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1 **Systematic review of the methods used in economic evaluations of targeted**
2 **physical activity and sedentary behaviour interventions**

3 Cochrane, M.^a, Watson, P.M.^a, Timpson, H.^b, Haycox, A.^c, Collins, B.^d, Jones, L.^c, Martin, A.^{d,e},
4 Graves, L.E.F.^a

5

6 ^aPhysical Activity Exchange, Research Institute for Sport and Exercise Sciences, Liverpool
7 John Moores University, 5 Primrose Hill, Liverpool, L3 2EX, UK

8 ^bPublic Health Institute, Faculty of Health, Education and Community, Liverpool John Moores
9 University, 3rd Floor, Exchange Station, Tithebarn Street, Liverpool, L2 2QP, UK

10 ^cManagement School, University of Liverpool, Chatham Street, Liverpool, L69 7ZH, UK

11 ^dDepartment of Public Health and Policy, University of Liverpool, Waterhouse Building, Block
12 B, 2nd Floor Liverpool, L69 3BX, UK

13 ^eHCD Economics, The Innovation Centre, Keckwick Lane, Daresbury, Warrington, WA4 4FS,
14 UK

15

16 **Abstract**

17 The burden of noncommunicable diseases (NCD) on health systems worldwide is
18 substantial. Physical inactivity and sedentary behaviour are major risk factors for NCD.
19 Previous attempts to understand the value for money of preventative interventions targeting
20 physically inactive individuals have proved to be challenging due to key methodological
21 challenges associated with the conduct of economic evaluations in public health. A
22 systematic review was carried out across six databases (Medline, SPORTSDiscus, EconLit,
23 PsychINFO, NHS EED, HTA) along with supplementary searches. The review examines
24 how economic evaluations published between 2009-March 2017 have addressed
25 methodological challenges with the aim of bringing to light examples of good practice for
26 future studies. Fifteen economic evaluations from four high-income countries were retrieved;
27 there is a dearth of studies targeting sedentary behaviour as an independent risk factor from
28 physical activity. Comparability of studies from the healthcare and societal perspectives
29 were limited due to analysts' choice in cost categories, valuation technique and time horizon

30 differing substantially. The scarcity of and inconsistencies across economic evaluations for
31 these two behaviours have exposed a mismatch between calls for more preventative action
32 to tackle NCD and the lack of information available on how resources may be optimally
33 allocated in practice. Consequently, this paper offers a table of recommendations on how
34 future studies can be improved.

35

36

Keywords

37 Systematic Review; Economic Evaluation; Physical Activity; Sedentary Behaviour; Equity;
38 Public Health; Cost effectiveness analysis; Cost utility analysis

39

Introduction

40 **Background**

41 The burden of noncommunicable disease (NCD) on health systems is substantial.
42 Worldwide NCD is the main cause of death and disability (WHO, 2018a). Physical inactivity
43 is a major risk factor for NCD and the fourth leading cause of death globally. There is
44 therefore an urgent need to invest in preventative interventions, such as those targeting
45 individuals who do not meet the international guidelines of 150 minutes of moderate physical
46 activity per week (Kohl et al., 2012). Furthermore, sedentary behaviour, defined as any
47 waking behaviour where an individual is in a sitting, reclining or lying posture, has been
48 identified as a risk factor for NCD and all-cause mortality independent of achieving the
49 recommended physical activity guidelines. The level of physical activity found to attenuate
50 the risks associated with sedentary behaviour is 60 minutes of moderate physical activity per
51 day, which equates to 420 minutes per week (Ekelund et al., 2016). As over a third (35%) of
52 females and a quarter (26%) of males in high-income countries do not presently meet the
53 recommended weekly guidelines, a daily target of 60 minutes is unlikely to be attained
54 (WHO, 2018b). Inaction to invest in preventative interventions tackling detrimental levels of
55 physical inactivity and sedentary behaviour is expected to lead to greater levels of NCD and
56 inequity, productivity losses and a continued overwhelming demand for costly curative health
57 services (OECD, 2015).

58 As public resources are scarce, economic evaluations are important to prevent both national
59 and local policymakers from disinvesting in highly cost-effective physical activity and
60 sedentary behaviour interventions. Economic evaluations are also needed as not all public
61 health interventions represent good value for money (Owen et al., 2017). Compared to
62 population-level interventions, physical activity and sedentary behaviour interventions
63 targeting individuals who are not meeting the recommended international physical activity
64 guidelines are more likely to be: (a) funded by local-level commissioners; (b) evaluated by
65 researchers. This is likely to be due to the challenge of measuring outcomes in the general
66 population. For this reason, this review focuses on economic evaluations of targeted

67 interventions such as exercise referral schemes, brief advice in primary care and exercise
68 sessions.

69 Despite recommendations for economic evaluations to become routine within public health
70 interventions (Kelly et al., 2005) cost-effectiveness information on physical activity and
71 sedentary behaviour interventions remains scarce (Abu-Omar et al., 2017). One reason for
72 this lack of analysis may be due to the lack of guidance and multidisciplinary efforts to inform
73 analysts on how to conduct economic evaluations in the field of public health (Davis et al.,
74 2014). Economic evaluations of public health interventions are subject to four key
75 methodological challenges identified and described in former reviews (Alayli-Goebbels et al.,
76 2014; Hill et al., 2017; Weatherly et al., 2009) as: attribution of effects; measuring and
77 valuing outcomes; identifying intersectoral costs and consequences; and incorporating
78 equity. These four challenges are explain in the subsequent sections.

79 **Challenge 1: Attribution of effects**

80 Randomised controlled trials (RCTs) are the gold standard for evaluating the effectiveness
81 of an intervention. RCTs alone are however insufficient to inform long-term investment
82 decisions in health systems aiming to be sustainable. This is because conducting
83 experimental studies such as RCTs over many years or decades is likely to be resource
84 intensive from both the research funder and participant's perspective. Attrition from the trial
85 and insufficient funding is inevitable. Yet, the greatest health outcomes and cost savings
86 attributable to physical activity and sedentary behaviour interventions do not typically
87 manifest until decades after an intervention has taken place. Due to this long pay-back time
88 (Wanless, 2004), it is recommended economic evaluations link up trial-derived intermediate
89 or surrogate outcomes with additional sources of evidence (e.g. observational studies)
90 (Ramsey et al., 2015).

91 **Challenge 2: Measuring and valuing outcomes**

92 Previous physical activity studies have used different outcomes, or have classified the same
93 type of outcomes in different ways, which makes it challenging to meaningfully use cost-
94 effectiveness results and compare interventions (Abu-Omar et al., 2017). This is likely to be
95 because physical activity and sedentary behaviour interventions are associated with a broad

96 range of outcomes, many of which are not captured in evaluations that conduct just one type
97 of valuation analysis. Furthermore, many broader important and relevant outcomes such as
98 improved wellbeing or someone's ability to return to work are difficult to assign a monetary
99 value, as they do not have a market price (Weatherly et al., 2014).

100

101 **Challenge 3: Identifying intersectoral costs and consequences**

102 Many physical activity and sedentary behaviour interventions take place outside of the
103 healthcare setting, necessitating a time and equipment commitment from intervention
104 participants and providers (which has an opportunity cost). Moreover, physical activity and
105 sedentary behaviour interventions are complex, impacting on various sectors simultaneously
106 (Dahlgren & Whitehead, 1991). Therefore, it is important to consider the impact of these
107 interventions on other stakeholders including public sector agencies beyond the health
108 sector, private individuals and the voluntary sector (Weatherly et al., 2014; Weatherly et al.,
109 2009). Yet, as there is no universal definition for each perspective type, the costs and
110 consequences deemed relevant for inclusion in the analysis is primarily analyst-dependent
111 (Husereau et al., 2013).

112 **Challenge 4: Incorporating equity**

113 A key objective in public health is to reduce inequity, meaning inequalities that are
114 avoidable, but have not yet been avoided and are therefore unfair (Marmot & Allen, 2014).
115 By contrast, a key objective in economic evaluation is to maximise efficiency across the
116 whole population (Weatherly et al., 2014). If authors fail to acknowledge equity by not
117 adapting their existing economic analysis approach, it is not transparent which socio-
118 economic group have gained or lost out due to a resource allocation decision. Until the
119 recent publication by Cookson et al. (Cookson et al., 2017) recommendations on how to
120 incorporate equity have been limited within international and national guidelines for
121 economic evaluation (Husereau et al., 2013; NICE, 2014; Ramsey et al., 2015; Sanders et
122 al., 2016). Approaches for incorporating equity into the analysis described by Cookson et al.
123 (2017) include: equity impact analysis, equity constraint analysis and equity weighting
124 analysis.

125

126 **Aim**

127 In an attempt to learn how the four challenges outlined above have been addressed in
128 practice, this systematic review aims to provide an overview of the methods used in
129 economic evaluations of physical activity and sedentary behaviour interventions since 2009.
130 Alayli-Goebbels et al. (2014) and Weatherly et al. (2009) reviewed the methods reported in
131 economic evaluations of a range of public health areas including 17 and 26 physical activity
132 economic evaluations published up to 2005 and 2009, respectively, but the reviews found
133 little insight from the empirical evidence. Economic evaluation is a rapidly developing field
134 especially with the growth of decision-analytic modelling and the economic evaluation
135 reporting standards (Drummond et al., 2015; Ramsey et al., 2015). Accordingly there is a
136 strong rationale to provide an update on methods carried out since 2009.

137

138 **Methods**

139 **Information sources and search strategy**

140 A comprehensive search took place across six electronic databases that host reports from
141 the medical and economic field (Medline via Ovid; SPORTSDiscus, EconLit and PsycINFO
142 via EBSCOHost, NHS EED and HTA via the Cochrane Library). The database NHS EED
143 stores records up to April 2015, thus searches in this database went up to 2015 only.
144 Additional, supplementary searching was performed: key websites were searched for
145 studies that included specific free text terms: 'physical activity', 'sedentary behaviour',
146 'economic' and 'cost'; reference lists of two relevant systematic reviews (Gc et al., 2016; Wu
147 et al., 2011) were hand searched; and protocols that met the majority of the eligibility criteria
148 were used to search for completed studies via online searching and contacting the authors.
149 An example of the full electronic search strategy for Medline is provided in Appendix A
150 [INSERT LINK TO ONLINE FILE A, B, C, D & E]. This search was replicated for all databases,
151 with amendments made as appropriate to align terms with individual database index terms.

152 **Study selection**

153 The protocol for this review can be retrieved from the PROSPERO database for registered
154 systematic reviews (registration number CRD42017074382). Full economic evaluations of
155 interventions targeting individuals aged 16 years or over, who are defined as being
156 physically inactive or sedentary, were eligible for inclusion in the review. Population level
157 interventions were excluded as well as protocols. Eligible studies needed to capture physical
158 activity or sedentary behaviour at two or more time points to observe if a change in
159 behaviour has occurred. Comparators could be any alternative intervention including no
160 intervention. Interventions and comparators targeting multiple behaviours such as physical
161 activity and diet were excluded unless the multiple behaviours were physical activity and
162 sedentary behaviour. Both trial and model based economic evaluations were eligible. Letters
163 to editors and conference briefings were excluded. Both published and unpublished 'grey'
164 literature were included. Abstracts where the full text could not be retrieved were excluded.
165 Only English language studies were included due to the restricted language skills of the
166 reviewers available. Eligibility criteria was applied during both screening phases. The
167 present systematic review identifies and discusses studies published from January 2009 to
168 March 2017. In addition, a rapid systematic scoping search was performed in Medline to
169 understand whether new studies had been published in this area from March 2017 to
170 January 2019. Details on methods of the scoping search are not discussed below, rather
171 they are presented in Appendix B [INSERT LINK TO ONLINE FILE A, B, C, D & E].

172 **Screening**

173 During the title and abstract screening phase two reviewers (first author, seventh author)
174 screened 10% (n=612/ 6,123) of the studies and there was a disagreement rate of 2.94%
175 (n=18). Reviewers discussed the disagreements and resolved them without the need to
176 seek the expertise of a third reviewer. Reviewer one (first author) went on to screen the rest
177 of the studies, informed by the disagreement discussions. Similarly, during the full text
178 screening phase reviewer two (seventh author) screened 10% (n=15/ 153) of the studies.
179 There was disagreement for 33.33% (n=5) of the studies. The reviewers discussed the
180 disagreements and again a consensus was met without the need for a third reviewer. Figure
181 1 shows an overview of the study selection process.

182 **Data extraction**

183 A data extraction form was developed based on the items featured on the Consolidated
184 Health Economic Evaluation Reporting Standards (CHEERS) checklist (Husereau et al.,
185 2013). The form was piloted independently by two reviewers (first author, seventh author) on
186 two (10%) randomly selected studies. Following discussions the form was shortened, and
187 items relevant to the four methodological challenges, and key study characteristics were
188 retained. Following the piloting stage, the first reviewer extracted data for the remaining
189 studies. A template of the final data extraction form is provided in Appendix C [INSERT LINK
190 TO ONLINE FILE A, B, C, D & E]. It was not necessary to request additional information from
191 the study authors.

192 **Quality assessment**

193 Drummond's 10-item checklist was selected as it is one of the most widely used quality
194 assessment tools (Drummond et al., 2015). A component approach was used when applying
195 the checklist in Appendix D [INSERT LINK TO ONLINE FILE A, B, C, D & E]. This approach is
196 advocated in the PRISMA statement and entails assessing each item individually rather than
197 generating a summary score (Liberati et al., 2009). Two reviewers (first author, seventh
198 author) independently conducted the quality assessment for 10% (n=2/ 15) of the included
199 studies. Disagreement was limited to item 6 (Item 6: Were costs and consequences valued
200 credibly?) on the checklist, examples in Drummond et al. (2015) were consulted to
201 overcome these disagreements. Practical application of item 10 (Item 10: Did the
202 presentation and discussion of study results include all issues of concern to the users?) was
203 challenging due to the limited guidance, thus findings from this question are less informative.
204 Alayli-Goebbels et al. (2014) also experienced this barrier in an earlier version of the
205 checklist.

206 **Method of analysis**

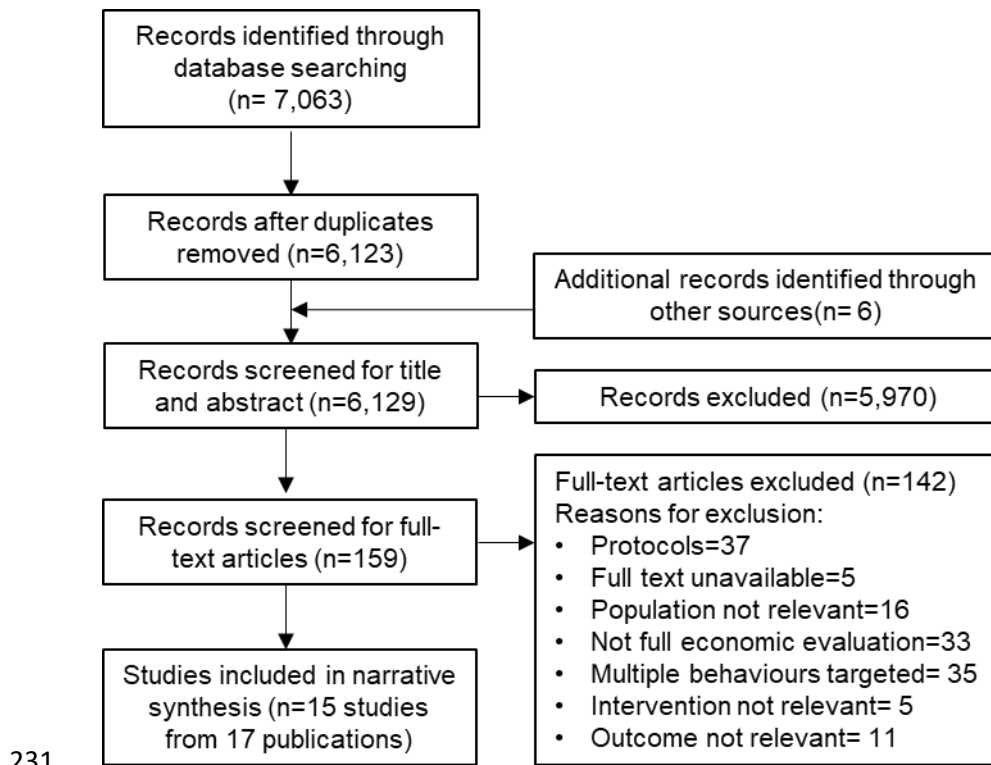
207 The published narrative synthesis framework by Popay et al. (2006) guided the analysis to
208 ensure a transparent and systematic approach was performed. The narrative synthesis in
209 this review goes beyond describing how authors have addressed each of the four challenges

210 by attempting to explain why specific approaches have been chosen. The analysis was an
211 iterative process. A priori analysis involved tabulating the data and producing bar charts on
212 key study characteristics: study design, time horizon, valuation technique, study perspective
213 and explicit/ implicit equity analysis. The same study characteristics were focused on in the
214 two former methodological reviews (Alayli-Goebbels et al., 2014; Weatherly et al., 2009).
215 The wider literature also indicated that the following contextual factors were important to
216 review when understanding an analyst's approach: intervention setting, country and year of
217 publication. Additional ad hoc analyses were performed where trends became apparent.
218 Lastly, the strength of the narrative synthesis and the conclusions derived from it were
219 considered by reflecting on the quantity of studies and results of the quality assessment.

220

Results

221 A total of 15 economic evaluations (17 publications) were included in the review (Figure 1).
222 Searching across Medline, SPORTSDiscus, EconLit, PsychINFO, NHS EED and HTA
223 databases retrieved 7,063 records. Supplementary searching retrieved six additional records
224 including: two records from hand searching on key websites, two from the reference list of a
225 systematic review (Gc et al., 2016), and a further two from searching for the completed
226 studies of two protocols (de Vries et al., 2013; Kolt et al., 2009) in Appendix E [INSERT LINK
227 TO ONLINE FILE A, B, C, D & E]. After removing duplicates 6,129 records remained of which a
228 further 5,907 records were removed as title and abstracts did not meet the eligibility criteria.
229 During the full text screening, 159 citations were examined in further detail, of which 142
230 studies were excluded. Reasons are outlined in Figure 1.



232 Figure 1. PRISMA flow diagram representing study selection process

233 **Study characteristics**

234 Of the 15 studies, ten were single trial-based economic evaluations and five were model-
 235 based; no studies were single trials that had extrapolated or modelled their results. Table 1
 236 provides an overview of study characteristics for the trial- and model-based studies
 237 respectively. Studies are arranged by country followed by year of publication. Interventions
 238 were set in primary care, community and the home, and setting did not appear to be related
 239 to intervention type or country. As shown in Table 1, no studies targeted sedentary
 240 behaviour as an independent risk factor from physical activity. The range of interventions
 241 was limited to the following types: physical activity programme/ on prescription in primary
 242 care (n=9); brief advice in primary care (n=2); home-based informational advice (n=1);
 243 physical therapy in a physical therapy setting (n=1); and fall prevention programme in both
 244 primary care and the home (n=1). The remaining study compared strategies for recruiting to
 245 physical activity interventions in primary care. The overall range of adult-based interventions
 246 matches the narrow range identified in a recent review of reviews focussing on the economic
 247 results of physical activity interventions (Abu-Omar et al., 2017). Studies came from four

248 high-income countries. More than half (n=8) of the 15 studies came from the UK, with the
249 remaining coming from New Zealand (n=3), the USA (n=2), and the Netherlands (n=2)
250 (Table 1).

251 **Quality assessment**

252 Overall, studies performed well against Drummond's 10-item quality assessment checklist
253 (Drummond et al., 2015) in Appendix D [INSERT LINK TO ONLINE FILE A, B, C, D & E].
254 Nevertheless, six studies scored 'No' on at least one item: two studies did not state their
255 perspective (item 1); three studies did not include all costs and consequences relevant to their
256 stated perspective (item 4); one study did not discount its costs and consequences (item 7);
257 and one study did not report their price source (item 6). Interpretation on whether item 4 was
258 met by any of the ten trial-based economic evaluations who captured costs and outcomes at
259 two years or less, is up for debate. It could be argued that not all important and relevant costs
260 and consequences can be identified for studies, which do not take a systems approach (e.g.
261 if they do not consider the impact on the wider system in which an intervention is being
262 implemented nor capture the long-term impact) (Rutter et al., 2017; Squires et al., 2016). In
263 order to align with other reviews which have used Drummond's checklist, the quality
264 assessment results for item 4 were based on the checklist's accompanying guidance
265 (Drummond et al., 2015). Costs and consequences identified, measured and valued are
266 discussed in greater depth in the subsequent sections.

267 Table 1: Overview of economic evaluations

| Trial-based economic evaluations | | | | | | | | |
|---|----------------------------|---------|--|-------------|--|--|--|---------------------|
| Study & Year of publication | Stated perspective | Country | Population targeted | Sample size | Intervention | Comparator | Setting | Valuation technique |
| Iliffe et al. 2014 | Health sector | UK | Inactive ≥65 years old who had fallen less than times in the previous 12 months | 100 | Falls Management Exercise Programme (Weekly group exercise class & 2 home-based exercise sessions) | Usual care (no intervention); Otago Exercise Programme | Primary care & community (as Home-based) | CEA |
| Edwards et al. 2013; Murphy et al. 2012 | Multi-agency public sector | UK | Sedentary, and over 16 years, with risk factors for coronary heart disease, or mild to moderate anxiety, depression or stress. | 798 | ERS (primary care) | Information leaflet only | Primary care | CUA |
| Boehler et al. 2011 | Health sector | UK | Inactive adults, 16 to 74 years old | 46 | Opportunistic recruitment strategy for physical activity interventions | Disease register strategy; Hypothetical no intervention strategy | Primary care | CEA |

| | | | | | | | | |
|----------------------|--------------|-------------|---|-----|---|--|--------------------------|----------|
| Shaw et al. 2011 | Not reported | UK | Inactive, adults (age not defined) | 79 | Individualised walking programme: a pedometer and a 30-min consultation | Individualised walking programme: a pedometer, but and 5 min brief advice | Primary care | CEA |
| Larsen et al. 2015 | Payer | USA | Inactive Latina women, 18-65 years old | 266 | Home print-based mail-delivered MVPA intervention linguistically and culturally adapted for Latinas | Wellness contact (information on health topics excluding MVPA) | Home-based | CEA |
| Young et al. 2012 | Societal | USA | Women, following coronary artery bypass surgery | 40 | Symptom management intervention delivered by telehealth device to improve the physical activity level | Usual care, 2 week follow up call by the primary providers and cardiac specialists | Community | CEA |
| de Vries et al. 2016 | Societal | Netherlands | Sedentary adults (or at risk of losing active lifestyle in near future) with mobility problems, ≥70 years old | 130 | Patient-centred physical therapy | Usual care for physical therapy, less patient-centred | Physical therapy setting | CUA |
| Maddison et al. 2015 | Not reported | New Zealand | ≥18 years old with diagnosis of IHD | 171 | Exercise prescription and behavioural support | Usual care (participation in usual Cardiac Rehabilitation) | Home-based | CEA; CUA |

| | | | | | | | | |
|-------------------|--------------------------------------|-------------|--------------------------------|-----|---|--|--------------|----------|
| | | | within previous 3-24 months. | | by mobile phone text messages and internet | e.g. education session and psychological support) | | |
| Leung et al. 2012 | Public health system and participant | New Zealand | Inactive adults, ≥65 years old | 330 | Pedometer-based prescription, focus was on step-related goals | Green prescription, focus was on physical activity time-related goals | Community | CEA; CUA |
| Elley et al. 2011 | Societal | New Zealand | Inactive, 40- 74 years old | 974 | Enhanced green prescription, 10 min of brief advice and a written exercise prescription with telephone support at 9 months and 30min face-to-face support at 6 months | Usual care from GP (not standard green prescription, usual care from GP not defined) | Primary care | CEA |

| Model-based economic evaluations | | | | | | | | |
|---|---|---------|-----------------------------------|---|-----------------------------|---|--------------|---------------------|
| Study & Year of publication | Stated perspective | Country | Population targeted | Model type & size of simulation cohort | Intervention | Comparator | Setting | Valuation technique |
| Campbell et al. 2015 | Health Sector | UK | Sedentary adults, ≥50 years old | Markov model (100,000 simulation cohort) | ERS (primary care) | Usual care (refers to Pavey et al. 2011's definition) | Primary care | CUA |
| Anokye et al. 2012; Anokye et al. 2014 | Health sector; Health sector and participant for CCA | UK | Inactive, ≥33 years old | Markov model (100,000 simulation cohort) | Brief Advice (primary care) | Usual care (no intervention) | Primary care | CUA (and CCA) |
| Anokye et al., 2011 | Health sector | UK | Sedentary adults, 40-60 years old | Decision tree model (1,000 simulation cohort) | ERS (primary care) | Usual care (refers to Pavey et al. 2011's definition) | Primary care | CUA |

| | | | | | | | | |
|----------------------|--|-------------|-----------------------------------|---|--|--|----------------|---------------|
| Pavey et al. 2011 | Health sector CUA; Partial-societal for CCA | UK | Sedentary adults, 40-60 years old | Decision tree model (1,000 simulation cohort) | ERS (leisure centre) | Usual care (no active ingredient- PA advice or leaflets) | Leisure-centre | CUA (and CCA) |
| Over et al. 2012 | Health sector | Netherlands | Inactive, 20-65 year olds | Markov model (100,000 simulation cohort) | GP pedometer prescription, counselling combined with pedometer use | Usual care (no intervention) | Primary care | CUA |

268 ERS: Exercise Referral Scheme; GP: General Practitioner; MVPA: Moderate-to-vigorous physical activity; CEA: cost-effectiveness analysis; CUA: Cost-utility

269 analysis

270 **Challenge 1: Attribution of effects**

271 Two thirds (n=10) of the studies in this review, all trial-based, did not compare the costs and
272 consequences of the comparator groups beyond the trial follow up period (Table 3). More
273 specifically, one study compared costs and consequences over a two-year period (Elley et
274 al., 2011), the remaining nine had a time horizon of 12-months or less. For six of these
275 studies, authors referred to their short time horizon as a limitation of their study (Boehler et
276 al., 2011; de Vries et al., 2016; Edwards et al., 2013; Larsen et al., 2015; Leung et al., 2012;
277 Shaw et al., 2011). For instance, it precluded the incorporation of any potential long-term
278 healthcare savings (Larsen et al., 2015). Just one study suggested future modelling
279 exercises could be used to address this challenge (Edwards et al., 2013). Yet, for Shaw et
280 al. (Shaw et al., 2011) a short-time horizon was justified as they reported there was
281 insufficient data to extrapolate their results over the participants' lifetime.

282

283 By contrast, all five model-based studies extrapolated a pooled trial-derived effectiveness
284 estimate over the rest of the participants' lifetime; bridging the gap between the short- and
285 long-term evidence (Table 2). Nevertheless, the assumptions underpinning the model-based
286 studies varied considerably. Two studies (Anokye et al., 2011; Pavey et al., 2011) made
287 large assumptions unsupported by evidence about the duration of the effect, assuming that
288 any short-term change in physical activity observed in the trials 6-12 months after the
289 intervention, would be long-lasting. Over et al. (2012) employed a different approach by
290 extrapolating an effect estimate, observed at 18 weeks, over a 40-year time horizon (the life
291 expectancy of the participants). The authors assumed that only 25% of the effect recorded at
292 18 weeks would remain over the 40-year time horizon; they too reported that their
293 assumptions were unsupported by evidence. These findings demonstrate how studies will
294 vary according to the assumptions made. It is therefore important that end-users of cost-
295 effectiveness results check they agree with the assumptions that underpin the economic
296 evaluation.

297

298 Assumptions underlying the two other model-based studies (Anokye et al., 2012; Campbell
299 et al., 2015) were supported by three robust cohort studies. Campbell et al. (2015) replicated

300 Anokye et al.'s (2012) approach. More specifically, they linked the short-term change in
301 physical activity level observed in trial data, with Hu et al.'s (2007; 2003; 2005) cohort
302 studies that followed a group of active and inactive individuals for a duration of at least 10
303 years to predict how their activity levels and risk of disease changed over time. Anokye et al.
304 (2012) explain how their identification and use of the cohort studies has strengthened
305 previous modelling attempts in the field of physical activity. Campbell et al.(2015) reported
306 this approach has enabled more conservative assumptions to be made around changing
307 physical activity levels and disease development over time.

308

309 Table 2. Time horizon and types of outcomes compared to costs

| Trial-based economic evaluations | | |
|----------------------------------|--|---|
| Study & Year of publication | Time Horizon (trial follow up) | Types of outcomes compared to costs per valuation technique |
| Larsen et al. 2015 | Trial duration (12 months) | CEA: Cost per minute of increase in physical activity |
| Iliffe et al. 2014 | Trial duration (12 months) | CEA: Cost per participant reaching or exceeding 150 minutes of moderate-to-vigorous physical activity per week |
| Young et al. 2012 | Trial duration (3 months) | CEA: Cost per incremental change in daily estimated energy expenditure; CEA: Cost per the incremental change in minutes spent on moderate-to-vigorous activity |
| Elley et al. 2011 | Trial duration (24 months; 12 months) | CEA: Cost per participant achieving 150 minutes of moderate intensity activity per week |
| Boehler et al. 2011 | Trial duration (3 months) | CEA: Cost per participant achieving 150 minutes of moderate intensity activity per week |
| Shaw et al. 2011 | Trial duration (12 months) | CEA: Cost per additional person achieving the target of a weekly increase of $\geq 15,000$ steps. |
| Maddison et al. 2015 | Trial duration (24 weeks / [6 months]) | CEA: Cost per MET-hour of walking and leisure activity; CUA: Cost per short-term QALY gain |

| | | |
|---|-----------------------------------|---|
| Leung et al. 2012 | Trial duration (12 months) | CEA: Cost per 30 minutes of weekly leisure walking; CUA: Cost per short-term QALY gain |
| de Vries et al. 2016 | Trial duration (6 months) | CUA: Cost per short-term QALY gain |
| Edwards et al. 2013; Murphy et al. 2012 | Trial duration (12 months) | CUA: Cost per short-term QALY gain |
| Model-based economic evaluations | | |
| Study & Year of publication | Time Horizon (trial follow up) | Types of outcomes compared to costs per valuation technique |
| Campbell et al. 2015 | Lifetime | CUA: Cost per short-term QALY gain (mental health gain); Cost per QALYs associated with coronary heart disease, stroke, type 2 diabetes due to reduced risk for developing these health states |
| Anokye et al. 2012; Anokye et al. 2014 | Lifetime | CUA: Cost per short-term QALY gain (mental health gain); Cost per QALYs associated with coronary heart disease, stroke, type 2 diabetes due to reduced risk for developing these health states CCA: Same outcomes outlined below for Pavey et al.'s (2011) CCA |
| Anokye et al., 2011 | Lifetime | CUA: QALYs associated with coronary heart disease, stroke, type 2 diabetes due to reduced risk for developing these health states |

| | | |
|-------------------|----------|--|
| Pavey et al. 2011 | Lifetime | <p>CUA: Cost per short-term QALY gain (mental health gain); Cost per QALYs associated with coronary heart disease, stroke, type 2 diabetes due to reduced risk for developing these health states</p> <p>CCA: Mental health (anxiety), Mental health (depression), Metabolic diabetes, Colon cancer, Breast cancer , Lung cancer, Hypertension (cardiovascular), Coronary Heart Disease, Stroke, Musculoskeletal (Osteoporosis), Musculoskeletal (Osteoarthritis), Lower back pain, Rheumatoid arthritis, Falls prevention, Absenteeism at work, Injury (disbenefit), Disability</p> |
| Over et al. 2012 | Lifetime | CUA: QALYs associated with myocardial infarction, stroke, diabetes, colorectal cancer, breast cancer due to reduced risk for developing these health states |

310

311 RCT: randomised controlled trial; cRCT: cluster randomised controlled trial; CEA: Cost-effectiveness analysis; CUA: Cost-utility analysis; CCA: cost-
312 consequence analysis ; MET: Metabolic Equivalent of Task

313 **Challenge 2: Measuring and valuing outcomes**

314 No studies in this present review conducted a cost-benefit analysis (CBA), despite health
315 economists (Drummond et al., 2015) stating this approach is superior to cost-utility analysis
316 (CUA) (Drummond et al., 2015). Recent UK and US guidelines recommended that studies
317 report a broad range of outcomes alongside their economic analyses, through the use of
318 approaches such as CBA, cost-consequence analysis (CCA) or an impact inventory (NICE,
319 2014; Sanders et al., 2016). Two studies (Anokye et al., 2012; Pavey et al., 2011) included a
320 CCA conducted alongside a CUA. A broad range of health outcomes were included in their
321 CCA (Table 2) yet the only non-health outcome reported was absenteeism.

322 Two thirds (n=11) of the studies presented just one type of valuation technique, either a
323 CUA (n=5) or cost-effectiveness analysis (CEA) (n=6) (Table 2). Table 2 demonstrates
324 further how despite having the same aim to increase physical activity levels and same
325 valuation technique, the way results are presented to the end-user are inconsistent. Young
326 et al. (Young et al., 2012) performed two CEAs reporting on the 'cost per incremental
327 change in daily estimated energy expenditure' and 'cost per incremental change in minutes
328 spent on moderate-to-vigorous activity'. Three other studies (Boehler et al., 2011; Elley et
329 al., 2011; Iliffe et al., 2014) performed a different type of CEA reporting on 'cost per
330 participant achieving 150 minutes of moderate physical activity per week'. The most
331 common way to present the result of the valuation analysis was as 'cost per short-term
332 quality-adjusted life year (QALY) gain'. Nevertheless, this was reported for just under half
333 (n=7) of the economic evaluations: four trial-based (de Vries et al., 2016; Edwards et al.,
334 2013; Leung et al., 2012; Maddison et al., 2015) and three model-based (Anokye et al.,
335 2012; Campbell et al., 2015; Pavey et al., 2011) studies. All model-based studies
336 conceptualised the long-term gain in QALY in the same way, in terms of the QALYs gained
337 due to not developing coronary heart disease, stroke or type 2 diabetes, or experiencing
338 premature mortality. Over et al.'s (2012) analysis differed slightly, as they also included
339 colorectal and breast cancer.

340

341 Rationale for the inclusion and exclusion of trial-derived QALYs varied considerably. Shaw
342 et al. (Shaw et al., 2011) argued against the inclusion of trial-derived QALYs in their

343 analysis, explaining it would be unnecessarily restrictive since evidence already shows that
344 physical activity is associated with a reduction in NCD and premature mortality, which in turn
345 is associated with a much greater gain in QALYs than trial-derived QALYs. Three model-
346 based studies (Anokye et al., 2012; Campbell et al., 2015; Pavey et al., 2011) deemed it
347 appropriate to incorporate both short-and long-term gain in QALYs. They conceptualised the
348 short-term QALY gain as being a one-off gain in mental health, which they assumed would
349 be achieved as a result of becoming physically active for at least 90 minutes per week. They
350 assumed the one-off mental health benefit would last for just one year, which they claimed
351 was a conservative assumption. Campbell et al. (2015) reported that their cost-effectiveness
352 result was highly sensitive to the inclusion and exclusion of the one-off gain in mental health.
353

354 **Challenge 3: Identifying intersectoral costs and consequences**

355 The most commonly reported perspective was the health sector perspective (n=7) (Table 1
356 and 2). Six of the eight studies from the UK were from this perspective. In 2014, the UK
357 reference case was updated to recommend the public sector perspective when conducting
358 economic evaluations of public health interventions (NICE, 2014). The multi-agency public
359 sector perspective adopted by Edwards et al. (2013) reflects the start of this paradigm shift.
360 Two more recent UK studies (Campbell et al., 2015; Iliffe et al., 2014) did not adopt a public
361 sector perspective. Despite studies being conducted from the same perspective, the type of
362 costs identified as relevant varied within and across countries and intervention type. This
363 weakness was identified through the quality assessment (Item 4 on Appendix D [INSERT
364 LINK TO ONLINE FILE A, B, C, D & E]), as five studies (Boehler et al., 2011; de Vries et al.,
365 2016; Maddison et al., 2015; Shaw et al., 2011; Young et al., 2012) did not relate their costs
366 to a study perspective. More specifically, two studies did not report their perspective
367 (Maddison et al., 2015; Shaw et al., 2011) and three included a narrower range of costs and
368 consequences than would be expected for their stated perspective (Boehler et al., 2011; de
369 Vries et al., 2016; Young et al., 2012). For example, two studies stated their study was from
370 the societal perspective yet assessed only direct intervention costs and short-term
371 healthcare savings (de Vries et al., 2016; Young et al., 2012), which were the same costs as

372 studies which stated taking a health sector perspective (Table 1 and 2). Weatherly et al.
 373 (2009) also found that many studies included only a narrow range of costs within their stated
 374 study perspectives.

375 Figure 2. Cost categories identified across all 15 included studies

376 Figure 2 shows that seven cost categories were identified across all 15 included studies.
 377 Like the findings in this review, Alayli-Goebbels et al. (2014) found the most common type of
 378 cost reported was the intervention costs, followed by healthcare costs. Participant out-of-
 379 pocket expenses and productivity losses appeared in only a small proportion of studies in
 380 this review and Alayli-Goebbels et al.'s (2014) review. Although most studies looked at both
 381 the direct and indirect costs of the interventions, only Edwards et al. (2013) looked at the
 382 unintended productivity costs to the provider. More specifically, they examined whether the
 383 provider where the intervention was set (the leisure centre) experienced a loss in revenue,
 384 as a result of providing the intervention.

385 **Challenge 4: Incorporating equity**

386 The two former reviews found that authors did not routinely consider equity in their analysis
 387 (Alayli-Goebbels et al., 2014; Weatherly et al., 2009). Table 3 shows that all but one study
 388 (Shaw et al., 2011) included in the present review did consider equity. All but one study
 389 (Edwards et al., 2013) did this implicitly, conducting subgroup analyses of the cost-
 390 effectiveness result (n=6) or targeting the intervention at a population deemed in need of
 391 intervention (n=8). Edwards et al. (2013) were the only authors to explicitly discuss equity
 392 and to consider socio-economic status in their equity analysis. They did this by asking
 393 participants from areas of different levels of deprivation about how much they would be
 394 willing to pay to participate in the intervention of interest; thus informing the reader about
 395 participants' economic preferences. Notably this was an exploratory analysis and so the
 396 results were not incorporated in the CUA.

397 Table 3. Types of equity considered

| | | |
|--|----------------------|------------------------|
| | Campbell et al. 2015 | Pre-existing condition |
| | Pavey et al. 2011 | Pre-existing condition |

| | | |
|--|---|--|
| Subgroup analyses of cost-effectiveness result | Anokye et al. 2011 | Pre-existing condition |
| | Edwards et al. 2013; Murphy et al. 2012 | Medical diagnosis |
| | | Referral reason |
| | | Adherence to scheme |
| | | Gender |
| | | Inequalities |
| | | Age group |
| | Over et al. 2012 | Age group |
| Anokye et al.2012 ; Anokye et al. 2014 | Age group | |
| Intervention targeted at equity group | de Vries et al.2016 | Frail older adults with mobility problems |
| | Leung et al.2012 | Older adults |
| | Iliffe et al. 2014 | Older adults |
| | Boehler et al. 2011 | Older adults |
| | Maddison et al. 2015 | People with ischaemic heart disease |
| | Elley et al. 2011 | Females |
| | Young et al. 2012 | Females |
| | Larsen et al. 2015 | Latinas |
| Willing to pay question | Edwards et al. 2013; Murphy et al. 2012 | Socio-economic status (level of deprivation) |

398

399 Table 3 details the eight studies which targeted their intervention at a specific population
400 group as well as the six studies that performed subgroup analyses of their cost-effectiveness
401 result. Older adults was the most common equity subgroup targeted for intervention (Boehler
402 et al., 2011; de Vries et al., 2016; Iliffe et al., 2014; Leung et al., 2012), followed by females
403 (Elley et al., 2011; Young et al., 2012). The most common subgroup analyses were on pre-
404 existing condition/ medical diagnosis (Anokye et al., 2011; Campbell et al., 2015; Edwards et
405 al., 2013; Pavey et al., 2011) and age group (Anokye et al., 2012; Edwards et al., 2013;
406 Over et al., 2012). Edwards et al.(2013) carried out seven types of equity analyses, all other
407 authors conducted just one type. Furthermore, no studies attempted alternative equity
408 analyses, such as an equity constraint or equity weighing analysis (Cookson et al., 2017).

409 **New studies**

410 The results of the rapid systematic scoping search are presented in Appendix B [INSERT
411 LINK TO ONLINE FILE A, B, C, D & E]. In brief, four additional studies were identified as
412 meeting the inclusion criteria of this review. Notably, one study (Gao et al., 2018) was an
413 intervention targeting sedentary behaviour as an independent risk factor from physical
414 activity. Furthermore, two studies (Gao et al., 2018; Harris et al., 2018) were both trial-and
415 model-based economic evaluations, as the analysts had extrapolated their within-trial results
416 a lifetime horizon.

417 **Discussion**

418 This review identified 15 economic evaluations of interventions that targeted physically
419 inactive adults, and no economic evaluations of interventions that targeted sedentary adults
420 (where sedentary behaviour was addressed an independent risk factor from physical
421 activity). Like Abu-Omar et al's (2017) review of reviews which focuses on the results of
422 economic evaluations, this present review identified economic evaluations on a limited range
423 of physical activity interventions (Abu-Omar et al., 2017). Studies came from just four high-
424 income countries, with over half (n=8) coming from the UK. This points to an important
425 evidence gap in countries where economic evaluations are deemed appropriate. Examining
426 a country's traditional beliefs around personal responsibility, efficiency and equity can
427 explain why countries such as France and Germany are low users of economic evaluations
428 and can in part explain why no studies in this review originated from these countries (Torbica
429 et al., 2018). Regardless of cultural and institutional differences, globally health economists
430 agree economic evaluations of preventative interventions are expected to have an important
431 impact on future healthcare decision-making (ISPOR, 2018). In order to answer upcoming
432 complex public health challenges, researchers need to go beyond clinical effectiveness
433 methods and use a multidisciplinary suite of methods (Rutter et al., 2017) which includes
434 economic evaluation. A prerequisite for this is an understanding on how key methodological
435 challenges can be addressed.

436 **Challenge 1: Attribution of effects**

437 *Modelling exercises*

438 All ten trial-based economic evaluations in this review had a short time horizon; meaning
439 they did not attempt to extrapolate or model the long-term impact of the intervention which
440 could be used to inform longer term investment decision making. Any future reduction in
441 incidence of NCD and premature mortality, attributable to physical activity and sedentary
442 behaviour interventions, is unlikely to manifest until decades after the intervention has taken
443 place. Yet, evaluating these interventions over the wrong timeframe means these
444 interventions may appear ineffective or markedly less effective; they are at risk of not being
445 appropriately prioritised by policymakers (Rutter et al., 2017). Curative interventions that
446 rescue people from very poor health to better health will continue to be favoured, even if
447 they are less cost-effective overall. Alayli-Goebbels et al. (2014) had previously suggested
448 modelling as a way to extend the time horizon of trial-based studies, yet none of the ten trial-
449 based studies in this review performed any modelling exercises. The challenges which can
450 preclude extrapolation include the availability of data, and time and skills of the analyst
451 (Squires et al., 2016).

452 *Cohort studies*

453 Campbell et al. (2015) and Anokye et al. (2012) were the only two studies in this review to
454 identify additional evidence to link up their short- and long-term effect estimate. The three
455 other model-based studies claimed there was insufficient evidence to verify the accuracy of
456 their assumptions (Anokye et al., 2011; Over et al., 2012; Pavey et al., 2011). Notably, the
457 cohort studies which Campbell et al.(2015) and Anokye et al.(2012) draw on were published
458 several years prior to the publication of the three other model-based studies. This suggests
459 that the methodological challenge of 'attribution of effect' may be more dependent upon the
460 analysts' time and skills as opposed to the availability of data.

461

462 **Challenge 2: Measuring and valuing outcomes**

463 *Cost-effectiveness and cost-utility analyses*

464 This review found large inconsistencies in the types of outcomes measured and valued.
465 There is no agreed classification system for physical activity outcomes (Abu-Omar et al.,

466 2017) since the analysis of raw objective accelerometer data measuring objective physical
467 activity levels is still in its infancy. Presenting a limited range of results can reduce the
468 applicability of the study's findings to other policymakers. Authors' views also differed firstly
469 on whether short-term QALYs should be included in the economic analysis, secondly on
470 whether a short-term QALY gain represented a one-off gain in mental health or general
471 functional health. Presently, within the economic literature the responsiveness of the EQ-5D-
472 3L to detect important differences in the severity of health is being challenged, and had led
473 to the development of the EQ-5D-5L, which measures health on five levels as opposed to
474 just three (Glick et al., 2014). This review has shown that outcomes used in physical activity
475 studies are diverse; therefore, there is a need for analysts to agree on a consistent outcome
476 that best captures the objectives of a physical activity intervention.

477 *Cost-benefit and cost-consequence analyses*

478 No studies in this review performed a CBA and just two presented a CCA alongside their full
479 economic evaluation. There is a lack of CBAs in other public health areas. Hill et al. (2017)
480 and Alayli-Goebbels et al. (2014) identified a small proportion of studies (n=1 and n=8
481 respectively) who reported conducting a CBA, but due to insufficient reporting gained limited
482 insight into how these were performed such as how outcomes had been monetised (Alayli-
483 Goebbels et al., 2014; Hill et al., 2017). Likewise, four studies claimed to be CBAs in the
484 review by Weatherly et al.(2009), but after further assessment were re-classified as CCAs
485 (n=3) and a CEA (n=1). Although classified as a partial-economic evaluation, CCA is a
486 useful alternative to CBA since all relevant costs and consequences can be presented to the
487 reader in the form of an inventory, rather than simplified into a single outcome measure or
488 index as is the case in CEA and CUA, respectively. If an outcome is deemed relevant to the
489 reader, they can reanalyse the data quantified in the CCA. However, CCA puts more onus
490 on decision makers than CBA or CUA, as it does not roll outcomes into a summary measure
491 that can be compared to a decision rule. An example of a decision rule in the UK is: invest
492 where the incremental cost-effectiveness ratio is less than £30,000 per QALY (NICE, 2014).

493

494 **Challenge 3: Identifying intersectoral costs and consequences**

495 *Inconsistent perspectives*

496 The three most common perspectives stated were the health system, payer and societal
497 perspectives. These match the three most commonly reported perspectives in the broader
498 field of economic evaluation (Husereau et al., 2013). Only Edwards et al. (2013) conducted
499 their analysis from the public sector perspective, a perspective recently recommended in the
500 UK reference case (NICE, 2014). That said, Edwards et al. (2013) did not incorporate
501 participant costs in their CUA, only through an exploratory analysis. Only three studies
502 considered the cost to the participant, which is not surprising since the health sector
503 perspective was the most commonly stated perspective. Participant and voluntary sector
504 costs are deemed important, but previously have not been routinely captured (Weatherly et
505 al., 2009).

506 It was found that even economic evaluations stated the same perspective did not always
507 include the same costs and consequences. This is likely to be because there is a lack of
508 standard definitions for the various perspective types (Husereau et al., 2013). Even where
509 there are examples of standard definitions, such as those proposed by the Second US Panel
510 on Cost-Effectiveness in Health and Medicine (Sanders et al., 2016), not all economists
511 agree with their definitions, and furthermore the definitions may not be applicable to other
512 countries since there are distinct features of each health system (Torbica et al., 2018). For
513 instance, deciding what costs and consequences to capture within a societal perspective is a
514 normative question, requiring the analyst to make social value judgements (Drummond et
515 al., 2015). This is an important issue, since the exclusion of relevant consequences can lead
516 to an underestimation of cost-effectiveness whilst the exclusion of relevant costs can lead to
517 an overestimation of cost-effectiveness (Hill et al., 2017).

518 *Cost categories identified*

519 The cost categories identified in this review match the five cost categories (healthcare
520 services, intervention costs, patient and family costs, lost productivity costs, future costs)
521 identified as most relevant for inclusion in economic evaluations, by health economists who
522 recently took part in a cross-Europe Delphi study (van Lier et al., 2017). This suggests
523 analysts' choice in costs in this review align with analysts in the more general field of

524 economic evaluation. It should be noted however that there was a difference in one of the
525 categories, as family costs were not identified as a relevant cost category in the studies from
526 this present review. Just two trial-based studies included absenteeism in their study;
527 similarly only two of the model-based studies included it in their CCA. It continues to be
528 debated in the literature as to whether absenteeism is an outcome of cost-offset, and thus
529 whether it should be included in the numerator or denominator part of the incremental cost-
530 effectiveness fraction (Drummond et al., 2015).

531 **Challenge 4: Incorporating equity considerations**

532 *Presenting results by subgroups*

533 Equity impact analysis can be as straightforward as presenting cost-effectiveness results by
534 equity subgroups (Alayli-Goebbels et al., 2014; Hill et al., 2017; Weatherly et al., 2009). Six
535 studies in this review presented an equity impact analysis (Anokye et al., 2012; Anokye et
536 al., 2011; Campbell et al., 2015; Edwards et al., 2013; Over et al., 2012; Pavey et al., 2011).
537 The most common subgroup analysed was individuals with pre-existing medical conditions,
538 nevertheless this analysis was performed in just four studies (Anokye et al., 2011; Campbell
539 et al., 2015; Edwards et al., 2013; Pavey et al., 2011). Furthermore, only one study
540 (Edwards et al., 2013) conducted more than one type of equity subgroup analysis. These
541 findings suggest analysts are not performing equity analyses in a comprehensive nor
542 consistent manner. Weatherly et al. (2009) outlined socio-economic status as an important
543 under-researched equity issue in economic evaluations, however only one study in this
544 review researched socio-economic status by asking participants about their willingness to
545 pay for an intervention component (Edwards et al., 2013). Incorporating equity into decisions
546 on physical activity and sedentary behaviour interventions is especially important, since it is
547 amongst the lower socioeconomic groups where physical inactivity is greatest (OECD,
548 2015).

549 **New studies**

550 Overall, the four studies published since March 2017 did not change the narrative of this
551 review since there remains a dearth of economic evaluations in the field of physical activity
552 and sedentary behaviour. What the studies have demonstrated is that firstly, there is an

553 indication that health economic methods have begun to be applied to targeted sedentary
554 behaviour interventions (Gao et al., 2018). Secondly, that it is feasible and informative to
555 extrapolate beyond the trial (Gao et al., 2018; Harris et al., 2018).

556 **Strengths and limitations**

557 This is the first systematic review conducted since 2009 to review the methods used in
558 economic evaluations of interventions targeted at physically inactive individuals, and the first
559 systematic review to search for economic evaluations targeting sedentary behaviour as an
560 independent risk factor from physical activity. This review included comprehensive literature
561 searching and a rigorous methodology in line with the PRISMA guidelines (Moher et al.,
562 2009). Economic evaluations aim to inform resource allocation decisions (Drummond et al.,
563 2015). Previous reviews have demonstrated that key methodological challenges preclude
564 economic evaluations in the field of public health from achieving this aim (Alayli-Goebbels et
565 al., 2014; Weatherly et al., 2009). By focusing on physical activity and sedentary behaviour,
566 this review has been able to not just provide an overview on whether or not the four key
567 methodological challenges have been addressed in the last decade, but crucially explain in
568 greater depth the methods performed in those few studies where progress has been made.

569 More specifically, progress has been observed in the 14 studies which have considered
570 equity in their analysis (Table 3) and the small proportion of studies where either: the long-
571 term model presented has been informed by robust epidemiological evidence (Anokye et al.,
572 2012; Campbell et al., 2015); all important and relevant costs and consequences have been
573 outlined to the reader in the form of a cost-consequence analysis (Anokye et al., 2012;
574 Pavey et al., 2011); and/or a multi-sector perspective has been selected (Edwards et al.,
575 2013). An output from the narrative synthesis of this review is a number of recommendations
576 (as outlined in Table 4) explaining how analysts can continue to make progress towards
577 addressing the four methodological challenges. Although, the comprehensive search
578 strategy only goes upto March 2017, a rapid systematic scoping search is presented which
579 highlights four new empirical studies. Two of these studies (Gao et al., 2018; Harris et al.,
580 2018) support the recommendations emerging from this review in terms of linking up the
581 intermediate evidence with longer term policy relevant outcomes.

582 It was not within the scope of this research to review the methods used in population-level
583 interventions such as national policies or media campaigns. It would therefore be useful for
584 future reviews to explore how economic evaluations are being carried out within this area. In
585 addition, this review focuses on the methods conducted in full economic evaluations and so
586 there is scope to review the methods used in partial evaluations. Nevertheless, full economic
587 evaluations are deemed more informative than partial evaluations, and so it would have
588 been expected that analysts would conduct for instance, a CCA alongside their full economic
589 evaluation, as was done in two studies (Anokye et al., 2014; Pavey et al., 2011) in this
590 review.

591

592

Recommendations

593 Table 4 presents a list of recommendations for researchers and users of economic
594 evaluations from a variety of disciplines (health economics, public health, physical activity
595 etc) to refer to when designing, analysing and appraising economic evaluations of
596 targeted physical activity and sedentary behaviour interventions.

597

598 **Table 4. Recommendations for future economic evaluations**

| Challenge | Recommendation | Explanation |
|--|---|--|
| Challenge 1. Attribution of Effects | <i>Visual representations of disease pathways</i> | <p>It is necessary for public health researchers to invest time in reviewing the existing evidence base and develop novel modelling skills. Best practice guidelines state well established published models are preferred to those developed specifically for a trial (Ramsey et al., 2015). If skill and time permits, analysts can draw on the structure of the published models (Anokye et al., 2012; Campbell et al., 2015) identified in this review and adapt them according to the local decision-making context. All five models in this review presented a visual depiction of the disease pathway for physical activity. Authors from non-economic disciplines could build on the disease pathways presented in the model-based studies in this review, in order to help policymakers and those designing interventions to consider the long-term costs and consequences of investing or disinvesting in physical activity interventions. The visual could be as simple as a logic model, a visual tool recommended for public health interventions (Moore et al., 2015).</p> |
| | <i>Long-term objective data derived from cohort studies</i> | <p>Future investment and disinvestment decisions should be informed by economic evaluations which not only assess the short-term impact of interventions, but also impact on the medium- and long-term (Academy of Medical Sciences, 2016). As long-</p> |

| | | |
|--|---|--|
| | | <p>term RCTs of physical activity and sedentary behaviour interventions are likely to be impractical or unethical, evidence from non-experimental studies such as cohort studies could be drawn on to evidence the long-term impact of physical activity and sedentary behaviour interventions as done in two studies. In the hierarchy of evidence, cohort studies are recognised as being the next best alternative to RCTs (Murad et al., 2016). The popularity of wireless-enabled wearable activity monitors in high-income countries present researchers with an opportunity to conduct more cohort studies and collect objective data on behaviour change over a longer time period.</p> |
| <p>Challenge 2. Measuring and valuing outcomes</p> | <p><i>Quality of life measurement tools</i></p> | <p>Future research should aim to understand whether a short-term gain in QALY represents a one-off benefit in mental health due to becoming physically active. The EQ-5D tool, is the most commonly used tool to measure QALYs but only captures the functional health of an individual. Future studies could use other recently developed quality of life tools such as the ICECAP-A (Al-Janabi et al., 2012; Al-Janabi et al., 2013; Flynn et al., 2015), which has been designed to capture capability in a broader sense, beyond functional health. Another solution is for analysts to agree on a tool which crosswalks between physical activity outcomes and a summary tool like the EQ-5D. There is currently a mapping database of studies that map the EQ-5D tool to other</p> |

| | | |
|--|----------------------------------|---|
| | | outcomes measures (Dakin et al., 2018). No studies on the database have mapped a physical activity specific tool to the EQ-5D; future research should address this gap. |
| | <i>Cost-consequence analysis</i> | There is a need for further methodological developments in the monetisation of effects in CBAs (Drummond et al., 2015; Sanders et al., 2016). In the meantime, it is deemed more appropriate to conduct a good quality CUA which may be of a narrower perspective, than a poor quality CBA which captures a broader perspective (Hill et al., 2017; Weatherly et al., 2009). In order to report on multiple outcomes which extend beyond health, a CCA or impact inventory conducted alongside a full economic evaluation is recommended (NICE, 2014; Sanders et al., 2016). If the word limit in journals precludes authors from presenting a CCA in the main manuscript, they should present this information in the online supplementary material. |
| Challenge 3. Identifying intersectoral costs and consequences | <i>Multi-sector perspective</i> | Three studies in this review omitted costs which would typically be deemed relevant to their stated perspective, and two studies did not report their perspective. It is imperative for analysts to describe and justify the costs and consequences which they have deemed relevant for their chosen perspectives (Husereau et al., 2013). Inevitably different assumptions on what costs and consequences are included in the analysis leads to different results (Sanders et al., 2016). Furthermore, future studies should aim |

| | | |
|--|---|--|
| | | <p>to present at least two types of perspectives and conduct a CCA or impact inventory alongside their CUA or CEA in order to present the various relevant costs and consequences to the various relevant sectors (Alayli-Goebbels et al., 2014; Sanders et al., 2016; Weatherly et al., 2009). A multi-sector perspective where costs and consequences are presented in their disaggregated form (i.e. in a CCA) for each sector is preferred over stating a societal perspective (Drummond et al., 2015; Hill et al., 2017).</p> |
| | <p><i>Systems thinking approach</i></p> | <p>Absenteeism was the only non-health effect identified in the two CCAs in this review (Anokye et al., 2012; Pavey et al., 2011). During the design stage of future economic evaluations analysts could conduct multi-stakeholder and expert consultations to map out which costs and consequences are deemed relevant to physical activity and sedentary behaviour interventions (Squires et al., 2016). A systems thinking approach (Rutter et al., 2017; Squires et al., 2016) is recommended to ensure interventions' indirect and unintended costs and consequences on the whole system are considered, not just those experienced by the health sector or payer. Two recently published frameworks can help analysts apply a systems approach (Cylus et al., 2016; Squires et al., 2016).</p> |

| | | |
|--|----------------------------------|--|
| Challenge 4. Incorporating equity | <i>Equity impact analysis</i> | <p>Analysts should present costs and consequences explicitly in their disaggregated form for various equity groups, so policymakers can start to build a better picture on which population groups gain and lose from a specific decision (Hill et al., 2017). From here, analysts can conduct an equity impact analysis. This type of analysis is deemed easier than conducting equity constraint or equity weighting analysis (Hill et al., 2017). The equity effectiveness loop framework (Welch et al., 2008) and PROGRESS-Plus framework (O'Neill et al., 2014) are recommended to help analysts consider, in a structured way, which equity factors may be relevant to their study (Alayli-Goebbels et al., 2014; Welch et al., 2017).</p> |
| | <i>Participant's preferences</i> | <p>Other types of equity-related analyses not identified in this review, but which future studies could investigate, include the public's perspective on trading off efficiency with equity (in public services) (Weatherly et al., 2009). It is also recommended that future studies, specifically trial-based studies, capture economic information on time, travel and out-of-pocket expenses incurred by the participant. The APEASE criteria by Michie et al. (2014) could also help analysts to consider the acceptability and affordability of an intervention to various stakeholders. Inevitably, these two issues will contribute to the success of interventions aiming to change behaviour (Michie et al., 2014).</p> |

600

601

Conclusions

602 A focus on the key methodological challenges in economic evaluations is important, as they
603 can impact on the derived cost-effectiveness result, which ultimately can impact on a
604 policymaker's resource allocation decision. As economic evaluation is a rapidly developing
605 field (Drummond et al., 2015) this systematic review has provided an important update on
606 the most recent methods used in targeted physical activity interventions. The review has
607 also highlighted there is a scarcity of economic evaluations for targeted sedentary behaviour
608 interventions. Importantly, this review makes it explicit to policymakers and researchers from
609 the varied disciplines in which physical activity and sedentary behaviour falls under, that
610 there are still key methodological challenges that need further attention. This review has
611 highlighted that methodological choices vary widely not just between countries but also
612 within them. Ultimately, these analyst-based choices affect the results presented and
613 subsequent resource allocation decisions made. A recent consensus statement has called
614 for collaboration across the disciplines to develop guidance specific to the context of
615 economic evaluations of physical activity interventions (Davis et al., 2014). To date, no
616 guidelines have been developed to address this need. The examples of methodological
617 development identified from the studies in this review and the resulting review
618 recommendations can be used to inform future guidelines and their supplementary
619 materials.

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