

Response to Reviewers Comments

We thank the reviewers for their comments. The necessary changes have been made accordingly.

Reviewer #1

1. No “references” are given in the background section.

Ans: References in the background section have been added.

2. Please use latest updated guidelines to define and treat and mention treatment considerations for subclinical endocrine diseases.

Ans: Updated guidelines have been referred to and cited as follows:

For prediabetes –

- a) McGuire H, Longson D, Adler A, Farmer A, Lewin I. Management of type 2 diabetes in adults: summary of updated NICE guidance. *Bmj*. 2016 Apr 6;353.
- b) Cosentino F, Grant PJ, Aboyans V, Bailey CJ, Ceriello A, Delgado V, Federici M, Filippatos G, Grobbee DE, Hansen TB, Huikuri HV. 2019 ESC Guidelines on diabetes, pre-diabetes, and cardiovascular diseases developed in collaboration with the EASD: The Task Force for diabetes, pre-diabetes, and cardiovascular diseases of the European Society of Cardiology (ESC) and the European Association for the Study of Diabetes (EASD). *European heart journal*. 2020 Jan 7;41(2):255-323.
- c) Jonas DE, Crotty K, Yun JD, Middleton JC, Feltner C, Taylor-Phillips S, Barclay C, Dotson A, Baker C, Balio CP, Voisin CE. Screening for prediabetes and type 2 diabetes: updated evidence report and systematic review for the US Preventive Services Task Force. *JAMA*. 2021 Aug 24;326(8):744-60.
- d) ADA 2022 guidelines. American Diabetes Association. Introduction: Standards of Medical Care in Diabetes—2022. *Diabetes Care*. 2022 Jan;45(Supplement_1):S1-2.

For osteopenia –

- a) Kanis JA, Cooper C, Rizzoli R, Reginster JY. European guidance for the diagnosis and management of osteoporosis in postmenopausal women. *Osteoporosis International*. 2019 Jan;30(1):3-44.
- b) Shoback D, Rosen CJ, Black DM, Cheung AM, Murad MH, Eastell R. Pharmacological management of osteoporosis in postmenopausal women: an Endocrine Society guideline update. *The Journal of Clinical Endocrinology & Metabolism*. 2020 Mar;105(3):587-94.
- c) Arceo-Mendoza RM, Camacho PM. Postmenopausal osteoporosis: latest guidelines. *Endocrinology and Metabolism Clinics*. 2021 Jun 1;50(2):167-78.

For subclinical hypothyroidism –

- a) Guideline NG145 NI. Thyroid disease: assessment and management. (NICE guideline for subclinical hypothyroidism)

- b) Bekkering GE, Agoritsas T, Lytvyn L, Heen AF, Feller M, Moutzouri E, Abdulazeem H, Aertgeerts B, Beecher D, Brito JP, Farhoumand PD. Thyroid hormones treatment for subclinical hypothyroidism: a clinical practice guideline. *Bmj*. 2019 May 14;365.
- c) Poppe K, Bisschop P, Fugazzola L, Minziori G, Unuane D, Weghofer A. 2021 European thyroid association guideline on thyroid disorders prior to and during assisted reproduction. *European thyroid journal*. 2021 Feb 1;9(6):281-95.
- d) Biondi B, Cappola AR. Subclinical hypothyroidism in older individuals. *The Lancet Diabetes & Endocrinology*. 2021 Dec 22.

For MACS –

- a) European Society of Endocrinology Clinical Practice Guideline in collaboration with the European Network for the Study of Adrenal Tumors, 2016
- b) Delivanis DA, Athimulam S, Bancos I. Modern management of mild autonomous cortisol secretion. *Clinical Pharmacology & Therapeutics*. 2019 Dec;106(6):1209-21.
- c) Yozamp N, Vaidya A. Assessment of mild autonomous cortisol secretion among incidentally discovered adrenal masses. *Best Pract Res Clin Endocrinol Metab*. 2021 Jan;35(1):101491. doi: 10.1016/j.beem.2021.101491. Epub 2021 Feb 6. PMID: 33593680.
- d) Prete A, Subramanian A, Bancos I, Chortis V, Tsagarakis S, Lang K, Macech M, Delivanis DA, Pupovac ID, Reimondo G, Marina LV. Cardiometabolic Disease Burden and Steroid Excretion in Benign Adrenal Tumors: A Cross-Sectional Multicenter Study. *Annals of internal medicine*. 2022 Jan 4.

Reviewer #2

1. Although this is a narrative unsystematic review, it is still preferably to outline briefly the method used to identify the literatures included (eg. years, keywords, search methodology used). This can provide readers a good overview of the evidence included in this manuscript.
Ans: The relevant search strategy has been included in the article after the “background” section.

2. Table 3: Unit of blood sugar & HbA1C should be stated. The different definition of AHA, WHO should be referenced as in Table 2 for consistency.
Ans: The definitions have been referenced and the units have been stated.

3. Some typos are noted within the text.
Ans: Corrections have been done.

Company editor-in-chief:

1. Authors are required to provide standard three-line tables
Ans: Tables have been formatted accordingly
2. Authors are advised to apply a new tool, the RCA. RCA is an artificial intelligence technology-based open multidisciplinary citation analysis database. In it, upon obtaining search results from the keywords entered by the author, "Impact Index Per Article" under "Ranked by" should be

selected to find the latest highlight articles, which can then be used to further improve an article under preparation/peer-review/revision

Ans: RCA tool has been used as per suggestion and mentioned in the search strategy as well.