Qualitative study of barriers to clinical trial retention in adults with recently diagnosed type 1 diabetes

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ABSTRACT

Objectives Regular physical exercise may preserve β cell function in newly diagnosed adults with type 1 diabetes (T1D). However, clinical trials to test this theory require the recruitment and retention of adults with new-onset T1D, which can be challenging. We sought to determine the overall experiences of newly diagnosed adults with T1D in an exercise study, to understand issues that influence the retention of trial participants in such studies.

Design Qualitative methodology using individual face-to-face (n=6) and telephone interviews (n=14). Interview transcripts were thematically analysed using the framework method.

Setting The study took place at five participating UK hospitals.

Participants Twenty participants, aged 19–55 years, in the Exercise for Type 1 Diabetes study were interviewed to explore their study experiences and identify motivators and deterrents towards the study. Participants in control and intervention arms were interviewed, as were people with T1D who had completed (n=16) and withdrawn (n=4).

Results Participants revealed barriers and facilitators to retention; the majority were generalisable to clinical trials of people with newly diagnosed T1D. Coming to terms with a diagnosis of T1D, lack of time, work pressures, level of health professional support, volume, clarity and consistency of information and feedback and a desire for knowledge about their condition were all cited as influencing factors to trial retention.

Conclusions To our knowledge, this is the first qualitative study to examine the experience of being involved in an exercise trial by people with T1D. Findings suggest appointments could be shorter, available outside of working hours and planned longer in advance; study information should be clear, consistent and in electronic and paper formats; questionnaires need minimising; healthcare support and feedback needs providing regularly; thought is required around how to support non-exercising arm participants. These considerations may improve participant retention rates in new-onset T1D studies.

INTRODUCTION

Type 1 diabetes (T1D) is a chronic autoimmune condition characterised by immune-mediated destruction of insulin-producing pancreatic β cells.1 Significant numbers of β cells are present at the time of diagnosis,2 and preservation of β cell function is associated with significant benefits for people with T1D. Interventions that can preserve residual β cell function in new-onset T1D are needed.

In animal models of T1D and type 2 diabetes (T2D), healthy humans and in people with impaired glucose tolerance with T2D, regular exercise has been shown to preserve β cell function.3 These findings have not been tested in people with T1D, and there is thus a need for a prospective clinical trial to test the hypothesis that exercise preserves β cell function in people newly diagnosed with T1D.

The Exercise for Type 1 Diabetes (EXTOD) study was a pilot study undertaken to explore whether exercise can preserve β cell function in adults newly diagnosed with T1D. In designing this study, we had to bear in mind that the incidence rate of T1D is low and recruitment of people with T1D to clinical trials is challenging.1-6 Other studies have shown recruitment rates as low as 17%.4 This means that retaining people with newly diagnosed T1D who are recruited to studies is even more important. In studies of immunotherapeutic agents for T1D, dropout rates are 12%–14%7 8; however, higher rates have
been reported for exercise studies. In a meta-analysis of unsupervised exercise programmes for people with T2D, 20% of studies had a dropout of >20%, 32% a dropout of 10%–20% and 48% a dropout of <10%.9

Barriers to participation in clinical trials are well documented,10 although no studies have looked at barriers to recruitment in people with T1D. There are very few studies that have looked at how to improve retention rates of people with T1D in a clinical randomised controlled trial (RCT). Those studies that have been done have concentrated on strategies to improve retention rather than barriers to retention.11 No studies have looked at barriers to retention in people newly diagnosed with T1D. We wished to address this important deficit by qualitatively exploring the experiences of people with newly diagnosed T1D who participated in a recently completed exercise and T1D study.12 Here we report our findings from a qualitative study of barriers to clinical trial retention in adults with recently diagnosed T1D. We have not reported other findings from the EXTOD study, which are reported separately.13 14

METHODS
Setting, access and recruitment
Study participants were from the EXTOD study, whose protocol had been described previously.12 In brief, all people aged between 16 and 60 years, diagnosed with T1D in the previous three months, from 19 UK hospital sites were invited to participate. The EXTOD study explored the barriers and benefits of exercise in adults with newly diagnosed T1D. The hypothesis being tested by the EXTOD study was that exercise preserved β cell function in adults recently diagnosed with T1D. EXTOD had two phases. Phase I consisted of a qualitative study to determine attitudes and barriers to exercise in people with newly diagnosed T1D who participated in a recently completed exercise and T1D study.13 Phase II was a pilot RCT to assess uptake, intervention adherence, dropout rates and rate of uptake in the usual care group during a 12-month exercise intervention (where the participants came from for this study).

At the time of recruitment to the pilot RCT (phase II) study, all people with T1D received participant information leaflets and provided informed consent to potentially take part in an interview to explore their experience of being involved in the pilot RCT. Of the 60 participants who took part in the pilot RCT, 20 participants from five participating sites (Birmingham, Leeds, Bristol, Gloucester, Taunton) were later selected using purposive sampling to ensure variety and diversity in terms of their key study characteristics.15 Participants were sampled in relation to their age, gender, study arm (intervention/control) and study status (completed/withdrawn) to ensure that an even spread of participants across the key characteristics were sampled. Similarly the sites were selected to allow a purposeful sampling of geographical areas participating in the EXTOD study (teaching hospitals vs district general hospitals). Selected participants were sent a letter at the end of the pilot RCT, informing them that they would be contacted by the EXTOD team. This was followed, a week later, by a phone call from a member of the EXTOD team (nurse or doctor) to check they were happy to be interviewed. If willing, they were then telephoned by the researcher (CH), who has a nursing background and is an experienced qualitative researcher (though she had not previously undertaken any research in this area), to arrange a suitable time and date for her to undertake the interview. CH had had no contact with participants prior to this and participants were unaware of the researcher’s background. All participants agreed to be interviewed.

Patient involvement
The research question was derived from people with T1D attending clinic and asking clinicians about any benefits and barriers to exercise as they were aware that much work had been undertaken on this topic in people with T2D, but not in people with T1D. This led to the formulation of the research question and an application for funding to undertake this research. People with T1D were involved in the study design from the outset as the researchers presented the study proposal to them and asked for any comments relating to it (approximately seven people with T1D were involved). Issues relating to the conduct of the study, such as the potential burden of the exercise intervention to participants, were also discussed and any feedback was incorporated into the study design. People with T1D also contributed to the study conduct by sitting on the study management committee and helping with study oversight, including helping to develop approaches to improve study recruitment (three people with T1D were involved). Study findings for phase II of the study were fed back to participants through an informal feedback evening where the findings were presented. This was well received by the participants. In addition, a summary of findings was posted to participants.

Data collection
Participants at the hospital where the researcher was based (Birmingham) were given the option of being interviewed face-to-face at the hospital or by telephone. The remaining participants were all interviewed by telephone due to financial, time and travel constraints. All interviews were carried out with only the participant and CH present.

Interviews were carried out by CH using a structured topic guide (table 1) that was developed in consultation with the EXTOD researchers. The interviews lasted between 20 and 50 min. No repeat interviews were conducted. Areas for discussion included levels of health professional support, information provided about diet and exercise and issues relating to recruitment and follow-up. All interviews were digitally recorded and transcribed verbatim by a local transcription company.
Data analysis

Data analysis was carried out during the interview period to enable the interview process to be shaped by the emerging data themes and to identify when saturation had been reached. For example, some participants in the early interviews spoke of how they had struggled to
come to terms with their new T1D diagnosis. As a result, subsequent interviews explored this area in more depth, focusing on the impact this had on their experience of participating in the EXTOD study. An inductive approach to data analysis was taken and themes were derived from the interview dataset. The predetermined topic areas set out in the topic guide (table 1) such as ‘health professional support’ and ‘recruitment and follow-up’ guided the analysis process as much of the data collected focused on these topic areas, enabling the researchers to elicit information that was relevant to the study question. Data were coded by CH and were inserted into a framework analysis matrix Excel spreadsheet, to enable data ordering and synthesis, while retaining the meaning and feeling of the interviewees’ words. Themes were developed by CH by reading and rereading the transcripts and these themes were then agreed during discussions with research team members (CH, PN and SG). The interviews were then analysed using thematic analysis. Transcripts were not returned to participants to comment on during this process.

**RESULTS**

**Participants**

In total, 20 participants were interviewed; 6 face-to-face and 14 by telephone (table 2). Participants were recruited from all five participating sites; 11 were men, ages ranged from 19 to 55 years and all were of white British ethnic origin. Thirteen had been allocated to the intervention group and seven to the usual care comparator. Sixteen participants had completed EXTOD, while four had withdrawn for reasons including work commitments, a cancer diagnosis and difficulties accepting their diabetes diagnosis.

**Themes**

The interviews yielded data on five main themes, and these help formulate the barriers and facilitators to trial participation (Box 1). These themes were study paperwork, feedback, barriers to continued participation, coming to terms with diagnosis of T1D and effect of allocated arm.

**Study paperwork**

Two main subthemes emerged within the ‘study paperwork’ theme; these were study information and study questionnaires.

**Study information**

The volume, clarity and consistency of information provided were important in determining how well informed participants felt about their diabetes and its management. Though a few participants felt they were given too much information, most felt that although there was a lot of information, it was useful, relevant and necessary so soon after diagnosis. Generally the information was cited as interesting, manageable and straightforward.
Box 1 Facilitators and barriers to continued clinical trial participation in people with type 1 diabetes mellitus

<table>
<thead>
<tr>
<th>Facilitators</th>
<th>Barriers</th>
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<tbody>
<tr>
<td>► Consistency and continuity of health professional support</td>
<td>► Time</td>
</tr>
<tr>
<td>► Clear, detailed and relevant information about diabetes and its management</td>
<td>► Work pressure</td>
</tr>
<tr>
<td>► Availability of both paper and electronic information and documentation</td>
<td>► Travelling to appointment</td>
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<tr>
<td>► Reduction in volume of study documentation</td>
<td>► Length of visits</td>
</tr>
<tr>
<td>► Flexible access to trial facilities outside of normal working hours</td>
<td>► Length of study</td>
</tr>
<tr>
<td>► Appointments scheduled in advance to allow for planning around work and social lives</td>
<td>► Coming to terms with diagnosis</td>
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<tr>
<td>► Early feedback of trial findings</td>
<td>► Being able to maintaining exercise levels</td>
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<tr>
<td>► Being allocated to the control (non exercising arm)</td>
<td>► Being allocated to the control (non exercising arm)</td>
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Others spoke of how they valued the repetition of some information as it meant they could fully absorb it.

To be honest I thought it was perfect … I was sort of drip-fed information throughout the year really, and it was very good … After … The shock of being told you’re Type 1 diabetic … I found it very good and the team was extremely helpful. EXTOD 12 (male, Taunton)

Most participants were happy receiving paper information, finding it easy to read and acknowledging that not everyone had internet access. Others, especially younger participants, stated they would have preferred digital information, such as apps, to access information faster. One participant suggested having a centralised webpage specific to the hospital’s diabetes unit, with links to different diabetes resources on it. Participants supplemented the trial information with other resources, such as the ‘Carbs and Cal’ book. Others used digital resources, such as the ‘Carbs and Cal’ or ‘Fitness Pal’ apps and online information.

Being a bit of a technology geek, I went with the app which is the Carbs and Cals app … It will give you a particular breakfast, like Alpen … and then it will show you … the size of portion that you’re supposed to have, and how much insulin you’re supposed to take. EXTOD 2 (male, Birmingham)

Study questionnaires

Most participants felt the study paperwork was too long and time-consuming, ‘switching off’ from it as a result, as it got in the way of work and personal lives. Much paperwork was questionnaires relating to diet, exercise and quality of life that were completed at the study visit. Participants felt that although the paperwork was generally easy to complete and understand, it could also be repetitive, confusing and contradictory, containing irrelevant, non-specific questions, which were hard to relate and respond to.

I remember always getting given the questionnaire sheet when I was having my bloods done. And I could just never fill it in. It used to be like – I used to say to my dad can you ask my questions and then I could do it, but it did go on for quite a while … And it was the same the second time I did it as well, and I was like I don’t know what to put for these questions again. It seemed like I was doing the same thing. EXTOD 19 (female, Birmingham)

Feedback

Some participants voiced disappointment at receiving little trial feedback, stating that the opportunity for feedback had incentivised them to join the trial, as a way of finding out about their diabetes and also that they wanted to learn more about the long-term trial outcomes.

It would be good to have some follow-up information … I wasn’t quite sure how my fitness level was affected … And I never got to find out if it actually improved. EXTOD 1 (female, Birmingham)

Barriers to continued participation

In total, 17 of the 58 participants who were randomised into the original EXTOD study withdrew before the end of the study. We thus sought to understand why this might have happened.

Participants described practical barriers as the most likely reasons for dropping out. These included time and work pressures, dislike of blood tests, travelling to appointments, the long study duration, volume of visits and moving away.

I’m not bothered for taking blood or doing injections or anything like that … But I can remember feeling bothered by that at the time. I think because I was very ill, really, still, and still quite fragile … I can remember that upsetting me … Because it was a study and because I didn’t have to do it … If you volunteered to have a blood test and it takes a few times for them to take the blood, it’s kind of like putting yourself through something difficult that you didn’t really need to do. EXTOD 6 (female, Leeds)

The long study duration also deterred many participants, who felt it was too long to fully commit to, meaning they became less vigilant at attending appointments or completing study documentation due to the repetition involved.

It’s quite long. I think that was quite daunting. It turned out to not really be an issue, but because it’s something that I could kind of blend into my lifestyle...
quite a lot. Because I suppose it didn’t really require any prolonged visits to the clinic or anything because it was mostly just things that I could do myself. But initially that was a bit daunting because it’s a year. EXTOD 16 (male, Bristol)

For most, committing to the EXTOD study had proved difficult, due to the time it required off work. This let to difficulties for participants in ensuring they could always attend appointments and sometimes proved costly due to having to take unpaid leave for study visits.

The time that you’re having to have off work … Often it’s unpaid leave so obviously that can be quite difficult. EXTOD 18 (female, Leeds)

These time pressures had led to two of the interviewed participants withdrawing from EXTOD prior to completion. Solutions to these problems suggested by participants—who had both withdrawn and remained in the study—included offering more flexible appointment times and planning clinic visits further in advance so there was more time to plan around them.

I left … Because I didn’t have enough time … The study is of such a long duration and I just found that too challenging with work … If you had appointments at eight o’clock in the evening every time you needed me then so be it … I have to do a hell of a lot of juggling in order to fit that sort of stuff in. EXTOD 17 (male, Bristol)

Coming to terms with diagnosis of T1D

Some participants spoke of how they had struggled to come to terms with their T1D diagnosis, with one participant citing this as their reason for withdrawing from EXTOD. For this participant, her difficulties coping emotionally with her diagnosis had prompted her doctors to advise her to withdraw.

Don’t think it kind of really sank in as to what I’d been diagnosed with … It had kind of hit me and I wasn’t really dealing with having it … I wasn’t taking my insulin and checking my levels as much … The doctors … Felt that it was best that I was taken off it. EXTOD 4 (female, Birmingham)

Effect of allocated arm

Some participants, who had been allocated the exercise intervention, spoke of difficulties maintaining the level of exercise expected of them due to lack of motivation or the extra time it took.

I think a year in it seemed to get a little bit ‘Oh God I’ve been doing this for a year now’ … A year’s a long time … Because towards the end as well, it was like you had to come in and then I had to do the gym thing, and it was kind of like … ‘oh this is getting really laborious’. EXTOD 2 (male, Birmingham)

Others, in the usual care comparator group, spoke of how this disincentivised them from remaining in EXTOD as they were not receiving the exercise benefits they had hoped for.

The sort of people that … Take part … Are often similar to me in kind of quite wanting to do the exercise, and presumably about half of the time you get randomised into not doing the exercise. And I don’t know whether that’s off-putting as well. EXTOD 16 (male, Bristol)

DISCUSSION

This is the first study to examine the experiences of people with T1D taking part in an exercise intervention trial. The findings have highlighted issues that should be considered when designing clinical trials involving this population group. Clinical trials which reflect the needs, wants and preferences of participants can lead to improvements in retention rates, statistical power and, in the longer term, to studies with more credible findings. The findings suggest that timing is an important consideration for many participants. Many may be offered trial entry shortly after their condition’s onset and may be experiencing multiple health and lifestyle changes and struggling to come to terms with their diagnosis of T1D.

This highlights the need for sensitive communication of information from health professionals when introducing clinical trials to adults with T1D as they may be experiencing tensions between not wanting to ‘become’ their new illness or allow it to govern them, while reconciling that they need to be adaptable to their illness if they are going to feel well again.19 20 The disappointment of some participants at being allocated the usual care group also highlights the need for a clear explanation of equipoise and other clinical trials terminology, so participants can make informed treatment decisions, minimising the potential for withdrawal rates after randomisation.21 Additional measures that could help with this are offering the intervention at the end of the study or randomising more to the exercise arm than the control arm, to enable more participants access to the intervention.

The study has illustrated the need for the transmission of clear, relevant and useful information to clinical trial participants. Participants showed preferences for a wide range of delivery modes, including paper formats, websites, apps and internet forums. The use of appropriate, accessible media to convey information effectively and engage with participants is dependent on individual preferences and may be influenced by factors such as age, gender and income.22 The popularity of apps and multimedia technologies among participants indicates that these modes of information delivery should be incorporated specifically into trials with T1D participants, who are likely to be a younger, more technologically minded population group than trials with T2D participants. Future trial designs could offer paper and electronic
information so participants can use their preferred resource. Similarly, when collecting data, a range of methods could be employed, including paper documentation, spreadsheets, phone apps, emails and websites. Demonstrating versatility and flexibility in data collection techniques may help motivate participants to complete study documentation that they might otherwise omit. The importance of providing regular study feedback to participants should also be considered, through letters, emails and presentations, so that their contribution is acknowledged and valued.23

Finally, the interview findings highlighted that practicalities such as work pressures, time commitments and geographical location influenced participants’ ability to commit to the trial. T1D onset usually occurs at a young age, with most people being diagnosed before 35, 24 As a result, people with T1D are likely to have busy lives incorporating work, family and social engagements, making committing to a clinical trial difficult. This is verified by the most common reason cited for the high dropout rate (29%) from EXTOD being time and work commitments. To rectify this, more flexible study visit times could be offered, with the option of attending outside normal working hours, such as evenings and weekends. Additionally, the provision of a timetable of scheduled appointments at the study outset would allow participants to plan for them. Improved hospital transport links could also facilitate ease of study attendance.25 By incorporating a flexible approach to these practical barriers, participants will experience less difficulty in complying with study visits, improving retention rates. Consideration should also ensure that the exercise intervention can be integrated into people’s lifestyles, without adding to their pressurised schedules. This could be done by providing a range of exercise options, in a range of locations and within a realistic timeframe, to minimise the likelihood of participants becoming overburdened by the extra commitment they have taken on.

Many of the challenges to clinical trial retention in people with T1D are similar to the challenges facing the general population who are recruited to clinical trials.26-30 The issues and challenges participants faced, such as the long study duration, difficulties completing documentation and time pressures, are also likely to be challenges for the general clinical trials population, including T2D. However, this study has identified that although many of these trial-related issues are not specific to T1D populations, the reasons for these issues are likely to be different. For example, people with T1D or T2D may report that time pressures prevent them from committing to a clinical trial. However, while for the T1D participant time pressures may stem from work priorities, the T2D participant time pressures may stem from regularly caring for grandchildren. By understanding more about the reasons behind these barriers to trial recruitment and retention, trials can be designed to accommodate and facilitate the wants and needs of these different groups. The average age of participants entering clinical trials is >50 years of age.26-31

While there is growing evidence that T1D is diagnosed throughout adult life,32,33 the mean age of the randomised EXTOD participants was 32 years, suggesting that trial retention considerations should focus on needs of younger adult populations. This younger age group is likely to face more difficulties in adhering to rigid time schedules and appointments due to added work, family and social pressures. This is important for clinicians and researchers to acknowledge, allowing them to design T1D studies that will facilitate trial recruitment and retention, by offering more flexible appointment times, out-of-hours services and realistic, manageable exercise schedules. These practical solutions are important considerations for retaining participants in clinical trials and must be valued if rich and full trial datasets are to be obtained. These considerations may also have some relevance for T2D populations, with the average age of onset for this disease getting younger.33,34

When considering study design, thought must be given to how to increase recruitment, as well as how to make the study experience appealing to participants once they have consented. By paying attention to the participant’s perspective, the chances of retaining participants for the study duration will increase, improving trial outcomes and participant satisfaction with trial entry.

**Strengths and limitations**

The interview participants were selected from five UK hospitals, increasing the transferability of the findings to a range of settings. Participants were purposively sampled, stratifying them according to age, gender, randomisation arm and whether they had completed or withdrawn from the study. This enabled a diverse and comprehensive collection of narratives to be gathered and analysed, enhancing the trustworthiness of the findings. However, although 12% of EXTOD participants recruited were non-white, none of the participants recruited to the qualitative study were non-white. This lack of representation is due to our exclusion of ethnicity as a criterion in our purposive sampling. Although we did not purposefully exclude non-white participants from the sample, the likely reason for their lack of inclusion is because T1D is far more common in white populations than other ethnic groups.35 Additionally, many of the geographical locations we recruited from had a white population of >84%.36-38 Despite this, including ethnicity in our purposeful sample could have ensured that non-white populations were represented. In addition, due to the interview participants being situated around the UK, most interviews were conducted by telephone. However, where possible, participants were given the choice of being interviewed face-to-face or by telephone, allowing them to choose the setting they found most relaxing, thus increasing opportunity for open dialogue between interviewees and researchers. The fairly even split between participants choosing face-to-face and telephone interviews suggests that providing a choice of setting may increase recruitment and retention to trials, giving participants the opportunity to select their most comfortable environment.
CONCLUSION
To our knowledge, this is the first qualitative study to examine T1D participants’ experiences of being involved in a clinical trial. Although people may be initially motivated to enter clinical trials for reasons such as altruism and a desire for information, practical factors such as work and time constraints, study duration and financial difficulties often act as deterrents for remaining on trials. Though these issues in themselves are not unique to the T1D population, the reasons for these issues are likely to be different. These reasons need considering when designing T1D clinical trials, to ensure that appropriate modifications are built into the trial design to enable people with T1D to participate with minimal disruption to their lives.

The study findings have highlighted that differences do exist between T1D participants and the general clinical trials population. First, despite T1D increasingly being diagnosed in adult life, the younger adult age of people with T1D at recruitment may make it harder for them to commit to clinical trials due to increased work, family and social pressures; this was verified by study participants. Second, the study has indicated that using multimedia technology might benefit T1D participants, who are used to handling information electronically. It has highlighted that to increase retention to T1D trials improvements to trial design are required. This can be done through providing flexible access to services, clear and relevant study information, documentation and feedback, as well as consistent healthcare support.

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Author note The COREQ reporting guidelines were followed.

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REFERENCES


