Abstract  

Objectives: To assess feasibility and effect of multimodal prehabilitation in patients with severe life-limiting intermittent claudication and complex infrainguinal disease.  

Design: Case series of patients who underwent a 12-week prehabilitation program.  

Setting: Outpatient clinic of a public tertiary hospital  

Participants: Patients with a diagnosis of severe life-limiting intermittent claudication (Fontaine stage IIb and III) with complex infrainguinal disease or previous failed bypass attempts (N=5) who were referred to the prehabilitation clinic by a vascular surgeon.  

Interventions: Patients underwent a baseline assessment that included quality of life questionnaires and functional capacity tests. After baseline assessment, they received a 12-week prehabilitation program that consisted of (1) a supervised exercise session 1 time per week; (2) home-based exercise prescription; (3) nutritional counseling; (4) smoking cessation; and (5) psychosocial intervention. Adherence to all components was recorded as well as the occurrence of any adverse event. After completion of the 12-week program, patients were reassessed.  

Main Outcome Measure: Feasibility of prehabilitation measured by adherence to the different components of the program and occurrence of adverse events.  

Results: All 5 patients completed the program. No serious adverse events occurred during the length of prehabilitation. Median adherence to each prehabilitation component was 91.7% (interquartile range [IQR], 33.5%) for supervised training, 91.7% (IQR, 40%) for home-based exercise, and 75% (IQR, 50%) for nutrition. Three of the 5 patients underwent psychosocial intervention and all who were active smokers enrolled in the smoking cessation program. Functional capacity measured with the 6-minute walk distance improved by 70 m (IQR, 99 m), and disease-specific quality of life measured with the Vascular Quality of Life Questionnaire improved by 25%.
Peripheral arterial disease (PAD) affects up to 10% of the worldwide population and the prevalence increases to 20% among patients 70 years of age and older. Individuals with PAD are often sedentary and experience decreased functional capacity, severe disability, poor quality of life, and increased risk of premature mortality. Lifestyle modifications, especially smoking cessation, exercise, and nutrition are critical for patients with PAD and should be encouraged before contemplating surgery. Despite their importance in preventing disease progression, there is a lack of implementation of all these elements in the clinical setting.

Multimodal prehabilitation, understood as a multidisciplinary optimization strategy that includes structured exercise, nutrition, smoking cessation, and psychological support, has been shown to improve surgical outcomes in different types of surgeries, and it is postulated as one of the most promising approaches to optimize a patient’s functional capacity before and after surgery. There is currently no data on the effect of prehabilitation in PAD population. This multidisciplinary holistic approach could potentially improve patient-centered outcomes in patients with severe life-limiting intermittent claudication who have expected complex revascularization surgery.

Methods

Patient selection and diagnosis of PAD

This study was approved by the Hospital Research Ethics Board. Five consecutive patients with a diagnosis of severe life-limiting PAD and complex infrainguinal disease described as Trans-Atlantic Inter-Society Consensus II D lesions or previous failed bypass attempts were referred to the prehabilitation clinic by vascular surgeons at a tertiary hospital between the years 2018 and 2019. Patients agreed to participate in the program and signed the informed consent. Moderate to severe PAD diagnosis was made clinically by functional limitations in walking activity or pain at night, anatomically by imaging demonstrating arterial stenosis below the inguinal, and hemodynamically by an ankle brachial index \( \leq 0.90 \). All patients were medically optimized and had stable PAD condition.

Assessments and intervention

Baseline assessment included sociodemographic and anthropometric data, disease-specific quality of life questionnaires (Vascular Quality of Life Questionnaire [VascuQol]-25 and Walking Impairment Questionnaire), presence of anxiety or depression (Hospital Anxiety and Depression Scale [HADS]10), and functional capacity (6-minute walk test,11 Gardner-Skinner’s test12). Onset of claudication during the walk test was also recorded. After baseline assessment, patients started a 12-week multimodal prehabilitation program, summarized in Table 1 and fully detailed in the supplemental appendix S1 (available online only at http://www.archives-pmr.org/). Patients were provided with a home-based exercise prescription notebook which they were expected to fill out every time the exercises were performed. Weekly compliance to home-based exercise as well as adherence to nutritional and psychosocial recommendations were recorded in a compliance sheet (see supplemental appendix S1, available online only at http://www.archives-pmr.org/) by the exercise specialist during the weekly supervised visit. During that visit, the exercise specialist also revised any limitations the patient might have had and progressed the exercises if necessary. Any exercise-related adverse events occurring during the supervised session were recorded. A reassessment was performed after completion of the 12-week prehabilitation program.

Analysis

Statistical analyses were performed using SPSS Statistics. Baseline and 12-week data were compared using the related-samples Wilcoxon signed rank test.

Results

A total of 5 patients with a median age of 76 years (range, 57-80y) underwent the 12-week prehabilitation program. Four patients had severe intermittent claudication (Fontaine stage IIb); 1 patient had rest pain (Fontaine III) and was using a wheelchair to move, because pain severely impaired his walking and he was not able to walk >50 m without needing to stop. All 5 patients had cardiovascular risk factors such as hypertension, dyslipidemia, and diabetes. There were 2 active smokers and 3 former smokers. Baseline functional capacity, disease-specific quality of life, and other parameters collected are shown in Table 2.

Median adherence to each prehabilitation component was 91.7% (interquartile range [IQR], 33.5%) for supervised training, 91.7% (IQR, 40%) for home-based, and 75% (IQR, 50%) for nutrition. Three of the 5 patients underwent psychosocial intervention and attended the follow-up visits. All participants who were active smokers (n=2) achieved smoking cessation by the end of the program with the smoking cessation program. No cardiovascular or other exercise-related events were observed during supervised exercise training, although 1 patient presented with a hypertensive peak (blood pressure, 210/90 mmHg) with symptomatology consisting of headache and dizziness upon arrival to one
exercise session. He was immediately referred to the emergency department and did not perform the exercise session on that day. During the program, a reduction in the insulin requirements was possible in 2 patients owing to improvement in glycemic control.

Baseline, 12-week, and changes in functional capacity, quality of life, and emotional status are summarized in table 1 and figure 1. The increase in functional capacity and onset of pain was clinically significant in all participants. Furthermore, the patient who used a wheelchair to ambulate owing to inability to walk for >50 m without needing to stop at baseline became fully functional and stopped using the wheelchair after completion of the 12-week program. Health-related quality of life and walking impairment measured by the VascuQol and Walking Impairment Questionnaire improved, with increases that almost doubled the baseline value in some cases.

Discussion

This is the first study evaluating the effects of a multimodal prehabilitation intervention in patients experiencing intermittent claudication with complex infrainguinal disease. Our findings suggest that a 12-week program may improve functional capacity, quality of life, and anxiety. Despite that international vascular surgery societies have underlined the importance of exercise, nutrition, and smoking cessation for PAD prevention and control,9,13,14 there has been a lack of implementation of these interventions or they have been only partially implemented. Structured exercise for patients with PAD has been proven to be effective at increasing walking capacity and delaying onset of pain and has been shown to be as effective as medication and endovascular revascularization with respect to increasing pain-free walking distance.15 Nevertheless, very few studies have formally addressed quality of life in patients with PAD undergoing an exercise intervention and none have examined the effect of prehabilitation on quality of life or functional outcomes in PAD.8,16

The feasibility of multimodal prehabilitation has been demonstrated in these 5 cases of patients with severe life-limiting claudication and complex infrainguinal disease or with previous failed bypass attempts. The improvements in functional capacity, health-related quality of life, and emotional status shown in this case series are promising. This is

| Table 1 Description of the 12-week multimodal prehabilitation program for patients with PAD |
| ----------------------------------------------- | ----------------------------------------------- | ----------------------------------------------- |
| **Exercise**                                   | **Supervised**                                 | **Home-based**                                 |
| - 12-wk program                               | 1 time/wk                                      | 3 times/wk                                      |
| - Goal progression every week if program is well tolerated | | |
| **Nutrition**                                 | **Nutritional education**                      | **Nutritional intervention**                   |
| - 1 first visit                               | - Balanced meals                               | - Nutritional assessment                       |
| - 1 follow-up visit                           | - Correct portion size                         | - Ensure balanced macronutrient intake         |
| **Psychosocial**                              | - Timing and spacing of meals                  | - Weight management                            |
| - If score in HADS-A > 6 or HADS-D > 8       | - Mindful eating                               | - Cognitive reframing                          |
| **Smoking cessation**                         | - Protein importance and sources               | - Relaxation and deep breathing exercises      |
|                                               |                                               | - Anxiety coping strategies                    |
|                                               |                                               | - Disease teaching and motivational interviewing|
|                                               |                                               | - Nicotine replacement therapy                 |
|                                               |                                               | - Progress tracking with exhaled carbon monoxide analyzer |

Abbreviations: HADS-A, Hospital Anxiety and Depression Scale—Anxiety; HADS-D, Hospital Anxiety and Depression Scale—Depression.
especially relevant given the complexity of these patients, with higher rates of unsuccessful surgery and, in some cases, ending up with worse outcomes than before surgery. Therefore, nonsurgical interventions aimed at improving postoperative outcomes and delaying or avoiding the need for surgery in this population are crucial.

We believe that the reason for the good results attained rely on the synergy created when combining all the components (structured aerobic and resistance exercise, psychosocial support, nutritional counseling, smoking cessation). This holistic approach might have helped in breaking the enrooted poor health behavior and reluctance to change that is characteristic of this population. Furthermore, adherence to all components was higher compared with other trials with behavioral interventions in this population.\textsuperscript{17,18} As the literature shows, supervised exercise is better than unsupervised; however, in a clinical setting, asking a patient to come to a hospital facility 3 or 4 times per week might drop the adherence to the program owing to the economic and transportation burden for patients and caregivers. The mixed approach (supervised and home-based) of the prehabilitation program helped patients to remain adherent to the program by using the supervised sessions to track progress and make sure that patients felt accompanied during their exercise “journey,” whereas the home-based portion kept the program low-cost and made it more

| Measure                          | Baseline | 12-Week | Change  | P Value |
|----------------------------------|----------|---------|---------|---------|
| **Functional capacity**          |          |         |         |         |
| 6-MWT distance, m               | 285 (69) | 362 (145)| +70 (99) | <.05    |
| Onset of pain, s                 | 98 (49)  | 145 (161)| +31 (136)| <.05    |
| Gardner-Skinner, s               | 97 (193) | 182 (363)| +90 (172)| <.05    |
| **Quality of life**              |          |         |         |         |
| VascuQol, total                  | 3.47 (1.93)| 4.84 (1.10)| +0.87 (1.24)| <.05    |
| Pain domain                      | 2.75 (2.13)| 4 (2.0)   | +1.0 (1.5) | .257    |
| Social domain                    | 4.5 (2.25)| 5.5 (1.5) | +1.0 (1.75)| .104    |
| Activity domain                  | 2.75 (1.68)| 4.25 (1.65)| +1.45 (1.45)| <.05    |
| Symptoms domain                  | 3.5 (2.78)| 6 (1.38)  | +1.03 (2.13)| .608    |
| Emotional domain                 | 3.29 (1.58)| 4.63 (1.59)| -0.86 (1.12)| <.05    |
| WIQ, total                       | 18.0 (10) | 45.2 (26.2)| +27.2 (24) | <.05    |
| WIQ distance                     | 5.9 (21.8) | 37.1 (36.7)| +18.6 (28.8)| <.05    |
| WIQ speed                        | 14.1 (10.7) | 39.1 (26.5)| +28.3 (18.4)| <.05    |
| WIQ stairs                       | 29.2 (20.8) | 54.2 (54.1)| +20.9 (35.4)| .104    |
| **Anxiety and depression**       |          |         |         |         |
| HADS-A                           | 9 (6.5)  | 4 (4)   | -3 (3.5) | <.05    |
| HADS-D                           | 7 (10.5) | 3 (4.5) | -4 (6)  | .144    |

NOTE. Data are expressed as median (IQR). Change from baseline to 12 weeks is expressed as median of differences (IQR).
Abbreviations: 6-MWT, 6-minute walk test; HADS-A, Hospital Anxiety and Depression Scale—Anxiety; HADS-D, Hospital Anxiety and Depression Scale—Depression; WIQ, Walking Impairment Questionnaire.
Multimodal prehabilitation appears to be feasible in patients with PAD experiencing severe life-limiting intermittent claudication with complex infragenual disease or previous failed bypass attempts. Multimodal prehabilitation could be an excellent tool to increase their quality of life and functional capacity, although further studies are needed to prove the safety and effect of this multimodal approach on quality of life, functional capacity, and postoperative outcomes, as well as its role in delaying or sparing the need for surgery in some cases.

Conclusions

Multimodal prehabilitation appears to be feasible in patients with PAD experiencing severe life-limiting intermittent claudication with complex infragenual disease or previous failed bypass attempts. Multimodal prehabilitation could be an excellent tool to increase their quality of life and functional capacity, although further studies are needed to prove the safety and effect of this multimodal approach on quality of life, functional capacity, and postoperative outcomes, as well as its role in delaying or sparing the need for surgery in some cases.

Supplier

a. SPSS Statistics for MacOS, version 23.0; IBM Corp.

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