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

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### Article

**Citation** (please note it is advisable to refer to the publisher's version if you intend to cite from this work)

**Cochrane, MA, Watson, PM, Timpson, H, Haycox, A, Collins, B, Jones, L, Martin, A and Graves, LEF (2019) Systematic review of the methods used in economic evaluations of targeted physical activity and sedentary behaviour interventions. Social Science and Medicine. ISSN 0277-9536**

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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.

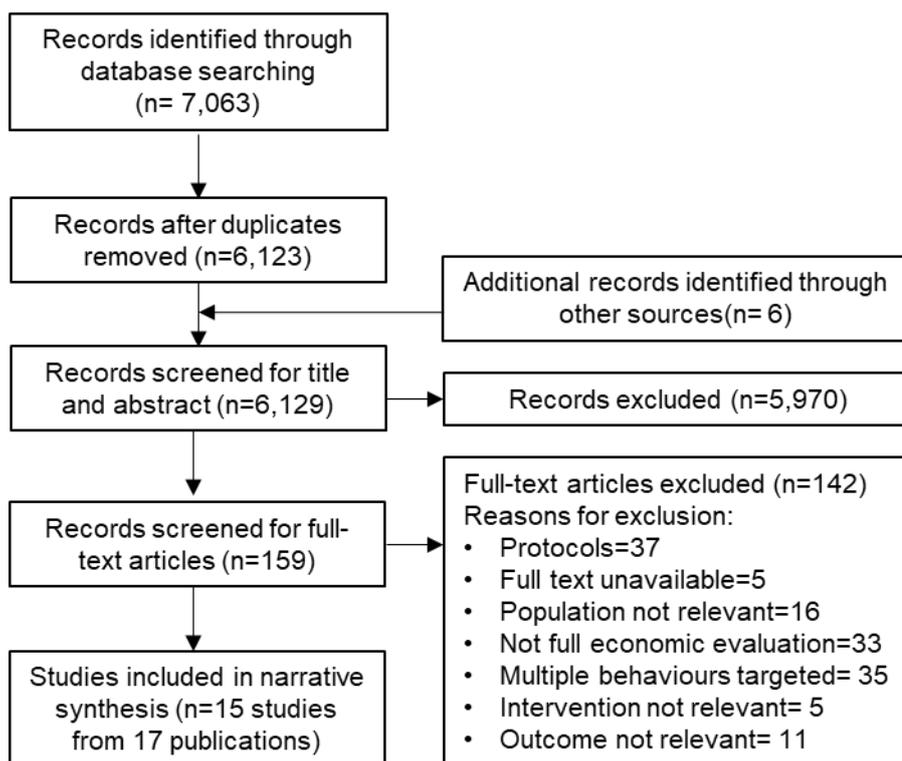
206 **Method of analysis**

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.



231

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 $\geq 65$ 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
<b>Challenge 1.</b> <b>Attribution of Effects</b>	<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><b>Challenge 2.</b> <b>Measuring and valuing outcomes</b></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.
<b>Challenge 3. Identifying intersectoral costs and consequences</b>	<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>

<b>Challenge 4.</b>  <b>Incorporating equity</b>	<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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