BMJ Open Virtual reality intervention to improve apathy in residential aged care: protocol for a multisite non-randomised controlled trial

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ABSTRACT

Introduction Apathy is a prevalent neuropsychiatric symptom for older adults residing in aged care. Left untreated, apathy has been associated with accelerated cognitive decline and increased risk of mortality. Reminiscence therapy is commonly used in aged care and has demonstrated to reduce apathy. Traditional methods of reminiscence use physical objects and more recently technology including tablets and laptop computers have demonstrated potential. Virtual reality (VR) has successfully been used to treat psychological disorders; however, there is little evidence on using VR for behavioural symptoms such as apathy in older adults. Using VR to deliver reminiscence therapy provides an immersive experience, and readily available applications provide access to a large range of content allowing easier delivery of therapy over traditional forms of therapy. This study aims to identify changes in apathy after a reminiscence therapy intervention using head-mounted displays (HMDs).

Methods and analysis Participants will be allocated to one of three groups; reminiscence therapy using VR; an active control using a laptop computer or physical items and a passive control. A total of 45 participants will be recruited from residential aged care (15 in each group). The three groups will be compared at baseline and follow-up. The primary outcome is apathy, and secondary outcomes include cognition and depression. Side effects from using HMDs will also be examined in the VR group. Primary and secondary outcomes at baseline and follow-up will be analysed using linear mixed modelling.

Ethics and dissemination Ethics approval was obtained from the University of South Australia Human Research Ethics Committee. The results from this study will be disseminated through manuscript publications and national/international conferences.

Trial registration number ACTRN12619001510134.

INTRODUCTION

The presence of apathy can be disabling for residents in long-term aged care facilities.1 Having apathy results in diminished goal-directed behaviour.2 Therefore, individuals with apathy are more likely to withdraw from care,3 therapeutic activities4 and have reduced self-care behaviours.5 In aged care facilities, neuropsychiatric symptoms occur in approximately 90% of people,6 and the prevalence of apathy has been reported to vary between 44% and 84%.7–10 It is common for apathy to go undiagnosed or less likely to be treated.11,12 However, patients with apathy can have almost a twofold increased risk of dementia.3

Despite the prevalence and consequences of having apathy, pharmacological treatment is limited and can be difficult due to differing factors including apathy subtypes, stage of neurodegeneration and age that can influence efficacy.13,14 Non-pharmacological treatments are a safe alternative, and there is a broad consensus that non-pharmacological approaches for treating apathy, in particular, personalised therapy using information and communication technologies should be used.15

Reminiscence therapy is an approach used in aged care that can be individualised and has demonstrated potential.16 One of the main strengths of reminiscence therapy is that it is a person-centred approach that emphasises the importance of personal identity.17 Loss of independence caused by moving into an aged care facility can cause a person’s identity to deteriorate and may increase apathy.18 A meta-analysis examining the use of reminiscence for improving psychological...
well-being in older adults found an overall medium effect size of 0.54. A more recent systematic review and meta-analysis examining reminiscence therapy for people with dementia included 22 studies of which two studies examined apathy. One study was included in the meta-analysis and found no difference at follow-up in apathy using the Apathy Inventory Scores after 3-month follow-up, mean difference 1.40 95% CI (−1.30 to 4.10); participants were classified as having Alzheimer disease and were non-institutionalised. The second study included in the review but not the meta-analysis found improvement after 3-month follow-up using the Apathy Evaluation Scale (AES) ($Z=−3.10$, $p=0.002$), in this study participants were institutionalised with mild-to-moderate dementia. Other benefits from reminiscence therapy found in this review were improvements in mood, cognition and communication, although results were inconsistent. This demonstrates that reminiscence therapy can work but calls for consistency in methods and alternative forms of delivery to assist with improving outcomes. Technology is now commonly used with reminiscence therapy in the form of touchscreen tablets or laptop computers, providing access to a multitude of content. The use of technology allows for personalisation and provides the ability to reduce the amount of time required to prepare for sessions. Videos can provide sound and audio, increasing the realism of the reminiscence experience. Using immersive technologies can be a way of providing personalised therapy to improve outcomes.

Virtual reality (VR) using head-mounted displays (HMDs) is a fully immersive technology that has been successfully used to treat conditions including post-traumatic stress disorder, anxiety and pain during medical procedures. Research using VR to treat apathy in older adults is limited. A single-session study measuring apathy using HMDs found improvements in apathy ($Z=−2.818$, $p=0.005$), this study included both group and individual sessions, used a generic library of videos and participants were limited to those with no or minimal cognitive impairment. Other studies using VR and looking at apathy have also been single-session studies, therefore, not examining changes in apathy. A recent feasibility study found that participants with varying levels of cognitive decline ranging from minimal-to-moderate impairment could tolerate the use of VR with HMDs in agreement with previous research. The VR reminiscence experience in this study was specifically tailored for each participant. Improvements in verbal fluency were found after the intervention in participants with higher levels of apathy at baseline ($r=0.719$, 95% CI 0.327 to 0.900, $p=0.003$), and 35% of participants in the study did experience temporary and minor side effects from using VR.

There is increasing evidence of acceptability of using HMDs in older adults. However, users of VR can experience symptoms of motion sickness. A recent systematic review and meta-analysis has highlighted the lack of research in examining side effects in older adults from using VR, and although older adults reported lower symptomatology than younger samples, these findings were based on a small number of studies and excluded clinical samples. Additionally, research with older adults does not always assess side effects from using VR. There is a need to increase our understanding of side effects from using VR in older adults, particularly with the increased use of HMDs for therapy in older clinical populations.

This study will deliver reminiscence therapy to older adults aged ≥65 years living in residential aged care to reduce apathy. Reducing apathy may assist with delaying the rate of cognitive decline and attenuate the expected functional decline that occurs in long-term care in maintaining a person’s overall health and well-being. The benefits of therapy are not limited to individuals as it may also provide more engaged residents for residential aged care facilities. Three groups will be compared: reminiscence therapy using VR (intervention group); reminiscence using a laptop computer or physical items (active control group) and usual care (passive control group). The primary hypothesis is that the VR group will have lower apathy scores than the traditional reminiscence group after the intervention. The intervention and active control groups will also be compared against the passive control group to establish the effectiveness of reminiscence therapy overall. The secondary outcomes include cognition and depression. For the VR group only, side effects from using HMDs will also be examined. During therapy for both the VR intervention and active control groups, heart rate variability, galvanic skin response and speech will be examined as exploratory outcomes, additional exploratory outcomes for all three groups include loneliness, quality of life and physical activity.

**METHODS AND ANALYSIS**

*Design*

This is a multisite non-randomised controlled trial. The Standard Protocol Items: Recommendations for Interventional Trials guidelines were used to set out the structure of this protocol. The study will include 45 adults aged ≥65 years, living in residential aged care. Assessments will be performed at baseline and follow-up approximately 2 weeks apart for all three groups. Activity monitors are fitted prior to baseline and on completion of follow-up. Therefore, all participants are in the trial for approximately 3 weeks. The VR and active control groups will receive a reminiscence intervention, and the passive control will receive usual care during the trial period.

*Patient and public involvement*

There was no patient and public involvement in the study design.

*Setting*

Data collection will be conducted across three residential aged care facilities operated by an aged care provider in South Australia. Selected sites are in the northeast or...
inner northern suburbs within the metropolitan area of Adelaide, South Australia. Site allocation was not randomised and determined by the aged care provider. All three facilities provide personal and healthcare services including dementia support, respite and palliative care.

**Eligibility criteria**

For all three groups, participants are eligible if they are aged 65 years or older, men or women and proficient English speakers. Participants also need to be willing to undertake follow-up assessments. For the VR group, participants need to be able to tolerate wearing the HMD and have vision that can be corrected with the use of their current glasses, their glass frame also need to be able to fit into the HMD.

For all three groups, participants are excluded if: they have known learning disabilities, their score on the Psychogeriatric Assessment Scale as assessed by the aged care facility is 16 or higher indicating severe cognitive impairment, have significant neurological disorders and other conditions including agitation and aggression at a level that would make assessment difficult. Finally, those who have issues with confusion/disorientation or who may become distressed due to confusion/re time and place.

**Intervention groups**

The study consists of three groups, see figure 1. The VR group undertakes reminiscence therapy using VR. The active control undertakes reminiscence therapy using a laptop computer and/or physical items including books, magazine or photos, and the passive control receives their usual daily care.

As participants are blinded to the other groups, each group is allocated to a separate residential aged care site. Many participants socialise at mealtimes or with group activities and this will minimise awareness of other groups. With the VR group it is first established if participants can see to a satisfactory level in the HMD prior to the consent process. This is done by displaying an image in the HMD that the participant is familiar with and asking the participant to assess the clarity of the image. If the quality is acceptable and it is established that the participant can tolerate the use of the HMD, then the consent process is undertaken.

**Study stages**

The study is divided into three stages, baseline, intervention and follow-up.

**Baseline**

Baseline measures take approximately 1 hour and include demographics, apathy, depression, quality of life, a loneliness measure and a cognitive assessment, see table 1. For the VR and the active control groups, a further 1 hour (approximately) is required. This is to conduct an interview about the participant’s history according to reminiscence therapy guidelines for establishing content to be used in the intervention sessions.

**Figure 1** Participant allocation and flow chart of study procedure. VR, virtual reality.

**Table 1** Timeline with measures and assessments used throughout the study

| Assessments                                      | Baseline | Intervention |
|--------------------------------------------------|----------|--------------|
| Activity                                         | X        |              |
| Apathy Evaluation Scale                          | X        |              |
| Addenbrooke’s Cognitive Examination III          | X        |              |
| Quality of life                                  | X        |              |
| Loneliness                                       | X        |              |
| Geriatric Depression Scale                       | X        |              |
| Simulator Sickness Questionnaire                 | X        | X            |
| Heart rate and galvanic skin response            | X        | X            |
| Speech tasks                                     | X        |              |
| Session record                                   | X        | X            |
| Staff questionnaire                              | X        |              |

VR, virtual reality.
During the reminiscence sessions for both intervention groups there is continual conversation, and feedback is requested after the first session to guide the content for the following sessions. The therapy may need to be flexibly tailored depending on the individual’s unique life experience and reaction to the therapy. For example, some participants may have increased positive reactions to music or memories at specific stages of their life. The use of objective measures has been recommended for assessing apathy in clinical trials including actigraphy.39

Galvanic skin response, heart rate40 and more recently, speech analysis41 and their association with apathy have also been examined. Therefore, to measure rest and physical activity, all three groups have the GENEActiv wrist-worn accelerometer (Activinsights, Kimbolton, Cambridgeshire, UK) fitted for a period of 48 hours at baseline, this is repeated after follow-up. The intervention will examine other objective measures of apathy as exploratory outcomes including speech, galvanic skin response and heart rate variability.

Intervention

The intervention consists of three sessions spaced at a minimum of 1 day apart within a 2-week period. All intervention sessions are carried out individually. Both the VR and active control groups have the Empatica E4 (Empatica, Boston, USA) fitted for the duration of the session to measure heart rate variability and galvanic skin response. Event markers on the Empatica E4 are set for each task during the session to distinguish between five separate periods. These include:

1. Pre reminiscence—Speech task 1: Voice recording of the participant’s response to a question (1 min).
2. Pre reminiscence—Speech task 2: Voice recording of the participant reading from a wordless picture book (up to a maximum of 5 min).
3. The reminiscence component (20 min).
4. Post reminiscence—Speech task 3: Voice recording of the participant’s response to a question (1 min).
5. Post reminiscence—Speech task 4: Voice recording of the participant reading from a wordless picture book (up to a maximum of 5 min).

In the VR group, if a participant experiences symptoms of VR sickness that creates discomfort, the session is immediately stopped. If symptoms subside, the intervention continues; if after a second attempt symptoms continue, the participant will be requested to withdraw from the study. All content is to be carefully selected avoiding fast moving scenes, using high-quality footage with smooth motion and stabilised video. Participants are to remain seated during the VR session as this will assist to avoid VR sickness and reduce risk of falls. In the VR group, motion sickness symptoms are measured before and after the reminiscence experience. The passive control group does not undertake any intervention and receives their usual care between baseline and follow-up.

Follow-up

The follow-up session is performed the day after the final intervention session and all baseline measures are repeated. Participants in all three groups are not prevented from participating in their normal lifestyle activities during the research.

Primary outcome

AES clinician version

The presence of apathy will be assessed using the AES clinician version.42 Trained researchers rate each item based on both verbal and non-verbal responses from the participant. The AES contains 18 items measured on a scale ranging from ‘not at all’ to ‘slightly’, ‘somewhat’ and ‘a lot’. Scores range from 18 to 72, with higher scores indicating a higher level of apathy. The clinician version of the AES has found to have a test–retest reliability of 0.88 and good internal consistency (α=0.90).42

Secondary outcomes

Addenbrooke’s Cognitive Examination III (ACE-III)

Cognition will be assessed using the ACE-III, this test is a widely used cognitive screening tool that assesses attention, fluency/language, verbal memory and visuospatial function.43 The ACE-III is scored out of 100 with higher scores indicating higher levels of cognition. There are three different versions of the ACE-III for use in repeated measures testing, two versions are used in this study. The ACE-III has reported good internal consistency (α=0.88).44

Geriatric Depression Scale (GDS) Short Form

The GDS is a tool for detecting a person’s level of depression specifically for older populations.45 This scale has a 92% sensitivity and 89% specificity when compared with diagnostic criteria and good internal consistency (α=0.80).46 The GDS consists of 15 items and is scored out of 15. Higher scores indicate likelihood of depression with a score equal to or greater than 10 suggesting high risk of depression.45

Exploratory outcomes

The Quality of Life in Alzheimer’s Disease (QOL-AD)

The QOL-AD Scale will assess a participant’s current level of quality of life.47 Originally developed for people with Alzheimer’s disease, the QOL-AD has also been used for those without dementia and in residential aged care settings. The QOL-AD is a self-report measure consisting of 13 items rated on a 4-point scale. Good internal consistency has been reported (α=0.82) for the QOL-AD.48

Three-Item Loneliness Scale

The level of loneliness will be assessed using the Three-Item Loneliness Scale. This scale is a shortened version of the Revised UCLA Loneliness Scale.49 The scale comprises three questions and is rated on a 3-point scale. Scores range from 3 to 9 with higher scores indicating increased levels of loneliness. The Three-Item Loneliness Scale has reported acceptable internal consistency (α=0.72).49
after the intervention, the VR group will spend more time in non-passive control groups. It is also expected that both intervention groups will spend more time in non-sedentary, light and moderate activity. Decreased daytime motor activity has been found in individuals with apathy compared with those without apathy. It is expected that after the intervention, the VR group will spend more time pursuing non-sedentary activities than the active and passive control groups. It is also expected that both intervention groups will spend more time in non-sedentary behaviour after the intervention compared with baseline than the passive control group. All data will be processed using a customised script in RStudio.

Heart rate variability and galvanic skin response
Physiological measures including heart rate variability and galvanic skin response will be measured using the Empatica E4 wrist-worn wireless device (Empatica, Boston, USA). The Empatica E4 will be fitted during intervention sessions 1 and 3 to analyse changes throughout the intervention for both the VR and active control groups. Heart rate variability will be derived from the interbeat interval recorded from the Empatica E4. Increased heart rate variability response has been associated with positive emotions. It is expected that a fully immersive experience provided to participants in the VR group will result in an increased heart rate variability response compared with the active control group during the reminiscence sessions.

Galvanic skin response will be measured through skin conductance to examine physiological responses during the reminiscence sessions. Exposure to autobiographical material has found to increase skin conductance response. It is expected that a fully immersive experience provided to participants in the VR group will result in higher physiological responses compared with the active control group during the reminiscence sessions. All data will be processed in MATLAB (MathWorks, Natick, MA, USA) using customised scripts.

Speech
Participants will be requested to perform four speech tasks including responding to a question and reading from a wordless picture book at the start and end of intervention sessions 1 and 3. For the first speech task, two different questions will be used and counterbalanced. For example, ‘describe your typical Sunday’ and ‘what did you do yesterday’. The response will be timed for 60s. For the second speech task, reading from a wordless picture book will be limited to a maximum time of 5 min. A series of four books from the same author with similar content (a different book for each time) will be used and counterbalanced. All speech tasks will be recorded using a digital recorder with a lapel microphone attached to the participant at sternum level. Prosodic, formant, source and temporal aspects of speech will be examined. It is expected that there will be a higher activation of speech markers after the reminiscence sessions compared with before sessions and that activation of speech markers will be higher in the VR group compared with the active control group. Data will be analysed using Praat.

Simulator Sickness Questionnaire (SSQ)
Side effects from using VR will be measured using the SSQ, which is the most commonly used questionnaire in VR research. Higher scores indicate higher side effects. The SSQ includes three subscales including nausea, oculomotor and disorientation. Good internal consistency of the SSQ has been reported (α=0.87).

Activity levels
Actigraphy will be measured using a GENEActiv wristband. The GENEActiv is placed for 48 hours at baseline and is repeated for 48 hours at the end of intervention after follow-up session for all three groups. Variables from GENEActiv will be categorised into levels of sedentary, light and moderate activity. Decreased daytime motor activity has been found in individuals with apathy compared with those without apathy. It is expected that after the intervention, the VR group will spend more time pursuing non-sedentary activities than the active and passive control groups. It is also expected that both intervention groups will spend more time in non-sedentary behaviour after the intervention compared with baseline than the passive control group. All data will be processed using a customised script in RStudio.
body-worn accelerometer that measures and tracks movement in all environments.

**Empatica E4**
Heart rate variability and galvanic skin response will be measured using Empatica E4 wristband (Empatica, Boston, USA). This is a medical-grade wristband that measures real-time physiological data.

**Sample size**
The required sample size was calculated using G*Power statistical analysis V3.1.9.7. A prior sample size was calculated. The approximate sample size was calculated with an analysis of variance for repeated measures (large effect size, power=0.80 and α=0.05). The large effect size was based on outcome of apathy using reminiscence therapy supported by internet based videos. The approximate sample size required was calculated to be n=36 (12 per group); therefore, to account for attrition, a total of 45 participants (15 per group) will be recruited.

**Recruitment**
A research nurse employed by the aged care provider has been assigned to recruit participants; in his/her absence, the primary investigator will undertake recruitment. Potentially suitable participants are identified by senior staff at the residential aged care facility in accordance with inclusion/exclusion criteria and a list is provided to the research nurse and primary investigator. Each group is allocated to a separate residential aged care site, this will assist with ensuring that adequate participant enrolment is achieved and that residents are blinded to the other groups. No honorarium will be offered for participation.

**Blinding and data collection**
The research team members conducting baseline and follow-up measures are blinded to group allocation. Research team members administering outcome measures will not have access to group allocation in the electronic database. Participants are blinded to the presence of the other groups assisted by allocating each group to a separate residential aged care site.

The reminiscence interview and intervention sessions are carried out by the primary researcher and an additional member of the research team who is not conducting baseline and follow-up measures.

**Statistical methods**
Data will be analysed to examine distributions and check for missing values and outliers. Descriptive statistics will be used to summarise the baseline characteristics. This is a per-protocol analysis, participants will be replaced if possible. The primary outcome at baseline and follow-up will be analysed using linear mixed modelling. Fixed factors will be group (VR, active control, passive control) and time (baseline, follow-up), with intercept of participant as a random factor. For comparisons between the two intervention groups with the usual care group, Helmert contrasts will be used where the contrasts will compare (1) combined intervention groups with the usual care group and (2) the two interventions. Additional analysis will be performed including only participants that meet the criteria for a diagnosis of apathy using a cut-off score of 37.5. Further secondary analysis will be conducted to examine the influence of potential covariates including depression and cognition at baseline. Secondary and exploratory outcomes will be analysed as per the primary outcome.

**ETHICS AND DISSEMINATION**

**Informed consent**
Participants that are interested and meet the exclusion/inclusion criteria are provided with written and verbal information about the study. A dedicated research nurse employed by the residential aged care facility or the primary researcher obtains informed consent (see online supplemental file 1). All participants are given the opportunity to discuss participation with family members or other responsible person close to the participant. Consent is continually monitored during the research by asking participants if they want to continue at the start and end of each session.

**Data management**
Each participant will be given a unique identification number that is stored in the electronic database. Research Electronic Data Capture, a secure web-based software platform, will be used for storing all data. All data are stored using anonymous codes. Codes with paper data are in lockable storage and can only be accessed by research staff involved in the project. Data will be stored for a minimum of 5 years from study completion.

**Harms**
Potential for harms is minimal. A risk associated with using HMDs is VR sickness characterised by side effects that are like those experienced in motion sickness from air, sea or land travel. If a participant does experience any side effects, they will be monitored by the researchers until symptoms subside. If symptoms persist longer than the research session, the participant will be referred to the nurse on duty and will be monitored by staff at the aged care facility until symptoms subside.

The process of reminiscence therapy has the potential to raise memories that are distressing. The use of reminiscence therapy will focus on memories that are positive to the participant, negative memories will be avoided. This will avoid causing undue stress to the participant due to memories that may be distressing. It is possible that even positive memories can cause emotional reactions. If any distress does occur, validation, reassurance and distraction approaches will be used depending on the response.

If a participant scores 10 or higher on the GDS at baseline or follow-up, staff at the residential aged facility will...
be made aware and will follow-up and offer support, as necessary.

Data monitoring
Due to the relatively small scale of the trial, there will be no data monitoring committee, interim analyses and auditing trial conduct.

Protocol amendments
Any protocol amendments will be submitted to the University of South Australia Human Research Ethics Committee for approval. The primary investigator will update the trial registry after amendments have been approved.

Access to data
Study staff including the primary investigator and research assistants will have access to raw data files. Access to data for each study staff member is limited to functions they are responsible for. Investigators outside of the study may request access to datasets through the corresponding author.

Dissemination
The results from this study will be disseminated through manuscript publications and national/international conferences. All study investigators will be eligible for authorship depending on contributions to the manuscripts, the use of professional writers is not intended.

Contributors
DS and TL: conception of the work. DS: drafted the manuscript. DS, HADK, MC and TL: revised the work critically for important intellectual content and have read and approved the manuscript.

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Supplemental material
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