Paper Title

Citation for published version (APA):

Citing this paper
Please note that where the full-text provided on King's Research Portal is the Author Accepted Manuscript or Post-Print version this may differ from the final Published version. If citing, it is advised that you check and use the publisher's definitive version for pagination, volume/issue, and date of publication details. And where the final published version is provided on the Research Portal, if citing you are again advised to check the publisher's website for any subsequent corrections.

General rights
Copyright and moral rights for the publications made accessible in the Research Portal are retained by the authors and/or other copyright owners and it is a condition of accessing publications that users recognize and abide by the legal requirements associated with these rights.

•Users may download and print one copy of any publication from the Research Portal for the purpose of private study or research.
•You may not further distribute the material or use it for any profit-making activity or commercial gain
•You may freely distribute the URL identifying the publication in the Research Portal

Take down policy
If you believe that this document breaches copyright please contact librarypure@kcl.ac.uk providing details, and we will remove access to the work immediately and investigate your claim.
Strategies to address the shortcomings of commonly used advanced chronic heart failure descriptors to improve recruitment in palliative care research: A parallel mixed-methods feasibility study

Pauline M Kane1, Fliss EM Murtagh1, Karen R Ryan2,3, Mary Brice4, Niall G Mahon3, Brendan McAdam5, Regina McQuillan2,5, Geraldine O’Gara5, Caroline Raleigh5, Cecelia Tracey3, Christine Howley3, Irene J Higginson1 and Barbara A Daveson1; on behalf of BuildCARE*

Abstract

Background: Recruitment challenges contribute to the paucity of palliative care research with advanced chronic heart failure patients.

Aim: To describe the challenges and outline strategies of recruiting advanced chronic heart failure patients.

Design: A feasibility study using a pre–post uncontrolled design.

Setting: Advanced chronic heart failure patients were recruited at two nurse-led chronic heart failure disease management clinics in Ireland.

Results: Of 372 patients screened, 81 were approached, 38 were recruited (46.9% conversion to consent) and 25 completed the intervention. To identify the desired population, a modified version of the European Society of Cardiology definition was used together with modified New York Heart Association inclusion criteria to address inter-study site New York Heart Association classification subjectivity. These modifications substantially increased median monthly numbers of eligible patients approached (from 8 to 20) and median monthly numbers recruited (from 4 to 9). Analysis using a mortality risk calculator demonstrated that recruited patients had a median 1-year mortality risk of 22.7 and confirmed that the modified eligibility criteria successfully identified the population of interest. A statistically significant difference in New York Heart Association classification was found in recruited patients between study sites, but no statistically significant difference was found in selected clinical parameters between these patients.

Conclusion: Clinically relevant modifications to the European Society of Cardiology definition and strategies to address New York Heart Association subjectivity may help to improve advanced chronic heart failure patient recruitment in clinical settings, thereby helping to address the paucity of palliative care research this population.

Keywords

Heart failure, palliative care, recruitment, patient-reported outcome measures

1Department of Palliative Care, Policy & Rehabilitation, Cicely Saunders Institute, Cicely Saunders International, King’s College London, London, UK
2St Francis Hospice, Dublin, Ireland
3Mater Misericordiae University Hospital, Dublin, Ireland
4St Christopher’s Hospice, London, UK
5Beaumont Hospital, Dublin, Ireland

*Project BuildCARE is about Building Capacity, Access, Rights and Empowerment for patients to improve the palliative care patients receive. Project BuildCARE is supported by Atlantic Philanthropies.

Corresponding author:
Pauline Mary Kane, Department of Palliative Care, Policy & Rehabilitation, Cicely Saunders Institute, Cicely Saunders International, King’s College London, Bessemer Road, London SE5 9PJ, UK.
Email: pauline.m.kane@kcl.ac.uk
Introduction

Chronic heart failure (CHF) is a progressive condition with a comparable mortality rate to cancer, predominantly affecting older adults with multi-morbidity. It is a condition with a risk of sudden or premature death and can necessitate therapeutic decision-making between life-prolonging treatments and quality of life. Older CHF patients with multi-morbidity particularly require a holistic patient-centred approach to care to facilitate therapeutic decision-making around quality of life versus life prolongation. Many CHF patients experience palliative needs. A palliative care needs assessment presents one way to integrate palliative care into CHF management and potentially improve patient care.

Recruitment challenges contribute to the paucity of CHF palliative care needs assessment research. Recruitment is difficult for many reasons: advanced CHF is difficult to define; it has unclear inconsistent descriptive nomenclature; it has a variable progression rate and unpredictable prognosis; there is high patient attrition due to illness burden and high mortality; and gate-keeping often occurs.

This article describes recruitment challenges experienced in a feasibility study of a palliative-specific patient-reported outcome measure (PROM) intervention and how these challenges were addressed. It provides recruitment solutions for future studies involving CHF patients.

Methods

Study design

This parallel mixed-methods feasibility study involved a pre–post uncontrolled longitudinal study design. This study examined the feasibility and acceptability of the palliative-specific PROM intervention to improve identification and management of palliative symptoms and concerns in advanced CHF patients. As part of this feasibility study, we examined our ability to recruit participants.

Setting

Study settings were CHF disease management clinics in two tertiary referral centres in Ireland led by heart failure nurses (HFN) and supported by CHF physicians.

Inclusion criteria development

Inclusion criteria (Figure 1) were developed by the study lead (P.M.K.), the HFNs and the research team. The criteria were informed by the advanced CHF definition in the 2012 European Society of Cardiology (ESC) Heart Failure Association position statement and by the ESC palliative care position statement. Although not included in the 2012 ESC definition, a definition of Heart Failure with preserved Ejection Fraction (HFpEF) as per the 2016 ESC definition was included in the criteria at the HFNs’ recommendation as this constituted over 50% of their patient population. Of note, HFpEF is included in the 2016 ESC definition which was not published at the time of this study. Once recruitment commenced, P.M.K. and the HFNs frequently reviewed the inclusion criteria. Subsequently, the requirement for a heart failure–related hospitalisation in the preceding 6 months was removed because it was not appropriate in a
setting where nurse-led clinics work to avoid heart failure–related admissions.

**Ethics**

Ethical approval was obtained from the Research Ethics Committees: Ref: 13/70, Ref: 1/378/1579 and Ref: BDM/13/14-25. Written informed consent was obtained from participants.

**Intervention**

The intervention involved administration of the Integrated Palliative care Outcome Scale (IPOS) together with HFN training. Training involved patient-centred education, the role of PROMs to facilitate patient-centred care, training in IPOS interpretation and use in consultation. The IPOS is a 20-item questionnaire measuring patient’s physical symptoms, psychological, emotional, spiritual, information and support needs.

**Strategies to address recruitment challenges**

To increase engagement, HFNs and service users helped develop study participant documentation. Posters advertising the study were placed in study sites. Screening logs were developed with HFN input. HFNs reviewed the clinical roster daily to identify eligible patients and completed weekly logs recording: numbers of screened and eligible patients, number approached, number recruited, reasons why patients were deemed ineligible, were not approached or declined participation.

Patients were given information on three occasions before written consent was requested. HFNs provided summary cards outlining key study points. They provided patients with documentation and obtained permission for the researcher to explain the study in depth. Potential participants returned signed consent forms. The researcher arranged a telephone appointment to complete the first set of questionnaires, which otherwise acted as a patient reminder for those who had not returned consent forms.

**Analysis**

Analysis of recruited patients was undertaken to determine whether the inclusion criteria modifications identified the same population (advanced CHF patients). We applied the Meta-analysis Global Group in Chronic Heart Failure (MAGiC) heart failure mortality risk score to

---

Figure 1. Adaptation of the European Society of Cardiology’s advanced chronic heart failure definition for the study inclusion criteria. CHF: chronic heart failure; HF: heart failure; NYHA: New York Heart Association; LVEF: left ventricular ejection fraction; PCWP: pulmonary capillary wedge pressure; RAP: right atrial pressure; BNP: brain natriuretic peptide; NT: N-terminal; 6-MWT: 6-minute walk test; VO2: oxygen consumption; CRT: cardiac resynchronisation therapy.

---

**Patient Inclusion Criteria for Feasibility Study**

- Patients with advanced chronic heart failure:
  - Severe symptoms (NYHA III/IV)*
  - Episodes with clinical signs of fluid retention and/or peripheral hypoperfusion
  - Objective evidence of severe cardiac dysfunction*:
    - Systolic dysfunction and left ventricular ejection fraction of ≤35% (HFrEF) or
    - Heart failure symptoms in the presence of preserved systolic function ≥40% (HFpEF)
  - Severe impairment of functional capacity demonstrated by an inability to exercise
  - History of 1 heart failure hospitalisation in the previous 6 months
  - Presence of all these features despite optimal therapy
  - Attending/referred to the heart failure clinical nurse specialist clinic ≥ 18 years of age*
  - Sufficient cognitive function to complete the questionnaires*

*essential inclusion criteria
calculate the median 1-year mortality risk for each recruited patient\textsuperscript{21} to determine participant homogeneity. The Kruskal–Wallis Test was applied to selected clinical markers to test for difference between participants assigned different New York Heart Association (NYHA) classes.

**Results**

**Recruitment**

Patients were recruited from September 2014 until February 2015 at the nurse-led clinics. Of 372 patients screened, 81 were approached, 38 were recruited (46.6\% conversion to consent\textsuperscript{22}) and 25 completed the IPOS. Figure 2 shows the modified CONSORT flow diagram.\textsuperscript{23}

**Recruited patients’ characteristics**

Table 1 reports the demographic and clinical data for participants. To ensure a representative sample of advanced CHF patients, comparison was made with data from Pocock et al.’s\textsuperscript{21} study. This demonstrated comparability with deceased patient data (Table 1).

A review of recruitment activity showed that Site A recruited low numbers. Two issues were identified. First, patients were deemed too unwell to be approached for recruitment. Second, patients frequently declined to participate. On exploration, a difference was identified in the manner in which Site A and Site B applied NYHA classification. Site A patients classified as NYHA III frequently had lower functional ability than Site B patients classified as NYHA III. Where Site B classified patients as NYHA III, Site A classified comparable patients as NYHA II. This resulted in Site A identifying a more unwell population of NYHA III patients than Site B limiting recruitment. To ensure recruitment homogeneity, eligibility criteria were modified to allow Site A to recruit patients who were currently classified as NYHA II, who had been NYHA III/IV in the previous 6 months. This resulted in an increase in the total median number approached monthly from 8 to 20, and in the median numbers recruited monthly, from 4 to 9 (Figure 3).

The MAGGIC mean integer mortality risk score was 27.9 ± 4.9 (median: 29.0) with a mean 1-year mortality proportional risk of 22.2 ± 9.2 (median: 22.7). A Kruskal–Wallis Test demonstrated no statistical difference in creatinine levels between patients assigned NYHA II (n=8) or assigned NYHA III/IV (n=17), \( \chi^2 (20, n=25)=17.75, p=0.63 \) between the two sites (Table 2). A Kruskal–Wallis Test demonstrated no statistical difference in 1-year mortality risk between patients assigned NYHA II (n=8) or assigned NYHA III/IV (n=17), \( \chi^2 (12, n=25)=17.75, p=0.12 \) between the two sites (Table 2). Median scores for creatinine and 1-year mortality were 117 and 14.7 for the NYHA II group and 122 and 24.8 for the NYHA III/IV group.
Kruskal–Wallis Test revealed a statistically significant difference in NYHA classification across sites (Site A, \( n = 10 \); Site B, \( n = 15 \)), \( \chi^2 (1, n = 25), 16.94, p = 0.0001 \) (Table 3). The median NYHA class for Site B was 3, that is, NYHA III, while in Site A, it was 2, that is, NYHA II. There was no statistical significant difference in systolic blood pressure, Charlson comorbidity index,\(^{24}\) creatinine or 1-year mortality risk\(^{21}\) between patients recruited at each site (Table 3).\(^{21}\)

**Discussion**

This study highlights that commonly used advanced CHF descriptors do not comprehensively identify advanced CHF patients, necessitating a different approach in palliative care study inclusion criteria development. The 2012 ESC advanced CHF definition\(^{15}\) which we used to inform our inclusion criteria (Figure 1) was too narrow to capture fluctuating health status.\(^{25}\) Patients who were initially too unwell to be approached were too well post-treatment to fulfil inclusion criteria based on this definition. This study demonstrates that the subjectivity of the application of NYHA classification and heterogeneity of this population mean using NYHA as a stand-alone inclusion criterion risks excluding patients from studies and reducing generalisability. Restricting participation to NYHA III/IV excludes more stable patients classified as NYHA II at the time of study recruitment, yet these patients may have palliative care needs.

The NYHA classification was designed to classify heart disease patients based on functional capacity, in the absence of more objective evidence.\(^{26,27}\) Its lack of interrater reliability,\(^{28}\) poor reproducibility and validity have been acknowledged.\(^{29}\) Yet it continues to play a role in inclusion criteria\(^{30,31}\) and as a determinant for palliative care access.\(^{31}\) To address NYHA limitations, the American College of Cardiology Foundation (ACC)/American Heart Association (AHA) CHF staging system describes CHF in stages of disease progression,\(^{32}\) while contextualising NYHA class to provide a more objective and reliable approach to identification (Figure 4).\(^{32}\)
CHF is characterised by symptom volatility meaning NYHA class can change over short time periods affecting eligibility for study inclusion criteria applied at a single time-point. We extended recruitment to patients classified as NYHA II, with a NYHA III/IV history. There was no statistical difference in clinical parameters between these patients and those recruited as class NYHA III/IV. Limiting recruitment to NYHA III/IV excludes symptomatically stable NYHA II patients who have been NYHA III/IV previously and are at risk of
deterioration from underlying heart disease or multi-morbidity. A limitation of this study is that it may not reflect CHF patients without access to nurse-led clinics. Additionally, we did not conduct quality assurance of the HFNs’ assessment of eligibility and information provision to patients.

**Conclusion**

The recently updated 2016 ESC definition better reflects the interplay of comorbidities with CHF but more emphasis is required on a palliative care approach throughout the disease trajectory. Despite its shortcomings, NYHA classification is widely used. Strategies to address NYHA subjectivity including the ACC/AHA CHF staging system may improve advanced CHF patient recruitment and advance palliative care research.

**Acknowledgements**

The views expressed are those of the authors and not necessarily those of the National Health Service, the NIHR or the Department of Health. CLAHRC South London is part of the NIHR, and a partnership between King’s Health Partners, St. George’s University of London and St. George’s Healthcare NHS Trust. The authors thank all collaborators and advisors including service users.

**Declaration of conflicting interests**

The author(s) declared no potential conflicts of interest with respect to the research, authorship and/or publication of this article.

**Funding**

The author(s) disclosed receipt of the following financial support for the research, authorship and/or publication of this article: This paper presents independent research funded by BuildCARE, and part-funded by the National Institute for Health Research (NIHR) under the Programme Grants for Applied Research Programme (RP-PG-1210-12015 C-CHANGE) and the NIHR Collaboration for Leadership in Applied Health Research & Care (CLAHRC) through CLAHRC South London. BuildCARE is supported by Cicely Saunders International (CSI) and Atlantic Philanthropies, and led by King’s College London, CSI, UK.

**References**


11. McMurray JIV, Adamopoulos S, Anker SD, et al. ESC Guidelines for the diagnosis and treatment of acute and chronic heart failure 2012: the Task Force for the diagnosis and treatment of acute and chronic heart failure 2012 of the European Society of Cardiology. Developed in collaboration with the Heart Failure Association (HFA) of the ESC. *Eur Heart J* 2012; 33: 1787–1847.


