

# NENC Lipids & FH Pathways & Guidelines:- Interpretation and Application

AHSN-NENC QOF Targets  
The Durham Centre  
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# Disclosures

- Research:- Sanofi, Novartis, Daiichi Sankyo
- Collaborative Working Projects :- Amarin, Novartis, GENinCode
- Advisory:- Amgen, Novartis
- Speaker Fees:- Lilly, Sanofi, Daiichi Sankyo, Amarin, Novartis
- Meeting Sponsorship:- NovoNordisk, Lilly, Sanofi, Boehringer Ingelheim, Amarin

# Why develop a Regional Lipid Guideline?

- Historically subregional guidelines (FATS/SLiMS/NEATS)
- Based on NICE
  - Guidelines (CG 71; 181)
  - Technology Appraisals (TA 385;393;394; )
- Increasingly developing regional approach to Lipid management
  - FH Genetic testing
  - Primary Care FH searches
  - Lipid Optimisation
- Useful to have a single regional resource that we can signpost to for issues pertaining to Lipid Management

# What have we learnt?

- Avoided “too much detail” = Secondary Care Guideline
- Single Sheet Summary – stick it on a wall
- Flowcharts – more of the same
- Colour coded sections
  - Ease of access
  - Ease of update
- Developed guidelines to cover the gaps
- Incorporated NICE approved AAC Pathways
- Googleable

# Contact with Reality

- Feedback mostly positive
- “Too Complicated”; “Don’t have time to wade through that”
- TA 694; TA 733; TA 805
  
- Added :-
- Problem based Signposting to relevant section
- Updated Secondary Prevention beyond standard therapy
- Frailty Guidance
- Inclisiran FAQ
- 2023 Version with updates AAC pathway awaiting NTAG approval

# Northern England Evaluation and Lipid Intensification guideline



Section Description	Primary Secondary prevention	Statin Intolerance	Severe Hypercholesterolaemia	Severe Hypertriglyceridaemia	Pregnancy	FH in Children and Young People	Supplementary information
Section Guideline			<a href="#">Simon Broome criteria for diagnosis of Familial Hypercholesterolaemia</a>		<a href="#">Lipid management and medication issues in pregnancy</a>		<a href="#">Frailty Guidelines</a> <a href="#">Common drug interactions</a> <a href="#">Lipid Clinic referral criteria</a> <a href="#">Lipoprotein (a)</a> <a href="#">Regional Lipid clinics</a> <a href="#">Inclisiran FAQs</a>
Flow charts	<a href="#">National Guidance for lipid management</a>  <a href="#">Secondary prevention treatment beyond standard therapy</a>	<a href="#">Statin intolerance flow chart</a>	<a href="#">Assessment pathway</a>	<a href="#">Assessment pathway</a>		<a href="#">Assessment pathway</a>	



## Result based reference guide and link to relevant guideline detail

### Alternative lipid lowering options - beyond Atorvastatin

Medication	Notes	When to prescribe	Section link
Rosuvastatin	Water soluble statin	First line alternative medication	<a href="#">Red section</a>
Ezetimibe		Additional if non HDLc above target or alternative if statin intolerant	
Ezetimibe + Bempedoic acid	Available as a single tablet	Consider in non HDLc above target on Ezetimibe monotherapy	
Evolocumab/ Alirocumab	Secondary care prescription only	Secondary prevention: single event LDLc>4mmol/l; two or more events LDLc>3.5mmol/l; Familial Hypercholesterolaemia LDLc>5mmol/l	<a href="#">Green section</a>
Inclisiran	Primary or Secondary care prescription	Secondary prevention; LDLc ≥2.6mmol/l	

### Guidance related to Interpretation of lipid profile

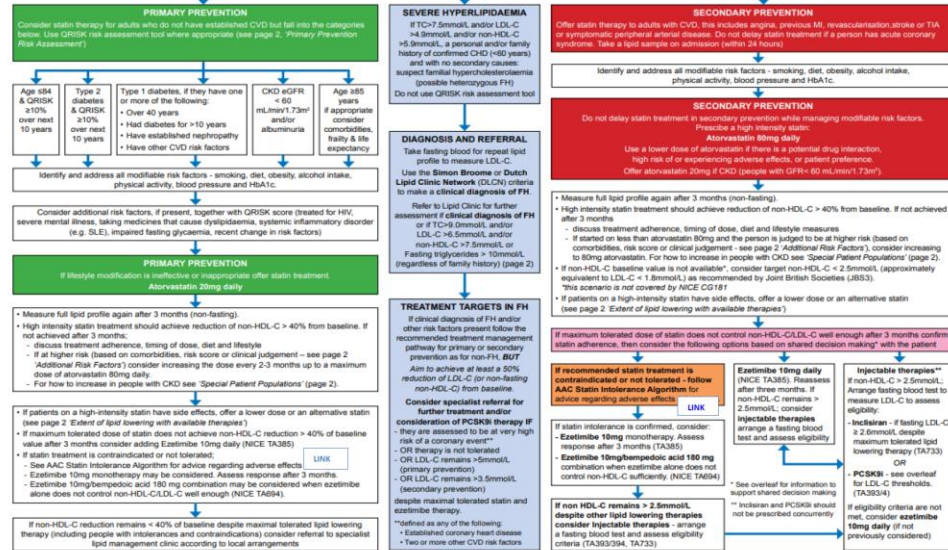
Parameter	Value	Clinical scenario	Section link
non HDLc	>5.9mmol/l	Assessment for severe hypercholesterolaemia	<a href="#">Peach section</a>
	≥2.5mmol/l	Need to intensify lipid lowering in secondary prevention patients	<a href="#">Green section</a>
Triglycerides	>5mmol/l	Measure fasting lipid profile	<a href="#">Yellow section</a>
Apolipoprotein B100		Measured if fasting triglycerides >10mmol/l	
Total cholesterol	>7.5mmol/l	+ Personal or first degree relative history of IHD/CVD event <60yrs <b>OR</b> + First /second degree relative with total cholesterol >7.5mmol/l or LDL>4.9mmol/l <b>Consider Familial Hypercholesterolaemia</b>	<a href="#">Peach section</a>
LDLc (fasting lipid profile)	>4.9mmol/l		
Lipoprotein (a)		Consider measuring if personal or first degree relative with history of IHD/CVD <60yrs or lipoprotein (a)>200nmol/l in a first degree relative	<a href="#">Blue section</a>

# AAC NICE Approved 1ry/2ry Pathway

## Summary of National Guidance for Lipid Management for Primary and Secondary Prevention of CVD

### INITIAL CONSIDERATIONS:

- Measure non-fasting full lipid profile (total cholesterol, HDL-C, non-HDL-C, triglycerides) and HbA1c as part of an initial baseline assessment.
- Consider secondary causes of hyperlipidaemia and manage as needed.
- Ensure appropriate baseline and follow up tests as detailed on page 2. Measure BMI.
- Identify and exclude people with contraindications/drug interactions.
- If non-fasting triglyceride above 4.5mmol/L see page 2.



**MANAGEMENT**

This guidance applies to new patients and may also be taken into consideration for those already on statins at their annual review. If 40% reduction of non-HDL-C not achieved, offer high intensity statin. Discuss with people who are stable on a low- or medium-intensity statin the likely benefits and potential risk of side effects if changed to a high-intensity statin when they have a medication review and agree with the person whether a change is needed.

Ezetimibe, alirocumab, evolocumab or inclisiran can be added when patients' LDL-C levels are not lowered enough with the maximally tolerated dose of statin. Bempesic acid with ezetimibe is an option when statins are contraindicated or not tolerated, and when ezetimibe alone does not control LDL-C well enough. Do not offer a fibrate, nicotinic acid, bile acid binder or omega-3 fatty acids alone or in combination with statin, for the prevention of CVD (Check NICE CG181 for exceptions).

**EXTENT OF LIPID LOWERING WITH AVAILABLE THERAPIES**

Statin dose regimen	Approximate reduction in LDL-C				
	0	10	20	40	80
Fluvastatin		21%	27%	33%	37%
Pravastatin		20%	24%	29%	42%
Simvastatin	27%	24%	26%	37%	42%
Atorvastatin	37%	41%	46%	56%	62%
Rosuvastatin	38%	43%	48%	59%	62%
Atorvastatin + Ezetimibe 10mg	49%	52%	57%	67%	67%

Low intensity statins will produce an LDL-C reduction of 20-30%.  
Medium intensity statins will produce an LDL-C reduction of 31-40%.  
High intensity statins will produce an LDL-C reduction above 40%.  
Simvastatin 80mg is not recommended due to risk of muscle toxicity.

**PRIMARY PREVENTION RISK ASSESSMENT**

QRISK3 is the current version of the QRISK calculator. <https://qrisk3.org/>

- Do not use this risk assessment tool for people with established CVD or those who are at high risk of developing CVD because of FH or other inherited disorders of lipid metabolism.
- Do not use a risk assessment tool to assess CVD risk in people with type 1 diabetes, or eGFR less than 60 mL/min/1.73 m<sup>2</sup> and/or albuminuria.
- Consider people aged ≥ 85 at increased risk of CVD because of age alone particularly people who smoke or have raised BP.

**Additional Risk Factors**

Note, standard CVD risk scores including QRISK may underestimate risk in people who have additional risk because of underlying medical conditions or treatments. These groups include the following groups of people:

- severe obesity (BMI ≥ 40kg/m<sup>2</sup>) increases CVD risk
- treated for HIV
- serious mental health problems
- taking medicines that can cause dyslipidaemia such as antipsychotic medication, corticosteroids or immunosuppressant drugs
- autoimmune disorders such as ... and other systemic inflammatory disorders
- non-diabetic hyperglycaemia
- significant hypertriglyceridaemia (fasting triglycerides 4.5-9.9mmol/L)
- recent risk factor changes e.g. quit smoking, BP or lipid treatment
- Consider socio-economic status as an additional factor contributing to CVD risk.

**IF QRISK < 10% over the next 10 years - Give lifestyle advice and ensure regular review of CVD risk in line with guidance.**

**SPECIAL PATIENT POPULATIONS**

**Type 1 Diabetes**

While NICE recommends offering statins to patients with Type 1 diabetes as detailed in the algorithm, it also states to consider statins in all adults with type 1 diabetes.

**Chronic Kidney Disease**

Offer atorvastatin 20mg for the primary or secondary prevention of CVD to people with CKD (eGFR less than 60 mL/min/1.73m<sup>2</sup> and/or albuminuria). Increase the dose if a greater than 40% reduction in non-HDL-C is not achieved and eGFR is 30 mL/min/1.73m<sup>2</sup> or more. Agree the use of higher doses with a renal specialist if eGFR is less than 30 mL/min/1.73m<sup>2</sup>.

**MONITORING**

Repeat full lipid profile in non-fasting. Measure liver transaminase within 3 months of starting treatment and then within 3 months of every additional up titration and then again at 12 months, but not again unless clinically indicated.

**TRIGLYCERIDES**

Refer to lipid clinic for urgent specialist review if a result of excess alcohol or poor glycaemic control. At risk of acute pancreatitis. Repeat the TG measurement with a fasting test after an interval of 3 days, but within 2 weeks) and review for potential secondary causes of hypertriglyceridaemia. Seek specialist advice if the TG concentration remains > 10mmol/L. At risk of acute pancreatitis.

**STATIN INTOLERANCE**

Statin intolerance is defined as the presence of clinically significant adverse effects from statin therapy that are considered to represent an unacceptable risk to the patient or that may result in adherence to therapy being compromised. For people who are intolerant of the recommended statin treatment see the NICE AAC statin intolerance algorithm, available on the NICE AAC page.

**ACCELERATED ACCESS COLLABORATIVE** **NHS**



# Basically.....

- Measure full fasted lipid profile and assess risk
- Address Secondary causes / Lifestyle issues
- Primary Prevention
  - Atorvastatin 20 mg OD
  - Aim for 40% reduction from baseline NonHDL-C
  - Up titrate and add Ezetimibe if not to target
- Secondary Prevention
  - Atorvastatin 80 mg OD (start with 20mg if eGFR < 60 or interaction)
  - Aim for NonHDL-C < 2.5 mmol/L [or LDL < 1.8 mmol/L]
  - Consider additional therapy if eligible

# Grouping of Statins

Dose mg / day	Reduction in LDL				
	5	10	20	40	80
Fluvastatin			21%	27%	33%
Pravastatin		20%	24%	29%	
Simvastatin		27%	32%	37%	42% MRHA
Atorvastatin		37%	43%	49%	55%
Rosuvastatin	38%	43%	48%	53%	
Atorvastatin + Ezetimibe		52%	54%	57%	61%

# NICE QOF Indicators

**CHOL 001** = 14 Points                      70-95%

% CHD, PAD, CVA/TIA or CKD prescribed statin or alternative lipid lowering therapy

**CHOL 002** = 16 Points                      20-35%

% CHD, PAD, CVA/TIA with NonHDL < 2.5 or LDL < 1.8 mmol/l

**DM022** = 4 Points                              50-90%

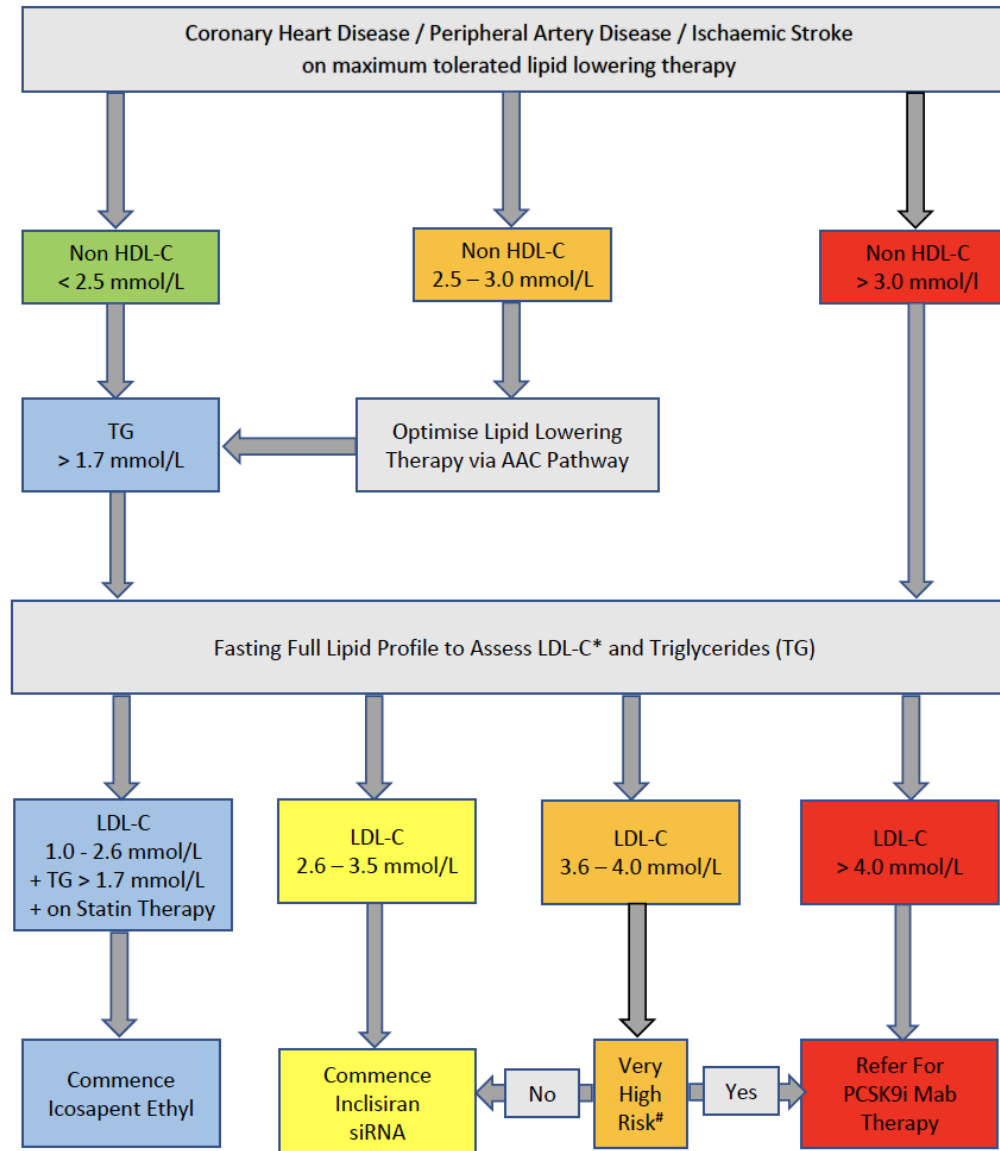
% Patients with DM >40yr without CVD or mod/severe Frailty prescribed statin

(exclude T2DM with QRisk < 10%)

**DM023** = 2 Points                              50-90%

% Patients with DM with CVD (excl haemorrhagic stroke) prescribed statin

# Secondary Prevention Beyond Standard Therapy



# Measure Fasting LDL if CVD + NonHDL not to target

- Non HDL < 2.5 mmol/L
  - To Target; Consider Vazkepa if TG > 1.7mmol/L and LDL 1.0 – 2-6 mmol/L
- Non HDL 2.5 – 3.0 mmol/L
  - Optimise Statin / Ezetimibe therapy
  - Consider Vazkepa if TG > 1.7mmol/L and LDL 1.0 – 2-6 mmol/L
- NonHDL ≥ 3.0 mmol/L
  - LDL > 4.0 mmol/L → Refer for PCSK9i MaB
  - LDL > 3.5 mmol/L → Refer for PCSK9i MaB if Very High Risk
  - Start Inclisiran if High Risk
  - LDL > 2.6 mmol/L → Start Inclisiran
  - LDL = 1.0 – 2.6 mmol/L → Consider Vazkepa if TG > 1.7mmol/L



# Statin Intolerance

- Affects up to 10% of patients
- Significant nocebo effect :- reinforced by health professionals
- More likely in certain subgroups (female, hypothyroid, alcohol excess)
- Ask, reassure, advise
- Check Creatinine Kinase if symptoms suggestive of myalgia
  
- Aim for max tolerated dose (low better than no)
- Re-try at lower dose after 4-6 week wash out period
- Switch to Rosuvastatin 10 mg
- Consider lower intensity statin
- Use Ezetimibe as statin sparing agent or as monotherapy
- Consider addition of Bempedoic acid to Ezetimibe if NonHDL

# Additional Sections

- Severe Hypercholesterolaemia
  - Simon Broome criteria
  - Pathway
- Severe Hypertriglyceridaemia
  - Use of Apo B100 alongside Full Fasted Lipid Profile
- Pregnancy
- FH in Children and Young People
- Supplementary
  - Frailty
  - Common Drug Interactions
  - Lipid Clinic Referral Criteria
  - Lipoprotein (a)
  - Regional Lipid Clinics
  - Inclisiran FAQ



# FH Diagnosis

Simon Broome Criteria

1. Total Chol > 7.5 or LDL > 4.9
2. Tendon Xanthoma
3. Family history of Premature MI\*
4. Family history of hypercholesterolaemia@

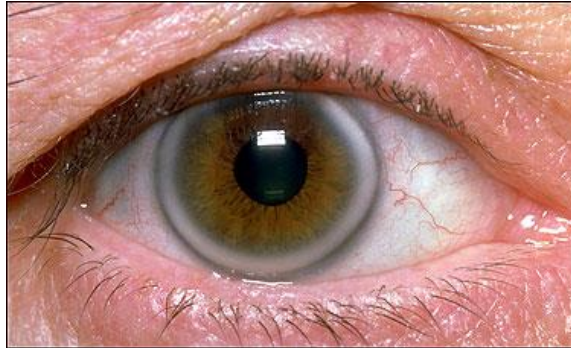
\* <60 yr if 1<sup>o</sup> Rel or < 50 yr 2<sup>o</sup> Rel

@ Total Chol > 7.5 or LDL > 4.9 in 1<sup>o</sup> / 2<sup>o</sup> Rel

1+2 = Definite FH

1+ 3 or 4 = Probable FH

Dutch Lipid Clinic Network score  $\geq 6$  used to determine whether or not to do genetic testing



# Referring FH

- Use Simon Broom Criteria to determine whether “could” have FH
  - 1 in 250 patients have FH = 0.4%
  - Population with Chol > 7.5 mmol/l = 4%
  - Hypercholesterolaemia common in women > 50 yrs
- Don't use DLCS; determines whether eligible for genetic testing
  - LDL > 6.4 mmol/L + family history premature MI / LDL > 5.5 mmol/L
  - LDL > 4.9 + personal MI < 60 yr + family history
  - Bilateral TX
  - TG < 2.3 mmol/L
- Refer via A+G if
  - > 30 yr with Chol > 9.0 mmol/L or NonHDL > 7.5 mmol/l or LDL > 6.5 mmol/L
  - < 30 yr with Chol > 7.5 mmol/L or NonHDL > 6.0 mmol/l or LDL > 4.9 mmol/L

# Criteria for Lipid Clinic Referral

- All Regional Lipid Clinics offer A+G
- Refer if
- Clinical Diagnosis of FH according to SB criteria
  - Relatives who require genetic testing (FH Nurses)
  - Children with FH (Paediatric Clinic)
- Total Chol > 9.0 mmol/L or NonHDL-C > 7.5 mmol/L
- TG > 10 mmol/L (Urgently if > 20 mmol/L)
- Other Lipid disorders (Remnant / FCH / Familial HyperTG)
- Patients with CVD and NonHDL-C > 4 mmol/L
- Patients for PCSK9i Mab therapy

# Where can I find it?

- NTAG Website

<https://ntag.nhs.uk/wp-content/uploads/2022/09/NEELI-edit-v2022.2-FINAL-NTAG.pdf>

- 2023 update awaiting NTAG approval
- Update existing weblinks

Any Questions?

