# Special considerations in stroke prevention

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March 7, 2024

# Outline

Vascular Protection Clinic

Review of stroke risk factors

Focus on management

- Hypertension
- Dyslipidemia
- Carotid disease
- Atrial fibrillation

# Vascular Protection Clinic

- Perth hospital site
- Staff:
  - 2 General internists
  - 1 Nurse
  - 1 Dietician
  - 1-2 Volunteers
- Schedule:
  - First appointment Wednesday afternoons
  - Follow up appointments telephone

### Ontario Triage Algorithm for Stroke Prevention Clinic Referrals

Patients with TIA or Non Disabling Stroke Symptoms

Acute Ataxia, Vision, or Hemibody MASH Symptoms within 48hrs seen MASH symptoms in ED OR Motor or Speech symptoms Sensory symptoms between 48hrs greater than 2 weeks between 48hrs and 2 wks and 2 wks Has Stroke Evaluation been completed? Any urgent findings? i.e. new stroke on imaging, untreated atrial fibrillation/flutter, untreated symptomatic >50% carotid stenosis, or other (thrombosis/dissection/stenosis) No Are other High Risks\*\* Present?

Symptoms LESS likely to be TIA/Stroke OR Discharged inpatient New Acute TIA / Stroke Symptoms

Within 48 hrs and no ED Visit Advise to go to CT-capable ED Immediately

#### Stroke Symptoms - MASH

likely  $\Pi\!A$  or minor stroke (transient, fluctuating) & if persistent, increase triage scale or admit

MOTOR (Unilateral weakness: face or arm or leg)

ACUTE ATAXIA or VISION CHANGE (monocular or hemifield vision loss or diplopia)

SPEECH (dysarthric or dysphasia/aphasia)

HEMIBODY SENSORY (unilateral numbness: face/arm or arm/leg)

#### Stroke Evaluation

- Head Imaging
- Vascular Imaging
- Cardiac Monitoring (ECG or Holter or Loop)
- Antiplatelet or Anticoagulation started

#### Triage Scale

Time frame to be seen at SPC from date of referral sent.

- Mithin 24hrs (ED or SPC Fast Track) (emergent)
- B Within I Week (urgent/high)
- Within 2 Weeks (*moderate)*
- Mithin 1 Month (*low)*
- Within 3 Months (discharge inpatients / less likely to be TIA/stroke but still may need attention)

Adapted from Northwestern Ontario Regional Stroke Network & Thunder Bay Regional Health Sciences Centre Source: Canadian Stroke Best Practice Recommendations (2020; 2017) & References (See Appendix)

Updated by Ontario Secondary Stroke Prevention Task Group (May, 2022)

- \* If untreated atrial fibrillation/flutter-implement plan for anticoagulation ASAP
- \*\* See over for other high risks

#### Other High Risks\*\* to Consider: Higher priority for those based on:

- Symptoms
  - Longer duration of symptoms
  - MASH stroke symptoms occurring > 2 weeks [with time the priority diminishes (e.g., 2 weeks vs > 3 months)]
  - Warrants OT/PT/SLP assessment
- Other vascular conditions
- Previous TIA/stroke
- Pregnancy including post-partum
- Cancer
- Vascular risk factors
- Already on Antiplatelet/Anticoagulation therapy
- Blood pressure reading is high (e.g., initial triage diastolic blood pressure ≥ 110 mmHg as per Canadian TIA Score)
- Abnormal blood work (e.g., Glucose  $\geq$  15 mmol/L &Platelet count  $\geq$  400 x 10 $^{9}$ /L as per Canadian TIA Score)
- High Canadian TIA Score ≥ 9
- Other considerations:
  - Lifestyle risks
  - o Age (younger)
  - Ethnicity
  - Family history

#### Not likely to be a TIA

- · Transient symptoms lasting only seconds
- Seizure
- · Isolated transient loss of consciousness or syncope
- · Transient global amnesia
- · Isolated non-vertiginous dizziness
- Vague generalized weakness without loss of power

#### OR

No other focal neurological findings

Defer back to referral source or primary care physician for follow up or as per internal processes



Brain imaging

CT or CTA +/- MRI



Carotid imaging

CTA
Carotid doppler





Echocardiogram

Trans-thoracic +/- Trans-esophageal



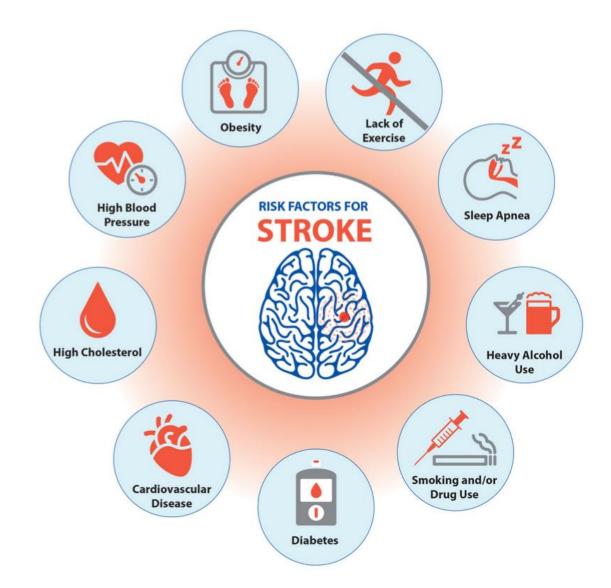
Holter monitors

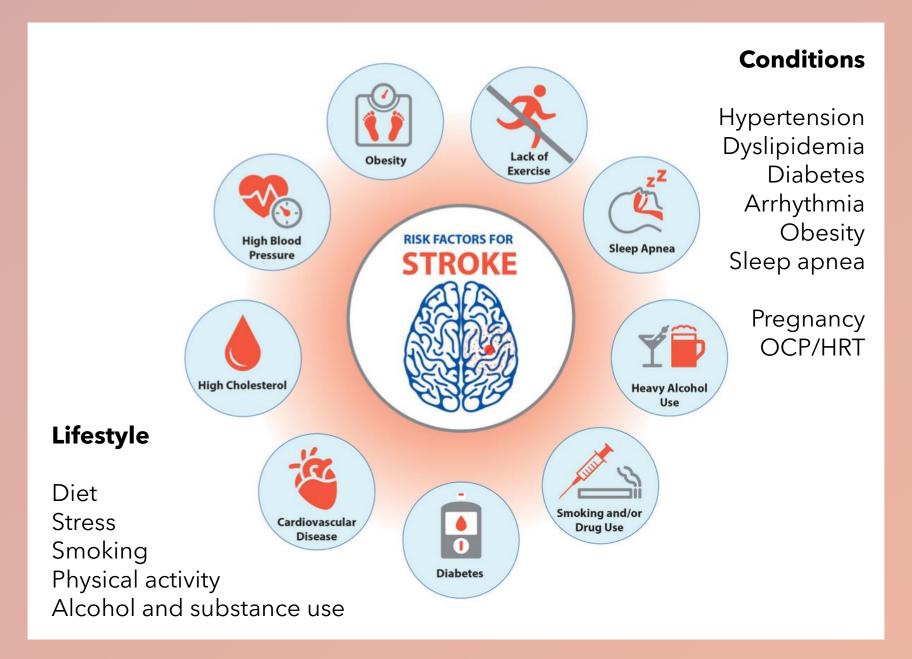
48 hour 14 day

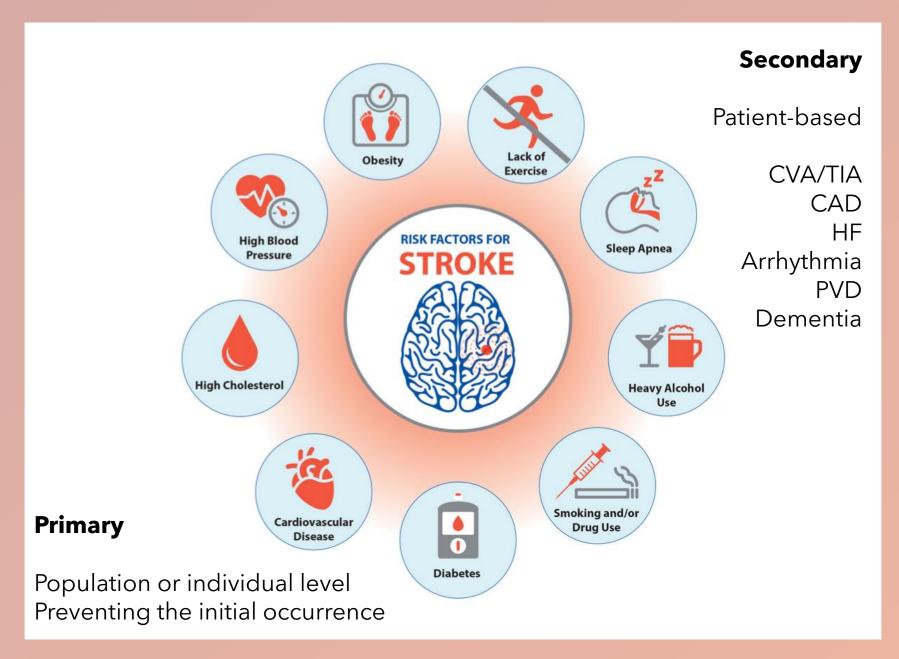


Blood work

Lipid panel HbA1c CBC, lytes, creat, ALT, CRP







# Targets for lifestyle modification

#### **Diet**

- Balanced diet
- Limited processed foods
- Limiting sodium
- DASH or Mediterranean diets

#### **Exercise**

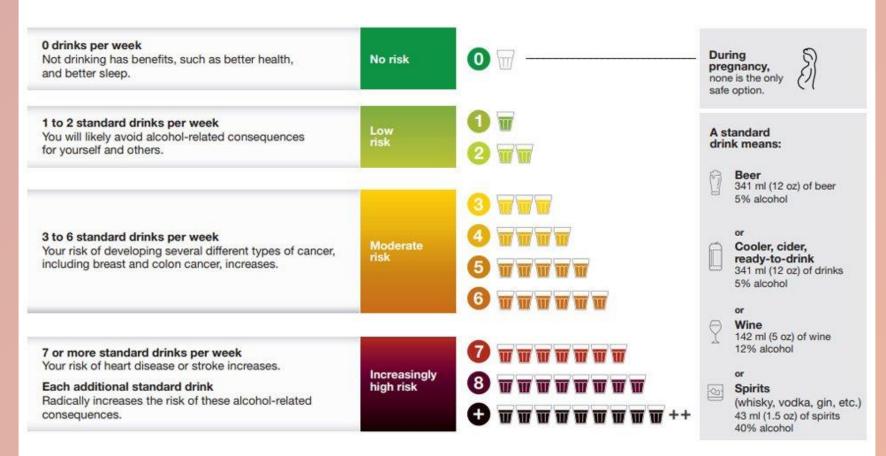
- At least 150 minutes per week
- In bouts of 10 minutes or more
- Moderate- to vigorous intensity aerobic physical activity

### Smoking / Substances

Cessation

### Alcohol consumption per week

Drinking alcohol has negative consequences. The more alcohol you drink per week, the more the consequences add up.



https://www.ccsa.ca/canadas-guidance-alcohol-and-health



# Hypertension

Most prevalent CV risk factor in Canada

## Figure 1

# Managing Hypertension

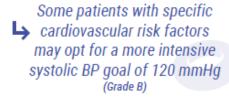
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#### 1 When to start a drug and what to aim for?

The threshold to start a drug may be different from the blood pressure goal for a patient on drug therapy

	Primary Prevention & Lower Risk	Secondary Prevention & Higher Risk	Diabetes
Who is included	<ul> <li>No history of heart disease, heart attack, heart failure, or stroke</li> </ul>	<ul> <li>History of heart attack/stroke         or</li> <li>10-year Framingham         CV risk score &gt;15%</li> </ul>	<ul><li>Type 1</li><li>Type 2</li></ul>
When to start a drug (threshold)	• >160/100 mmHg	<ul> <li>&gt;140/90 mmHg</li></ul>	<ul> <li>&gt;130/80 mmHg</li></ul>
	(Grade A)	(SBP Grade C; DBP Grade A)	(Grade C)
What to aim for (goal)	<ul><li>&lt;140/90 mmHg</li></ul>	<ul> <li>&lt;140/90 mmHg</li></ul>	<130/80 mmHg
	(Grade A)	(Grade A)	(Grade C)

Table is based on the 2018 Hypertension Canada guidelines. SBP (systolic), DBP (diastolic). Grade A evidence: strong evidence; Grade B evidence: moderate evidence; Grade C evidence: weak evidence.





Help patients choose a threshold and goal based on their preferences, medical history, and frailty



Consider waiting if there is a short-term cause of hypertension (e.g., pain, stress, trauma)



Table 5. Blood pressure thresholds for initiation of antihypertensive therapy and treatment targets in adults

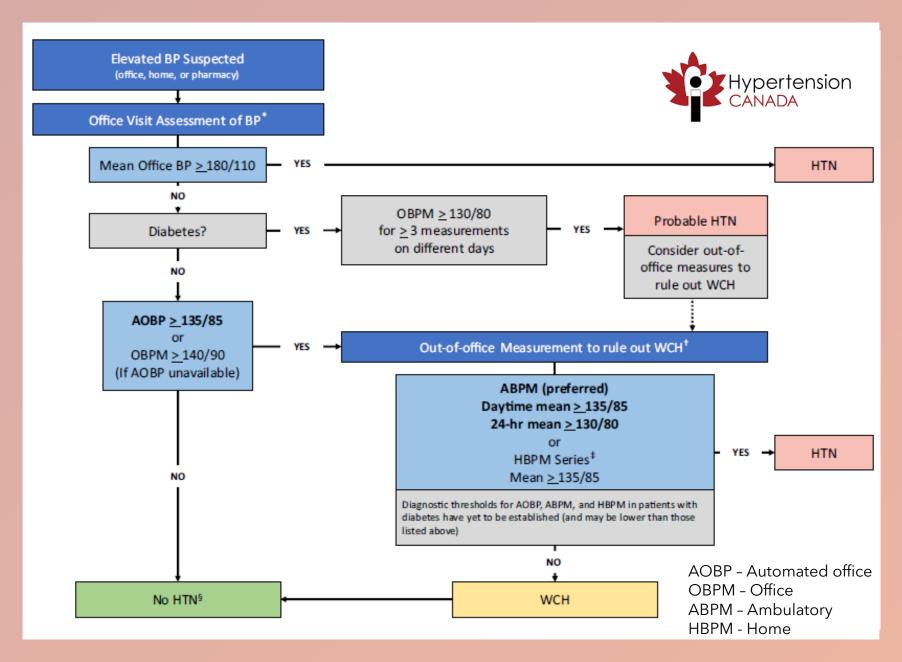
Patient population	BP threshold (mm Hg) for initiation of antihypertensive therapy	BP target (mm Hg) for treatment
Low risk (no target	SBP ≥ 160 (Grade A)	SBP < 140 (Grade A)
organ damage or cardiovascular risk factors)	DBP ≥ 100 (Grade A)	DBP < 90 (Grade A)
High risk of cardiovascular disease*	SBP ≥ 130 (Grade B)	SBP < 120 (Grade B)
Diabetes mellitus	SBP ≥ 130 (Grade C)	SBP < 130 (Grade C)
	$DBP \ge 80 \text{ (Grade A)}$	DBP < 80 (Grade A)
All others	$SBP \ge 140$ (Grade C) $DBP \ge 90$ (Grade A)	SBP < 140 (Grade A) DBP < 90 (Grade A)

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

### **High risk**

- Clinical or subclinical CV disease
- CKD / proteinuria
- 10y CV risk >15%
- Age >75y

<sup>\*</sup>See Table 6; on the bass of automated office blood pressure measurement.



## 2023 ESH Guidelines for the management of arterial hypertension



Hypertension disease	Other risk factors, HMOD, CVD or CKD	BP (mmHg) grading			
staging		High-normal SBP 130–139 DBP 85–89	Grade 1 SBP 140–159 DBP 90–99	Grade 2 SBP 160–179 DBP 100–109	Grade 3 SBP ≥ 180 DBP ≥ 110
	No other risk factors <sup>a</sup>	Low risk	Low risk	Moderate risk	High risk
Stage 1	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	HMOD, CKD grade 3, or diabetes mellitus	Moderate to high risk	High risk	High risk	Very high risk
Stage 3	Established CVD or CKD grade ≥4	Very high risk	Very high risk	Very high risk	Very high risk

<50 years	60–69 years	≥70 years	
<2.5%	<5%	<7.5%	
2.5 to <7.5%	5 to <10%	7.5 to <15%	Complementary risk estimation in Stage 1
≥7.5%	≥10%	≥15%	with SCORE2/SCOR2-OP

FIGURE 4 Cardiovascular risk according to grade and stage of hypertension.

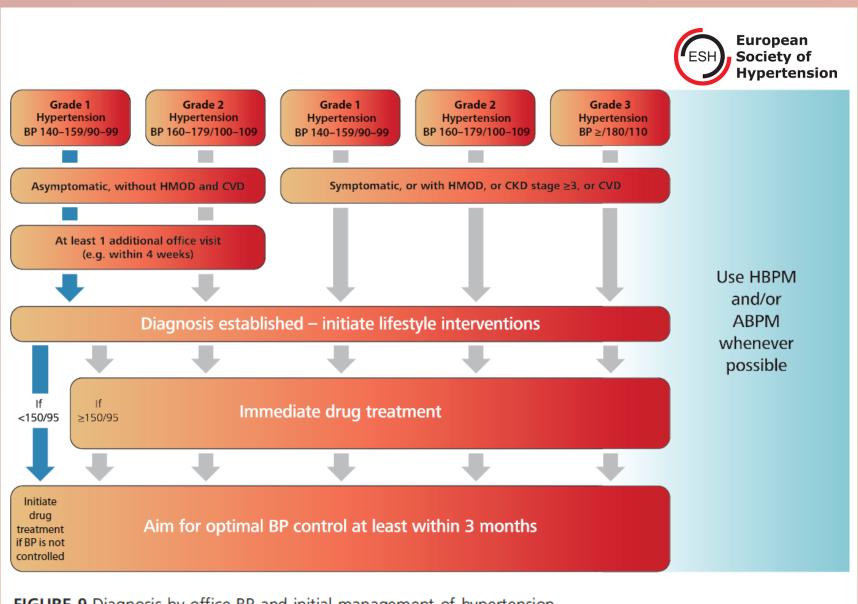


FIGURE 9 Diagnosis by office BP and initial management of hypertension.



# 2023 ESH Guidelines for the management of arterial hypertension

The Task Force for the management of arterial hypertension of the European Society of Hypertension

Endorsed by the International Society of Hypertension (ISH) and the European Renal Association (ERA)

Treatment BP targets

Adults 18–79 y, primary SBP/DBP goal <140/90 mm Hg; if treatment well tolerated target SBP/DBP <130/80 mm Hg but not SBP <120 mm Hg or DBP <70 mm Hg. For adults with isolated systolic hypertension, target SBP lowering, albeit cautiously.

In adults ≥80 y, target SBP/DBP <140/90 mm Hg, if well tolerated.

Recommendations are like previous ESH Guideline



#### Table 7. Considerations in the individualization of pharmacological therapy in adults

	Initial therapy	Second-line therapy	Notes and/or cautions		
Hypertension without other compelling indications					
Diastolic hypertension with or without systolic hypertension	Monotherapy or SPC. Recommended monotherapy choices include thiazide/thiazide-like diuretics (with longer-acting diuretics preferred), β-blockers, ACE inhibitors, ARBs, or long-acting CCBs. Recommended SPC choices include combinations of an ACE inhibitor with CCB, ARB with CCB, or ACE inhibitor/ARB with a diuretic (consider statins in selected patients)		Not recommended for monotherapy: α-blockers, β-blockers in those 60 years of age or older, ACE inhibitors in black people. Hypokalemia should be avoided in those prescribed diuretics. Combination of an ACE inhibitor with an ARB is not recommended		
Isolated systolic hypertension without other compelling indications Diabetes mellitus	Thiazide/thiazide-like diuretics, ARBs, or long-acting dihydropyridine CCBs	Combinations of first-line drugs	Same as diastolic hypertension with or without systolic hypertension		
Diabetes mellitus with microalbuminuria,* renal disease, cardiovascular disease, or additional cardiovascular risk factors	ACE inhibitors or ARBs	Additional use of a dihydropyridine CCB is preferred over a thiazide/ thiazide-like diuretic	A loop diuretic could be considered in hypertensive chronic kidney disease patients with extracellular fluid volume overload		
Diabetes mellitus not included in the above category	ACE inhibitors, ARBs, dihydropyridine CCBs, or thiazide/ thiazide-like diuretics	Combination of first-line drugs. If combination with ACE inhibitor is being considered, a dihydropyridine CCB is preferable to a thiazide/ thiazide-like diuretic	Normal urine microalbumin to creatinine ratio < 2.0 mg/mmol		
Cardiovascular disease					
Coronary artery disease	ACE inhibitors or ARBs; β-blockers or CCBs for patients with stable angina	When combination therapy is being used for high-risk patients, an ACE inhibitor/dihydropyridine CCB is preferred	Avoid short-acting nifedipine Combination of an ACE inhibitor with an ARB is not recommended. Exercise caution when lowering SBP to target if DBP is ≤ 60 mm Hg, especially in patients with LVH		
Recent myocardial infarction	$\begin{array}{l} \beta\text{-Blockers and ACE inhibitors (ARBs} \\ \text{if ACE inhibitor-intolerant)} \end{array}$	Long-acting CCBs if β-blocker contraindicated or not effective	Nondihydropyridine CCBs should not be used with concomitant heart failure		

Table 7. Considerations in the individualization of pharmacological therapy in adults

	Initial therapy	Second-line therapy	Notes and/or cautions
Heart failure	ACE inhibitors (ARBs if ACE inhibitor-intolerant) and β-blockers. Aldosterone antagonists (mineralocorticoid receptor antagonists) may be added for patients with a recent cardiovascular hospitalization, acute myocardial infarction, elevated BNP or NT-proBNP level, or NYHA class II-IV symptoms	ACE inhibitor and ARB combined. Hydralazine/isosorbide dinitrate combination if ACE inhibitor and ARB contraindicated or not tolerated. Thiazide/thiazide-like or loop diuretics are recommended as additive therapy; dihydropyridine CCB can also be used. A combined ARB/neprilysin-inhibitor is recommended (in place of an ACE inhibitor or ARB) in symptomatic patients with hypertension and HFrEF according to standard guideline-based therapies	Titrate doses of ACE inhibitors and ARBs to those used in clinical trials Carefully monitor potassium and renal function if combining any of ACE inhibitor, ARB, and/or aldosterone antagonist
LVH	ACE inhibitor, ARB, long-acting CCB, or thiazide/thiazide-like diuretic	Combination of first-line agents	Hydralazine and minoxidil should not be used
Past stroke or TIA	ACE inhibitor and a thiazide/thiazide- like diuretic combination	Combination of first-line agents	Treatment of hypertension should not be routinely undertaken in patients with acute stroke unless extreme BF elevation. Combination of an ACE inhibitor with an ARB is not recommended
Nondiabetic chronic kidney disease Nondiabetic chronic kidney disease with proteinuria <sup>†</sup>	ACE inhibitors (ARBs if ACE inhibitor-intolerant) if there is proteinuria Diuretics as additive therapy	Combinations of first-line agents	Carefully monitor renal function and potassium for those receiving an ACE inhibitor or ARB. Combinations of an ACE inhibitor and ARB are not recommended

ACE, angiotensin converting enzyme; ARB, angiotensin receptor blocker; BNP, brain natriuretic peptide; BP, blood pressure; CCB, calcium channel blocker; DBP, diastolic blood pressure; HFrEF, heart failure with reduced ejection fraction < 40%; LVH, left ventricular hypertrophy; NT-proBNP, N-terminal pro B-type naturietic peptide; NYHA, New York Heart Association; SBP, systolic blood pressure; SPC, single-pill combination; TIA, transient ischemic attack.

<sup>\*</sup>Microalbuminuria is defined as persistent albumin to creatinine ratio > 2.0 mg/mmol.

<sup>&</sup>lt;sup>†</sup>Proteinuria is defined as urinary protein > 150 mg in 24 hours or albumin to creatinine ratio > 30 mg/mmol in 2 of 3 specimens.

# Dyslipidemia

Primary versus secondary prevention



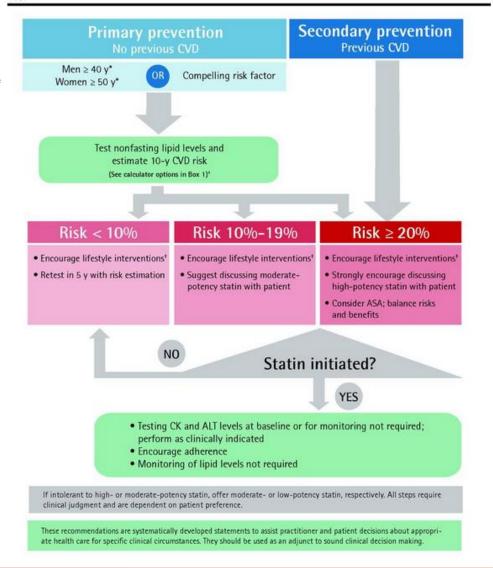
The official journal of the College of Family Physicians of Canada

#### Simplified lipid guidelines

Prevention and management of cardiovascular disease in primary care

(2015)

**Figure 1. Lipid algorithm:** For primary or secondary prevention; excludes those with familial hypercholesterolemia.



# Dyslipidemia



# 2021 Canadian Cardiovascular Society Guidelines for the Management of Dyslipidemia for the Prevention of Cardiovascular Disease in Adults

- New treatment targets
  - Non-HDL and ApoB
- New therapies
  - PCSK9 inhibitors and IPE

# Dyslipidemia



### **Statin indicated conditions**

Atherosclerotic cardiovascular disease (ASCVD)	Other
Coronary artery disease	Diabetes
Stroke / TIA	CKD
Peripharal arterial disease	LDL≥5 mmol/L
Carotid disease	Familial hypercholesterolemia
AAA	



#### Table 1. Who to screen for dyslipidemia in adults at risk

#### Who to screen

# Men 40 years of age or older; women 40 years of age or older (or postmenopausal)

 Consider earlier in ethnic groups at increased risk such as South Asian or indigenous individuals

#### All patients with any of the following conditions, regardless of age

- Clinical evidence of atherosclerosis
- Abdominal aortic aneurysm
- Diabetes mellitus
- Arterial hypertension
- Current cigarette smoking
- Stigmata of dyslipidemia (corneal arcus, xanthelasma, xanthoma)
- Family history of premature CVD\*
- · Family history of dyslipidemia
- CKD (eGFR ≤ 60 mL/min/1.73 m<sup>2</sup> or ACR ≥ 3 mg/mmol)
- Obesity (BMI ≥ 30)
- Inflammatory diseases (RA, SLE, PsA, AS, IBD)
- HIV infection
- Erectile dysfunction
- COPD
- · History of hypertensive disorder of pregnancy

#### How to screen:

- Lipid panel
- HbA1c / FPG
- eGFR
- Lipoprotein A\*\*
- ApoB
- Urine ACR

#### PRIMARY PREVENTION<sup>†</sup>

Low-Risk\* FRS <10% Intermediate-Risk\* FRS 10-19.9% and

LDL-C ≥3.5 mmol/L **or** Non-HDL-C ≥4.2 mmol/L **or** ApoB ≥1.05 g/L **or** 

Men ≥50 yrs and women ≥60 yrs with one additional risk factor: low HDL-C, IFG, high waist circumference, smoker, or HTN **or** 

with presence of other risk modifiers: hsCRP ≥2.0 mg/L, CAC >0 AU, family history of premature CAD, Lp(a) ≥50 mg/dL (100 nmol/L) High-Risk\* FRS ≥20%



Statin therapy not recommended for most low-risk individuals; exceptions include: (a) LDL-C ≥5.0 mmol/L (or ApoB ≥1.45 g/L or non-HDL-C ≥ 5.8 mmol/L) – see Figure 2; or (b) FRS is 5%-9.9% with LDL-C ≥3.5 mmol/L (or non-HDL-C ≥4.2 mmol/L or ApoB ≥1.05 g/L), particularly with other CV risk modifiers (eg, FHx, Lp(a) ≥50 mg/dL [or ≥100 nmol/L] or CAC >0 AU) as the proportional benefit from statin therapy may be similar to other treated groups.



#### **Health Behaviour Modifications:**

- · Smoking cessation
- Diet: It is recommended all individuals adopt a healthy dietary pattern.
- Exercise: It is recommended adults accumulate at least 150 mins/week of moderate-vigorous intensity aerobic physical activity.

#### Monitor

- · response to statin Rx
- · response to add-on lipid-lowering Rx
- · health behaviour changes

#### Discuss health behaviour modifications

#### INITIATE STATIN TREATMENT

If LDL-C ≥2.0 mmol/L or ApoB ≥0.8 g/L or non-HDL-C >2.6 mmol/L on maximally tolerated statin dose

YES

NO

YES

YES

Discuss add-on therapy with patient: Evaluate reduction in CVD risk vs. cost/access and side effects

ADD-ON

(BAS as alternative)<sup>¶</sup>

#### STATIN INDICATED CONDITIONS

#### LDL ≥5.0 mmol/L

(or ApoB ≥1.45 g/L or non-HDL-C ≥5.8 mmol/L) (familial hypercholesterolemia or genetic dyslipidemia)

#### Most patients with diabetes:

- Age ≥40y
- Age ≥30y & DM x≥15y duration
- · Microvascular disease

#### **Chronic Kidney Disease**

 Age ≥50y and eGFR <60 mL/min/1.73 m<sup>2</sup> or ACR >3 mg/mmol

### Atherosclerotic Cardiovascular Disease (ASCVD):

- Myocardial infarction (MI), acute coronary syndromes (ACS)
- Stable angina, documented coronary artery disease using angiography
- Stroke, TIA, documented carotid disease
- Peripheral arterial disease, claudication, and/or ABI < 0.9</li>
- Abdominal aortic aneurysm (AAA) -abdominal aorta >3.0 cm or previous aneurysm surgery



Review/Discuss health behavioural modifications (refer to Figure 1)

#### INITIATE STATIN TREATMENT

If LDL-C ≥2.5 mmol/L If LDL-C ≥2.0 mmol/L or If LDL-C ≥1.8 mmol/L or (or <50% reduction) or ApoB ≥0.80 g/L or ApoB ≥0.70 g/L or NO ApoB ≥0.85 g/L or non-HDL-C ≥2.6 mmol/L on non-HDL-C ≥2.4 mmol/L on maximally tolerated statin dose maximally tolerated statin dose† non-HDL-C ≥3.2 mmol/L YES YES YES Discuss intensification of Discuss add-on therapy with patient: Evaluate reduction in CVD risk vs. cost/access and side effects therapy with patient ADD-ON INTENSIFICATION ADD-ON Ezetimibe first-line Ezetimibe or (BAS° as alternative -PCSK9 inhibitor Refer to Figure 3 add-on to other drugs) Monitor NO NO · response to add-on lipid-lowering Rx · healthy behaviour modifications · response to statin Rx



#### Patients with Atherosclerotic Cardiovascular Disease (ASCVD) Receiving maximally tolerated statin dose If LDL-C is ≥1.8 mmol/L or if ApoB ≥0.70 g/L\*\* or If TG is ≥1.5 to 5.6 mmol/L if non-HDL-C >2.4 mmol/L Consider LDL-C >2.2 mmol/L or LDL-C 1.8-2.2 mmol/L or Icosapent ethyl 2000 mg BID† ApoB >0.80 g/L or ApoB 0.70-0.80 g/L or non-HDL-C >2.9 mmol/L or non-HDL-C 2.4-2.9 mmol/L high PCSK9i benefit patient\* \*May also be considered for patients without ASCVD but with DM requiring medication treatment in patient ≥50 years of age, and ≥1 additional CV risk factor (from REDUCE-IT169): men ≥55 y and women ≥65 y; · cigarette smoker or stopped smoking within 3 months: hypertension (≥140 mmHg systolic OR ≥90 mmHg diastolic) or Consider Consider HDL-C ≤1.04 mmol/L for men or ≤1.3 mmol/L for women; ezetimibe ± PCSK9 inhibitor PCSK9 inhibitor ± ezetimibe hsCRP >3.0 mg/L; Renal dysfunction: eGFR >30 and <60 mL/min;</li> · Retinopathy: \*Patients shown to derive largest benefit form intensification of statin therapy with PCSK9 inhibitor therapy are identified in Table 3.

"At low levels of LDL-C or non-HDL-C, measurement of apoB is more accurate than other markers.

#### Cardiovascular Risk Reduction with Icosapent Ethyl for Hypertriglyceridemia

Micro- or macroalbuminuria;

ABI <0.9 without symptoms of intermittent claudication)</li>

Deepak L. Bhatt, M.D., M.P.H., P. Gabriel Steg, M.D., Michael Miller, M.D., Eliot A. Brinton, M.D., Terry A. Jacobson, M.D., Steven B. Ketchum, Ph.D., Ralph T. Doyle, Jr., B.A., Rebecca A. Juliano, Ph.D., Lixia Jiao, Ph.D., Craig Granowitz, M.D., Ph.D., Jean-Claude Tardif, M.D., and Christie M. Ballantyne, M.D. for the REDUCE-IT Investigators\*

Article Figures/Media Metrics January 3, 2019 N Engl J Med 2019; 380:11-22 DOI: 10.1056/NEJMoa1812792 38 References 1906 Citing Articles Letters 7 Comments



# PEER simplified lipid guideline 2023 update

Prevention and management of cardiovascular disease in primary care

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#### **Abstract**

**Objective** To update the 2015 clinical practice guideline and provide a simplified approach to lipid management in the prevention of cardiovascular disease (CVD) for primary care.

### **PEER Simplified Lipid Guideline 2023: Summary**

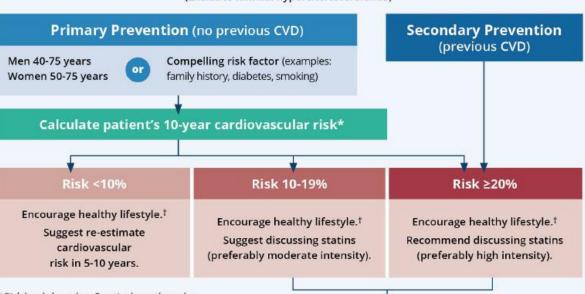
Simplified approach

Shared decision making

Reduce unnecessary testing

### **Treatment Algorithm**

(Excludes familial hypercholesterolemia)



- Risk levels based on Framingham, the only 10-year calculator validated in Canada.
- † Lifestyle includes smoking cessation, physical activity and the Mediterranean diet

CVD = cardiovascular disease

EPA = eicosapentaenoic acid

PCSK9 = proprotein convertase subtilisin-kexin type 9

Statin Intensity				
Statin (mg)	Low	Moderate	High	
Atorvastatin	5	10-20	40-80	
Pravastatin	10-20	40-80	1070	
Rosuvastatin	2.5	5-10	20-40	
Simvastatin	5-10	20-40	0.70	

#### Statin initiated?

 Suggest re-estimating cardiovascular risk in 5-10 years, sooner if risk factors change.

- No repeat lipid testing.
- No baseline creatine kinase or alanine transaminase unless clinically indicated.

For secondary prevention, if additional cardiovascular risk reduction is desired beyond maximum statin dose:

- · Recommend discussing ezetimibe or PCSK9 inhibitors.
  - Due to adverse events, suggest EPA ethyl ester (icosapent) only after ezetimibe or PCSK9 inhibitor considered.

# An assessment by the Statin Muscle Safety Task Force: 2014 update

# Statin intolerance

# Box 1 Factors that may increase the risk of statin induced myopathy

Advanced age (>80 years old)

Female sex

Low body mass index

Multisystem diseases (for example, diabetes mellitus)

Diseases affecting kidney or liver function

Hypothyroidism (untreated)

Drug interactions, especially with drugs that are inhibitors or substrates of the cytochrome P450 pathway (for example, fibrates, nicotinic acid, calcium channel blockers, ciclosporin, amiodarone, thiazolidinediones, macrolide antibiotics, azole antifungals, protease inhibitors, warfarin)

Vigorous exercise

Excess alcohol

Intercurrent infections

Major surgery or trauma

Diet (excessive grapefruit or cranberry juice)

Genetic factors (for example, polymorphisms of the cytochrome P450 isoenzymes or drug transporters, inherited defects of muscle metabolism, traits that affect oxidative metabolism of fatty acids)

- new-onset or increased symptoms of myalgia (muscle aches, stiffness, cramping, soreness, and tenderness) that were unassociated with recent exercise;
- · symptoms that persisted for at least 2 weeks;
- symptoms that resolved within 2 weeks of stopping the study drug; and
- symptoms that reoccurred within 4 weeks of restarting the medication.

### An assessment by the Statin Muscle Safety Task Force: 2014 update

# Statin intolerance

#### **Table 1** Spectrum of statin-associated muscle adverse events

- Myalgia—unexplained muscle discomfort often described as "flu-like" symptoms with normal CK level. The spectrum of myalgia complaints includes:
  - Muscle aches;
  - Muscle soreness;
  - Muscle stiffness;
  - Muscle tenderness; and
  - Muscle cramps with or shortly after exercise (not nocturnal cramping).
- Myopathy—muscle weakness (not attributed to pain and not necessarily associated with elevated CK).
- Myositis—muscle inflammation
- Myonecrosis—muscle enzyme elevations or hyperCKemia
  - Mild >3-fold greater than baseline untreated CK levels or normative upper limit that are adjusted for age, race, and sex.
  - Moderate ≥10-fold greater than untreated baseline CK levels or normative upper limit that are adjusted for age, race, and sex.
  - Severe ≥50-fold above baseline CK levels or normative upper limit that are adjusted for age, race, and sex.
- Myonecrosis with myoglobinuria or acute renal failure (increase in serum creatinine ≥0.5 mg/dL (clinical rhabdomyolysis).

CK, creatine kinase.

Rheumatology International (2023) 43:383–390 https://doi.org/10.1007/s00296-022-05230-0



#### CASE BASED REVIEW



Challenges in the diagnosis and management of immune-mediated necrotising myopathy (IMNM) in a patient on long-term statins

# Dyslipidemia in the elderly

Considerations in patients older than 75 y

- 15. For primary prevention in patients older than 75 y, we recommend against lipid testing and the assessment of risk using a CVD risk calculator
- 16. We **suggest against** the routine initiation of statin therapy for primary prevention in patients older than 75 y. However, it may be reasonable to discuss the benefits and risks of statin therapy for primary prevention in some patients older than 75 y whose overall health status is good
- 17. In patients older than 75 y who have had a cardiovascular event, we **recommend** clinicians discuss the benefits and risks with patients and encourage the initiation of statin therapy
- 18. In patients already taking and tolerating a statin, we recommend against stopping the statin or reducing the dose just because patients have aged beyond 75 y
- 19. We **recommend against** altering statin prescribing for cognitive concerns

# Dyslipidemia in the elderly

**Table 2.**Ongoing Studies of Statin Therapy in the Elderly

	SCOPE-RCT	SITE	STAREE
Country	South Korea	France	Australia
Age	>75 years	>75 years	>70 years
Type of prevention	Primary	Primary	Primary
Study Arms	Moderate dose vs high dose statin	D/C statin vs not D/C statin	Atorvastatin 40 mg/day vs placebo
Primary endpoint	Statin-Associated Muscle Symptoms	Incremental cost per QALY gained, mortality	Death, dementia or disability
Secondary endpoints	Fatal and nonfatal CV events	New events: cardiovascular, cognitive, diabetes	Fatal and nonfatal CV events, diabetes, dementia, hospitalization, QALY, cost- effectiveness
Duration	6 years	3 years	7 years
Number of enrollees	2,234	2,430	18,000
Year of completion	2024	2021	2023

# Carotid artery disease

The clock is ticking!



# Carotid Artery disease

## Imaging modality

- Carotid doppler (CD), CT-A or MR-A
- If intervention considered based on CD, need further imaging

### Intervention

- CEA > CAS
- CAS may be considered in patients who are poor surgical candidates



# Carotid Artery disease

### Indication for revascularization

- Symptomatic event with <u>ipsilateral</u> carotid artery stenosis
- Recommended: Men with 50-99% and women with 70-99%
- May be considered: Women 50-69%

### Timing

- As early as possible\*\* and ideally 14 days from event
- Men 50-69%: greatest benefit within 14 days from event, attenuated beyond that time window



# Carotid Artery disease

Asymptomatic or remotely symptomatic

- Aggressive medical management
- Highly selected patients may be considered for intervention, for 60-99% stenosis
  - Acceptable surgical risk, >5yr life expectancy
  - Women 60-99% unclear benefit
  - Less than 3% risk of peri-op morbidity and mortality (surgeon)
  - Significant improvement in risk factor modification since trials for asymptomatic stenosis interventions
- CEA > CAS, though CAS if not operative candidate

#### **Screening for Carotid Artery Stenosis**

Carotid artery stenosis, the buildup of plaque in the major arteries of the neck, can lead to stroke and death. Medications to treat risk factors such as high cholesterol, high blood pressure, and diabetes can reduce the risk of these poor outcomes.





#### **Population**

Adults without a history of transient ischemic attack, stroke, or other neurologic signs or symptoms



#### USPSTF recommendation

The USPSTF recommends against screening for asymptomatic carotid artery stenosis in the general adult population.

Asymptomatic Carotid stenosis

Choosing Wisely®

# Screening for Carotid Artery Stenosis in Asymptomatic Adult Patients

#### Recommendation

Don't screen for carotid artery stenosis (CAS) in asymptomatic adult patients.

- There is good evidence that for adult patients with no symptoms of carotid artery stenosis, the harms of screening outweigh the benefits.
- · Screening could lead to non-indicated surgeries that result in serious harms, including death, stroke and myocardial infarction.



### Atrial fibrillation with CVA/TIA

- Oral anticoagulation is strongly recommended over antiplatelet
- DOAC preferred over warfarin
- If already on warfarin, with INR time in range >70%, reasonable to continue. \*patient preference
- Stroke/TIA while on anticoagulation
  - Adherence
  - Confirm dosing / INR in control
  - Medication interactions?
  - Other potential stroke etiologies?
  - Risk factor modification



## References

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### **OBESITY** IN ADULTS

A clinical practice guideline



**BMI IS NOT AN ACCURATE** TOOL FOR IDENTIFYING OBESITY-RELATED COMPLICATIONS

Obesity comp body fat impai Effects:

▼ health

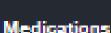
People with o experience v and stigma

Weight blas obesity do not or are not coop



#### Psychological 8 2 2

- Cognitive approach to behaviour change
- Manage sleep, time and stress
- Psychotherapy if appropriate



 For weight loss and to help maintain weight 1033

#### Bariatric surgery

 Surgeon-patient discussion.

#### THE PATIENT JOURNEY IN OBESIT



### PERMISSION

"Would it be all right if we discussed your weight?"

Asking permission

- · Shows compassion and empathy
- · Builds patient-provider trust



**Psychological** 

Manage sleep,

time and stress

· Psychotherapy if appropriate

· Cognitive approach

to behaviour change









#### Medications

#### For weight loss and to help maintain weight

#### Bariatric surgery

 Surgeon-patient discussion



#### ASSESS THEIR STORY

- · Goals that matter to the patient Obesity classification
- (BMI and waist circumference)
- Disease severity (Edmonton Obesity Staging System)

Treating the root causes of weight gain is the foundation of obesity management

Focus on patient-centred health outcomes weight loss alone



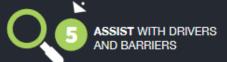






AGREE ON GOALS

Collaborate on a personalized, sustainable action plan



### **Obesity in adults: a clinical** practice guideline





# OBES

A clinical pr









Treating the root causes of weight g is the foundation o obesity management

Focus on patient-centred health outcomes versus weight loss alone

#### Key messages for health practitioners

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- Pharmacological treatments are an effective and scalable approach to treating obesity. As with any chronic disease, such as type 2 diabetes (T2DM) or hypertension, pharmacotherapy is an important pillar in the management of obesity.
- The focus of obesity management should be the improvement of health parameters (metabolic, mechanical, mental, and/or quality of life [QoL]), not solely weight reduction, and should include outcomes that the patient identifies as important. Obesity is defined by body mass index (BMI) in clinical trials, which itself does not adequately reflect the burden of
- There are four medications indicated for long-term obesity management in Canada as adjuncts
  to health-behaviour changes: liraglutide (Saxenda®), naltrexone/bupropion (Contrave®) in a
  combination tablet, orlistat (Xenical®) and semaglutide (Wegovy®). All four medications are
  effective in producing clinically significant weight loss and health benefits greater than placebo
  over a duration of at least one year.
- The individual response to pharmacotherapy for obesity management is neterogeneous.
   Efficacy (both for weight and management of obesity-related health issues), mechanism of action, safety, potential side effects/tolerability, contraindications, medication interactions, mode of administration and cost are important considerations in choosing the most appropriate obesity pharmacotherapy.
- Obesity medications are intended as part of a long-term treatment strategy. Clinical trials of pharmacotherapy for obesity management consistently demonstrate regain of weight when treatment is stopped.
- Medications that are not approved as pharmacotherapy for obesity management should not be used for this purpose.

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nical

# Stroke in Women

#### Stroke risk factors in women

#### Common risk factors

Less favorable for women

Comparison of women	to men
Prevalence	
Hypertension in older age <sup>19</sup> Migraine <sup>58</sup> Obesity <sup>64,65</sup>	Greater (women > men)
Association with stroke risk	*
Hypertension <sup>29-31</sup> Diabetes mellitus <sup>29,37</sup> Atrial fibrillation (AF) <sup>39,40</sup> Migraine <sup>63</sup> Smoking <sup>29,69</sup>	Greater (women > men)
Treatment rate	
Oral anticoagulant therapy <sup>42,43</sup> Statin therapy <sup>56,57</sup>	Lesser (women < men)
Procedural complication rate	9.
Catheter ablation of AF <sup>45</sup> Left atrial appendage closure <sup>46,47</sup>	Greater (women > men)

#### Women-specific risk factors

Unique to women

Pregnancy & adverse pregnancy outcomes

Hypertensive disorders of pregnancy

Preterm delivery

Gestational diabetes

Exogenous estrogen

Oral contraceptive

Oral postmenopausal hormone therapy

Lifetime endogenous estrogen exposure

Early & late menarche

Early menopause

- Natural menopause
- Surgical menopause (oophorectomy with/without hysterectomy)

Stroke in Women: A Review Focused on Epidemiology, Risk Factors, and Outcomes

#### Stroke outcomes in women

#### Less favorable for women than men



#### Crude mortality4,124

Adjusted mortality | (after adjusting for confounders such as age, stroke severity, pre-stroke status, risk factors including atrial fibrillation):

It means that women's higher mortality is attributable to advanced age, greater stroke severity, worse pre-stroke status, and higher prevalence of atrial fibrillation.

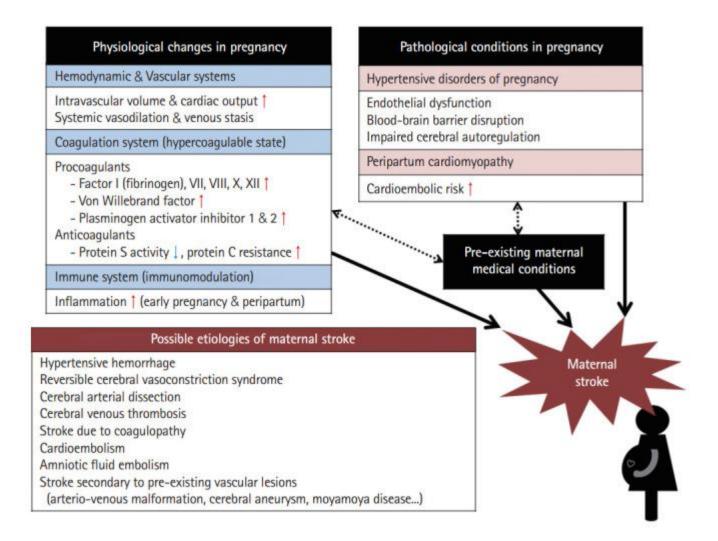
Functional recovery4,10,125-128

Quality of life4,10,125-128

Post-stroke depression 126,129 1

Post-stroke cognitive impairment 130,131

Stroke in Women: A Review Focused on Epidemiology, Risk Factors, and Outcomes



Stroke in Women: A Review Focused on Epidemiology, Risk Factors, and Outcomes