Insomnia is associated to depressive symptoms in patients with chronic heart failure

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ABSTRACT

Introduction: Insomnia and depressive symptoms are common among patients with chronic heart failure (HF). Aim: The aim was to describe the prevalence of insomnia and depressive symptoms, as well as to examine the association between insomnia and depressive symptoms in patients with HF. Method: A cross-sectional descriptive study including 212 patients with HF. All patients responded to questionnaires regarding sleeping difficulties (Uppsala Sleep Inventory-Chronic Heart Failure), daytime sleepiness (Epworth Sleepiness Scale) and depressive symptoms (The Mental Health Scale). Results: Patients with depressive symptoms (34%) had, compared to those without, significantly more major complaints of insomnia as indicated by the sub-types; non-restorative sleep (66% vs. 32%, p < 0.0001), difficulty in maintaining sleep (35% vs. 16%, p = 0.003), difficulty in initiating sleep (29% vs. 14%, p = 0.009) and early morning awakenings (25% vs. 10%, p = 0.004). The odds ratio (OR) to suffer from depressive symptoms; were for; non-restorative sleep 5.2 (CI 95%, 2.2 - 12.3), difficulties maintaining sleep 2.5 (CI 95%, 1.2 - 4.9), difficulties in initiating sleep 2.2 (CI 95%, 1.1 - 4.4) and early morning awakenings 2.4 (CI 95%, 1.1 - 5.4). When categorising insomnia into three severity groups, 1) non insomnia, 2) mild insomnia, and 3) severe insomnia, the OR for depressive symptoms for the mild insomnia and severe insomnia group were 2.2 (CI 95%, 1.1 - 4.2) and 7.4 (CI 95%, 2.4 - 22.8) respectively, compared to the non insomnia group. Conclusion: Insomnia is independently associated to depressive symptoms. Assessment of depressive symptoms and insomnia in patients with HF is important since treatment could be targeted to depressive symptoms only and/or to the sleep disturbance.

Keywords: Congestive Heart Failure; Insomnia

Disorders; Depressive Symptoms

1. INTRODUCTION

Disturbed sleep measured objectively, as well as subjectively, has been recognized as a common problem among patients with HF [1-3]. One study found that 36% of 223 patients with HF reported not getting enough sleep and 21% reported excessive daytime sleepiness [2]. Both the quality, as well as the continuity of sleep are important since these independently are associated to functional performance and to the physical and mental quality of life [4]. Reasons for this may be related to insomnia [3]. Insomnia is according to the Diagnostic and Statistical Manual of Mental Disorders described as a complaint of difficulty initiating sleep, difficulty maintaining sleep and/or a non-restorative sleep [5]. Primary insomnia means that the disturbance is due to unknown causes while secondary insomnia suggests medical, psychiatric or abuse being the causes for the disturbance [5]. Many studies also include early morning awakenings in the definition of insomnia [6]. As much as 20% - 50% of patients with HF may suffer from insomnia [1,2,7], and complaints of difficulties maintaining sleep or insufficient sleep have been found to be the two most commonly reported disturbances [1,2]. Compared to non-insomniac HF patients, as well as to a normal population, HF patients complaining of insomnia also rate a poorer quality of life [2].

In HF secondary insomnia can be caused by physical aspect such as dyspnoea, palpitations or nocturia [8]. Pain or sleep disordered breathing can be other causes [9], however, the relationship between insomnia and sleep disordered breathing is unclear, since some studies report an association [10] whereas others not [11,12]. Depression can be a psychiatric cause for insomnia [13]. The diagnosis of depression (Table 1) is made upon the presence of at least one of the two core symptoms, i.e.
Table 1. Core and secondary symptoms of depression according to American psychiatric association [5].

<table>
<thead>
<tr>
<th>Core symptoms</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Depressed, sad mood most of the day.</td>
<td></td>
</tr>
<tr>
<td>Markedly diminished interest or pleasure in all or almost all activities</td>
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</tr>
<tr>
<td>nearly every day.</td>
<td></td>
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</tbody>
</table>

<table>
<thead>
<tr>
<th>Secondary symptoms</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Weight loss or weight gain, or decrease or increase in appetite.</td>
<td></td>
</tr>
<tr>
<td>Insomnia or hypersomnia.</td>
<td></td>
</tr>
<tr>
<td>Psychomotor agitation/retardation.</td>
<td></td>
</tr>
<tr>
<td>Fatigue or loss of energy.</td>
<td></td>
</tr>
<tr>
<td>Feelings of worthlessness or inappropriate guilt.</td>
<td></td>
</tr>
<tr>
<td>Diminished ability to think or concentrate or ambivalence.</td>
<td></td>
</tr>
<tr>
<td>Thoughts of death or suicidal ideation.</td>
<td></td>
</tr>
</tbody>
</table>

sadness and loss of pleasure (anhedonic symptoms), together with at least four or more secondary symptoms [5]. Studies in aged non HF populations have shown mood disturbances to be one of the most important factors associated with sleep disturbances [14]. The link between sleep disturbances and depression is however complicated. Sleep disturbances can in some cases be a precursor to depression and in other cases, a consequence of depression [13]. Although data indicate that approximately 40% of the HF population might suffer from depression or depressive symptoms [15], only one study to our knowledge have investigated the association between insomnia and depression in patients with HF. Redeker et al. [16] reported that patients with difficulties initiating sleep and/or difficulties maintaining sleep were five time more likely to be depressed. A deeper knowledge of the associations between insomnia and depressive symptoms is needed to design patient centred interventions to improve sleep and psychological wellbeing in patients with HF. The aim of this study was therefore to describe the prevalence of insomnia and depressive symptoms, as well as to examine the association between insomnia and depressive symptoms in patients with HF.

2. METHODS

2.1. Design and Setting

This study used a cross-sectional descriptive design. Patients were recruited from one medical ward (in a university hospital), two heart failure clinics (one university and one county hospital) and five primary health care centres in the southeast of Sweden.

2.2. Study Population

The template is used to format your paper and style the text. All margins, column widths, line spaces, and text fonts are prescribed; please do not alter them. You may note peculiarities. For example, the head margin in this template measures proportionately more than is customary. This measurement and others are deliberate, using specifications that anticipate your paper as one part of the entire journals, and not as an independent document. Please do not revise any of the current designations.

2.3. Uppsala Sleep Inventory-Chronic Heart Failure

Insomnia was measured with the well-validated Uppsala Sleep Inventory-Chronic Heart Failure (USI-CHF) [2, 17]. The USI-CHF consists of 26 items of which four focuses on insomnia (i.e., difficulty initiating sleep, difficulty maintaining sleep, non-restorative sleep and early morning awakenings). The items about insomnia are to be answered on a five-point likert-type scale; from no problems (1), small problems (2), some problems (3), great problems (4), to very great problems (5). In the statistical processing and analysis of the insomnia variables of the USI-CHF scores of 1 - 3 were considered to represent “no complaints” and scores 4 and 5 “major complaints”.

2.4. Epworth Sleepiness Scale

Excessive daytime sleepiness was measured with the well-validated Epworth Sleepiness Scale (ESS) [18,19]. The 8 items focuses the expectation of “dozing” in a variety of situations. The probability ratings are zero (0), slight (1), moderate (2), or high (3) in the hypothetical situations. The ratings can be summarized to a total score of 24, with a cut-off value of >10 indicating excessive daytime sleepiness.

2.5. Mental Health Scale

In the present study depressive symptoms was measured by the Mental Health Scale (MH-5), from the 36-Item Short Form Health Survey [20]. The 5 items focuses the anhedonic aspect of depression (i.e., sadness and loss of pleasure) and the scores can range from 0 (the worst) to 100 (best mental health). With a cut-off point of ≤60 the MH-5 seems valid and reliable to use in the screening for depressive symptoms indicating a high probability of a diagnosis of major depression [21-23]. The MH-5 can be said to assess a depressive symptomatology which might indicate the presence of different types of depressive disorders such as subsyndromal or minor depression, dysthymia or major depression [15]. Due to this the term
depressive symptoms is used in this study. Hence, participants with a score ≤60 on the MH-5 scale were therefore labeled as having depressive symptoms and those with a score >60 were labeled as having no depressive symptoms.

2.6. Data Collection

HF nurses and primary health care nurses acted as coordinators at each center. Eligible patients with HF were found through a screening process on wards and heart failure clinics at the two hospitals and through the Diagnosed Related Groups register at each of the five primary health care centers. The medical records were scrutinized by an experienced heart failure nurse, together with one of the nurse-coordinators and a physician. All participants gave their informed written consent. The study protocol was approved by the ethics committee at the faculty of health sciences, University of Linköping, Sweden, and is in accordance with the provisions of the Helsinki declaration.

2.7. Statistical Processing and Analysis

Descriptive statistics were used to describe differences between the HF population with and without depressive symptoms. Categorical variables were analyzed with the Chi-square test. Continuous variables, if normally distributed, were analyzed with the Student’s t-test (2-tailed) and presented as means and standard deviation (SD). The Mann-Whitney non-parametric test were used on skewed continuous variables, and presented as the median (md) and the 25th 75th percentiles. To determine the interrelationship between difficulties initiating sleep, difficulties maintaining sleep, early morning awakenings and non restorative sleep a Spearman’s rank correlation analysis was done. Binary regression modeling was used to analyze the independent relation between depressive symptoms and the insomnia variables. The model was adjusted for covariates that in Table 2 had a univariate p value < 0.15 (i.e. NYHA class). The significance level was set at 0.05. Statistical computing was done with the SPSS 18.0.

Table 2. Characteristics of HF patients (n = 212) and univariate differences between patients with and without DS.

<table>
<thead>
<tr>
<th></th>
<th>Total group</th>
<th>No DS n = 140</th>
<th>DS n = 72</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sex, male, n (%)</td>
<td>129 (61)</td>
<td>87 (41)</td>
<td>42 (20)</td>
<td>0.59</td>
</tr>
<tr>
<td>Age, mean, (SD)</td>
<td>76 ± 9</td>
<td>76 ± 8</td>
<td>75 ± 11</td>
<td>0.40</td>
</tr>
<tr>
<td>Living with a partner, n (%)</td>
<td>89 (42)</td>
<td>55 (39)</td>
<td>34 (38)</td>
<td>0.27</td>
</tr>
<tr>
<td>MH-5 score, md, (Q25 - Q75)</td>
<td>76 (56 - 88)</td>
<td>84 (76 - 92)</td>
<td>52 (40 - 56)</td>
<td>0.001</td>
</tr>
<tr>
<td>NYHA-class</td>
<td></td>
<td></td>
<td></td>
<td>0.006</td>
</tr>
<tr>
<td>II, n (%)</td>
<td>86 (40)</td>
<td>65 (46)</td>
<td>21 (29)</td>
<td></td>
</tr>
<tr>
<td>III, n (%)</td>
<td>114 (54)</td>
<td>70 (50)</td>
<td>44 (61)</td>
<td></td>
</tr>
<tr>
<td>IV, n (%)</td>
<td>12 (6)</td>
<td>5 (4)</td>
<td>7 (10)</td>
<td></td>
</tr>
<tr>
<td>Aetiology</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>IHD, n (%)</td>
<td>94 (44)</td>
<td>61 (44)</td>
<td>33 (46)</td>
<td>0.75</td>
</tr>
<tr>
<td>MI, n (%)</td>
<td>87 (41)</td>
<td>56 (40)</td>
<td>31 (43)</td>
<td>0.67</td>
</tr>
<tr>
<td>HT, n (%)</td>
<td>73 (34)</td>
<td>50 (36)</td>
<td>23 (32)</td>
<td>0.58</td>
</tr>
<tr>
<td>Diabetes, n (%)</td>
<td>53 (25)</td>
<td>36 (26)</td>
<td>17 (24)</td>
<td>0.74</td>
</tr>
<tr>
<td>Other, n (%)</td>
<td>29 (14)</td>
<td>19 (14)</td>
<td>10 (14)</td>
<td>0.95</td>
</tr>
<tr>
<td>Blood pressure</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>SBP, mean, SD</td>
<td>131 ± 22</td>
<td>133 ± 22</td>
<td>127 ± 21</td>
<td>0.04</td>
</tr>
<tr>
<td>DBP, mean, SD</td>
<td>71 ± 13</td>
<td>72 ± 12</td>
<td>70 ± 13</td>
<td>0.31</td>
</tr>
<tr>
<td>Sleep variables</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>DIS, n (%)</td>
<td>41 (19)</td>
<td>20 (14)</td>
<td>21 (29)</td>
<td>0.009</td>
</tr>
<tr>
<td>DMS, n (%)</td>
<td>48 (23)</td>
<td>23 (16)</td>
<td>25 (35)</td>
<td>0.003</td>
</tr>
<tr>
<td>EMA, n (%)</td>
<td>32 (15)</td>
<td>14 (10)</td>
<td>18 (25)</td>
<td>0.004</td>
</tr>
<tr>
<td>NRS, n (%)</td>
<td>93 (44)</td>
<td>45 (32)</td>
<td>48 (66)</td>
<td>0.0001</td>
</tr>
<tr>
<td>EDS, n (%)</td>
<td>46 (22)</td>
<td>27 (19)</td>
<td>19 (26)</td>
<td>0.23</td>
</tr>
</tbody>
</table>

IHD—Ischaemic Heart Disease; MI—Myocardial Infarction; HT—Hypertension; SBP—Systolic Blood Pressure; DBP—Diastolic Blood Pressure; DIS—Difficulty Initiating Sleep; DMS—Difficulty Maintaining Sleep; EMA—Early Morning Awakenings; NRS—Non Restorative Sleep; EDS—Excessive Daytime Sleepiness.
3. RESULTS

3.1. Participants

The mean age of the patients was 76 years, 61% males and 42% lived with a partner (Table 2). A total of 94% were in NYHA classes II-III. Ischaemic heart disease (44%) mostly explained by myocardial infarction (41%) and hypertension (34%) were the most common causes of HF (Table 2). A total of 82% of the patients had their diagnosis of HF based on radiographic evidence and/or echocardiography.

3.2. Prevalence of Depressive Symptoms

Of the patients 34% (n = 72) suffered from depressive symptoms. Regarding age, partnership, sex and aetiology of HF, there were no significant differences between those with and without depressive symptoms (Table 2). A higher proportion of the non-depressed patients were in NYHA class II and a higher proportion of those with depressive symptoms were in NYHA class III (p = 0.006). Those with depressive symptoms had a lower systolic blood pressure (p = 0.04) compared to those without depressive symptoms (Table 2).

3.3. Insomnia, Excessive Daytime Sleepiness and Depressive Symptoms

The most common insomnia type was non restorative sleep, which was reported as a major complaint by 44% of the patients. Major complaints of difficulties maintaining sleep, difficulties initiating sleep and early morning awakenings were reported by 23%, 19% and 15% of the patients, respectively. Patients with depressive symptoms had significantly more major complaints of non restorative sleep (p < 0.0001), difficulties maintaining sleep (p = 0.003), difficulties initiating sleep (p = 0.009) and early morning awakenings (p = 0.004) than to those without depressive symptoms (Table 2). Fully one fifth (22%) reported excessive daytime sleepiness. There was no significant difference in excessive daytime sleepiness reported between the patients with and without depressive symptoms (Table 2).

3.4. Association between Insomnia and Depressive Symptoms

To examine which of the insomnia variables that independently was associated to depressive symptoms, all variables were included in the same regression model. However in this analysis none of the insomnia variables became significant. This can probably be explained because of an interrelationship between the insomnia variables (Table 3). Therefore separate models including the insomnia variables were calculated with adjustments for NYHA classes, with NYHA class II as the referent. In the unadjusted model all insomnia variables had a significant relation to depressive symptoms. Major complaints of non restorative sleep more than five fold the OR to have depressive symptoms. The OR to have depressive symptoms for major complaints of early morning awakenings almost increased by three fold, while the OR for major complaints of difficulties initiating sleep and difficulties maintaining sleep increased more than two fold respectively (Table 4). After adjustment for NYHA class, all insomnia variables were still significantly related to depressive symptoms. In the adjusted model major complaints of non restorative sleep more than five fold the odds to suffer from depressive symptoms. Reports of major complaints of early morning awakenings, difficulties initiating sleep and difficulties maintaining sleep respectively more than doubled the odds for having depressive symptoms. However, NYHA class by itself had a significant association to depressive symptoms only in the models for difficulties initiating sleep and difficulties maintaining sleep (Table 4).

Since many patients suffer from more than one type of insomnia, we studied the relationship between the amount of insomnia subtypes and depressive symptoms (i.e., the higher degree of insomnia the higher proportion of patients with depressive symptoms). In this model insomnia was categorized into three groups; the first group included those who did not report to suffer from a major complaint in any of the insomnia groups (non insomnia), the second group (mild insomnia) included those that had a major complaint in one or two of the insomnia groups; and the third group (severe insomnia) consisted of those who had a major complaint in three or four of the insomnia groups. The percentages of patients with depressive symptoms increased proportionally from 24% in the no insomnia group to 74% in the severe insomnia group (Figure 1). Compared to the non insomnia group, the patients with mild and severe insomnia had more than two fold and seven fold increased risk respectively to suffer from depressive symptoms. In the adjusted model,

Table 3. Interrelationship between the insomnia subtypes in patients with HF:

<table>
<thead>
<tr>
<th></th>
<th>DIS</th>
<th>DMS</th>
<th>EMA</th>
<th>NRS</th>
</tr>
</thead>
<tbody>
<tr>
<td>DIS</td>
<td>-</td>
<td>0.4**</td>
<td>0.260**</td>
<td>0.349**</td>
</tr>
<tr>
<td>DMS</td>
<td>0.449**</td>
<td>-</td>
<td>0.338**</td>
<td>0.395**</td>
</tr>
<tr>
<td>EMA</td>
<td>0.260**</td>
<td>0.338**</td>
<td>-</td>
<td>0.320**</td>
</tr>
<tr>
<td>NRS</td>
<td>0.349**</td>
<td>0.395**</td>
<td>0.320**</td>
<td>-</td>
</tr>
</tbody>
</table>

DIS—Difficulty Initiating Sleep; DMS—Difficulty Maintaining Sleep; EMA—Early Morning Awakenings; NRS—Non Restorative Sleep; EDS—Excessive Daytime Sleepiness. **p < 0.01.
Table 4. Odds ratios (OR) for the association between the insomnia variables and depressive symptoms (DS) in patients with HF.

<table>
<thead>
<tr>
<th></th>
<th>Unadjusted OR (CI 95%)</th>
<th>p</th>
<th>Adjusted OR (CI 95%)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>DIS</td>
<td>2.5 (1.2 - 4.9)</td>
<td>0.011</td>
<td>2.2 (1.1 - 4.4)</td>
<td>0.03</td>
</tr>
<tr>
<td>NYHA</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>II</td>
<td>ref</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>III</td>
<td>1.7 (0.9 - 3.2)</td>
<td>0.09</td>
<td></td>
<td></td>
</tr>
<tr>
<td>IV</td>
<td>3.9 (1.1 - 13.9)</td>
<td>0.034</td>
<td></td>
<td></td>
</tr>
<tr>
<td>DMS</td>
<td>2.7 (1.4 - 5.2)</td>
<td>0.003</td>
<td>2.5 (1.2 - 4.9)</td>
<td>0.007</td>
</tr>
<tr>
<td>NYHA</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>II</td>
<td>ref</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>III</td>
<td>1.7 (0.9 - 3.3)</td>
<td>0.08</td>
<td></td>
<td></td>
</tr>
<tr>
<td>IV</td>
<td>4.1 (1.1 - 14.7)</td>
<td>0.02</td>
<td></td>
<td></td>
</tr>
<tr>
<td>EMA</td>
<td>2.9 (1.3 - 6.4)</td>
<td>0.005</td>
<td>2.4 (1.1 - 5.3)</td>
<td>0.03</td>
</tr>
<tr>
<td>NYHA</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>II</td>
<td>ref</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>III</td>
<td>1.8 (0.9 - 3.3)</td>
<td>0.07</td>
<td></td>
<td></td>
</tr>
<tr>
<td>IV</td>
<td>3.05 (0.2 - 11.2)</td>
<td>0.09</td>
<td></td>
<td></td>
</tr>
<tr>
<td>NRS</td>
<td>5.9 (2.5 - 13.9)</td>
<td>0.001</td>
<td>5.2 (2.2 - 12.3)</td>
<td>0.001</td>
</tr>
</tbody>
</table>

DIS—Difficulty Initiating Sleep; DMS—Difficulty Maintaining Sleep; EMA—Early Morning Awakenings; NRS—Non Restorative Sleep; EDS—Excessive Daytime Sleepiness.

NYHA class did not have any significant relation to depressive symptoms (Table 5).

4. DISCUSSION

The main findings in this study were that one third of the HF patients suffered from depressive symptoms. Insomnia was common, with non restorative sleep reported by 32% of those without and by 66% of those with depressive symptoms. The association between insomnia and occurrence of depressive symptoms was strong. The OR for HF patients with severe insomnia to suffer from depressive symptoms was more than seven folded.

4.1. Prevalence of Depressive Symptom and Insomnia

The prevalence of depressive symptoms in this study was 34%, which is comparable with the reported median prevalence of 42% found in studies that have used dimensional depression screening tools [15]. Of the patients in the present study 44% complained of non restorative sleep, while 23%, 19% and 15% of the patients complained of difficulties maintaining sleep, difficulties initiating sleep and early morning awakenings, respectively. Patients with depressive symptoms also reported more insomnia. Principe-Rodriguez et al. [7] found that 31% of the HF patients presented symptoms of insomnia, but the different insomnia types were not described. Another study found 37%, 35% and 36% of the patients reporting difficulties initiating sleep, difficulties maintaining sleep and early morning awakenings [1]. This is higher rates than in our study. However, the 44% who

Table 5. Odds ratios (OR) for the association between severity of insomnia and depressive symptoms (DS).

<table>
<thead>
<tr>
<th>Insomnia</th>
<th>Unadjusted OR (CI 95%)</th>
<th>p</th>
<th>Adjusted OR (CI 95%)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>*Non insomnia, (n = 130)</td>
<td>ref</td>
<td></td>
<td>ref</td>
<td></td>
</tr>
<tr>
<td>**Mild insomnia, (n = 62)</td>
<td>2.5 (1.3 - 4.7)</td>
<td>0.006</td>
<td>2.2 (1.1 - 4.2)</td>
<td>0.01</td>
</tr>
<tr>
<td>***Severe insomnia, (n = 19)</td>
<td>8.9 (2.9 - 26.8)</td>
<td>0.001</td>
<td>7.4 (2.4 - 22.8)</td>
<td>0.001</td>
</tr>
<tr>
<td>NYHA</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>II</td>
<td>ref</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>III</td>
<td>1.5 (0.8 - 2.9)</td>
<td>0.19</td>
<td></td>
<td></td>
</tr>
<tr>
<td>IV</td>
<td>2.6 (0.7 - 10.0)</td>
<td>0.15</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

*The group includes patients not suffering of insomnia (i.e. DIS, DMS, EMA, NRS); **The group includes patients suffering from major complaint in one or two of the insomnia variables; ***The group includes patients suffering from major complaint of three or all of the insomnia variables.
reported non restorative sleep in our study are at the same level as those who reported that they felt tired or fatigued in the morning at least 3 - 4 times/week in Prin-
cipe-Rodriguez et al. [7], and the 45% who reported in-
sufficient sleep in the study of Redeker and Stein [1]. In
total this indicates, that about 15% - 30% of the patients
with HF suffer from insomnia and that almost 50% still
are sleepy and fatigued when they wake up in the morn-
ing. The high prevalence of insomnia found also suggests
that a large proportion of these patients also are at risk
for suffering from depressive symptoms.

4.2. The Association between Depressive
Symptoms and Insomnia

We found a strong association between insomnia and
depressive symptoms. The OR to suffer from depressive
symptoms for patients complaining of difficulties initiat-
ing sleep, difficulties maintaining sleep and early morn-
ing awakenings increased by more than two fold. These
rates corresponds almost the figures found by Leesman-
Leege et al. [24] who reported the OR between sleep
disturbance and depressive symptoms to be three folded.
However, in the study of Redeker et al. [1] the OR be-
tween difficulties initiating sleep and/or difficulties
maintaining sleep and early morning awakenings was five
folded [16]. In our study a five folded increased risk was
found for the association between non restorative sleep
depressive symptoms. One explanation for this may
be explained by that fact that non restorative sleep might
include feelings of fatigue and tiredness, which both are
symptoms of HF without the co-existence of a sleep dis-
turbance. However, NYHA class, a measure of functional
capacity, had no significant relation to depressive symp-
toms in the separate non restorative sleep model. More-
over the relationships between insomnia and depressive
symptoms persisted after that NYHA class was taken
into consideration. This corresponds to the study of Re-
derker et al. [16] who also reported the relationship be-
tween depression and insomnia to persist after adjust-
ments for clinical variables such as NYHA class and
ejection fraction. In another study patient’s perception of
their sleep explained 30% of the variance in mental
health compared to the 15% explained by NYHA-class,
age, gender and co-morbidity [25]. Hence, insomnia in
HF patients may not only be a marker for severity of HF.
Furthermore, in the present study we also analysed the
association between severity of insomnia and depressive
symptoms. Those with severe insomnia had a seven fold
increased risk to suffer from depressive symptoms. This
has not before been reported in patients with HF high-
lights the importance of interventions such as cognitive
behavioural therapy (CBT) that can be used to target
both insomnia and depressive symptoms. In older adults
with arthritis and insomnia, CBT improved sleep as well
as pain [26]. In patients with HF, CBT has been found to
improve depressive symptoms and health-related quality
of life [27,28]. However, the numbers of CBT studies
performed on patients with HF are few and none have
examined of the effects on sleep disturbances and de-
pressive symptoms.

In future studies it is of importance to consider that
there are several different factors that can cause difficul-
ties initiating sleep, difficulties maintain sleep and early
morning awakenings (i.e. secondary insomnia). Theo-
retically, insomnia can be described as the combination
of sleep interfering and sleep interpreting processes [29].
The sleep interfering processes include different types of
arousals (i.e., physiological, emotional and cognitive). A
physiological arousal may be explained by an elevated
baseline level of the stress-response system, as well as a
slow return to the baseline level after being aroused. Emo-
tional and cognitive arousals may be explained by a
chronic tendency to worry, an elevated response to
stressful events, problems, or increased dysphoric mood.
The sleep interpreting processes imply that the insomnia-
acs’ perceptions of their own sleep are misinterpreted
[29]. Sleep interfering processes leading to negative
thoughts and anxiety, i.e., emotional and cognitive arousal
in patients with HF can be can be associated with prob-
lems to perform activities in daily life, thoughts about HF
including deterioration, medication and an unknown life
situation [8,16]. Dysrhythmia, coughing and dyspnoea or
have also been described causing a sense of panic during
the night. Nocturia or inability to sleep in a supine posi-
tion can also cause nightly awakening and be possible
sleep interfering processes [8,30].

However, not all patients with a disturbed sleep will
develop insomnia and/or depressive symptoms. In an
elderly non HF population factors describing a psycho-
logical vulnerability (i.e., somatisation, intrusive and in-
resistible thoughts and worries) were associated to in-
somnia [31]. In another study of non-HF patients showed
that insomniacs and good sleepers did not differ in the
frequency of experienced daily stressors. The difference
was in how they coped and perceived the stressors in
daily life [32]. Poor psychological functioning in elderly
people with and without HF has been found to mediate
the association between insomnia and health related
quality of life [33]. Thus, an HF patient who complains
of insomnia should therefore be carefully evaluated for
presence of depressive symptoms with common and
validated specific instruments for insomnia and depres-
sion [15,34,35]. Treatment of insomnia in clinical prac-
tice often includes sleep-medications. However, in eld-
ery with or without HF sleep-medications should be used
carefully. This is because of the risk for polypharmacy
and side-effects, such as sedation, drowsiness or somno-

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lence [36]. Moreover, sleep medications have in elderly, independent of insomnia, been found to cause depression [37]. CBT (e.g., reduction of dysfunctional believes about sleep) can therefore be preferable. CBT studies in HF populations are missing but in other populations face to face CBT interventions have been successfully [38,39]. In a cardiovascular setting the accessibility of face to face CBT is limited due to lack of competence in CBT among the personnel. CBT provided by the internet has shown promising results to reduce insomnia as well as depressive symptoms [40-42].

5. STUDY LIMITATIONS

Because this study had a cross-sectional design it is not possible to determine causal relationship between insomnia and depressive symptoms. Identification of depression can be made by categorical and/or dimensional instruments. Categorical instrument can be used by a trained interviewer to decide whether the symptom profile corresponds to an inclusion or exclusion of a diagnosis of depression. In contrast, dimensional instruments are not diagnostic tools. These are based on self-reports and ranks the patients symptom profile on a continuum, and if above a certain threshold, there is a high probability for a diagnosis of depression [15]. The use of a dimensional depression screening tool (i.e., the MH-5) does not allow us to assess a clinical diagnosis of depression. However dimensional depression screening tools are widely accepted and frequently used [15,43]. Another problem with dimensional depression screening tools is that these might indicate false depressed cases due to the fact that some symptoms such as loss of appetite, fatigue and sleeplessness occur in depression as well as in HF. However, the MH-5 measures the anhedonic symptoms of depression only [20], which might reduce the problem with a false strong relationship between insomnia and depressive symptoms in the present study. The ESS is a self-assessment tool in which the individual evaluates his/her expectation of “dozing” in a variety of situations. A problem can be individual variations in relation to age, gender, habits, or other aspects that affects napping. However, validation studies using differential item functioning (i.e., analyses focusing on if a subject’s response to the item is affected by other aspects than that which the test is intended to assess) have found only minor effects of age in three of the items included in the ESS. The mentioned items were, inactive in a public place (item 3), passenger in a car (item 4), and item 8 focusing on “dozing” when a car has stopped in traffic [44].

Another limitation is that at the time of inclusion data regarding existence and/or treatment of SDB, as well as if the patients were prescribed antidepressant or hypnotic medication were not collected.

6. CONCLUSION

This study found that insomnia as indicated by difficulties initiating sleep, difficulties maintaining sleep, early morning awakenings and non restorative sleep was independently associated to depressive symptoms in patients with HF. The study also revealed that the higher severity of insomnia, the higher was the risk of the patient to suffer from depressive symptoms. The relation between insomnia and depressive symptoms is complex and probably bi-directional. Assessment of sleep disturbances and depressive symptoms in patients with HF is therefore important, since treatment could be targeted to depressive symptoms only and/or to the sleep disturbances. Therefore more research is needed to study the relation between as well as the treatment of sleep disturbances and depressive symptoms in patients with HF.

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