

The effect of a clinical pharmacist discharge service on medication discrepancies in patients with heart failure

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Abstract *Objective* Heart failure patients are regularly admitted to hospital and frequently use multiple medication. Besides intentional changes in pharmacotherapy, unintentional changes may occur during hospitalisation. The aim of this study was to investigate the effect of a clinical pharmacist discharge service on medication discrepancies and prescription errors in patients with heart failure. *Setting* A general teaching hospital in Tilburg, the Netherlands. *Method* An open randomized intervention

study was performed comparing an intervention group, with a control group receiving regular care by doctors and nurses. The clinical pharmacist discharge service consisted of review of discharge medication, communicating prescribing errors with the cardiologist, giving patients information, preparation of a written overview of the discharge medication and communication to both the community pharmacist and the general practitioner about this medication. Within 6 weeks after discharge all patients were routinely scheduled to visit the outpatient clinic and medication discrepancies were measured. *Main outcome measure* The primary endpoint was the frequency of prescription errors in the discharge medication and medication discrepancies after discharge combined. *Results* Forty-four patients were included in the control group and 41 in the intervention group. Sixty-eight percent of patients in the control group had at least one discrepancy or prescription error against 39% in the intervention group (RR 0.57 (95% CI 0.37–0.88)). The percentage of medications with a discrepancy or prescription error in the control group was 14.6% and in the intervention group it was 6.1% (RR 0.42 (95% CI 0.27–0.66)). *Conclusion* This clinical pharmacist discharge service significantly reduces the risk of discrepancies and prescription errors in medication of patients with heart failure in the 1st month after discharge.

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Impact of findings on practice

- The studied clinical pharmacist discharge service reduces the percentage of heart failure patients with one

- or more discrepancies or prescription errors by almost a half (68% vs. 39%).
- The information about medication at discharge for heart failure patients needs to be optimized for community pharmacists, for general practitioners, and for the patients.

Introduction

Heart failure is a chronic, progressive disease characterized by frequent hospital admissions and high mortality rates [1]. The primary goals of improving disease management in patients with heart failure are optimization of the pharmacological therapy and improving adherence to medication and lifestyle. Medication such as angiotensin-converting enzyme inhibitors and β -blockers are well established in the treatment of heart failure, reducing mortality and readmissions [2]. Despite pharmacotherapy, outcomes for patients remain poor and frequent hospitalisations remain necessary [3]. Hospital admissions usually lead to changes in medication use. Most changes are intended, for example adjusting dosage during hospital admission. Other changes, however, are unintended as is the case with prescription errors. In addition, medication discrepancies after hospital discharge can occur by insufficient patient education or insufficient communication to the general practitioner (GP) or the patient his community pharmacist about the intentional changes [4]. A possible consequence of these discrepancies and prescription errors can be readmission [3].

A recent review demonstrated that pharmacist care in the treatment of heart failure greatly reduces the risk of all-cause and heart failure hospitalization, particularly if the pharmacist was a member of a multidisciplinary team [5]. Yu et al. have systematically compared the disease management programmes for older patients with heart failure. This review showed that a program must be multifaceted and should consist of an in-hospital phase of care, intensive patient education, exercise and psychosocial counselling, self care supportive strategy, optimization of medical regimen, and ongoing surveillance and management of clinical deterioration [6]. There is also evidence that a pharmacist intervention for outpatients with heart failure can improve adherence to cardiovascular medications and decrease health care cost [7]. However, the study of Holland et al. [8] did not find a reduction in hospital admissions when a community pharmacist led the intervention in contrast to the results found when a specialist nurse led the intervention. These studies all looked at readmission and hospitalization. None of the studies have focused on the reduction of medication discrepancies after hospital discharge, which may be an indicator of readmission.

Several studies have been published which describe methods to optimize medication use after hospital discharge, but the most effective method has not been established yet [9–11]. A possible useful strategy is providing patients with more information about the (side)effects of their medication and explaining the changes made in pharmacotherapy during their admission. Al-Rashed et al. [10] showed that pharmaceutical counselling before discharge, together with a medication and information discharge summary and a medicine reminder chart lead to better drug knowledge and compliance and a reduction of readmissions. Providing the patient with a copy of the drugs prescribed on discharge, i.e. a full overview of the current medication which can be incorporated into the patient his medication record at the community pharmacy, also seems to be effective. Discharging 19 patients with such information to take to their community pharmacist could result in the prevention of one unintentional discrepancy having a definite adverse effect [11].

Research on the prevention of medication discrepancies has mainly taken place in the USA and UK. Within these countries the availability of hospital pharmacists is higher than within the countries of the European continent. This makes extrapolation of the results to the European situation difficult and projects can not be easily implemented in European hospitals because of this manpower problem. Several solutions are possible for the manpower problem. First of all, the use of alternative personnel may be an option as was shown in two Dutch studies on medication reconciliation [12, 13]. A second solution may be to focus the attention of the clinical pharmacist on high risk patients during these interventions. Heart failure patients are such high risk patients because they use a large number of medicines and are frequently admitted to hospital [1].

Aim of the study

The aim of this study was to investigate the effects of a multifaceted clinical pharmacist discharge service on the number of medication discrepancies after discharge in heart failure patients. A secondary aim was to make an estimation of the effect of the service on non-adherence.

Method

Design

An open randomized intervention study was performed comparing an intervention group provided with the clinical

pharmacist discharge service, with a control group provided with regular care by doctors and nurses.

The study protocol was approved by a medical ethics committee (Medische-Ethische Toetsing Onderzoek Patiënten en Proefpersonen, Tilburg, The Netherlands).

Setting and study population

The study was conducted at the department of cardiology of a teaching hospital in Tilburg, the Netherlands between May 2007 and July 2008.

Eligible patients were adults (aged over 18 years) admitted with a diagnosis of heart failure and prescribed five or more medicines (from any class) at discharge. We excluded patients living in a nursing home or unable to give informed consent, due to mental incapacity or terminal illness.

All patients who provided written informed consent were randomised using a random number table, to receive intervention or regular care.

Regular care

Patients in the control group received regular care, consisting of verbal and written information about their drug therapy from a nurse at hospital discharge. The discharge prescription was made by the physician and given to the patient to hand over to the GP.

Intervention

First of all, the intervention consisted of a clinical pharmacist identifying potential prescription errors in the discharge medication and discussing them with the cardiologist. This resulted in the final discharge medication. Furthermore, patients in the intervention group received verbal and written information about (side)effects of, and changes in, their in hospital drug therapy from a clinical pharmacist upon hospital discharge. In addition to this, the clinical pharmacist made a discharge medication list which contained additional information related to dose adjustments and discontinued medication. After it had been approved by the physician, the discharge medication list was faxed to the community pharmacy and given as written information to the patient with the instruction to hand it over to the GP.

All patients (both regular care and intervention) collected medication at their community pharmacy and received usual routine management by their cardiologist after discharge. This included an outpatient visit within 6 weeks after hospital discharge and an additional visit to the heart failure nurse if necessary.

Data collection

The following patient characteristics have been collected: age, sex, education (primary school or higher education), living situation (single or cohabitating), chronic co-morbidity (Chronic Disease Score, CDS [14]), routine check ups at the heart failure unit before admission, length of admission, number of medicines at discharge, living conditions after discharge (i.e. living in a nursing home, in a residential home for elderly people or at home with additional care), patient or pharmacy control over medication (i.e. patient is in control or the patient receives a “week box”, pre-filled by the pharmacy; a week box contains all the medication for a week arranged by day and hour of intake) and New York Heart Association (NYHA) class upon discharge.

In addition, medication was classified by ATC code and the “source” of each drug, i.e. start or discontinuation during admission, dose adjustment or preadmission medication, was noted.

During the first follow up consultation after discharge with the cardiologist or heart failure nurse at the clinic, an estimate of adherence was made with the “Brief Medication Questionnaire—Regimen Screen” (BMQ), a validated tool for screening for adherence consisting of seven questions. This tool requires patients to list all medication taken in the past week and subsequently for each medicine listed four questions about the use of the medicine are asked, as well as three general questions about medication use. For each item patients received a score of “1” if their initial or spontaneous report indicated potential non-adherence with the current regimen for the target medication (i.e. when the specific question was answered with ‘yes’) and a score of “0” if this reports indicated no non-adherence (i.e. when the question was answered with ‘no’). The maximum score is 7 and a score of 1 or higher is an indication for potential non-adherence. Table 1 shows the questions of the BMQ [15].

In addition, the patient’s medication was checked for discrepancies and for prescription errors. Discrepancies were discussed with the patient and the cardiologist or heart failure nurse. A discrepancy was defined as a deviation in medication use compared to the medication on the discharge prescription. Discrepancies were classified as: re-start of discontinued medication, discontinuation of prescribed discharge medication, use of higher or lower dose, more or less frequent use than prescribed and incorrect time of taking medication.

A prescription error was defined as an error which occurs in the process of prescribing medication, namely dosing errors, dosage form errors, contra-indications, drug-drug interactions and double-medication. All prescription

Table 1 Questions of the BMQ-Regimen Screen [15]

Question	Score ^a
Did patient fail to list the prescribed drugs in the initial (spontaneous) report?	Yes = 1; no = 0
Did patient stop or interrupt therapy due to late refill or other reason?	Yes = 1; no = 0
Did patient report any missed days or doses?	Yes = 1; no = 0
Did patient reduce or cut down the prescribed amount per dose?	Yes = 1; no = 0
Did patient take any extra doses or more medication than prescribed?	Yes = 1; no = 0
Did patient report “do not know” in response to any questions?	Yes = 1; no = 0
Did patient refuse to answer any questions?	Yes = 1; no = 0

^a Score of ≥ 1 indicates potential non-adherence

Table 2 NCC MERP Classes

Class	Content
A	Circumstances or events that have the capacity to cause error
B	An error occurred but the error did not reach the patient
C	An error occurred that reached the patient but did not cause patient harm
D	An error occurred that reached the patient and required monitoring to confirm that it resulted in no harm to the patient and/or required intervention to preclude harm
E	An error occurred that may have contributed to or resulted in temporary harm to the patient and required intervention
F	An error occurred that may have contributed to or resulted in temporary harm to the patient and required initial or prolonged hospitalization
G	An error occurred that may have contributed to or resulted in permanent harm
H	An error occurred that required intervention necessary to sustain life
I	An error occurred that may have contributed to or resulted in the patient’s death

errors identified by the clinical pharmacist and agreed upon by the cardiologist were collected.

The clinical relevance of the discrepancy or prescription error was assessed by making use of the NCC MERP-index [16]. This index categorizes medication errors (class A–I), using an algorithm, see Table 2 (briefly, class A: no error, class B, C and D: error, but no harm, class E, F, G and H: error and harm, class I: error and death). Discrepancies and prescription errors in class E or higher (i.e. errors resulting in harm) are considered as clinically relevant. Three pharmacists and a cardiologist assessed the clinical relevance; for those discrepancies they disagreed on they met to reach consensus.

End points

The primary end point in this study is the total sum of the percentage of prescription errors and discrepancies after hospital discharge. The estimate of adherence as indicated by the BMQ was chosen as a secondary end point. Patients with a score of ≥ 1 were considered to be potentially nonadherent [15].

Data analysis

The program PS sample size (version 2.1.31) was used to determine the sample size [17]. The sample size was

calculated at 62 patients per group based on $\alpha = 0.05$, a power of 0.8, an estimated frequency of the end point in the control group of 30% [3, 4, 11, 18, 19] and an expected reduction to 10% [10, 20].

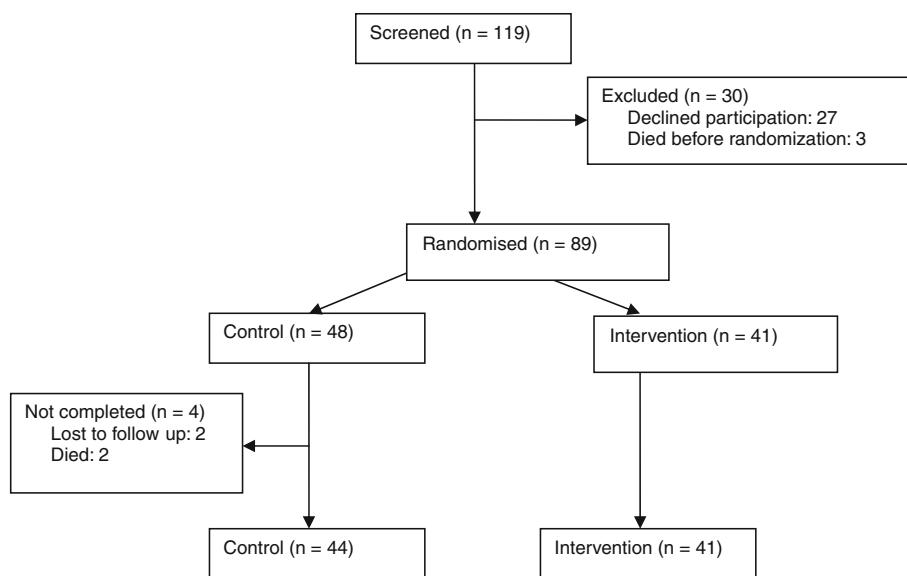
All data were processed in Microsoft Access 2003 and analysed with SPSS version 16.0.

The average and standard deviation were determined for continuous variables and the percentage was calculated for categorical variables. The differences between the intervention and the control group were analysed by the two sample *t* test for continuous variables and by the Chi-square test for categorical variables. A *P* value of ≤ 0.05 was considered to be significant.

For analysis of the primary and secondary end point the relative risk (RR) was calculated with a confidence interval (CI) of 95%. For the primary endpoint this was performed both on the medication level (number of medications as denominator) and on the patient level (% of patients with one or more discrepancy or prescription errors), and for the secondary endpoint on the patient level.

Results

We approached a total of 119 patients to participate after screening them for eligibility. The 85 patients who agreed were randomised; 44 patients in the control group and 41

Fig. 1 Study participant flowchart**Table 3** Patient characteristics

	Control (n = 44)	Intervention (n = 41)	P value
Age (years ± sd)	72 ± 10	74 ± 12	>0.05 ^a
Sex (% male)	75	59	>0.05 ^b
Time to follow-up (days ± sd (range))	23 ± 10 (6–40)	24 ± 12 (7–48)	>0.05 ^a
Education (% primary school only)	39	42	>0.05 ^b
Single or cohabitating (% single)	41	46	>0.05 ^b
Chronic co-morbidity (CDS ± sd)	8 ± 3	8 ± 3	>0.05 ^a
Treatment at the heart failure clinic before admission (% no)	75	78	>0.05 ^b
Length of admission (days ± sd)	12 ± 8	13 ± 7	>0.05 ^a
Medication at moment discharge (number ± sd)	9 ± 3	10 ± 4	>0.05 ^a
Living conditions after discharge			
At home (%)	66	66	>0.05 ^b
At home with additional care (%)	14	22	
Residential home for elderly people (%)	20	12	
Patient control over medication (% yes)	82	81	>0.05 ^b
NYHA class at discharge			
I/II (%)	52	56	>0.05 ^b
III (%)	48	39	
IV (%)	0	5	

^a Tested with t test^b Tested with Pearson Chi-square

patients in the intervention group (see Fig. 1). Patient characteristics are represented in Table 3. The characteristics of both groups did not differ.

Sixty-eight percent of patients in the control group had at least one discrepancy or prescription error against 39% in the intervention group (RR 0.57 (95% CI 0.37–0.88)). The percentage of medications with a discrepancy or prescription error in the control group was 14.6% and in the intervention group it was 6.1% (RR 0.42 (95% CI 0.27–0.66)) (see Table 4).

Prescription errors are most common in both groups. These are followed by re-start of discontinued medication and use of higher dose of medication in the control group. In the intervention group prescription errors are followed by discontinuation of prescribed medication and use of a lower dose of medication (see Table 5).

In the intervention group as well as the control group the highest percentage of the total number of prescription errors and discrepancies fell into class B of the NCC MERP-index (36% vs. 53%). The percentage of class E or

Table 4 Discrepancy/prescription error

	Intervention	Control	Total
Number of patients with a discrepancy/prescription error	16	30	46
Number of patients without a discrepancy/prescription error	25	14	39
Total	41	44	85
Relative risk (95% CI)	0.57 (0.37–0.88)		
Number of medications with a discrepancy/prescription error	25	62	87
Number of medications without a discrepancy/prescription error	382	363	745
Total	407	425	832
Relative risk (95% CI)	0.42 (0.27–0.66)		

Table 5 Classification and examples of discrepancies and prescription errors

	Example	Control, no (%)	Intervention, no (%)
Re-start of discontinued medication	Before hospitalization a patient was prescribed verapamil. During hospitalization this was stopped. After discharge he was still taking verapamil because “nobody told me to stop” ^C	9 (15)	2 (8)
Discontinuation of prescribed medication	Not filling a prescription for a loop diuretic, because “I already use a diuretic (hydrochlorothiazide)”. Readmission within 2 weeks ^C	6 (10)	6 (24)
Use of higher dose of medication	Discharge prescription and written information for patient: bumetanide 1 mg tablet, once a day 2 mg The pharmacy delivered tablets of 2 mg. The patient took 2 tablets of 2 mg instead of one ^C	7 (11)	2 (8)
Use of lower dose of medication	Discharge prescription: paracetamol 1,000 mg four times a day. Patient used 500 mg four times a day ^I	1 (2)	4 (16)
Use more frequent	Discharge prescription: lactulose if necessary; the label of the pharmacy prescribed twice a day ^C	2 (3)	–
Use less frequent	Discharge prescription: lactulose once a day; the patient took it if necessary ^I	6 (10)	2 (8)
Incorrect time of taking	Discharge prescription: calcium at bedtime due to interaction with ferrofumarate. Patient took both tablets at the same time ^I	1 (2)	2 (8)
Prescription error—dosage error	Discharge prescription: acenocoumarol 1 mg once a day. This should be: according to scheme ^{C, I}	30 (48)	7 (28)
Prescription error—dosage form error	Discharge prescription: thiamine injections. This should be tablets ^C		
Total		62 (100)	25 (100)

^C Control; ^I intervention

higher was 32% in the intervention group and 29% in the control group (see Table 6).

No difference was found in the estimate of adherence between both groups: 79.5% in the control group had a BMQ score ≥ 1 (potentially non-adherent) versus 78.0% in the intervention group (RR: 1.07 (95% CI 0.47–2.44)).

Discussion

This study has investigated the effect of a discharge service by a clinical pharmacist on the occurrence of discrepancies and prescription errors in a population of heart failure

patients. The percentage of patients with one or more discrepancies or prescription errors has been lowered by almost a half (68% vs. 39%).

International studies show that supervision of and providing heart failure patients with information at the time of discharge as well as postdischarge support reduces the number of readmissions and improves the patient's quality of life [21–23]. It was not possible in this study to measure the number of readmissions caused by incorrect medicine use because it was considered unethical to leave discrepancies uncorrected during the check up at the outpatients' clinic. However, by classifying discrepancies and prescription errors into classes of seriousness an estimate can

Table 6 Clinical relevance and examples of the discrepancies and prescription errors

NCC MERP index	Example	Control, no (%)	Intervention, no (%)
A	Discharge prescription: zopiclon at bedtime. On advise of the pharmacy this prescription stopped ¹	–	1 (4)
B	Discharge prescription: acenocoumarol 1 mg once a day. This should be: according to scheme ^{C, 1}	33 (53)	9 (36)
C	Before hospitalization a patient was prescribed nitroglycerin patch. During hospitalization this was stopped. After discharge she was still using nitroglycerin ^C	9 (15)	7 (28)
D	Discharge prescription: amiodarone 200 mg three times a day. This should be reduced to 200 mg once a day after a week ^C	2 (3)	–
E	Before hospitalization, a patient was prescribed lisinopril/hydrochlorothiazide. During hospitalization hydrochlorothiazide was stopped. After discharge the patient still used the combination tablet ^{C, 1}	15 (24)	7 (28)
F	Not filling a prescription for a loop diuretic, because “I already use a diuretic (hydrochlorothiazide)”. Readmission ^C	3 (5)	1 (4)
G		–	–
H		–	–
Total		62 (100)	25 (100)

^C Control; ¹ intervention

be made of the possible consequences. The percentage of discrepancies and prescription errors in class E or higher (error and harm) is similar in both groups but the absolute numbers decrease by half in the intervention group.

The percentage of patients with at least one discrepancy or prescription errors (68% vs. 39%) was consistent with other general reports describing medication discrepancies in 14.1–59.6% of patients at discharge [4, 24–26]. Although a variety of factors contribute to the occurrence of medication discrepancies, prescribers often fail to routinely compare a patient's inpatient medication list with his or her preadmission list at the time of prescribing and may not communicate medication information effectively at the time of discharge.

The fact that the percentage of patients who are potentially non-adherent is similar in both groups is not surprising. It is a known fact that a combination of interventions spread out over more than 3 weeks is needed to improve medication adherence [7, 27].

This study is limited in several aspects. First, medication discrepancies arising at the moment of admission were not included in the study. Results of the study by Bolas et al. [28] show that preparation of an accurate medication record at admission by a community liaison pharmacist reduces the number of these discrepancies. A combination of admission and discharge consultations could have led to a further decrease in the number of discrepancies.

Second, the interventions in this study were done by one clinical pharmacist in one hospital only, which limits the generalisability. Yet, the study is one of the few European studies that provides information on a clinical pharmacist discharge service outside the UK and the situation in this single hospital is likely to be similar to many other European

hospitals with respect to the limited number of clinical pharmacists. As the study focuses on a specific high risk group, these limited resources may be used in an effective way when implementing the intervention as described in this study.

Third, less patients were included in the study than the calculated group size. Nonetheless, a statistically significant effect of the intervention could be demonstrated. The results need to be interpreted in the light that the predetermined sample size was not achieved.

Finally, the BMQ only provides an indication for potential non-adherence. Therefore it is not possible to distinguish between good or poor levels of adherence.

Strengths of the study are its randomized design and the assessment of the discrepancies by a multidisciplinary team. Other studies often solely rely on the assessment by pharmacists and they may assess the clinical relevance of discrepancies different than doctors [29].

Future studies should be performed investigating clinical pharmacist discharge services in multicenter settings, preferably using readmissions as a clinical endpoint. Such studies should also pay attention to aspects as patient satisfaction and quality of life.

Conclusion

Information about (side)effects and changes in the drug therapy given by a clinical pharmacist combined with a written overview of the discharge medication and communication to both the community pharmacist and the GP reduces discrepancies and prescription errors within a population of heart failure patients. No effect on non-adherence was found in this short follow-up study.

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