Subclinical Hypothyroidism in Primary Care

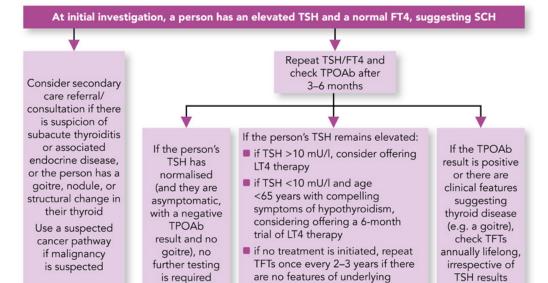
Medscape UK X Guidelines Primary Care Hacks

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What is Subclinical Hypothyroidism?

- SCH is commonly observed in primary care and affects up to 10% of the population, with highest prevalence in women and older people[1]
- SCH is a biochemical state where TSH is elevated (usually <10 mIU/l) but with normal
- Many individuals with SCH are asymptomatic. A recent cross-sectional study found that SCH did not confer a symptom burden particularly in people age >65 years, and treatment was not recommended;[3] **treat** the person, not the number!
- SCH often normalises without intervention within 5 years. 44 The risk of progression from SCH to overt hypothyroidism is around 2–5% per year; the presence of TPOAb and higher TSH levels increase this risk

Managing Subclinical Hypothyroidism^[5]



Managing Subclinical Hypothyroidism

- Most individuals with SCH do not require treatment; a recently published clinical practice guideline gave a strong recommendation against thyroxine for all adults with SCH (at least 2 consecutive tests with or without mild-to -moderate symptoms)^[6]
 - o important exceptions are women who are trying to conceive or individuals with TSH >20
 - o additionally, this recommendation may also not apply to individuals with severe symptoms or young adults aged <30 years.

Hypothyroidism, SCH, and Pregnancy

- Hypothyroidism is common in women of childbearing age and occurs in 2-3% of all pregnancies^{[7}
 - o suboptimal thyroid function (including SCH, although evidence is weak^[8]) in pregnancy is associated with adverse outcomes
 - o refer all women with SCH or overt hypothyroidism who are planning a pregnancy, or who are pregnant, to endocrinology
 - ocheck TFT results before conception if possible; if levels are not within the euthyroid range, advise delaying conception and using contraception until stabilised on LT4 treatment
 - ensure the patient understands that there is likely to be an increased need for LT4 treatment during pregnancy, and LT4 dose must be adjusted as early as possible in pregnancy to reduce the chance of adverse outcomes
 - discuss dosing urgently with endocrinology; as a rule, hypothyroid women require a 25-50% increase in thyroxine dose on conception; increase thyroxine dose by 25 mcg if usual dose ≤100 mcg, or by 50 mcg if ≥100 mcg
 - early pregnancy vomiting, antacids, and iron therapy may also reduce thyroxine bioavailability
 - QIA: as a clinical team, discuss preconception counselling strategies for women with SCH or hypothyroidism. Consider using the British Thyroid Foundation leaflet, Your guide to pregnancy and fertility in thyroid disorders.

Hypothyroidism in Primary Care: Practical Pointers Screening

There is no evidence for population screening for thyroid disease. However, offer an annual TSH test to those:

thyroid disease

- owith Down's syndrome or Turner's syndrome
- otaking lithium, amiodarone, or rifampicin
- owho have received radioiodine treatment or neck radiotherapy
- who have undergone subtotal thyroidectomy
- owith T1D or PAI (Addison's disease)
 - $^{\circ}$ the British Thyroid Association recommends that people living with T2D should be screened for thyroid disease at diagnosis only.

Initiation and Titration of LT4[10]

- The BNF recommends for those aged 18–49 years, initially commence 50–100 mcg od taken on an **empty stomach at least 30–60 minutes before food, caffeine-containing liquids, or other drugs** (particularly antacids, PPIs,
 - oadjust dose 25-50 mcg every 3-4 weeks according to response; usual maintenance dose is 100-200 mcg od
- The BNF recommends for those aged >50 years and those with cardiac disease or severe hypothyroidism, start 25 mcg od and adjust by 25 mcg every 4 weeks according to response; usual maintenance dose is
- Aim to resolve symptoms and signs of hypothyroidism and to maintain serum TSH and FT4 levels within or close to the normal reference range
- A longitudinal study found that long-term outcomes remain similar across a wide range of 'normal' TSH levels,^[11] supporting **individualisation with LT4** dosing regimens and TSH targets
 - of fine-tuning of TSH levels inside the reference range can therefore be considered for some individuals. Consider aiming for the lower half of the reference range for those whose symptoms persist
- Note: deterioration of symptoms with LT4 treatment can be suggestive of coexisting PAI and can lead to adrenal crisis if steroid replacement is not introduced[[]
 - 25% of those with PAI have hypothyroidism
- There is no high-quality evidence for the use of desiccated thyroid extract, triiodothyronine, or dietary supplementation for hypothyroidism.

Treatment-refractory Hypothyroidism[13]

- Explore non-compliance with LT4 and reinforce correct administration
- Exclude drug interactions for both prescribed and over-the-counter medications
- GI conditions causing malabsorption (e.g. coeliac disease, *H. pylori* gastritis, inflammatory bowel disease) may reduce absorption of LT4 in the gut ocheck bloods to screen for malabsorption—FBC; vitamin B12 and folate; ferritin; calcium; albumin; and coeliac serology
- Weight gain and pregnancy may increase LT4 requirements.

- Medscape. Hypothyroidism. Available at: emedicine.medscape.com/article/122393-overview (accessed 6 June 2023).
- Biondi B, Cappola A, Cooper D. Subclinical hypothyroidism; a review. J Am Med Soc 2019; 322 (2): 153–160.
- McCahon D, Haque M, Parle J et al. Subclinical thyroid dysfunction symptoms in older adults: cross-sectional study in UK primary care. Br J Gen Pract 2020; 70 (692): 3.
- Meyerovitch J, Rotman Pikielny P, Sherf M et al. Serum thyrotropin measurements in the community: five-year follow-up in a large network of primary care physicians. Arch Intern Med 2007; 167 (4): 1533-153.
- NICE. Hypothyroidism: scenario: subclinical hypothyroidism (non-pregnant). Available at: cks.nice.org.uk/topics/hypothyroidism/management/subclinical- hypothyroidism-non-pregnant (accessed 6 June 2023).
- Bekkering G, Agoritsas T, Lytvyn L et al. Thyroid hormones treatment for subclinical hypothyroidism: a clinical practice guideline. Br Med J 2019; 365: 12006. 6.
- NICE. Hypothyroidism: scenario: preconception or pregnant. Available at: cks.nice.org.uk/topics/hypothyroidism/management/preconception-or-pregnant (accessed 6
- 8.
- Wiles K. Management for women with subclinical hypothyroidism in pregnancy. *Drug Ther Bull* 2019; **57**: 22–26. The Association of Clinical Biochemistry. British Thyroid Association, British Thyroid Foundation. *UK guidelines for the use of thyroid function tests*. British Thyroid 9. Association, 2006. Available at: british-thyroid-association.org/sandbox/bta2016/uk guidelines for the use of thyroid function tests.pdf (accessed 6 June 2023).
- British National Formulary. Levothyroxine sodium. Available at: bnf.nice.org.uk/drugs/levothyroxine-sodium/#indications-and-dose (accessed 6 June 2023). 10. 11. Thayakaran R, dderley N, Sansbury C et al. Thyroid replacement therapy, thyroid stimulating hormone concentrations, and long term health outcomes in patients with
- hypothyroidism: longitudinal study. *Br Med J* 2019; **366**: l4892. Murray J, Jayarajasingh R, Perros P. Deterioration of symptoms after start of thyroid hormone replacement. *Br Med J* 2001; **323** (7308): 332–333.
- McNally L, Ofiaeli C, Oyibo S. Treatment-refractory hypothyroidism. *Br Med* J 2019; **364**: I1579.

Abbreviations: BNF=British National Formulary; FBC=full blood count; FT4=free thyroxine; GI=gastrointestinal; od=once daily; LT4=levothyroxine; PAI=primary adrenal insufficiency; PPI=proton pump inhibitor; SCH=subclinical hypothyroidism; T1D=type 1 diabetes; T2D=type 2 diabetes; TFT=thyroid function test; TPOAb=thyroid peroxidase antibodies; TSH=thyroid-stimulating hormone; QIA=quality improvement activity.





