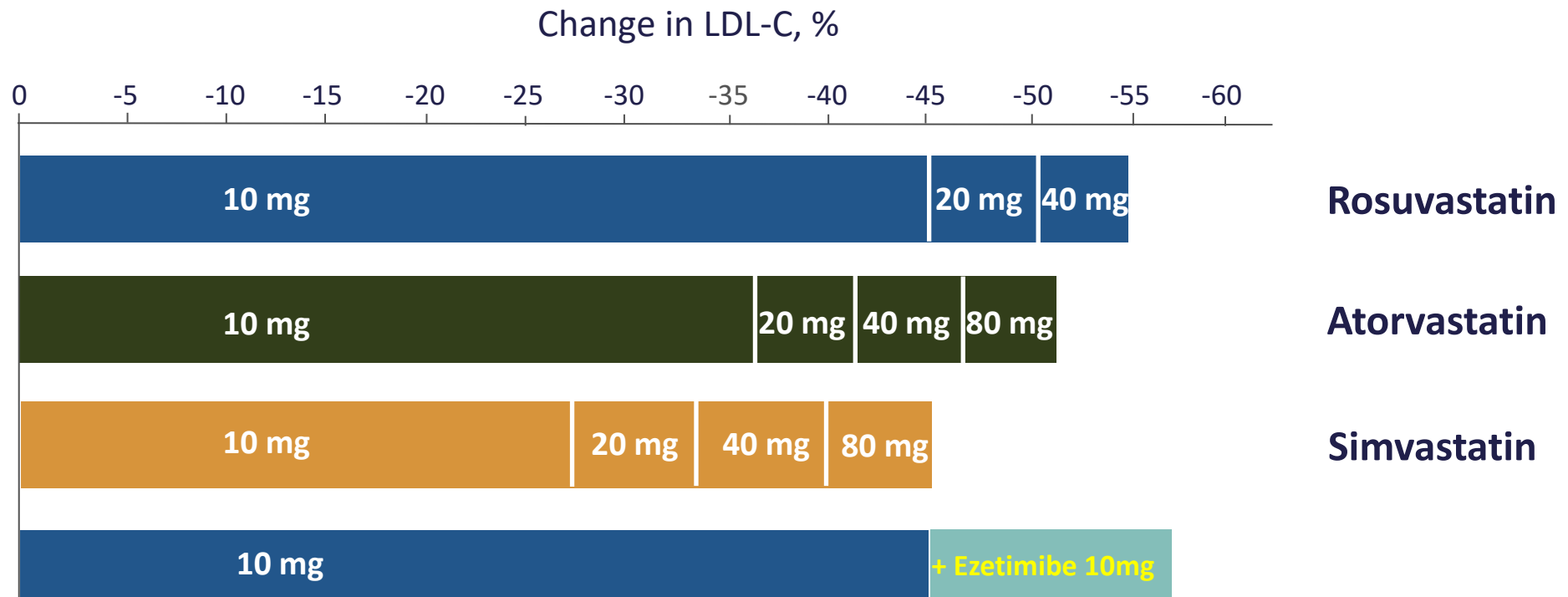
High-Intensity Statin Therapy	Moderate-Intensity Statin Therapy	Low-Intensity Statin Therapy
Daily dose lowers LDL on average by $\geq 50\%$	Daily dose lowers LDL on average by approximately 30-49%	Daily dose lowers LDL on average by $< 30\%$
Atorvastatin 40-80 mg Rosuvastatin 20-40 mg	Atorvastatin 10-20 mg Rosuvastatin 5-10 mg Simvastatin 20-40 mg Pravastatin 40-80 mg Lovastatin 40 mg Fluvastatin XL 80 mg Fluvastatin 40 mg BID Pitavastatin 2-4 mg	Simvastatin 10 mg Pravastatin 10-20 mg Lovastatin 20 mg Fluvastatin 20-40 mg

# Doubling Statin Dose will only Achieve ~6% Additional LDL-C Reduction



LDL-C, low-density lipoprotein cholesterol. Adapted from "FDA drug safety communication: New restrictions, contraindications, and dose limitations for simvastatin to reduce the risk of muscle injury." US Food & Drug Administration website. Accessed June 2020.

# Doubling Statin Dose will only Achieve ~6% Additional LDL-C Reduction



LDL-C, low-density lipoprotein cholesterol. Adapted from “FDA drug safety communication: New restrictions, contraindications, and dose limitations for simvastatin to reduce the risk of muscle injury.” US Food & Drug Administration website. Accessed June 2020.

# Clinical Case – pt TS

- 49-year old male
- No current treatments
- Weight gain of 7kg during pandemic
- Smoking 5 cigarettes a day
- Family history of hypertension (father) and CVD
- Current complaints: fatigue, otherwise asymptomatic
- BMI 27kg/m<sup>2</sup>
- Office bp: 143/82mmHg, HR 78/bpm

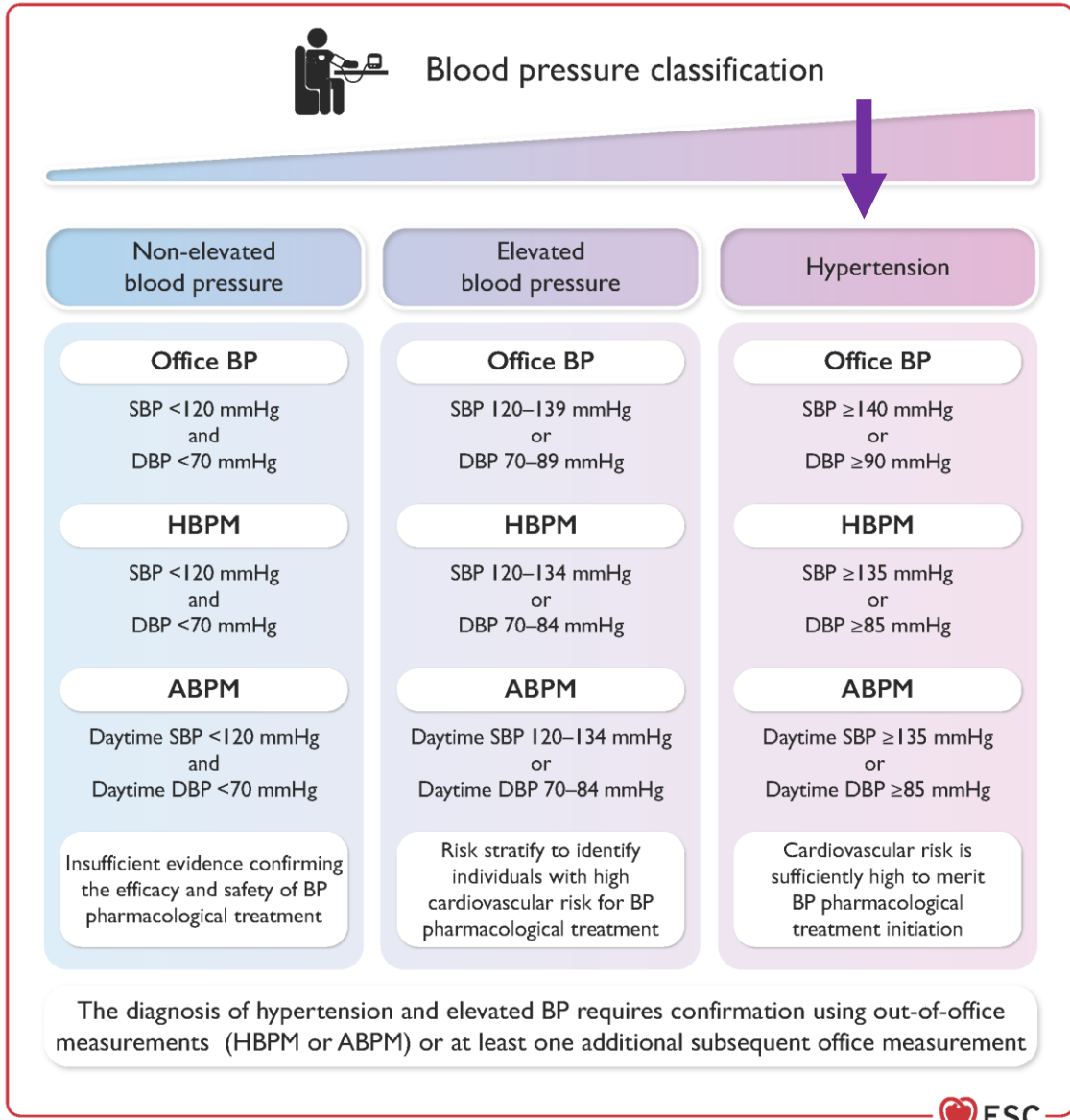


# Clinical Case – pt TS

- Fasting glucose 97mg/dL (5.4mmol/L)
- HbA1c 5.9%
- Normal serum Na<sup>+</sup> & K<sup>+</sup>
- eGFR >60mL/min/1.73m<sup>2</sup>
- Total cholesterol 7.1mmol/L
- HDL cholesterol 0.7 mmol/L
- TG 1.8 mmol/L
- LDL-c 5.5 mmol/L
- Non-HDL 6.4 mmol/L

# Blood pressure categories

- NEW**
- Simplified Blood pressure categories
  - New group of patients with elevated bp
  - No hypertension grades defined
  - For “Elevated bp”: risk stratify to identify high CV risk for BP pharmacological treatment



## SCORE2 & SCORE2-OP

10-year risk of (fatal and non-fatal) CV events in populations at moderate CVD risk



Women

Men

Non-smoking

Smoking

Non-smoking

Smoking

Non-HDL cholesterol

Systolic blood pressure (mmHg)  
**SCORE2-OP**

3.0-3.9  
4.0-4.9  
5.0-5.9  
6.0-6.9  
150 200 250

3.0-3.9  
4.0-4.9  
5.0-5.9  
6.0-6.9  
150 200 250

mmol/L  
mg/dL

3.0-3.9  
4.0-4.9  
5.0-5.9  
6.0-6.9  
150 200 250

3.0-3.9  
4.0-4.9  
5.0-5.9  
6.0-6.9  
150 200 250

160-179

37 39 40 42 41 43 44 46

Age (y)

37 45 53 62 37 45 53 61

140-159

35 36 38 39 39 40 42 43

85-89

36 43 51 59 35 43 51 59

120-139

32 34 35 37 36 38 39 41

34 41 49 57 34 41 48 57

100-119

30 32 33 34 34 35 37 38

32 39 47 55 32 39 46 55

160-179

27 28 30 31 34 35 37 39

30 35 41 47 34 40 46 53

140-159

24 25 27 28 30 32 33 35

80-84

27 32 37 43 31 36 42 48

120-139

21 22 24 25 27 28 30 31

25 29 34 40 28 33 38 44

100-119

19 20 21 22 24 25 27 28

22 26 31 36 25 30 35 40

160-179

19 20 21 23 27 29 30 32

75-79

24 27 31 35 31 35 39 44

140-159

16 17 18 19 24 25 26 28

21 23 27 30 27 30 34 38

120-139

14 15 15 16 20 21 22 24

17 20 23 26 23 26 29 33

100-119

12 12 13 14 17 18 19 20

15 17 19 22 19 22 25 29

160-179

13 14 15 16 22 23 25 26

19 21 23 25 28 31 34 36

140-159

11 11 12 13 18 19 20 22

15 17 18 20 23 25 28 30

120-139

9 9 10 11 15 16 17 18

70-74

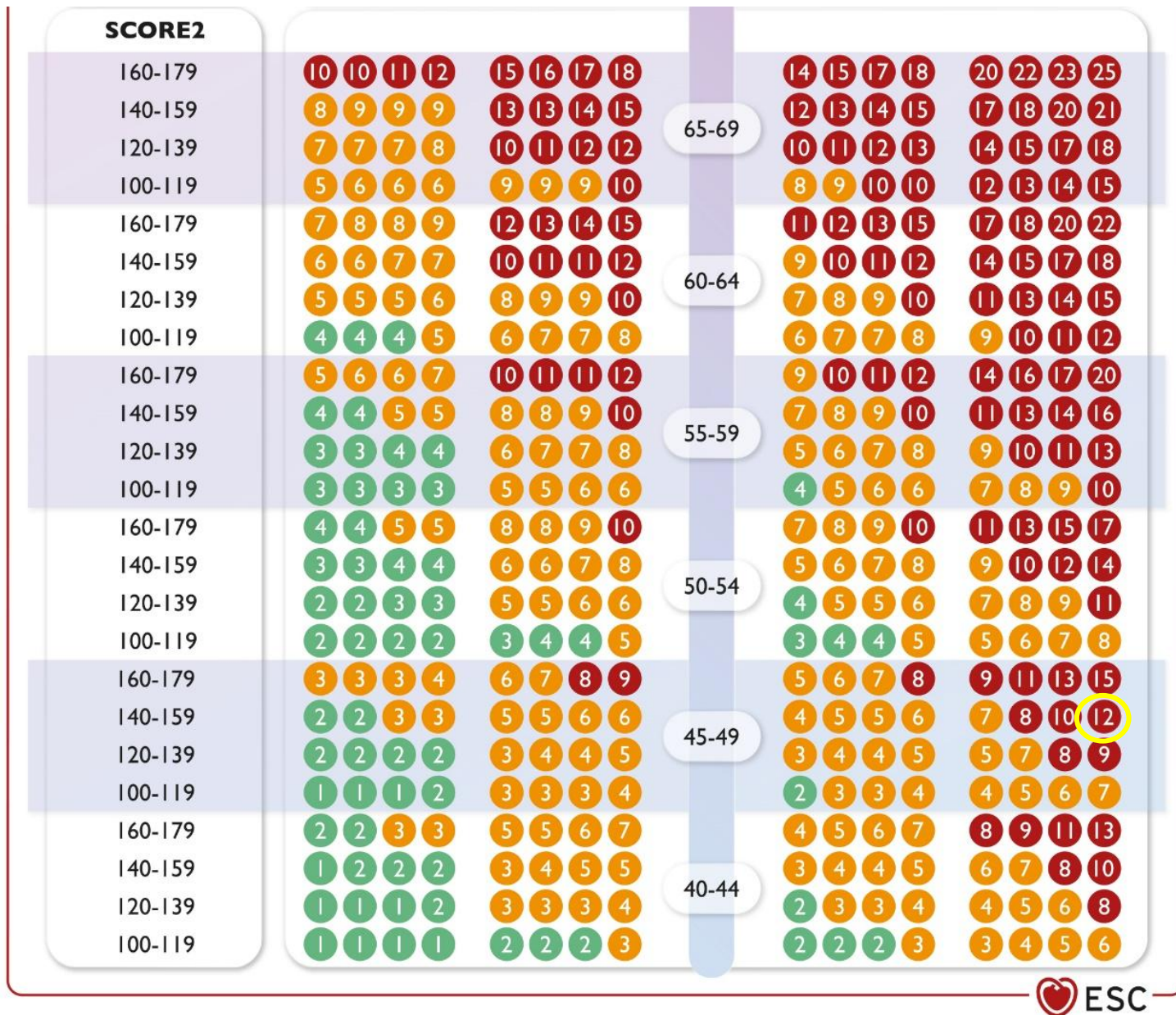
12 13 15 16 19 20 22 24

100-119

7 7 8 8 12 13 13 14

10 11 12 13 15 16 18 20

**SCORE2 and SCORE2-OP risk chart for fatal and non-fatal (MI, stroke) ASCVD Moderate CVD Risk (1)**

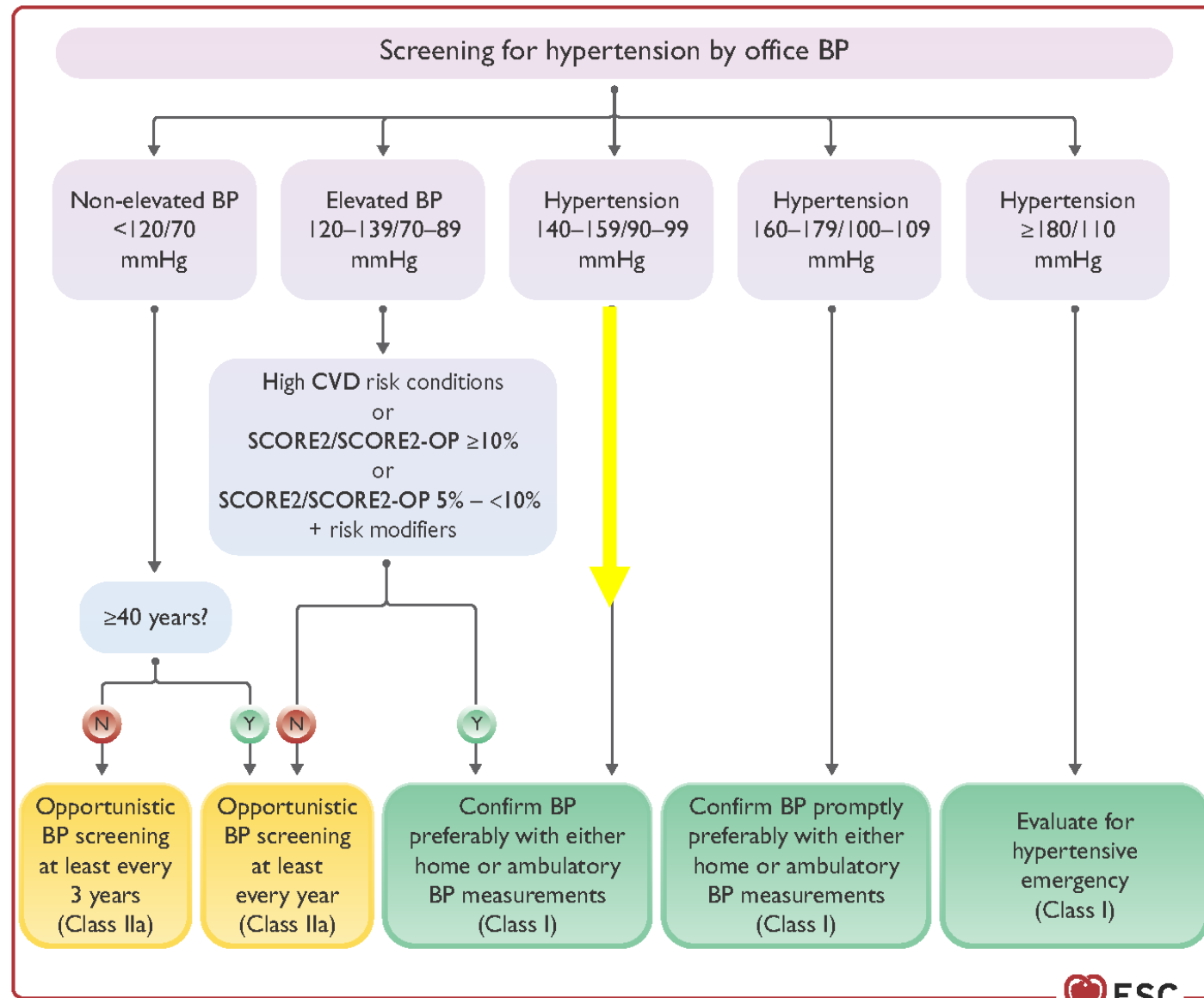


**SCORE2 and SCORE2-OP risk chart for fatal and non-fatal (MI, stroke) ASCVD**

**Moderate CVD Risk (2)**

← BP: 143/82mmHg  
Non-HDL: 6.4mmol/L

# Protocol for confirming hypertension diagnosis



# New recommendations (1)

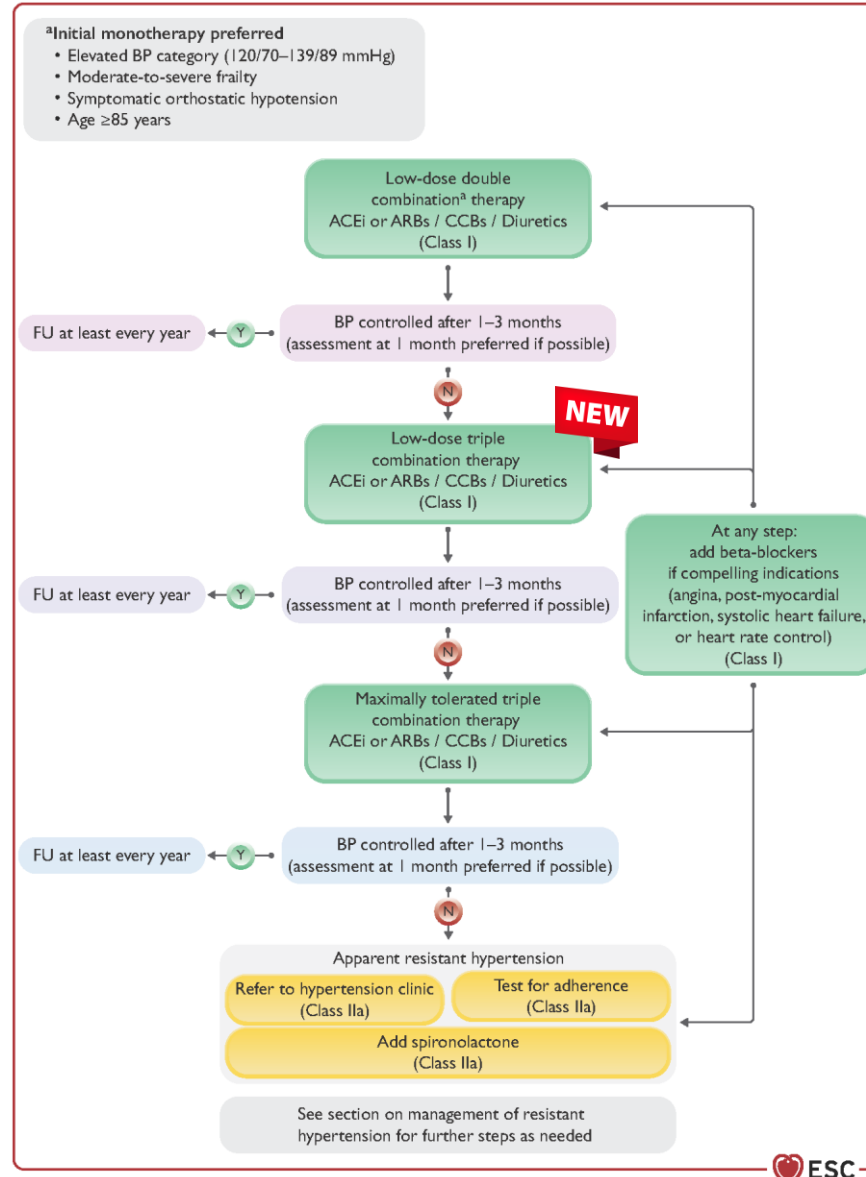
Recommendations	Class	Level
<b><i>Measuring blood pressure</i></b>		
It is recommended to measure BP using a validated and calibrated device, to enforce the correct measurement technique, and to apply a consistent approach to BP measurement for each patient.	I	B
Out-of-office BP measurement is recommended for diagnostic purposes, particularly because it can detect both white-coat hypertension and masked hypertension. Where out-of-office measurements are not logistically and/or economically feasible, then it is recommended that the diagnosis be confirmed with a repeat office BP measurement using the correct standardized measurement technique.	I	B
Most automated oscillometric monitors have not been validated for BP measurement in atrial fibrillation; BP measurement should be considered using a manual auscultatory method in these circumstances, where possible.	Ila	C

# Comparison of office, home, and ambulatory blood pressure measurement thresholds for elevated blood pressure and HTN



	Office BP (mmHg)	Home BP (mmHg)	Daytime ABPM (mmHg)	24 h ABPM (mmHg)	Night-time ABPM (mmHg)
Non-elevated BP	<120/70	<120/70	<120/70	<115/65	<110/60
Elevated BP	120/70– <140/90	120/70– <135/85	120/70– <135/85	115/65– <130/80	110/60– <120/70
Hypertension	≥140/90	≥135/85	≥135/85	≥130/80	≥120/70

# Practical algorithm for pharmacological blood pressure lowering



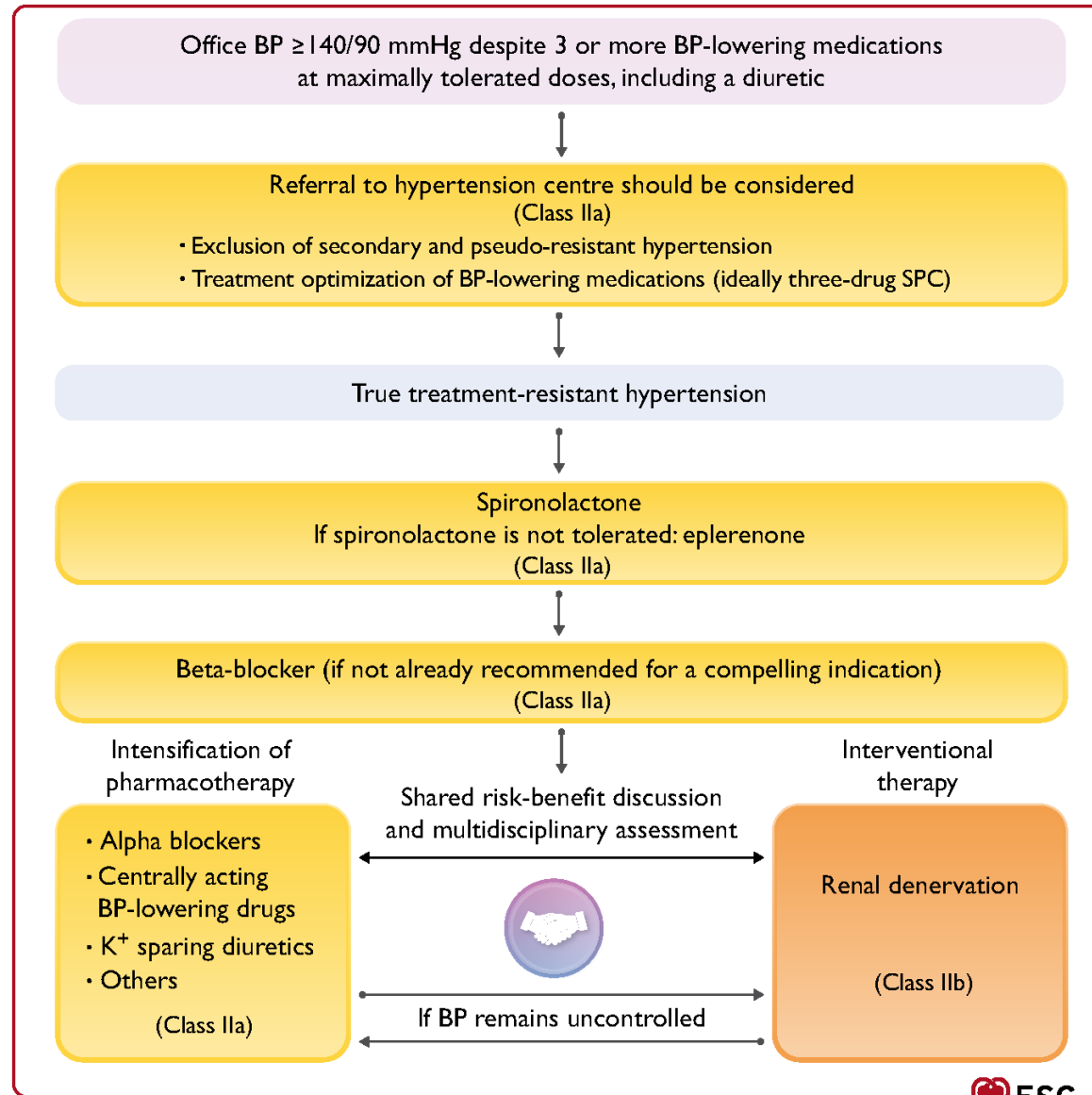
← Treatment initiation with Dual combination

← Earlier use of triple combination

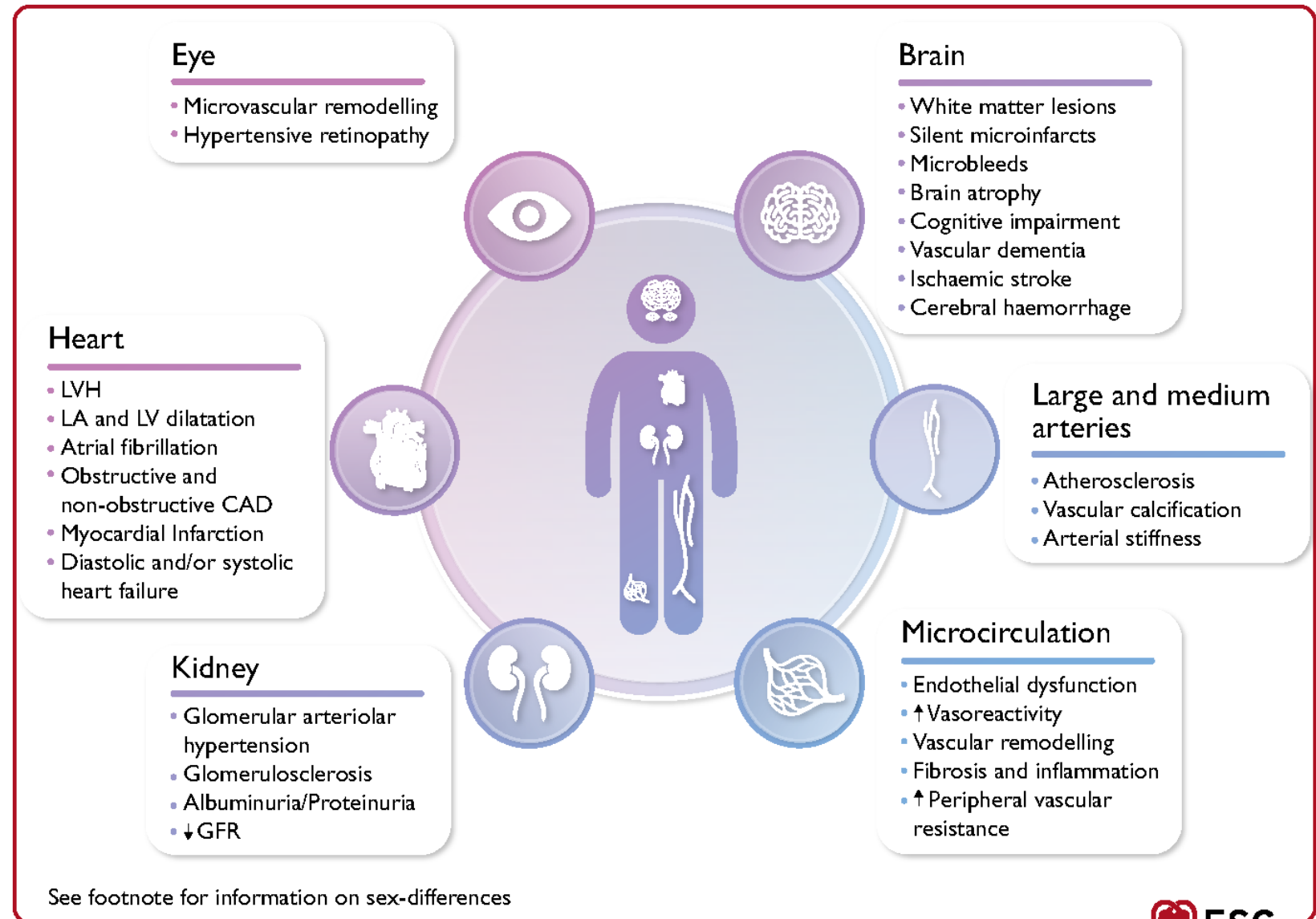


# Management of resistant hypertension

In patients with true resistant hypertension, the fourth line treatment should include the MRA spironolactone (or eplerenone), or a BB (if MRA not tolerated or not effective)




# Persistently elevated blood pressure and hypertension lead to hypertension-mediated organ damage and cardiovascular disease



# Clinical Case – pt TS

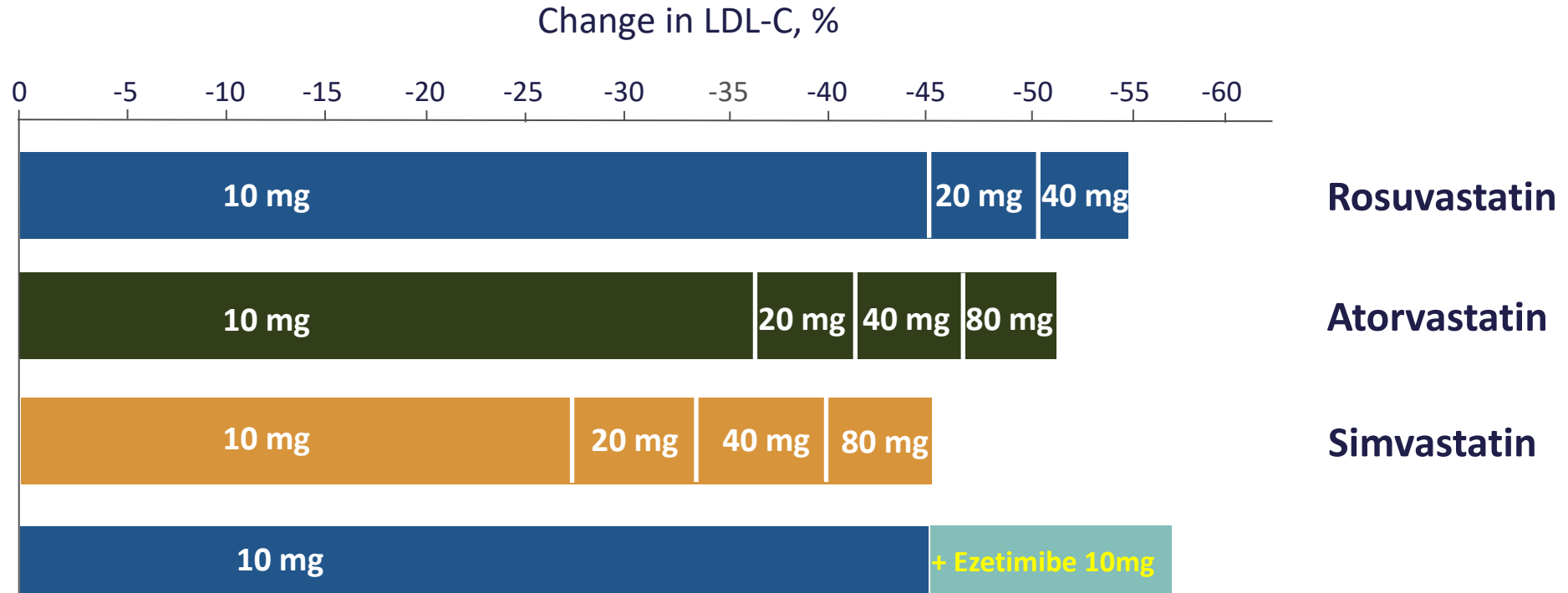
- Fasting glucose 97mg/dL (5.4mmol/L)
- HbA1c 5.9%
- Normal serum Na<sup>+</sup> & K<sup>+</sup>
- eGFR >60mL/min/1.73m<sup>2</sup>
- Total cholesterol 7.1mmol/L
- HDL cholesterol 0.7 mmol/L
- TG 1.8 mmol/L
- LDL-c 5.5 mmol/L
- Non-HDL 6.4 mmol/L

# 2019 ESC/EAS Dyslipidemia Targets

<p>LDL-C goal </p> <p>LDL &gt; 1.4mmol/L</p>	<p><b>Very high risk</b></p>	<p>People with any of the following:</p> <ul style="list-style-type: none"> <li>• Documented ASCVD, either clinical or unequivocal on imaging</li> <li>• DM with target organ damage, or at least three major risk factors, or early onset of T1DM of long duration (&gt; 20 years)</li> <li>• Severe CKD (eGFR &lt; 30 mL/min/1.73 m<sup>2</sup>)</li> <li>• A calculated SCORE ≥ 10% for 10-year risk of fatal CVD</li> <li>• FH with ASCVD or with another major risk factor</li> </ul>
<p>LDL &gt; 1.8mmol/L</p>	<p><b>High risk</b></p>	<p>People with:</p> <ul style="list-style-type: none"> <li>• Markedly elevated single risk factors, in particular TC &gt; 8 mmol/L (310 mg/dL), LDL-C &gt; 4.9 mmol/L (190 mg/dL), or BP ≥ 180/110 mmHg</li> <li>• Patients with FH without other major risk factors</li> <li>• Patients with DM without target organ damage, with DM duration ≥ 10 years or another additional risk factor</li> <li>• Moderate CKD (eGFR 30-59 mL/min/1.73 m<sup>2</sup>)</li> <li>• A calculated SCORE ≥ 5% and &lt; 10% for 10-year risk of fatal CVD</li> </ul>
<p>LDL &gt; 2.6mmol/L</p>	<p><b>Moderate risk</b></p>	<ul style="list-style-type: none"> <li>• Young patients (T1DM &lt; 35 years; T2DM &lt; 50 years) with DM duration &lt; 10 years, without other risk factors</li> <li>• Calculated SCORE ≥ 1% and &lt; 5% for 10-year risk of fatal CVD</li> </ul>
<p>LDL &gt; 3 mmol/L</p>	<p><b>Low risk</b></p>	<ul style="list-style-type: none"> <li>• Calculated SCORE &lt; 1% for 10-year risk of fatal CVD</li> </ul>

ASCVD, atherosclerotic cardiovascular disease; BP, blood pressure; CKD, chronic kidney disease; CVD, cardiovascular disease; DM, diabetes mellitus; eGFR, estimated glomerular filtration rate; FH, familial hypercholesterolaemia; LDL-C, low-density lipoprotein cholesterol; SCORE, Systematic Coronary Risk Estimation; T1DM, type 1 diabetes mellitus; T2DM, type 2 diabetes mellitus; TC, total cholesterol. Adapted from Mach F, et al. *Eur Heart J* 2020;41(1):111-88.

# Doubling Statin Dose will only Achieve ~6% Additional LDL-C Reduction




LDL-C, low-density lipoprotein cholesterol. Adapted from "FDA drug safety communication: New restrictions, contraindications, and dose limitations for simvastatin to reduce the risk of muscle injury." US Food & Drug Administration website. Accessed June 2020.

# Clinical Case – pt EB

- 48 year old male
- Smoker / on no regular Rx
- Presented with STEMI late December 2023
- Normal LV ejection fraction
- History: Revascularization (PDA / RCA / LAD) in January 2023
- TC 7.5 LDL 4.5 TG 4.4

\*PDA=Posterior Descending Artery; RCA=Right coronary artery; LAD = left anterior descending artery

# 2019 ESC/EAS Dyslipidemia Targets

<p>LDL-C goal </p> <p><b>LDL &gt; 1.4mmol/L</b></p>	<p><b>Very high risk</b></p>	<p>People with any of the following:</p> <ul style="list-style-type: none"> <li>• Documented ASCVD, either clinical or unequivocal on imaging</li> <li>• DM with target organ damage, or at least three major risk factors, or early onset of T1DM of long duration (&gt; 20 years)</li> <li>• Severe CKD (eGFR &lt; 30 mL/min/1.73 m<sup>2</sup>)</li> <li>• A calculated SCORE ≥ 10% for 10-year risk of fatal CVD</li> <li>• FH with ASCVD or with another major risk factor</li> </ul>
<p>LDL &gt; 1.8mmol/L</p>	<p><b>High risk</b></p>	<p>People with:</p> <ul style="list-style-type: none"> <li>• Markedly elevated single risk factors, in particular TC &gt; 8 mmol/L (310 mg/dL), LDL-C &gt; 4.9 mmol/L (190 mg/dL), or BP ≥ 180/110 mmHg</li> <li>• Patients with FH without other major risk factors</li> <li>• Patients with DM without target organ damage, with DM duration ≥ 10 years or another additional risk factor</li> <li>• Moderate CKD (eGFR 30-59 mL/min/1.73 m<sup>2</sup>)</li> <li>• A calculated SCORE ≥ 5% and &lt; 10% for 10-year risk of fatal CVD</li> </ul>
<p>LDL &gt; 2.6mmol/L</p>	<p><b>Moderate risk</b></p>	<ul style="list-style-type: none"> <li>• Young patients (T1DM &lt; 35 years; T2DM &lt; 50 years) with DM duration &lt; 10 years, without other risk factors</li> <li>• Calculated SCORE ≥ 1% and &lt; 5% for 10-year risk of fatal CVD</li> </ul>
<p>LDL &gt; 3 mmol/L</p>	<p><b>Low risk</b></p>	<ul style="list-style-type: none"> <li>• Calculated SCORE &lt; 1% for 10-year risk of fatal CVD</li> </ul>



ASCVD, atherosclerotic cardiovascular disease; BP, blood pressure; CKD, chronic kidney disease; CVD, cardiovascular disease; DM, diabetes mellitus; eGFR, estimated glomerular filtration rate; FH, familial hypercholesterolaemia; LDL-C, low-density lipoprotein cholesterol; SCORE, Systematic Coronary Risk Estimation; T1DM, type 1 diabetes mellitus; T2DM, type 2 diabetes mellitus; TC, total cholesterol. Adapted from Mach F, et al. *Eur Heart J* 2020;41(1):111-88.

## 2019 ESC/EAS Guidelines for the management of dyslipidaemias: lipid modification to reduce cardiovascular risk

### Treatment goals for LDL-C

In secondary prevention for patients at very-high risk, an LDL-C reduction of  $\geq 50\%$  from baseline and an LDL-C goal of  $< 1.4$  mmol/L ( $< 55$  mg/dL) are recommended.

I

A

In primary prevention for individuals at very-high risk, an LDL-C reduction of  $\geq 50\%$  from baseline and an LDL-C goal of  $< 1.4$  mmol/L ( $< 55$  mg/dL) are recommended.

I

C

In patients at high risk, an LDL-C reduction of  $\geq 50\%$  from baseline and an LDL-C goal of  $< 1.8$  mmol/L ( $< 70$  mg/dL) are recommended.

I

A

### Pharmacological LDL-C lowering

It is recommended that a high-intensity statin is prescribed up to the highest tolerated dose to reach the goals set for the specific level of risk.

I

A

If the goals are not achieved with the maximum tolerated dose of a statin, combination with ezetimibe is recommended.

I

B

For secondary prevention in patients at very-high risk not achieving their goal on a maximum tolerated dose of a statin and ezetimibe, a combination with a PCSK9 inhibitor is recommended.

I

A

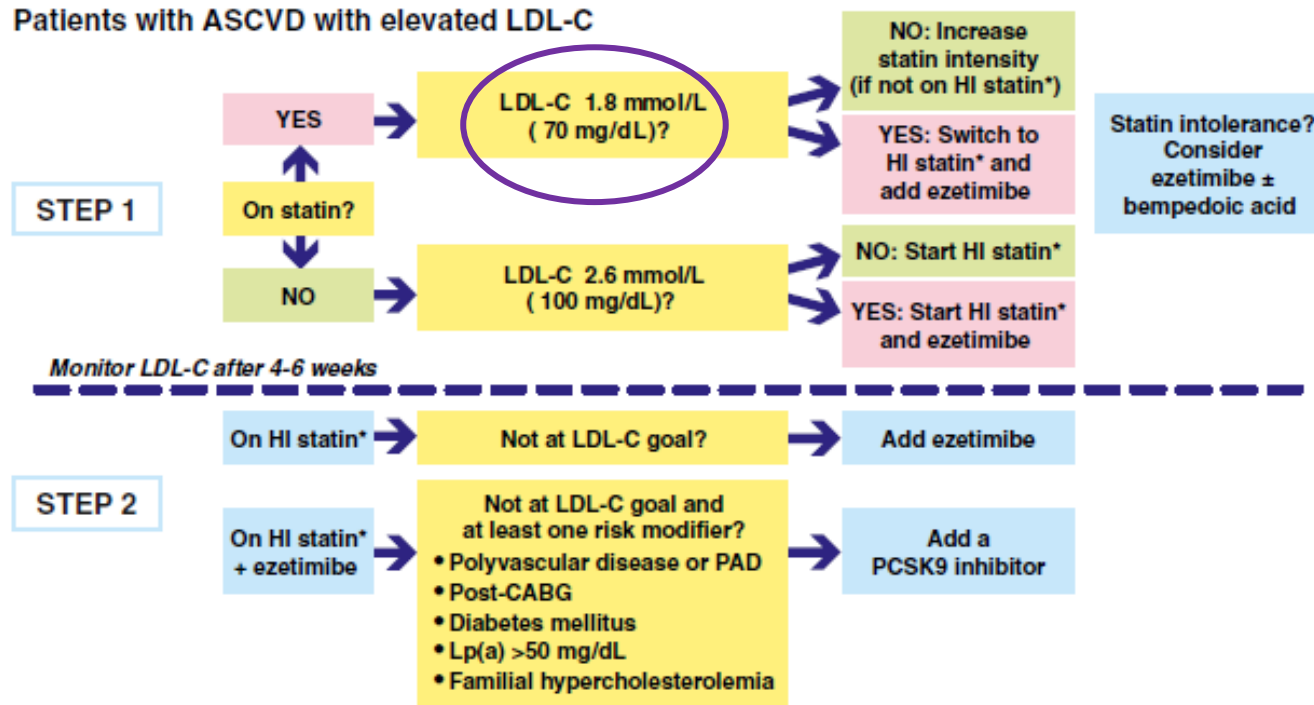
For very-high-risk FH patients (that is, with ASCVD or with another major risk factor) who do not achieve their goal on a maximum tolerated dose of a statin and ezetimibe, a combination with a PCSK9 inhibitor is recommended.

I

C



# New EAS 2021 Statement supporting upfront combinations of high-intensity statin/ezetimibe and fixed-dose combinations (FDCs)



\* HI statin: high-intensity statin or maximally tolerated statin therapy

**Upfront combinations**  
 2.1.3. *Why upfront combination treatment with a statin and ezetimibe?*  
 Patients with ASCVD, particularly those at enhanced risk with additional risk moderators, or FH without ASCVD and high LDL-C levels, are unlikely to attain LDL-C goal with intense statin monotherapy. Therefore, this Task Force recommends upfront combination high-intensity statin-ezetimibe treatment in the patients. This approach has particular advantages in avoiding repeated follow-up, allowing patients to be on target as early as possible, with favorable impact on cardiovascular outcome.

**In FDCs**  
 Proportion of patients at LDL-C goal by 3-fold [28]. The availability of a fixed combination of ezetimibe and high dose of a more efficacious statin will likely improve patient adherence.

## Paradigm shift in dyslipidemia management:

moving from a sequential treatment strategy to the upfront use of combinations

# Dose of Rosuvastatin on CV Morbidity & Mortality is important

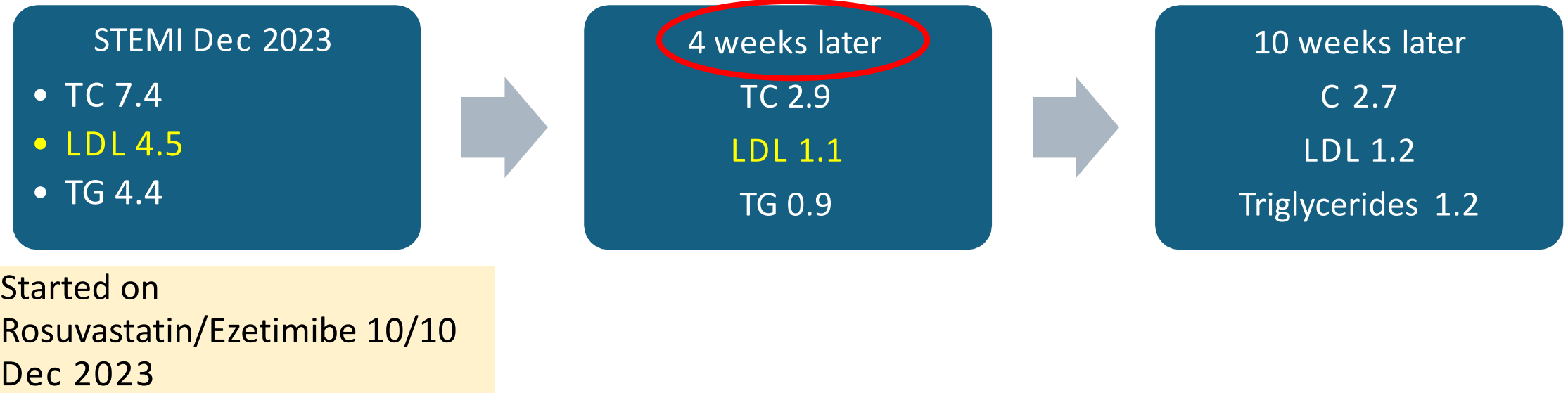
Study Name	Rosuvastatin Dose	Population	Primary Outcome
JUPITER Trial	20 mg daily	Individuals with elevated CRP levels and normal LDL levels	Significant reduction in cardiovascular events
HOPE-3 Trial	10 mg daily	Intermediate-risk individuals without cardiovascular disease	Reduced incidence of cardiovascular events
CORONA Trial	10 mg daily	Patients with systolic heart failure	No significant reduction in primary outcome; reduced hospitalizations
ASTEROID Trial	40 mg daily	Patients with coronary artery disease	Significant regression of atherosclerosis
METEOR Trial	40 mg daily	Individuals with low Framingham risk scores and evidence of subclinical atherosclerosis	Slowed progression of carotid intima-media thickness

In high & very high risk patients, an LDL-C reduction of at least 50% from baseline together with an LDL-C goal of <1.4 mmol/L) are recommended (both goals need to be achieved).

High-intensity statins at a dose of Rosuvastatin **10-20mg** are required to give CV protection.

# Clinical Case – pt EB

## Timeline



# Re-Testing

The expected LDL-C reductions in response to therapy are shown in *Figure 13*, and may vary widely among individuals. Therefore, monitoring the effect on LDL-C levels is recommended, with **assessment of LDL-C levels 4 - 6 weeks** after any treatment strategy initiation or change.



ESC

European Society  
of Cardiology

European Heart Journal (2021) 42, 3227–3337  
doi:10.1093/eurheartj/ehab484

ESC GUIDELINES

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**2021 ESC Guidelines on cardiovascular disease  
prevention in clinical practice**

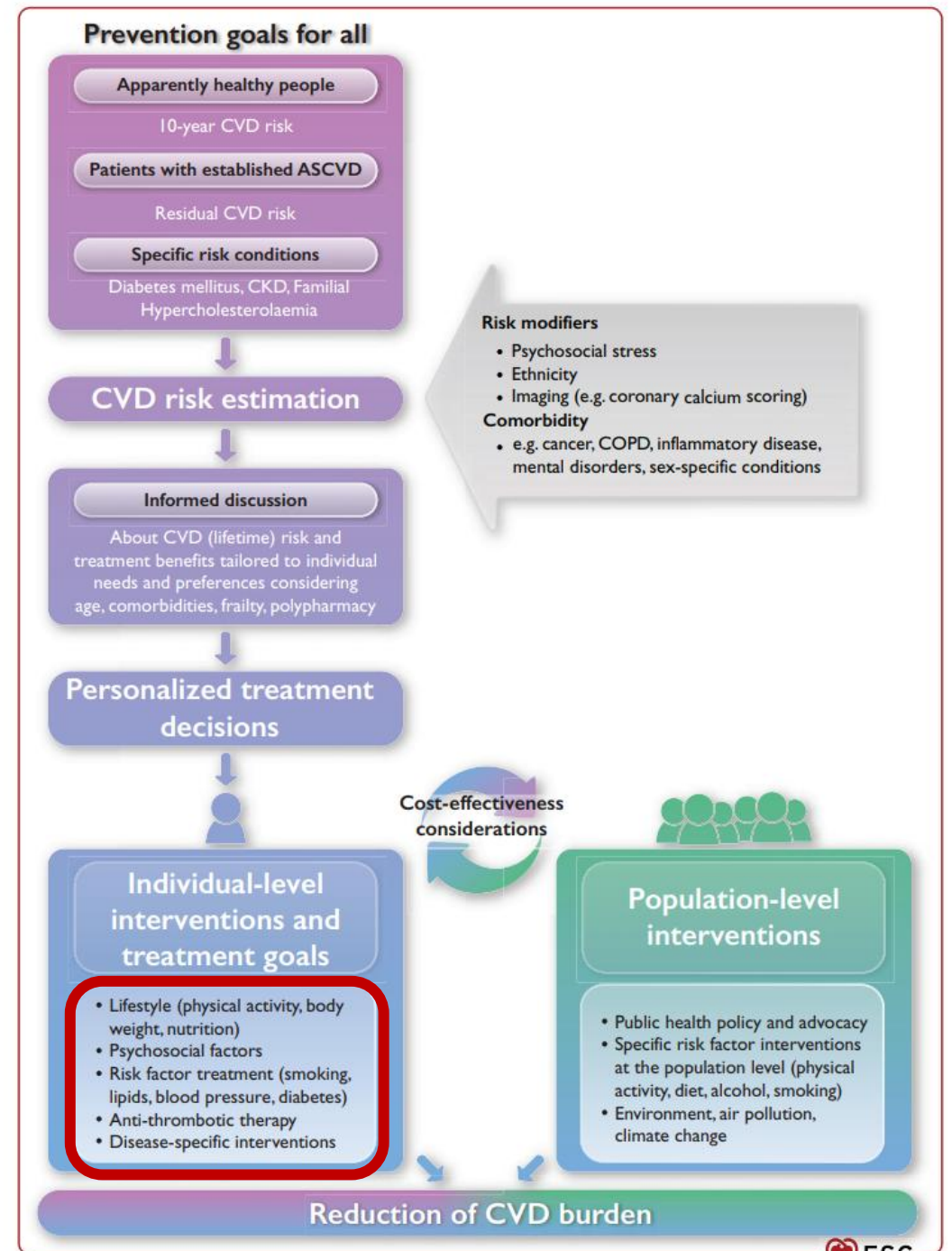
# Conclusions

- Dyslipidemia & hypertension are important variables in Cardiovascular Disease
- Different targets for different patient populations
- Dyslipidemia treatment: *the earlier, the lower, the longer...the better*
- Hypertension guidelines place a greater emphasis on individualized care and out-of-office monitoring, while introducing a more aggressive treatment target and new definitions for elevated BP.

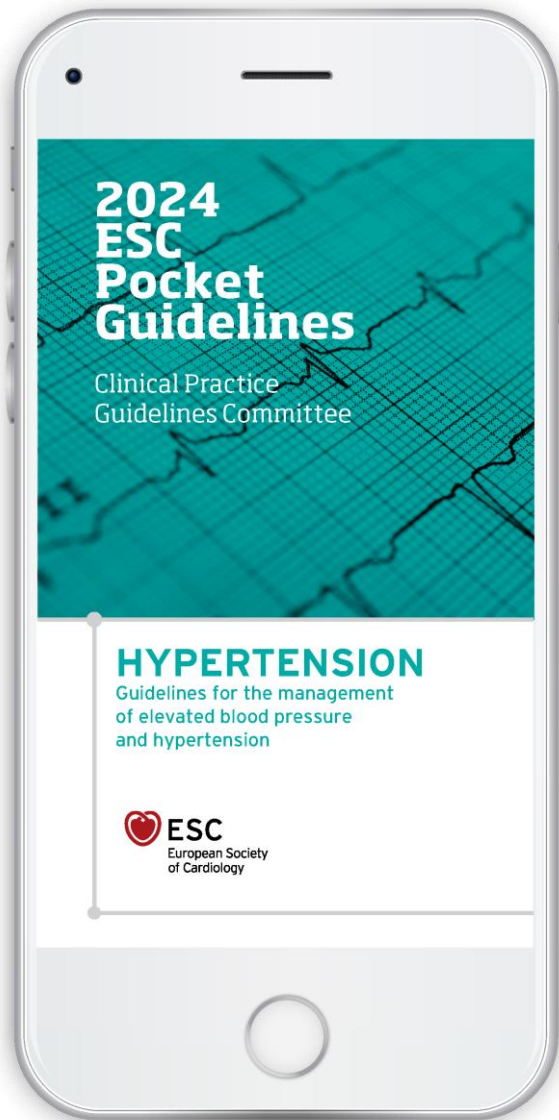
Back-up slides

# Risk Assessment

## 2021 ESC Guidelines on cardiovascular disease prevention in clinical practice



# ESC Pocket Guidelines App to access



- **All ESC Pocket Guidelines**

**AND**

- **Over 140 interactive tools**

- > Clinical decision support
- > Algorithms
- > Calculators
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- **Congress guidelines presentations**

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