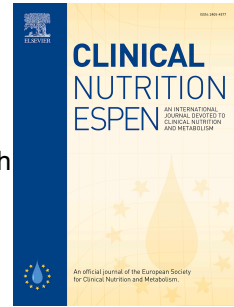


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Nutrigenomics-guided lifestyle intervention programmes: A critical scoping review with directions for future research

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Nutrigenomics-guided lifestyle intervention programmes: A critical scoping review with directions for future research

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Abstract:

Genetic testing is increasingly used in clinical practice to provide personalized information and recommendations about health risks and lifestyle habits at a relatively low cost. Research on the effectiveness of nutrigenomics-guided lifestyle interventions is growing. A scoping review approach was adopted to identify pertinent published studies on nutrigenomics-guided intervention programmes from 2007 to 2023. The review shows that despite the growing interest in nutrigenomics-guided lifestyle interventions, there are still few empirically supported studies, primarily based on developed countries. Furthermore, the findings on the impact of personalised genetic advice are mixed, leaving the field unclear. Existing studies have some empirical strength, contributing to further understanding of the relationship between food and gene expression. However, some limitations that affect the robustness of findings exist, such as a small sample size, insufficient monitoring of the data collection process, and a short follow-up period. Future research needs to address reliability concerns and provide more robust practical evidence.

Keywords:

Nutrigenetics; nutrigenomics; nutrition intervention; gene expression

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3

4 **Abstract:**

5 Genetic testing is increasingly used in clinical practice to provide personalized
6 information and recommendations about health risks and lifestyle habits at a relatively
7 low cost. Research on the effectiveness of nutrigenomics-guided lifestyle interventions
8 is growing. A scoping review approach was adopted to identify pertinent published
9 studies on nutrigenomics-guided intervention programmes from 2007 to 2023. The
10 review shows that despite the growing interest in nutrigenomics-guided lifestyle
11 interventions, there are still few empirically supported studies, primarily based on
12 developed countries. Furthermore, the findings on the impact of personalised genetic
13 advice are mixed, leaving the field unclear. Existing studies have some empirical
14 strength, contributing to further understanding of the relationship between food and
15 gene expression. However, some limitations that affect the robustness of findings exist,
16 such as a small sample size, insufficient monitoring of the data collection process, and
17 a short follow-up period. Future research needs to address reliability concerns and
18 provide more robust practical evidence.

19

20 **Keywords:**

21 Nutrigenetics; nutrigenomics; nutrition intervention; gene expression

22

23 Background:

24 The advancement in understanding the science of the interaction between individual
25 genetic variation, dietary intake and changes in gene expression, structure and
26 function (nutrigenomics and nutrigenetics) has led to a growing research interest in
27 nutrigenomics-guided lifestyle intervention [1, 2, 3, 4, 5]. Genetic testing is increasingly
28 used in clinical practice to provide personalized information and recommendations
29 about health risks and lifestyle habits at a relatively low cost [6, 7]. Many specialised
30 companies can now offer genetic testing services without the involvement of clinicians,
31 focusing on predicting the risk of developing complex diseases during one's life course
32 and then making nutritional recommendations on personal lifestyle changes [8]. The
33 genetic testing investigation can be focused on health-related outcomes such as
34 fitness, pharmacogenetics and nutrigenetics [3, 9, 10]. In nutrigenetics, genetic testing
35 could provide personalised nutrition recommendations for weight control, food
36 intolerance and sensitivity [3, 11].

37 Personalised nutrition recommendations offer great potential for optimising outcomes
38 of weight management intervention [12]. However, research lacks human intervention
39 studies [2, 12, 13]. Further, there is a positive consumer attitude towards genetic-
40 based nutritional advice, partly explaining the growing interest in this field [14].
41 Notwithstanding, consumers believe the potential benefits of nutrigenomics outweigh
42 the risks [15]. Other studies have also shown that the receptivity of genetic-based
43 dietary advice is higher, considering that a one-size-fits-all approach to weight
44 management and fitness is not optimal [12, 14, 16]. Hence, nutrigenomics-guided
45 lifestyle intervention programmes result in long-term adherence to dietary
46 guidelines/recommendations [16]. As such, there is potential for genetically guided,
47 actionable nutrition recommendations to help motivate changes in dietary behaviours
48 [8, 13, 16]. In a study on genetic testing and behaviour change, adequate dietary
49 intake is the most promising lifestyle component that could be motivated through
50 personalised genetic-based advice [2]. However, the effectiveness of genetic testing
51 in promoting changes in lifestyle habits has conflicting results, too [14, 17, 18, 19]. For
52 instance, changes in dietary fat quality due to personal genetic information affecting
53 health behaviour were short-lived [14,19]. Thus, further research is "required to
54 determine how to utilize genotype-based health information and how to efficiently
55 achieve sustainable long-term changes in the prevention of lifestyle-related diseases"
56 [18, p. 161].

57 Aim and Objectives

58 This scoping review aims to build context to study the effectiveness of personalized
59 nutrition intervention on body weight management among females [18-24 years old]
60 in Jeddah Kingdom of Saudi Arabia by critically evaluating existing studies on
61 nutrigenomics-guided lifestyle intervention programmes.

62 The research objectives of the study which guided the search of the literature are:

- 63 1. To investigate the impact of personalized genetic-based nutritional programmes
64 on weight management of obese individuals.
- 65 2. To determine the effectiveness of a nutrigenomics-guided lifestyle programme on
66 sustainable weight management.
- 67 3. To evaluate the strengths, constraints and receptivity of genetic-based nutritional
68 programmes on weight management.

69

70 **Method:**

71 The scoping review approach has been chosen as it helps to address the broad aim
72 of this study. [20] argue that scoping reviews are more flexible and allow for the
73 inclusion of a diverse range of study designs. In mapping and summarising evidence,
74 scoping reviews can also help to inform future research and contribute to policy
75 implications [21]. The scoping framework proposed by [20] has been adopted in this
76 research. This process includes identifying the research question, finding relevant
77 studies, selecting studies meeting inclusion criteria, and collating, summarising and
78 reporting the results.

79 *Search Strategy*

80 The Population, Intervention, Comparison and Outcome (PICO) format was used
81 firstly to define the PICO question and then to help plan our search strategy following
82 the Arksey and O'Malley methodological framework [20]. Table 1 shows the PICO
83 search elements with related keywords/phrases to aid the literature search.

84

85

86 Table 1: PICO elements

87

PICO ELEMENT	KEYWORDS/PHRASES	Key Terms	Search Number
P (Population)	Individual living with obesity and considering weight management intervention	Obese adults OR overweight AND weight management	S1
I (Intervention)	A personalized genetic-based nutritional programme	nutrigenomics OR Genetic-based OR nutrigenomics-based OR genotype-based AND intervention OR programme	S2
C (Comparison)	Non-genetic-based nutrition intervention	Non-genetic-based OR standard-based OR population-based	S3
O (Outcome)	Sustainable reduction of body weight	Body weight OR body mass index OR Fat composition OR body circumference AND reduction OR loss AND Sustained OR Long term	S4
Final search	S1+S2+S3+S4 = Results		

88

89 The intervention was defined as providing personalised genetic-based nutritional
90 information for weight management. The comparison is, therefore, against non-
91 genetic-based nutritional interventions/programmes or population/standard-based
92 interventions, which do not involve providing genetic information. The desired outcome
93 from the personalised genetic-based nutrition intervention is a sustainable reduction
94 of body weight (body mass index, body composition, body circumference).

95 The search using key terms from the PICO table was conducted via electronic
96 searches of databases, including PubMed and Medline, on the Westminster University
97 Library database. The PubMed MeSH search involved three main concepts:
98 nutrigenomics (genetic-based, genetics*), obesity (body weight/body mass index), and
99 weight loss*. The searches in the databases were structured using Boolean operators
100 ("OR" and "AND"). This was useful in broadening the results.

101

102 ***The inclusion Criteria***

103 The inclusion criteria detail the basis on which sources were considered for inclusion
104 in the scoping review to address the research objectives [21]. Utilising the PICO
105 framework, the inclusion criteria were developed as follows:

- 106 • Adult Individuals (18+) living with obesity and considering weight
107 management intervention.
- 108 • Weight management interventions involving genetic-based
109 (nutrigenomics guided) information/advice.
- 110 • Published literature on any research design

- 111 • Published literature in the English Language or translated into English.
112 • Studies in the period 2007 to 2023

113 ***The exclusion criteria***

- 114 • Animal studies (non-human studies)
115 • Studies not involving adults (less than 18 years)
116 • Studies not involving genetic-based information (i.e., standard or
117 population-based weight management intervention)
118 • Studies on genetic-based interventions not involving obesity/weight loss.
119 • Studies not published or translated into the English language.
120 • Studies published before 2007.

121 The search strategy aimed to identify published nutrigenomics-guided intervention
122 studies relevant to the research objectives from 2007 to 2023. As such, the literature
123 search aimed to identify and review empirical studies that demonstrate the impact of
124 genetic-based nutritional intervention in weight management. The examined studies
125 were not restricted to one age group but to all adults. In addition, the search for studies
126 was not limited to any region/country.

127 ***Data Extraction***

128 The relevant studies identified were transferred to Mendeley's referencing software,
129 which helped locate duplications across the searched databases. After removing the
130 duplications, 76 articles were placed for initial screening. The articles' titles and
131 abstracts were screened. This resulted in only 11 articles meeting the criteria. To
132 identify further studies not possibly captured in the database search, a manual check
133 of the reference lists of the included studies was conducted to determine any other
134 studies that meet the inclusion criteria. This resulted in 2 additional articles. In
135 reviewing the full article text, Microsoft Excel was used to chart the data by applying
136 the relevant aspects of the Critical Appraisal Skills Programme (CASP) checklist [22].
137 The search strategy that resulted in 13 relevant articles is presented using the
138 Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) flow
139 diagram in Figure 1.

140

141

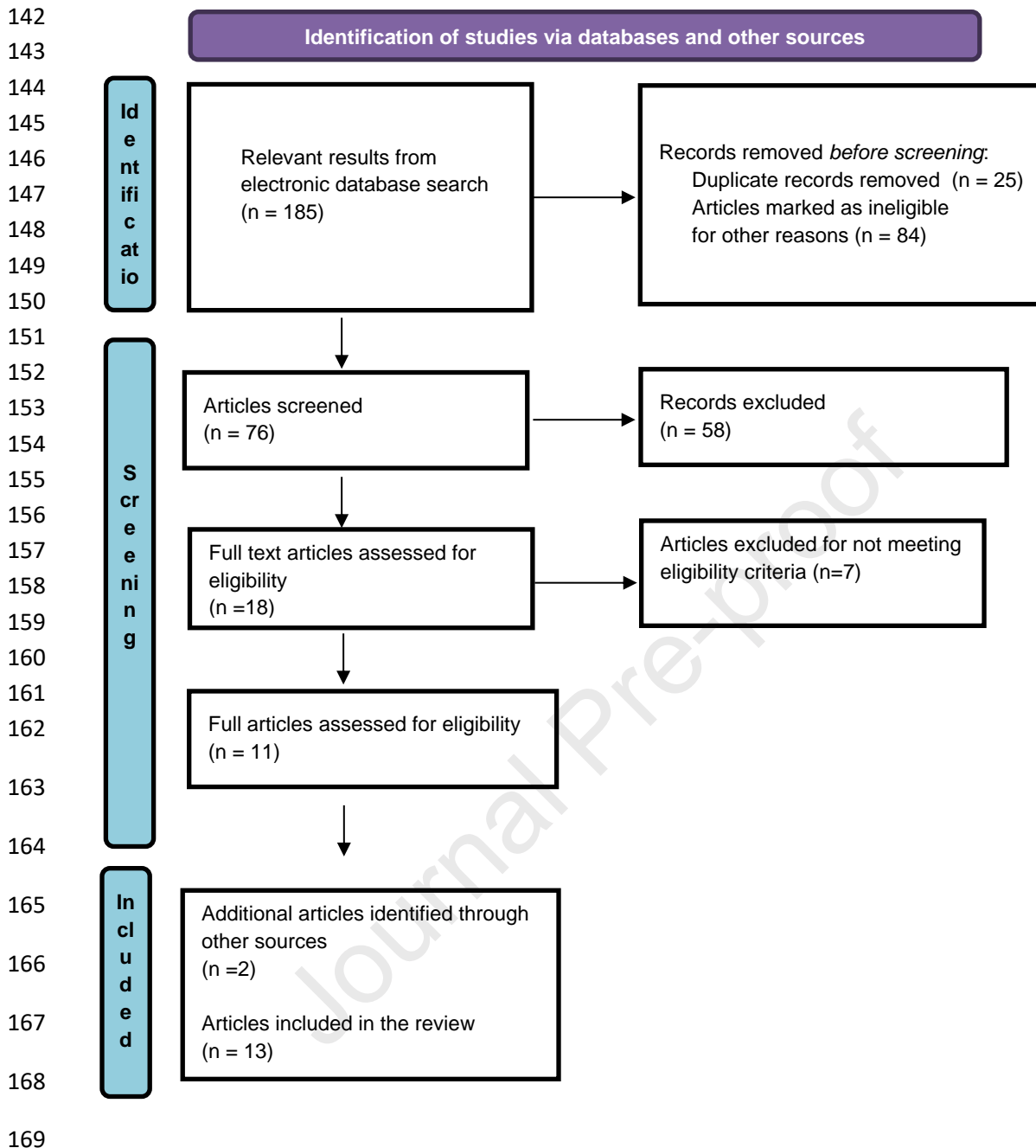


Figure 1: PRISMA flow chart showing the articles identified for critical review.

Results:

Table 3 below summarises the relevant studies on genetic-based personalised health interventions for obesity/overweight, satisfying the selection criteria developed following Arksey and O'Malley's framework. All studies were randomised clinical trials except 1 [23], a scoping review. The strengths and weaknesses of these studies have also been included. The authors' critical reflections on the findings are discussed in context with the existing body of knowledge.

179 **Table 3: Published interventions on weight reduction programmes [behavioural modification with Genetic-based**
 180 **intervention]**

Author/s (year)	Title	Aim	Participants (baseline; follow-up)	Intervention	Comparison group	Target condition (gene tested)	Follow-up	Lifestyle habit assessed	Outcome	Strength	Weakness
[13]	A comparative analysis: improved weight management using nutrigenetically tailored diet among Indians.	The study examined whether a nutrigenetically tailored diet could improve an individual's compliance with long-term weight management	106 (54 intervention groups, 52 comparisons)	Genetic based (nutrigenetic test)	Standard/population-based	Weight loss (FTO, AG, LIPC, MC4R, PPARGC1A, CD36, ADIPOQ, PPARG, CD36, MTHFR, APOA5)	30 days, 60 days, 90 days, 120 days	Body mass index and waist circumference,	The intervention group was more likely to maintain some weight loss (82%) than the comparison group (21%). Motivation and willingness to lose weight were also higher than the comparison group.	The study focussed on Indian participants in a non-western context. 15 variants in 10 genes associated with body weight and metabolism were tested.	Not all participants were obese. Only 69.8% of participants in the intervention group were obese. The reliability of the results was weak. A more detailed analysis of the results was needed.
[24]	A double-blinded, randomized, parallel intervention to evaluate biomarker-based nutrition plans for weight loss: The PREVENTOMICS study.	The study evaluated the efficacy of the PREVENTOMICS platform—which uses metabolomic and genetic information to classify individuals into different	Adults (18-65) (n=100, f=82)	Genetic-based (metabolome and genotype)	Standard/population-based	Obesity (Not specified)	Ten weeks	Fat mass, weight, waist circumference, lipid profile, glucose homeostasis markers, inflammatory markers,	The study found no differences between groups in the changes in body weight, body fat percentage, and waist circumference and no	The study examined the efficacy of personalised recommendation diets (based on genetic, nutritional, biochemical,	The study relies on the effectiveness of the platform (preventomcis). The study has a problem with the reliability of results since

		'metabolic clusters' and create personalized dietary plans—for improving health outcomes in subjects with overweight or obesity.						blood pressure, physical activity, stress and eating behaviour.	interactions with genotype or baseline insulin secretion.	physiological and behavioural factors). The approach used is different from that of other studies.	participants were not monitored but asked to self-report. The sample size is small, and the follow-up period was relatively short (10 weeks). Strong adherence to behavioural change takes time. The study examines too many aspects. Each of these aspects requires more than one observation.
[16]	Change in Weight, BMI, and Body Composition in a Population-Based Intervention Versus Genetic-Based Intervention: The NOW Trial.	To compare changes in body fat percentage (BFP), weight, and BMI between a standard intervention and	Adults (n = 140, f=38)	Genetic-based personalised lifestyle advice	Population-based lifestyle advice	Obesity (Body fat percentage, weight and BMI) (12 gene variants – FTO, UCP1,	3, 6 and 12 months	BFP, weight and BMI	The nutrigenomics group experienced significantly more significant reductions in per cent and absolute BFP at	The study provides strong evidence of change in BFP and BMI based on genetic-based	The sample size was relatively small compared to other studies (e.g. [27]). The sample size would be

		a nutrigenomics intervention.				TCF7L2, APOA2, ACE, MC4R, ADRB3, NRF2, GSTP1, NFIA-AS2, ACNT3)			the 3-month follow-up and per cent BFP at the 6-month follow-up compared with the standard group.	lifestyle evidence. Also, the number of genes tested was relatively higher than in other studies (e.g., [27]. See Appendix 1	estimated to be above 275. Also, the participants included in the study were already part of a weight management programme.
[25]	Exploring attitudes, subjective norms and perceived behavioural control in a genetic-based and a population-based weight management intervention: A one-year randomized controlled trial.	To determine the impact of providing genetically tailored and population-based lifestyle advice on key constructs of the Theory of Planned Behaviour (TPB)	Caucasian female adults (n = 140, f=70)	Genetic-based personalised lifestyle advice	Population-based lifestyle advice	Overweight/ body fat percentage (FTO)	3, 6 and 12 months	Attitudes, subjective norms and perceived behavioural control	Significant changes in attitudes, subjective norms, and perceived behavioural control tended to be short-term in the population-based group and long-term for the genetic-based group.	Provided some good empirical insight on the effect of personalised genetic data provision and applied a behavioural theory (TPB) Also, follow-ups were done at different levels, though 12 months is not a long-term change.	Observing attitudes, subjective norms or perceived behavioural control is affected by several factors, and it is hard to distinguish whether genetic-based advice was the sole or primary contributor in this case. The sample size is also tiny, and the focus was on

											one gene (FTO)
[23]	Assessing the effectiveness of actionable nutrigenomics and lifestyle genomics interventions for weight management in clinical practice: A critical, scoping review with directions for future research.	A scoping review was conducted to summarize and evaluate the current knowledge on the effectiveness of providing DNA-based lifestyle advice on weight-related outcomes to provide direction for future research.	N/A	N/A	N/A	Weight management	N/A	Weight management	Research in this area is promising but limited. Identified some limitations of prior studies: e.g., study designs, the nature of the recommendations provided to participants, small (underpowered) sample sizes, the use of self-reported weight/BMI data and lack of consideration of important confounding factors.	Provided an excellent scoping review of existing studies	Not an empirical primary study to show the impact of genetic-based intervention
[26]	Enhanced long-term dietary change and adherence in a nutrigenomics-guided lifestyle intervention compared to a population-based	To determine if a nutrigenomics-guided lifestyle intervention programme could be used to motivate greater dietary adherence and change in	Adults (n = 140)	Genetic-based personalised lifestyle advice	Standard population-based weight management intervention	Overweight/obesity (UCP1, FTO, TCF7L2, APOA2, PPAR γ 2 and MC4R)	3, 6 and 12 months. 24-hour recalls	Dietary adherence and change in dietary intake (short-term, moderate-term term)	Only the genetically guided intervention group significantly reduced their total fat intake from baseline to	The use of the theory of planned behaviour in exploring the impact of genetically based intervention.	The study was confined to participants already on a weight management programme (group lifestyle

	(GLB/DPP) lifestyle intervention for weight management: results from the NOW randomised controlled trial.	dietary intake short-term, moderate-term and long-term compared to the gold standard population-based weight management intervention (Group Lifestyle Balance (GLB)/Diabetes Prevention Programme (DPP)).						and long-term)	12-month follow-up. Long-term dietary adherence to total fat and saturated fat guidelines were also significantly more significant in the genetically guided group compared to the standard/population-based weight management group	The study considered short-term and long-term dietary changes and adherence to dietary guidelines.	balance programme). Only a few participants (i.e., 140)
[27]	Can genetic-based advice help you lose weight? Findings from the Food4Me European randomized controlled trial.	To determine whether the provision of FTO genotype information affected obesity-related traits across different levels of personalized nutrition, and between risk and non-risk FTO genotypes	Adults (n=583)	High-risk genetic results	Non-risk genetic result	Overweight/obese (FTO)	3 and 6 months	Nutrition, physical activity	High-risk FTO genotype group had significantly greater reductions in weight and WC compared with the control group (standard, nonpersonalized lifestyle advice);	The sample size in this study was relatively higher than other studies (e.g., [16]) The relative strength of the observed change in weight reduction was also higher.	The follow-up in this study was only 3 and 6 months. Thus, some long-term weight and waist circumference (WC) changes might not be observed.

[19]	Changes in physical activity following a genetic-based internet-delivered personalized intervention: randomized controlled trial	The purpose of this study was to determine if disclosing FTO risk had an impact on change in PA following a 6-month intervention.	Adults (b = 265; f=130)	High-risk genetic results	Non-risk genetic result	Overweight/obesity (FTO)	Six months	Physical activity	No significant change in subjective or objective physical activity with the provision of FTO genotype risk info	The sample size in this study was large compared to other studies. The study was more focused on fat mass and obesity-associated (FTO) genotype and provided empirical evidence.	The study only examined changes in physical activity without also considering the change in dietary adherence as these affect the predisposition to overweight. Also, the provision of genetic-based information/advice was web-based, which could affect the impact on participants.
[28]	Genetic susceptibility testing and readiness to control weight: Results from a randomized controlled trial	To test the hypothesis that adding obesity gene feedback (FTO) to simple weight control advice at a life stage with a raised risk of weight gain (university)	Young Adults (b = 1,016; f=279)	Genetic results	No genetic testing	Obesity (FTO)	One month	Nutrition (adherence to a variety of eating behaviours) and Physical activity	There was no significant change in nutrition and physical activity (pooled) between groups. Adding FTO feedback to weight control	The sample size in this study was relatively high (b=1,016 and f=279). Also, targeted young adults are more susceptible to	The follow-up in this study was too short (1 month only). Subsequent follow-ups were needed.

		increases readiness to control weight.							advice enhanced readiness to control weight, without evidence for genetic determinism, but had no more effect on behaviour than weight control advice alone.	weight gain (university level) The study examined both nutrition/dietary change and physical activity, which is good as these are two critical factors in obesity.	
[29]	Effects of a web-based personalized intervention on physical activity in European adults: a randomized controlled trial.	To investigate the impact of different levels of personalization on PA change, using phenotypic and genotypic information to tailor the PA advice	Adults (b-1480, f-1233)	High-risk genetic result	Non-risk genetic result	Overweight/obesity (FTO)	Six months	Physical activity	There is no evidence that personalized advice is more effective than conventional "one size fits all" guidelines to promote changes in PA in our Web-based intervention when PA was measured objectively.	The sample size in this study was high (b-1480), though this was all self-reported and web-based.	Focused more on physical activity without considering 'diet'. Both diet and physical activity are critical drivers of obesity.
[18]	An intervention study of individual, apoE genotype-based dietary and physical-activity	To assess the behavioral effects of receiving personal genetic information, use	Adults (b-151, f-130)	Genetic testing	No genetic testing	Overweight/obesity, cardiovascular disease (apoE)	Ten weeks, six months,	Diet/nutrition, alcohol consumption, physical activity	Personal genetic information affects health behaviour. Dietary fat	This study's sample size was relatively small compared to	The study did not focus specifically on obesity/overweight but on

	advice: impact on health behavior.	apoE genotypes to promote lifestyle changes.					12 months		quality improved more in the high-risk group than in the low-risk and control groups after personal, genotype-based health advice.	other studies, e.g. (29). Examination of dietary behaviour and physical activity was considered, in addition to alcohol consumption.	susceptibility to cardiovascular diseases in addition to overweight.
[30]	Differences in weight loss between persons on standard balanced vs nutrigenetic diets in a randomized controlled trial.	To determine whether more participants who followed a nutrigenetic-guided diet lost more significantly than 5% of their body weight than participants on a standard diet	Adults (b=51)	Nutrigenetic-guided diet	Standard balanced diet	Obesity (APOA2, ADIPOQ, FTO, KCTD10, LIPC, MMAB, PPARG)	Eight weeks, 24 weeks	Weight loss	There was no significant difference in the percentage of participants on the balanced diet vs the nutrigenetic-guided diet who lost 5% of their body weight. Both groups had difficulty adhering to the diets.	Study shows that adherence to diet is a challenge regardless of the information provided. However, weight loss is more when a nutrigenetic-guided diet is followed.	The study concentrates on age groups 46 and above. Also, physical activity was not incorporated. The sample size is too small relative to other studies.
[31]	Is the information on genetic determinants of obesity helpful or harmful for obese people?—A	To assess the positive and negative effects of informing obese people about the genetic	Adults (b-410, f-294)	Genetic testing and consultation	Consultation only	Obesity	Six months	Nutrition (restraint eating)	No negative effects (e.g., loss of self-efficacy/self-control, increase of	The sample size in this study was high, improving the	The study concentrates on 'feelings' about the state of participants following the

	randomized clinical trial.	etiology of being overweight.							body weight) were observed to inform obese people about the genetic etiology of being overweight. The consultation resulted in long-term improvement of negative mood if it included genetic information in the case of participants with a family history of obesity and if it included no genetic information in the case of obese people without a family history of obesity.	reliability of the findings. Gives a unique perspective on participants' attitudes or feelings about themselves following genetic information provision.	provision of genetic information about susceptibility to obesity. The study did not specifically address adherence to a healthy diet or change in physical activity. Feelings after consultation are subject to change.
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182 **Discussion:**

183

184 ***Critical observation***

185 The search for studies on nutrigenomics-guided lifestyle intervention programs on
186 weight management revealed that there are few directed studies despite the many
187 studies on the effectiveness of weight management intervention (i.e., with no genetic
188 information associated). In most cases, the focus on the provision of genetic
189 information was aimed at addressing other health issues (e.g., cardiovascular
190 diseases, hypertension, cancer), not specifically obesity/overweight health issues [1,
191 2, 7]. This scoping review focuses on nutrigenomics-guided studies in weight
192 management.

193 The findings show that most studies were based in developed countries where gene
194 services are becoming widely available. However, this does not necessarily limit the
195 conduct of nutrigenomics-guided lifestyle intervention programmes to geographical
196 regions. Specialised genetic testing services can also be provided to international
197 customers [11]. In such a case, a cost-benefit analysis, risk assessment, and ethical
198 consideration become even more necessary due to data protection law, potential cost
199 implications, and findings' reliability when samples are transported across long
200 distances between continents.

201 A general observation also highlights, in part, the effect of the COVID-19 pandemic
202 (from 2020). Only two studies were identified as useful/relevant post-COVID. This
203 suggests that the COVID-19 period caused a significant gap in the conduct of
204 randomised clinical trials on genetic-based intervention programmes. As such, a 'big
205 gap' exists to be filled in terms of empirically supported studies on the effectiveness of
206 genetic-based nutrition intervention.

207 ***Authors critical reflection on presented studies:***

208 There are mixed results concerning the effect of the provision of personalised genetic
209 information in obesity/overweight intervention programmes. In a study on the impact
210 of genetically based personalised lifestyle advice involving 140 adults in the baseline
211 phase and 38 adults in the follow-up phase, [16] investigated the changes in body fat
212 percentage (BFP), weight and body mass index (BMI) between a group with
213 genetically based personalised lifestyle advice and standard intervention group. Their
214 study tested 12 gene variants (FTO, UCP1, TCF7L2, APOA2, ACE, MC4R, ADRB3,
215 NRF2, GSTP1, NFIA-AS2, ACNT3) and had 3, 6 and 12 months follow-up periods.
216 [16] found a statistically significant change in a reduction in BFP (in per cent and
217 absolute terms) between the two groups. The group with genetic-based personalised
218 lifestyle advice outperformed the standard intervention groups. The positive effects of
219 genetic-based personalised lifestyle advice were also observed in the [27] study that
220 involved 583 participants with one group provided with FTO genotype information and
221 personalised nutritional advice and a standard group (non-risk FTO genotype). [27]

222 found that the high-risk FTO genotype group significantly reduced weight and weight
223 circumference.

224 Further, positive changes in attitude, nutritional adherence and physical activity in the
225 long term have been found in some studies when personalised genetic-based advice
226 is provided [25, 26]. This is significant considering that lifestyle changes for weight
227 management are preferred in the long term than the short term [32]. [25] explored
228 attitudes, subjective norms, and perceived behavioural control from the perspective of
229 planned behaviour theory. The study found significant changes in attitudes, subjective
230 norms and behavioural control that were long-term oriented for groups with
231 personalised genetic-based lifestyle advice compared to the standard/population-
232 based group that tended to be short-term oriented. In this respect, the provision of
233 personalised genetic information was vital in influencing the long-term behavioural
234 changes of participants, providing motivation to adhere to dietary guidance over a long
235 time. These results were further reinforced in [26] study that aimed to determine if a
236 nutrigenomics-guided lifestyle intervention programme could motivate greater dietary
237 adherence and change in dietary intake in the short-term, moderate-term and long-
238 term. The study found that dietary adherence and change in dietary intake were
239 significantly more significant in the long term when personalised genetic-based
240 information was provided to participants. In other words, genetic-based personalised
241 lifestyle advice positively influenced participants to adhere to dietary guidelines in the
242 long term. This is also consistent with the observation in the [18] study, in which
243 personalised genetic-based information improved the quality of dietary fat and health
244 behaviour. The health behaviour related to physical activity, dietary intake and alcohol
245 consumption, with participants most at risk based on their genotype making significant
246 changes.

247 On the contrary, other studies have found no significant difference in the impact of
248 personalised genetic-based information on weight management [19, 24, 28]. [24]
249 assessed the fat mass, weight, waist circumference, lipid profile, glucose homeostasis
250 markers, inflammatory markers, blood pressure, physical activity, stress and eating
251 behaviour of 100 participants. They found no statistically significant difference
252 between the group with genetic information and the control group with no information
253 about their genotype. Behavioural changes concerning physical activity were
254 examined in [19] and [28] studies, which found that participants did not change their
255 behavioural patterns despite being given personalised genetic information about their
256 risk susceptibility. For instance, [19] examined whether disclosing FTO risk had an
257 impact on change in physical activity following a 6-month intervention and found no
258 statistically significant change in subjective or objective physical activity despite the
259 provision of FTO genotype risk information. These observations were also found in
260 [29] and [28], as behavioural change concerning physical activity did not change
261 despite the provision of genetic information and the related risk profile. Adherence to
262 nutrition advice (dietary intake) for weight management did not change either, despite
263 participants being provided with personalised genetic-based nutritional advice in these

264 studies. Thus, [29] argue that 'one size fits all' guidelines are equally practical in weight
265 management even without genetic information. This perception can also be seen in
266 [31] study that disclosing genetic information and subsequent consultation did not
267 negatively affect some key psychological attributes (e.g., loss of self-efficacy or self-
268 control). In other words, participants seem to have accepted their obesity/overweight
269 predisposition, and no additional information motivated them to change their behaviour.

270 ***Strengths and weaknesses of studies***

271 Given the mixture of results, it is imperative that a critical evaluation of the strengths
272 and limitations of the reviewed studies is discussed. This helps to highlight not only
273 the existing gaps in the literature but also directs attention to areas for further
274 investigation. A significant contribution to the literature on the effect of personalised
275 genetic-based information has been provided by [16, 23, 25, 26]. These studies have
276 provided solid empirical evidence showing the positive impacts of genetic-based
277 nutritional advice. These studies provided strong empirical evidence and engaged well
278 with theoretical perspectives explaining the observed behavioural change. In particular,
279 the theory of planned behaviour was utilised in understanding the attitudes, subjective
280 norms and perceived behaviour controls in [25]. The supportive results of [27] show
281 that the relative strength of observed change in weight reduction was higher, and the
282 sample size (583 adults) was significantly large. Sample size impacts the statistical
283 inference of results, with results strengthened when sample sizes are large and
284 participants are randomised into groups [33]. In this respect, some strength of studies
285 [19, 27-29, 31] lie in their large sample size. A critical appraisal of sample size
286 determination shows that when population size is unknown, the estimated sample size
287 to achieve a 95% confidence level, 5% margin of error and 50% population proportion
288 of characteristic/attribute would be 385 (see www.calculator.net). In this respect, there
289 is a relative strength that emanates from the size of the sample sizes in [27], [28], [29]
290 and [31] studies that had 583, 1,016, 1,480 and 410 participants respectively. Similarly,
291 some criticism can be revealed regarding the reliability of findings in the studies with
292 small sample sizes. For instance, [16, 25, 26] studies all had small sample sizes
293 (i.e., 140 participants). However, as [33] argue, the population size impacts the sample
294 size requirement and affects the reliability of findings.

295 In genetic-based interventions, identifying the relevant genotype associated with the
296 target condition is necessary (Goodarzi, 2018). Identifying the relevant focus genotype
297 strengthens some of the studies reviewed. [19], for instance, tested for the FTO
298 genotype, similar to [28] and [27], while [18] examined the apoE genotype. Figure 2
299 below (appendix) highlights that the number of genes associated with obesity and
300 overweight are numerous, affecting different aspects of the health issue [34]. Thus,
301 some strength of studies lies in examining more than one genotype. For instance, [16]
302 examined 12 gene variants, [26] examined six gene variants, and [13] examined 15
303 variants in 10 genes, giving relative strength to these studies. On the contrary, the
304 study by [24] did not specify which genotype was examined. Nonetheless, considering

305 the many gene variants associated with the health issue of obesity, the reviewed
306 studies have a weakness in not expanding their focus to consider more gene variants.

307 Obesity prevention strategies require dietary changes and physical activities [35, 36].
308 Thus, there is a strength in some studies [18, 24, 27, 28] that assessed both aspects:
309 nutritional adherence and physical activity. On the contrary, the limitations can be
310 argued for studies that focussed on only one of the aspects, i.e., physical activity [19,
311 29] or dietary change [13, 26, 30, 31] as this gives an incomplete assessment in weight
312 management.

313 There is a further limitation in some studies [16, 25-27] concerning the population and
314 sample selection. The participants for these randomised controlled trials were all
315 drawn from an existing weight management programme. As such, the provision of
316 genetic information would be expected to provide additional motivation along the
317 continuum of positive behavioural change. [16, 23, 26] participants were drawn from
318 the Group Lifestyle Balance (GLB) Program, designed for non-diabetic, overweight
319 individuals aged 18 and older [37]. The program aims to achieve a 7% weight loss
320 through healthy eating and promoting 150 minutes of brisk physical activity each week
321 [37]. In this respect, any participants from this group would have committed to the
322 program's goals. [27] study participants were part of the Food4Me project, an EU-
323 funded research project to understand the relationship between food and gene
324 expression [38]. Thus, instead of participants already on a weight management
325 programme, it would have been insightful to see the impact of such information
326 provision to non-participants on a weight management programme. This would be
327 useful in identifying whether the provision of personalised genetic information and risk
328 susceptibility provided the incentive to overcome the inertia (resistance) for
329 behavioural change. Nonetheless, it could also be argued that the true impact of
330 personalised genetic-based nutritional advice is on whether it provided additional
331 momentum (imperative) to the existing path to behavioural change.

332 Some criticism can also be levelled against studies such as [24] and [27] for the limited
333 monitoring. These studies relied significantly on self-reporting and self-recruitment,
334 which was internet-based. As such, the rigorous monitoring process that helps
335 improve the reliability of findings of randomised clinical trials [33] is reduced. The
336 challenge lies in the provision of 'accurate information' and, thus, the importance of
337 monitoring or tracking processes in any randomised clinical trial. This would help
338 strengthen the findings and contribution to the field. This has been aptly observed by
339 the Food4Me project, which states that "there is a need to comprehensively analyse
340 the opportunities and challenges in the field of personalised nutrition" [38, p. 1]. This
341 remains a challenge in genetic-based randomised clinical trials [39].

342 Further, some studies [13, 24, 28] had very short follow-up periods, which arguably
343 does not give sufficient time to observe the effect of behavioural change. [24] and [28]
344 had follow-up periods of 10 weeks and one (1) month, respectively. The importance
345 of the observation period is demonstrated in [32] study, which found that twelve (12)

346 months for the weight loss programme was more effective than six (6) months as solid
347 adherence to behavioural change takes time. The study of [24] can also be criticised
348 for focusing on the effectiveness of a nutritional platform (PREVENTOM CIS) instead
349 of genetic-based information. Further, [25] study that explored attitudes, subjective
350 norms, and perceived behavioural control could be criticised because several factors
351 affect attitudes and norms [40]. Thus, it is hard to distinguish whether the provision of
352 genetic-based advice was the primary or sole contributor to the observed change.

353 **Conclusion:**

354 The scoping review has highlighted that the research landscape of nutrigenomics-
355 guided lifestyle intervention programmes is still growing. The evidence on the
356 effectiveness of nutrigenomics-guided lifestyle intervention programmes is mixed.
357 Thus, more research is needed to demonstrate whether the provision of personalised
358 genetic-based nutritional advice significantly influences health behavioural changes.
359 A key aspect of further research is considering the reliability/validity of the randomised
360 clinical trials and issues such as sample selection, follow-up periods, and monitoring
361 tools. Further research is warranted to incorporate physical activity and dietary
362 adherence, as these aspects are essential to sustain weight management.

363 Further, most studies have been based in developed countries, providing a research
364 gap to understand not only the attitudes or receptivity of nutrigenomics-guided lifestyle
365 intervention programmes but also their effectiveness and contribution to the unclear
366 (mixed) empirical evidence. This scoping review has identified a research gap through
367 re-directing the focus on emerging countries and also on young adults who may be
368 exposed to packed lifestyle-related risk factors for overweight/obesity. The limitations
369 arise mainly from the nature of a scoping review (unlike systematic reviews), in that
370 quality assessment of the included studies is not comprehensively undertaken.

371

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373

374 ***Authors' contributions***

375 Saba Aljasir and Ihab Tewfik designed the study and collected the data. All authors
376 analysed the data and prepared the manuscript. All authors read and approved the
377 final manuscript.

378

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382 ***Conflict of interests***

383 The authors declare that they have no competing interests.

384

385 ***Ethics approval and consent to participate***

386 Not applicable

387

388

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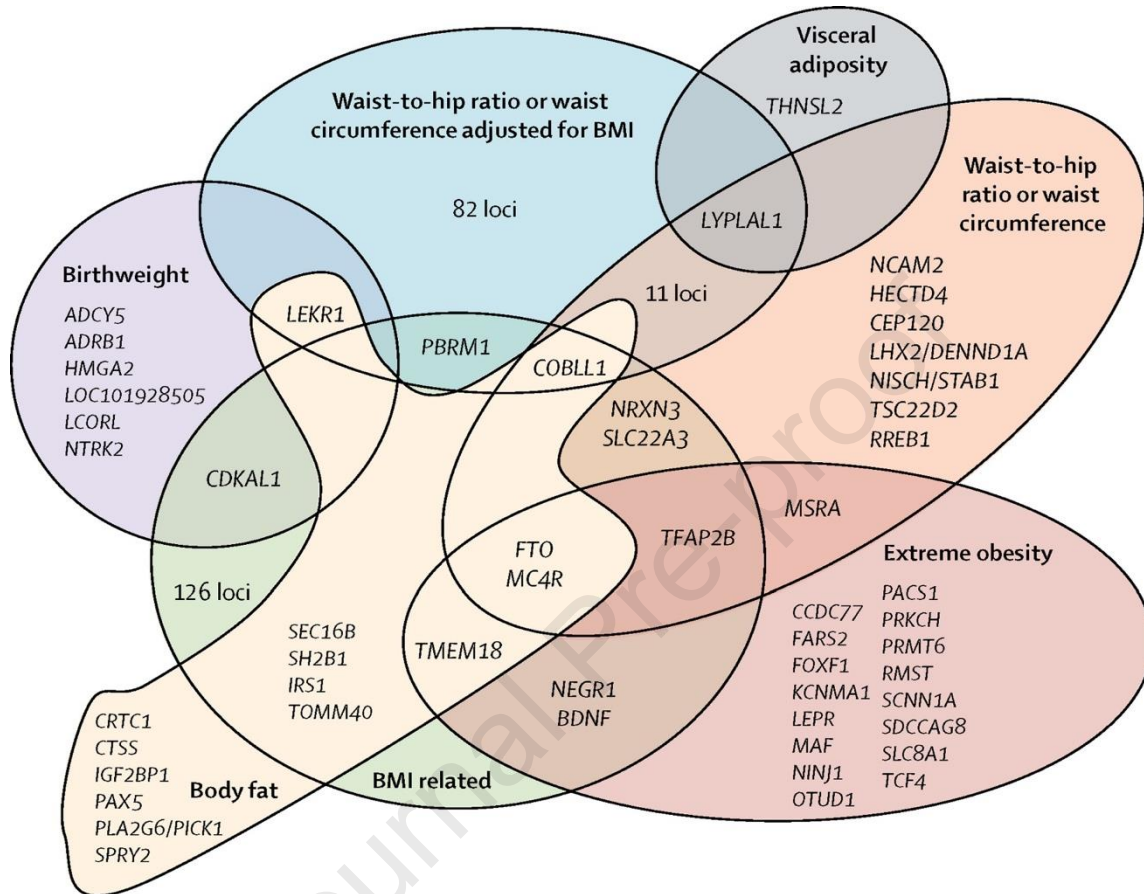
527 **List of abbreviation:**

528	BFP	-	Body Fat Percentage
529	BMI	-	Body Mass Index
530	22	-	Critical Appraisal Skills Programme
531	GLB	-	Group Lifestyle Balance
532	PICO	-	Population, Intervention, Comparison and Outcome

533 PRISMA - Preferred Reporting Items for Systematic Reviews and Meta-
534 Analyses

535 Appendix 1

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539

540 Figure 2: Selected genes associated with obesity. [Source: Goodarzi, 2018]

541

542 Appendix 2: PRIMA Checklist

543 Preferred Reporting Items for Systematic reviews and Meta-Analyses extension for
544 Scoping Reviews (PRISMA-ScR) Checklist

SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #
TITLE			
Title	1	Identify the report as a scoping review.	1
ABSTRACT			
Structured summary	2	Provide a structured summary that includes (as applicable): background, objectives, eligibility criteria, sources of evidence, charting methods, results, and conclusions that relate to the review questions and objectives.	1
INTRODUCTION			
Rationale	3	Describe the rationale for the review in the context of what is already known. Explain why the review	2

SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #
		questions/objectives lend themselves to a scoping review approach.	
Objectives	4	Provide an explicit statement of the questions and objectives being addressed with reference to their key elements (e.g., population or participants, concepts, and context) or other relevant key elements used to conceptualize the review questions and/or objectives.	2-3
METHODS			
Protocol and registration	5	Indicate whether a review protocol exists; state if and where it can be accessed (e.g., a Web address); and if available, provide registration information, including the registration number.	N/A
Eligibility criteria	6	Specify characteristics of the sources of evidence used as eligibility criteria (e.g., years considered, language, and publication status), and provide a rationale.	4-5
Information sources*	7	Describe all information sources in the search (e.g., databases with dates of coverage and contact with authors to identify additional sources), as well as the date the most recent search was executed.	4
Search	8	Present the full electronic search strategy for at least 1 database, including any limits used, such that it could be repeated.	3-4
Selection of sources of evidence†	9	State the process for selecting sources of evidence (i.e., screening and eligibility) included in the scoping review.	5-6
Data charting process‡	10	Describe the methods of charting data from the included sources of evidence (e.g., calibrated forms or forms that have been tested by the team before their use, and whether data charting was done independently or in duplicate) and any processes for obtaining and confirming data from investigators.	5-6
Data items	11	List and define all variables for which data were sought and any assumptions and simplifications made.	N/A
Critical appraisal of individual sources of evidence§	12	If done, provide a rationale for conducting a critical appraisal of included sources of evidence; describe the methods used and how this information was used in any data synthesis (if appropriate).	7-15, 16-20
Synthesis of results	13	Describe the methods of handling and summarizing the data that were charted.	7-15
RESULTS			
Selection of sources of evidence	14	Give numbers of sources of evidence screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally using a flow diagram.	6
Characteristics of sources of evidence	15	For each source of evidence, present characteristics for which data were charted and provide the citations.	7-15
Critical appraisal within sources of evidence	16	If done, present data on critical appraisal of included sources of evidence (see item 12).	7-15

Results of individual sources of evidence	17	For each included source of evidence, present the relevant data that were charted that relate to the review questions and objectives.	7-15
Synthesis of results	18	Summarize and/or present the charting results as they relate to the review questions and objectives.	7-15
DISCUSSION			
Summary of evidence	19	Summarize the main results (including an overview of concepts, themes, and types of evidence available), link to the review questions and objectives, and consider the relevance to key groups.	16-20
Limitations	20	Discuss the limitations of the scoping review process.	20
Conclusions	21	Provide a general interpretation of the results with respect to the review questions and objectives, as well as potential implications and/or next steps.	20
FUNDING			
Funding	22	Describe sources of funding for the included sources of evidence, as well as sources of funding for the scoping review. Describe the role of the funders of the scoping review.	N/A

546