1 Abstract

2 *Objectives:* To identify barriers to medication adherence in patients prescribed medicines for 3 the prevention of cardiovascular disease and map these to the Theoretical Domains 4 Framework (TDF), to produce a conceptual framework for developing a questionnaire-based 5 medication adherence tool. 6 **Methods:** A scoping review of barriers to medication adherence in long-term conditions was 7 conducted to generate an initial pool of barriers. After preliminary mapping to the TDF, these 8 barriers were presented to two focus groups of patients prescribed medicines for the 9 prevention of cardiovascular disease (n=14) to stimulate discussion. The group discussions 10 enabled the patients' interpretations of the adherence barriers to be determined, provided 11 validity from the patient perspective, and identified additional barriers unrepresented in the 12 scoping review. 13 **Key findings:** The preliminary pool of adherence barriers was identified from 47 studies 14 across a range of long-term conditions. The majority of TDF domains were represented by 15 these literature-identified barriers except 'social/professional role and identity' and 'behavioural regulation'. Barrier mapping was largely endorsed by focus group participants, 16 17 who also contributed additional barriers, including those relating to not having a 'system' in 18 place for managing their medicines and the negative emotions evoked by medicine taking. 19 **Conclusion:** The TDF enabled full exploration of adherence barriers including those relating 20 to emotions which have received limited attention in the literature. This work has provided a 21 conceptual framework for developing a questionnaire to identify an individual's adherence 22 barriers which may then be coupled with appropriate behaviour change techniques to deliver a 23 theory-based intervention tailored for individual need.

- 24 Keywords: Scoping review, Theoretical Domains Framework (TDF), focus group,
- 25 questionnaire, IMAB-Q

Introduction

27

28

29

30

31

32

33

34

35

36

37

38

39

40

41

42

43

44

45

46

47

48

49

An estimated 30 to 50% of patients with long-term conditions (LTCs) are non-adherent to their prescribed medicines. A large-scale meta-analysis, estimated adherence to medicines for the prevention of cardiovascular disease (CVD) to be 57% (95% CI 50-64%).² These medicines are prescribed for a range of LTCs including hypertension, dyslipidaemia and angina and are amongst the most commonly prescribed medicines in the UK.² Medication adherence is a complex health behaviour, influenced by a plethora of factors.³ Non-adherence can diminish treatment effects leading to increased morbidity and mortality⁴ plus wasted healthcare resources.³ Evidence suggests that a greater understanding of the barriers to adherence is needed to improve the effectiveness of adherence interventions.⁵ A plethora of theoretical models have been developed to explain the complexities of medication adherence, including those focused on the balance between patient perceived necessity and concerns about medicines⁶ and those focused on the importance of practitioner consultation style⁷. Though these models highlight important considerations for medication adherence research, the most recent Cochrane review highlights that meaningful progress with adherence research is still sub-optimal. ⁵ Theoretical models such as Social Cognitive Theory, the Health Belief Model and Self-regulation model have been applied to medication adherence interventions.⁸ However, a systematic review of theory-based interventions to improve medication adherence identified that none have successfully guided the development of an effective adherence intervention applicable to all long-term medications⁸. Psychology-based behaviour change techniques, such as motivational interviewing, show promise as effective adherence interventions. However, core training of the existing healthcare workforce is not designed to equip practitioners in selecting the most appropriate

50 behaviour change techniques (BCT) for improving adherence, according to identified individual adherence barriers. 10,11,12, 51 52 Developing an adherence tool which identifies a patient's barriers to adherence and guides the 53 practitioner to work with the patient to select the most appropriate BCTs may enable the 54 healthcare workforce to respond to the call for theory and evidence guided, individualised interventions, ^{13, 14,} which identify potential barriers to behaviour change. ^{8, 155,} 55 The Theoretical Domains Framework (TDF)^{16, 17} is a composite of health psychology theory 56 which offers a structured approach for exploring the determinants of individual behaviour.¹⁸ 57 The domains of the TDF have been linked to evidence-based BCTs, ^{19,20} leading to successful 58 use of the TDF to guide the intervention development for behaviour change.²¹ The TDF may 59 60 therefore be suitable for mapping adherence barriers and creating a conceptual framework. Literature describing application of the TDF to medication adherence²²⁻²⁵ represents notable 61 62 advancements in the field. However, each study focusses on medication adherence in a 63 specific disease rather than multiple LTCs. Most patients have multiple diseases for which 64 they are prescribed multiple medicines; routine practice consultations such as medication 65 reviews are therefore not focused on medication adherence in one specific disease state. Intervention implementation is supported by compatibility with routine practice, ²⁶ thus, an 66 adherence support tool applicable across a range of LTCs is a stronger candidate for effective 67 implementation into routine practice.²⁷ Exploration of barriers to adherence in medicines 68 69 prescribed for the prevention of CVD (which covers multiple LTCs) is therefore an intuitive 70 opportunity to broaden TDF-based adherence research towards multiple LTCs, whilst 71 minimising the confounding factors that could be introduced by considering all LTCs 72 collectively.

The current article presents the developmental work which underpinned the Identification of Medication Adherence Barriers Questionnaire (IMAB-Q); ²⁸ a TDF-based questionnaire to support practitioners in identifying non-adherent patient's and elucidating their individual reasons for non-adherence. It comprises a scoping review of barriers to adherence in LTCs, the initial mapping of these barriers to the TDF and the qualitative exploration of these barriers in patients prescribed medicines for the prevention of CVD, in order to develop a conceptual framework to inform questionnaire development.

Existing literature syntheses (e.g. ^{29,30}) report quantitative findings from intervention studies and non-modifiable adherence determinants such as age, gender and socioeconomic status. Modifiable determinants of adherence, relating to psychosocial and environmental barriers are

and non-modifiable adherence determinants such as age, gender and socioeconomic status. Modifiable determinants of adherence, relating to psychosocial and environmental barriers are often overlooked. These reviews also consider non-adherence in all conditions, yet important differences in adherence determinants exist between acute and LTCs.³ A broader evidence synthesis, narratively combining both quantitative and qualitative studies may therefore provide a better foundation for exploring adherence barriers. Scoping reviews are an appropriate method to 'map' relevant literature and address broad topics where differing study designs are available.³¹

Correct mapping of adherence barriers to a theoretical framework requires deep understanding which cannot always be elucidated from the literature. Qualitative exploration to supplement a literature review can provide this depth of understanding,³² enhance the utility of a scoping review and ensure meaningful mapping.

Methods

- The programme of work included four phases:
- 97 1. Scoping review of barriers to medication adherence in LTCs

98 2. Preliminary mapping of literature-identified barriers to the TDF 99 3. Focus groups with patients prescribed medicines for the prevention of CVD 100 4. Refinement of adherence barriers mapping 101 Phase 1 Scoping review 102 This phase aimed to generate a preliminary repository of barriers to medication adherence in 103 LTCs, for stimulating focus group discussions. 104 Search strategy The Embase, Medline and PsychINFO databases were accessed via the Ovid interface on 18th 105 106 September 2012, to undertake the search detailed in supplementary file 1. The search was 107 restricted to articles written in English and since 2005, as scoping searches indicated that prior 108 to this, psychosocial determinants of adherence were seldom explored. Abstracts were 109 screened against pre-defined inclusion and exclusion criteria. 110 Inclusion and exclusion criteria 111 Abstracts of any study design, reporting medication adherence barriers in LTCs were eligible 112 for inclusion. LTCs beyond those covered by 'CVD prevention' were included to ensure 113 breadth of the preliminary pool of adherence barriers before later refinement. 114 Abstracts were excluded if they: 115 Included participants with drug addiction or mental health problems (the nature of non-

6

adherence in this population is condition-specific)

Data collection and synthesis (charting)

116

Full texts were accessed where possible, but when unavailable, adherence barriers were extracted from abstracts. Adherence barriers were initially recorded using the exact terminology in the article. Once all barriers had been extracted, barriers with the same underpinning characteristic but presented differently due to specifics of context or variations in language were grouped, for example 'forgetting to take medicines' and 'not remembering doses' were grouped as one barrier related to forgetting medicines.

Phase 2 Mapping of adherence barriers to the TDF

Adherence barriers were mapped to one of the 12 domains of the original TDF.¹⁶ . Existing literature^{16, 17, 33} were utilised to interpret each of the TDF domains in the context of barriers to medication adherence. Preliminary mapping was discussed by the authors until consensus was achieved about which barriers belonged to each domain.

Phase 3 Focus groups with patients prescribed medicines for the prevention of CVD

- Focus groups with patients prescribed medication for CVD prevention were undertaken to:
- 1. Identify additional adherence barriers not elicited from the scoping review
- 2. Optimise the research team's understanding of identified barriers
- 3. Ensure appropriate mapping of barriers to the TDF

134 Participants and recruitment

Recruitment commenced post ethical approval from the University of East Anglia Faculty of Health ethics committee (reference number 2012/2013-04). The large pool of employees and students at the university were used as potential participants and gatekeepers to the wider non-university community for recruitment. Recruitment was via posters placed across campus, a weekly e-bulletin emailed to all staff and students, and university social media.

Advertisements were worded to extend recruitment beyond university students and staff, to include their friends and family, thus increasing the likelihood of recruiting a diverse population. Participants were offered a £10 high street shopping voucher for participation.

Inclusion and exclusion criteria

Adults (individuals aged 18 years or older) able to provide informed consent were eligible if prescribed medication for the prevention of CVD as defined in the literature.² Those who were unable to read or speak English, or receiving medication for the treatment of addiction or mental illness were excluded.

Procedures

Eligible members of the public expressing interest in participation were posted a study information leaflet, consent form and brief questionnaire to collect demographic information, plus the number of medicines prescribed and prescription charge exemption status. Returned consent forms and questionnaires were used to assign participants to one of two focus groups. Two focus groups, each with six to eight participants was deemed to be appropriate for generating sufficient data for the exploratory nature of this stage, whilst not over-burdening members of the public. Recruitment continued until each focus group had between six and eight participants representing a range of demographic characteristics.

158 Focus groups

Each focus group was audio-recorded, approximately two hours long, transcribed verbatim and moderated by the lead author with co-facilitation. The TDF-domains deemed applicable to medication adherence barriers (established in phase one) were divided across the two focus groups. Adherence barriers mapped to differing behavioural domains were considered in each

focus group but the 'emotions' domain was duplicated to investigate consistency of interpretation between participants of the two focus groups. This domain was selected for duplication across both focus groups as it was considered to be the domain most likely influenced by differing personal experience; we therefore aimed to explore how these personal experiences differed across the largest possible number of participants. Each behavioural domain was described to participants in turn, before discussing the literature-identified adherence barriers mapped to the domain. The initial mapping of barriers to each domain of the TDF is provided in supplementary file 4; this mapping therefore served as the topic guide for the focus groups. Participants were encouraged to share their experiences and thoughts, using the adherence barriers presented as prompts for discussion. For each behavioural domain, participants were asked if there were any additional adherence barriers that were not represented. Data analysis Primary data analysis was undertaken by the lead author then validated by the co-authors as recommended in the literature.³⁴ Data were analysed using a framework approach,³⁵based upon the domains of the TDF. Phase 4 Refinement of adherence barriers mapped to the TDF and summary Data from the focus groups were used to refine the mapping of adherence barriers, according to the participants' understanding of their meaning and relevance. Any additional barriers generated during the consultation exercises were also considered. **Results**

163

164

165

166

167

168

169

170

171

172

173

174

175

176

177

178

179

180

181

182

183

184

Phase 1

Scoping review

Forty-seven eligible studies (representing a range of LTCs) were identified, from which the preliminary pool of adherence barriers were extracted. Similar barriers were initially grouped into 17 themes, (as summarised in supplementary file 2) which included beliefs, cognitive and memory associated factors, knowledge-related factors and administration problems.

Phase 2 Mapping of adherence barriers to the TDF

185

186

187

188

189

190

191

192

193

194

195

196

197

198

199

200

201

202

203

204

205

206

207

208

The agreed interpretations of how each behavioural domain of the TDF relates to medication adherence barriers are provided in supplementary file 3. All adherence barriers were considered carefully, though some required a deeper level of consideration and discussion. An interesting example here is the adherence barrier 'experience of side effects' which was ultimately mapped to the 'beliefs about capabilities' domain of the TDF. This decision was reflective of the recognition that it is not the side effects per se that influence medication adherence, but more an individual's ability to appropriately cope with the medication side effect that determines their behaviour. The 'skills' domain was considered to encompass both physical skills, (e.g. medicines administration) and cognitive skills (e.g. processing and understanding instructions). A number of barriers such as 'being too busy' and 'having a chaotic lifestyle' related to competing goals; these barriers did not intuitively map onto any of the existing behavioural domains of the TDF. Guided by relevant literature, ³³ an additional behavioural domain termed 'goal conflicts' was created. The behavioural domains termed 'social/professional role and identity' and 'behavioural regulation' were excluded as no literature identified adherence barriers were mapped to these domains. The constructs associated with the 'behavioural regulation' domain are barriers and facilitators to behaviour¹⁶; as the study was focused on barriers to medication adherence, the 'behavioural regulation' domain was redundant. The 'nature of the behaviour' domain was also excluded; Michie et al^{16} explain that this domain is accorded to a different order as it describes the

dependent variable, in this case, taking medicines as prescribed.³⁶ It is therefore not treated as a domain of behaviour change, but its constructs such as habits, were considered throughout the mapping task. Of the original 12 domain TDF, the three domains of 'social/professional role and identity', 'behavioural regulation' and 'nature of the behaviour' were not therefore active in the context of medication adherence barriers and an additional 'goal conflicts domain' was generated yielding 10 active domains in the present study.

The adherence barriers initially grouped to each TDF domain are detailed in supplementary file 4. Barriers were well distributed across the 10 relevant domains, though the beliefs about capabilities, beliefs about consequences and social influences domains had the broadest range of adherence barriers. Some barriers, for example 'no medical insurance' were excluded as they were not relevant to the UK healthcare system.

Phase 3 Focus groups with medicine-taking members of the public

Interest in focus group participation was expressed by 32 members of the public; signed consent forms and demographic questionnaires were returned by 17 (54.8%) respondents, of whom, 14 (82.4%) were able to attend one of the two focus groups. Table 1 summarises the participant's descriptive characteristics. Across all participants, there was a relatively even gender split and a median (IQR) age of 62.0 (51.5, 75.5) years. The majority of participants were exempt from prescription charges and most were prescribed multiple medicines; the median (IQR) number was 3 (1.5, 6). Only three participants (21.4%) were students or employees of the university.

[Table 1 about here]

Participant discussions demonstrated an understanding of the TDF and agreement with the mapping process.. Participants discussed adherence barriers known through personal

232 experience as well as offering opinion on potential adherence barriers that others may 233 experience. 234 Focus group one 235 A summary of topics discussed is provided in supplementary file 5. Topics were discussed 236 across all six TDF domains presented in this focus group. Three adherence barriers, 237 undetected by the scoping review were discussed: 238 Not knowing about medicine delivery and repeat ordering systems – mapped to the 239 knowledge domain 240 Difficulties with identifying medicines, especially when the brands and packaging 241 regularly change – mapped to the skills domain 242 Hostility from GP receptionists which can prohibit medicine access – mapped to the social 243 influences domain 244 Focus group two 245 A summary of the topics of participant discussion is provided in supplementary file 6. Topics 246 were discussed across all five behavioural domains presented but the beliefs about 247 consequences domain was particularly stimulating of discussion. Adherence barriers 248 discussed by participants undetected by the scoping review were: 249 Negative emotions caused by feelings of getting a 'raw deal' with regards to medicines 250 supply, e.g. only getting one month's worth of medicines when others get three months' – 251 mapped to the emotions domain 252 Reduced motivation to adhere caused by questioning whether medicines represent 'good

value for money' – mapped to the motivation and goals domain

• 'Annoyance' about medicines taking when medicines have to be declared on insurance forms – mapped to the emotions domain.

The emotions domain was discussed in both focus groups, whilst there were similarities in the discussions on this topic between the two focus groups, differing personal experiences meant that in the second focus group, emotions related to 'annoyance' and 'getting a raw deal' were discussed which were not raised within the first focus group.

Phase 4 Refinement of adherence barriers mapped to the TDF

A summary of the re-mapping of adherence barriers from one TDF domain to another due to the additional perspectives identified from the focus groups is provided in supplementary file 7. Seventeen adherence barriers were re-mapped at this stage. Some barriers, for example knowing how to identify tablets or access them from packaging were moved from the knowledge domain to the skills domain. Additional understanding gained from the patients' perspective meant that these behaviours could be understood as an ability that can be acquired through practice (skill), rather than direct knowledge. Similarly, barriers such as feeling negative about medicines taking or burdened by this were originally conceived to relate to motivation and goals but understanding from the patient perspective enabled an appreciation of the genuine emotive aspects of these barriers.

Table 2 summarises the adherence barriers mapped to the domains of the TDF¹⁶ highlighting the wide range of adherence barriers captured.

[Table 2 near here]

Discussion

Use of the TDF¹⁶ to both organise literature-identified barriers to adherence and structure focus group discussions has facilitated their detailed analysis. It has identified ten active domains, each incorporating a range of determinants of medication adherence, such as those relating to emotions, which have previously received less attention in literature.²⁹ It is acknowledged that further relevant literature may have emerged since the conduct of the scoping review, however, its function was to act as a vehicle for prompting discussion in the focus groups. Given that the scoping review was designed to be supplemented by qualitative work and not intended to quantify the importance or prevalence of different barriers to adherence a full systematic review was inappropriate. The new adherence barriers and changes in mapping arising from the focus groups indicate that the methodological approach was appropriate for initiating and structuring the discussions. Recruitment through university advertisements for the focus groups may have introduced biases. However, participants represented a wide range of ages and medication regimen complexities. Furthermore, only three participants were university students or employees, of which only one was an academic. Whilst anecdotal evidence gathered from the focus group discussions means that we are confident that a wide range of educational and professional backgrounds were covered in our sample of focus group participants, characterisation of participants through formal data collection about educational level may have added further rigour. Additional information regarding whether adherence barriers suggested by focus group participants were based upon personal experience or supposition, may have been beneficial and provided readers with further contextual information. No relevant adherence barriers were identified for three of the TDF domains and a new domain termed 'goal conflicts' was added to capture adherence barriers that were not

276

277

278

279

280

281

282

283

284

285

286

287

288

289

290

291

292

293

294

295

296

297

299 reflected by the 2005 version of the TDF. The appropriateness of the adaptation is confirmed by the updated version of the TDF,¹⁷ which now incorporates goal conflicts. 300 301 Contrary to the present paper which mapped adherence barriers to all bar three of the TDF domains, Presseau et al.²² report that fewer TDF domains were relevant and did not map 302 303 adherence barriers to the skills, beliefs about capabilities, motivation and goals, 304 environmental context or emotions domains. Differing methodological approaches may 305 account for this as Presseau and colleagues sought to identify the most relevant domains 306 whereas the present article sought to explore the breadth of determinants. The latter approach 307 has allowed exploration of adherence barriers which are often overlooked. A further 308 difference is that Presseau and colleagues included the social/professional role and identity domain which was excluded from the present paper. Crayton et al.²³ also report redundancy 309 310 of this domain when exploring adherence determinants in stroke survivors, as do Voshaar et al. 24 with regards to adherence barriers and facilitators for disease-modifying anti rheumatic 311 312 drugs. In the present paper the social norms domain was used for barriers associated with not 313 identifying oneself as a medicines taker. These minor differences in mapping highlight that 314 despite robustly employed methods, there is still inherent subjectivity in TDF interpretation. The inherent subjectivity of the TDF mapping process means that a different theoretical map 315 316 could have been produced by other researchers, as highlighted by the work reported by 317 Presseau et al.²² The mapping decision being undertaken by a research team with expertise in 318 behavioural science and medication adherence plus refinement of this mapping based on 319 patient input provides some confidence in the final map. However, further validation of the 320 mapping decisions by an independent peer with expertise in these fields may have added 321 further rigour.

Crayton et al.²³ highlight that 'emotions', 'beliefs about consequences' and 'knowledge' appeared to be most influential TDF domains when mapping adherence determinants in stroke survivors. This finding is consistent with the qualitative explorations reported in this present paper. Voshaar et al.²⁴ also report mapping of adherence barriers across the range of TDF domains, with notable consistency in mapping compared to the work presented in the present paper. Both studies therefore support applicability of the work presented in the current article, beyond CVD prevention. The studies reporting mapping of adherence barriers to the TDF²²⁻²⁴ provide useful contextualisation of the present work and highlight the similarities of adherence barriers across a range of LTCs. However, the utility of each of these studies for adoption as routine practice is limited by their focus on specific diseases. The present paper presents the first TDF-based conceptual framework of medication adherence barriers across multiple LTCs, and is also the first paper to develop a framework based on both literature-identified and qualitatively explored adherence barriers. The focus groups in the present study, added richness to the data and, despite a large body of existing literature regarding adherence barriers, new barriers were identified spanning a range of TDF domains. An awareness of barriers such as a lack of knowledge about repeat prescription ordering services may be useful in supporting patients who wish to adhere but struggle with the management of their medicines. Likewise, the information yielded about the range of negative emotions associated with medicines taking, adds to our knowledge of the factors that may influence a patient's decisions to not adhere. Emotions, such as feelings of frustration and being 'short-changed', may represent modifiable determinants of adherence

322

323

324

325

326

327

328

329

330

331

332

333

334

335

336

337

338

339

340

341

342

343

344

worthy of further investigation as these are often overlooked^{29,37}. Practitioners seeking to

resolve non-adherence should be aware of the diverse plethora of factors that may influence adherence and mindful of the emotional components of medicines-taking behaviour.

The present work creates an evidence-based platform for developing novel, theory guided interventions to improve medication adherence. Whilst other theoretically informed adherence interventions have not always yielded improved outcomes, 15,37 this may be influenced by the lack of guidance regarding how these theories should be used for intervention design. The structured approach offered by the TDF and availability of work linking TDF domains to evidence based BCTs may address this difficulty. A programme of work to develop a novel adherence intervention, based on this conceptual framework will follow. Whilst theory guided litertaure 20 can be utilised to match BCTS to the domains of the TDF, much work is needed in understanding how these BCTs are appliacable to medicines-related consultations. Moreover, notable implementation work is necessary to explore how these BCTs are best delivered, from where and by whom.

Conclusion

This work provides the foundations for developing a patient questionnaire, grounded in the adherence barriers mapped to the TDF which will enable identification of an individual's barriers to adherence. As the focus groups were undertaken in the context of medicines prescribed for the prevention of CVD, it is intuitive to develop and trial a questionnaire in the same population. However, as the literature-identified barriers discussed in these focus groups were sourced from a variety of LTCs, it is likely that the adherence barriers will also be applicable to medication non-adherence in other LTCs. Further work is necessary to confirm this and to establish how adherence barriers vary for acute conditions.

References

- 369 1. World Health Organisation (WHO). Adherence to long term therapies: evidence for action. Geneva. 2003.
- Naderi SH et al. Adherence to drugs that prevent cardiovascular disease: Meta-analysis on
 376,162 patients. *The American Journal of Medicine*. 2012;125(9): 882-887
- Horne R et al. Concordance, adherence and compliance in medicine taking. Report for the
 National Co-ordinating Centre for NHS Service Delivery and Organisation R & D
 (NCCSDO). London. 2005.
- 376 4. Simpson SH et al. A meta-analysis of the association between adherence to drug therapy377 and mortality. *BMJ*. 2006; 333:15
- Nieuwlaat R et al. Interventions for enhancing medication adherence. *The Cochrane Library*. 2014.
- 380 6. Clifford S et al. Understanding different beliefs held by adherers, unintentional nonadherers, and intentional nonadherers: application of the necessity–concerns framework. *Journal of psychosomatic research*. 2008; 64(1), pp.41-46.
- 383 7. Bultman DC and Svarstad BL. Effects of physician communication style on client
 384 medication beliefs and adherence with antidepressant treatment. *Patient education and counseling*. 2000; 40(2), pp.173-185
- 8. Patton DE et al. Theory-based interventions to improve medication adherence in older adults prescribed polypharmacy: as sytematic review. *Drugs in Aging*. 2017. 34:97-113
- 9. Easthall C et al. A meta-analysis of cognitive-based behaviour change techniques as interventions to improve medication adherence. *BMJ open.* 2013. 1;3(8):e002749.
- 390 10. General Pharmaceutical Council (GPhC). Future Pharmacists: Standards for the initial
 391 education and training of pharmacists. 2011. Available from:
 392 https://www.pharmacyregulation.org/education/education-standards. Accessed 18th
 393 January 2018. Archived at https://www.webcitation.org/6wZ2ee7pd on January 18th 2018.
- Nursing and Midwifery Council (NMC). Standards for pre-registration nurising education.
 2010. Available from:
 https://www.nmc.org.uk/globalassets/sitedocuments/standards/nmc-standards-for-pre-registration-nursing-education.pdf. Accessed January 18th 2018. Archived at http://www.webcitation.org/6wZ31Gg3U on January 18th 2018
- 399 12. General Medical Council (GMC). Promoting excellence: standards for medical education and training. 2015. Available from: http://www.gmc-uk.org/Promoting_excellence_standards for medical_education_and training_0715.pdf
 402 61939165.pdf. Accessed January 18th 2018. Archived at http://www.webcitation.org/6wZ3G0t89 on January 18th 2018.
- 404 13. National Institute of Clinical Excellence (NICE). Behaviour change: individual approaches. London. 2014
- 406 14. National Institute of Clinical Excellence (NICE). Behaviour change at population,407 community and individual levels. London. 2007.
- 408 15. Craig P et al. Developing and evaluating complex interventions: the new Medical Research Council guidance. *BMJ*. 2008. 29;337:a1655
- 410 16. Michie S et al. Making psychological theory useful for implementing evidence based 411 practice: a consensus approach. *Quality and Safety in health care*. 2005;14(1):26-33.
- 412 17. Cane J et al. Validation of the theoretical domains framework for use in behaviour change and implementation research. *Implementation Science*. 2012; 7(37).
- 18. Francis JJ et al. Theories of behaviour change synthesised into a set of theoretical
 groupings: introducing a thematic series on the theoretical domains framework.
 Implementation Science. 2012; 7(1):35.
- 417 19. Abraham C, Michie S. A taxonomy of behavior change techniques used in interventions. *Health psychology*. 2008;27(3):379

- 419 20. Michie S et al. The Behavior Change Technique Taxonomy (v1) of 93 Hierarchically
 420 Clustered Techniques: Building an International Consensus for the Reporting of Behavior
 421 Change Interventions. *Annals of Behavioral Medicine*. 2013;46(1):81-95.
- 422 21. French SD et al. Developing theory-informed behaviour change interventions to
 423 implement evidence into practice: a systematic approach using the Theoretical Domains
 424 Framework. *Implementation Science*. 2012;7(1):38.
- 22. Presseau J et al. Identifying determinants of medication adherence following myocardial
 infarction using the Theoretical Domains Framework and the Health Action Process
 Approach. *Psychology & Health*. 2016;18:1-9.
- 23. Crayton E et al. Psychological determinants of medication adherence in stroke survivors:
 a systematic review of observational studies. *Annals of Behavioral Medicine*. 2016;31:1-3.
- 430 24. Voshaar M et al. Barriers and facilitators to disease-modifying antirheumatic drug use in
 431 patients with inflammatory rheumatic diseases: a qualitative theory-based study. *BMC* 432 *musculoskeletal disorders*. 2016;17(1):442.
- 433 25. McCullough AR et al. Defining the content and delivery of an intervention to Change AdhereNce to treatment in BonchiEctasis (CAN-BE): a qualitative approach incorporating the Theoretical Domains Framework, behavioural change techniques and stakeholder expert panels. *BMC health services research*. 2015;15(1):342.
 - 26. May C, Finch T. Implementing, embedding, and integrating practices: an outline of normalization process theory. *Sociology*. 2009;43: 535-554.
- 439 27. Durlak JA, DePre EP. Implementation matters: a review of research on the influence of
 440 implementation on program outcomes and the factors affecting implementation. *Am J Community Psychol.* 2008;4: 327-335.
- 28. The Identification of Medication Adherence Barriers Questionnaire (IMAB-Q).
 [homepage on the internet]. 2018. Available from:
 https://www.uea.ac.uk/pharmacy/research/imab-q/quest. Accessed 26th January 2018.
 Archived at https://www.webcitation.org/6wlIFZKVv on January 26th 2018.
- 29. Kardas P et al. Determinants of adherence: a review of systematic reviews. *Frontiers in pharmacology*. 2013;4(91).
- 30. van Dulmen S et al. Patient adherence to medical treatment: a review of reviews. *BMC Health Services Research*. 2007;7(1):1
- 31. Arksey H, O'Malley L. Scoping studies: towards a methodological framework.
 451 *International journal of social research methodology*. 2005;8(1): 19-32
- 452 32. Patton MQ. Qualitative research. John Wiley & Sons, Ltd. 2005

437

438

465

- 33. Taylor N et al. Development and initial validation of the Determinants of Physical
 Activity Questionnaire. *International Journal for Behavioral Nutrition and Physical* Activity. 2013;10(74).
- 456 34. Armstrong D et al. The Place of Inter-Rater Reliability in Qualitative Research: An Empirical Study. *Sociology*. 1997;31(3):597-606
- 458 35. Ritchie J, Spencer L. Qualitative data analysis for applied policy research. *The qualitative researcher's companion*. 2002;305-29.
- 36. Taylor N et al. Development and initial validation of the Influences on Patient Safety
 Behaviours Questionnaire. *Implementation Science*. 2013;8(81).
- 37. Munro S et al . A review of health behaviour theories: how useful are these for developing interventions to promote long-term medication adherence for TB and HIV/AIDS? *BMC* 464 *Public Health*. 2007;7:10

 Table 1:
 Summary of participant characteristics for consultation exercises

Participant characteristics	Measure	Consultation exercise one (n = 5)	Consultation exercise two (n=9)
Male gender	Number (%)	3 (60%)	5 (55.5%)
Age (years)	Median (IQR)	70.0 (45.5, 76.5)	62.0 (54.0, 75.5)
Exempt from prescription charges	Number (%)	3 (60%)	6 (66.7%)
Number of regularly prescribed medicines	Median, (IQR)	3 (1, 5)	2 (2, 6)
Employed by the university	Number (%)	2 (40%)	1 (11.1%)

Table 2: Summary of adherence barriers mapped to each domain of the original TDF

TDF Domain	Adherence barriers mapped to this domain
Knowledge	 Not knowing how to order prescriptions or about services that facilitate this process
	 Not knowing how to collect prescriptions or about services that facilitate this process
	 Having insufficient information about medicines e.g. how they work, why they were prescribed, side effects and benefits
	Not knowing how (and when) to take medicines as prescribed
Skills	 Physical inability to take medicines as prescribed e.g. swallowing difficulties and problems accessing medicines from packaging
	 Cognitive inability to take medicines as prescribed e.g. inability to read and/or understand instructions
	Inability to identify and differentiate between different medicines
	 Lack of organisational and forward planning skills (not having a system in place to help manage medicines)
Beliefs about	 Lack of confidence in ability to adhere and manage medicines e.g. feeling regimen is too complex
capabilities	 Lack of confidence to overcome difficulties with medicines taking e.g. experience of side effects
	Perceived inability to cope with medicines related changes
Beliefs about	Fear that medicines will be (are) harmful
consequences	Belief that medicines cannot be trusted

	 Doubting the efficacy of medicines 	
	 Not believing that there is a need for treatment 	
	Denial of illness or non-acceptance of diagnosis	
	 Decision making process justified belief about consequences (or lack of consideration of consequences) e.g. preference for alternative remedies 	
Motivation and	Not perceiving medicines taking as a priority	
Goals	• Lack of intention to adhere	
	Lack of motivation to adhere	
Goal Conflicts*	Cost of medicines (having to choose between paying for a prescription and something else)	
	• Having a busy lifestyle (e.g. work and travel) and other priorities (e.g. family commitments or meal times) which impede medicines taking at specific times	
	Being too busy to order and collect prescriptions/having other priorities which impede ordering and collecting medicines	
Memory, attention &	Forgetting to take medicines	
decision processes	Forgetting to order/collect medicines from pharmacy	
	• Lack of attention in medicines taking e.g. making errors or forgetting due to distractions	
Environmental	Problems with pharmacy/GP surgery e.g. not stocking medicines, lost prescriptions, failed orders etc.	
context and	Difficulties getting to pharmacy/GP surgery to collect prescriptions	
resources	Changes to environment or daily routine which impede medicines taking	
Social influences	Fear of judgement, discrimination or social stigma	
	Cultural and religious norms and expectations	
	• Lack of trust in prescriber	
	• Lack of social support	
Emotion	Experience of negative emotions associated with medicines taking e.g. frustration or embarrassment	
	Perceiving medicines taking as a negative reminder of illness/condition	
	Perceiving medicines taking as a burden	
Social/professional role & identity	No adherence barriers mapped to this domain	
Behavioural regulation	No adherence barriers mapped to this domain	
Nature of the behaviour	No adherence barriers mapped to this domain	

^{*} A newly created domain to reflect adherence barriers that did otherwise not fit