

Impact on the survival in occasional clusters of symptoms with coronary heart failure

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Introduction

Coronary heart failure (HF) is a serious fitness trouble international. Within the U.S.A. On my own, HF afflicts round 5.7 million humans. Bodily and emotional signs are a defining feature of HF. Patients with HF have stated experiencing an average of 15 physical and emotional signs. Incidence of a couple of signs adversely have an effect on patient lives by using growing mental distress and lowering first-rate of lifestyles.

Up to now, most researchers have focused on symptoms in isolation. But, sufferers with HF commonly revel in multiple symptoms at a time. The effect of multiple, concurrent signs and symptoms on outcomes may be multiplicative. Patients with more signs or a more degree of symptom burden have worse outcomes. Fatigue severity mediated the affiliation between pain and performance popularity in sufferers with most cancers. Accordingly, the investigation of multiple symptoms as an interconnected enjoy is important to acquire a greater whole photograph of affected person symptom stories.

The consideration of symptom clusters, described as two or extra interrelated signs and symptoms going on together offers a singular technique to symptom assessment and control. Patient's capacity to apprehend a exchange in situation that calls for urgent attention from their healthcare carriers may be facilitated by means of expertise about symptom clusters. Identifying the profiles of sufferers who're at risk for worse results based totally on symptom clusters may also help the development and shipping of powerful, individualized techniques for specific groups of patients.

Gender may play a position within the HF symptom enjoy due to the fact HF characteristics, which include etiology and occurrence of HF symptoms, vary among males and females. Ladies generally tend to offer with greater symptoms and signs and symptoms associated with HF (e.G., dyspnea, fatigue, decrease extremity edema, third coronary heart sound, jugular venous distension, and

rales) than guys. A few symptoms, together with fatigue, despair, and anxiety, are mentioned to be more standard and intense in girls than men. In addition, symptoms are a crucial determinant of outcomes, which includes mortality and pleasant of lifestyles. Given that women with HF tend to have worse consequences in comparison to men, it's far feasible that symptom experiences are one-of-a-kind among ladies and men. Consequently, evaluation of signs and symptoms between genders may also provide valuable facts for tailoring interventions for patients with HF.

The motive of this examine was to compare symptom clusters among men and women with HF, differences in affected person characteristics among symptom clusters, and the impact of those symptom clusters on outcomes. The particular targets have been to: (1) decide whether different symptom clusters had been found in ladies and men with HF, (2) compare patient traits of companies within symptom clusters, and (3) examine the effect of symptom clusters on occasion-unfastened survival defined as time to first cardiac occasion

Techniques

The records for this study were compiled from 3 potential, longitudinal studies that had similar inclusion and exclusion standards. Institutional evaluation Board approval turned into received for every take a look at. Eligibility of sufferers who have been stated the investigators through nurses and physicians at each site was showed by way of the investigators or educated research nurses the use of scientific document assessment. Sufferers who agreed to participate in the examine provided written informed consent and finished the questionnaire packets during a visit to the general medical research center. Sufferers had been observed for a mean of 361 days to acquire cardiac event-loose survival records.

Patients

A complete of 331 patients had been protected in this examine. Patients have been recruited from HF outpatient clinics related to six big community hospitals or instructional clinical centers in Kentucky, Georgia, and Indiana. Sufferers had been blanketed who: (1) had a showed analysis of HF with either preserved or non-preserved systolic feature, (2) have been receiving choicest scientific therapy, (three)

have been capable of study and talk English, and (4) had no apparent cognitive impairment that prevented completing the questionnaire packets and interview with research nurses. Patients had been excluded in the event that they had: (1) valvular heart ailment as an etiology of their HF, (2) a myocardial infarction in the previous three months, (three) been referred for coronary heart transplantation, or (4) important existence-threatening comorbidities along with give up-level renal or liver ailment. Exclusion criteria for this study have been decided on because HF attributable to valvular coronary heart sickness can be correctable not like HF from other etiologies. Further, sufferers who had a myocardial infarction recently, been referred for coronary heart transplantation or severe comorbidities are much more likely to be hemodynamically volatile and feature a shorter lifestyles expectancy.

Measures

Heart failure signs and symptoms

Symptoms were diagnosed using the Minnesota dwelling with coronary heart Failure Questionnaire (MLHF), which consists of 21 items assessing fitness-related great of existence in patients with HF. The device contains the following 8 objects measuring misery from HF-related symptoms which can be concept to persuade health-associated great of lifestyles: edema, dyspnea, fatigue/multiplied want to rest, fatigue/low power, sleep disturbances, traumatic, feeling depressed, and cognitive problems (issue concentrating or remembering things). Sufferers rated each item on a scale from zero (no misery) to 5 (very severe distress). On this examine the internal consistency of these 8 gadgets was proven by a Cronbach's alpha of 0.89.

Demographic and medical characteristics.

Demographic and scientific traits (e.g., age, gender, ethnicity, and frame mass index [BMI]) had been obtained the use of a demographic and scientific questionnaire. Total comorbidity scores have been received from the Charlson Comorbidity Index. The rankings of the Charlson Comorbidity Index have been weighted through taking into account the variety and seriousness of comorbid illnesses. The big apple heart association (NYHA) functional category turned into determined via educated research nurses through in-depth established patient interviews

Cardiac occasion-unfastened survival.

Cardiac occasion-unfastened survival became defined as time to first cardiac occasion which blanketed loss of life, rehospitalization, or ED visit due to cardiac reasons. The records had been received by way of month-to-month follow-up calls to patients or circle of relatives and by means of administrative assessment of medical statistics and public demise records.

Statistical Analyses

Descriptive information together with frequency distributions, manner, preferred deviations, and degrees were used to explain the demographic and scientific characteristics of sufferers. Cluster evaluation become used to perceive symptom clusters due to the fact this technique maximizes the homogeneity of variables within clusters at the same time as concurrently maximizing the heterogeneity between clusters. We used the hierarchical cluster agglomerative approach which begins with treating every variable as a separate cluster and then combines the variables into consecutively large clusters based totally on their similarity.

The Euclidean distance became used to measure the similarity of variables. Proximity between corporations of variables was measured the use of Ward's technique through which clusters have been joined by using minimizing the overall within-cluster blunders sum of squares. Ward's method was selected due to the fact it's far sensitive to outliers and powerful while figuring out clusters in comparison to other inter-group proximity measures.

The resulting clusters have been pictorialized with dendrograms, which illustrate the proximity of variables to each other. Semi-partial R-squared scores have been used to determine the diploma of homogeneity of variables within the clusters, with larger values reflecting much less similarity among clusters. To decide the most excellent range of clusters we used dendrograms, the pseudo-F statistic, and the pseudo-T squared statistic. To illustrate the validity of the diagnosed range of clusters, principal factor analysis become performed and the first and second essential thing ratings were plotted.

Based at the diagnosed symptom clusters, patients were divided in companies through the median

split of general scores of each symptom cluster, which were calculated with the aid of summing distress rankings of signs and symptoms in each cluster. Evaluation of variance (ANOVA) or the chi-square take a look at turned into used to examine differences in demographic and clinical characteristics amongst 4 affected person groups (i.g., low distress, physical misery, emotional/cognitive distress, and excessive distress). Publish hoc analysis turned into finished the usage of the Bonferroni adjustment. Hierarchical Cox proportional risks regression turned into used to determine whether general ratings of symptom misery inside symptom clusters expected time to first cardiac occasion (i.E., loss of life, rehospitalization, or ED go to due to cardiac reasons) after controlling for age, gender, overall comorbidity scores, BMI, and NYHA functional elegance.

Result

Sample traits

The suggest age of sufferers become 61years with various 24 to 87 years (table 1).

Characteristics	Total	Male (n=216)	Female (n=115)	p-value
	Mean (\pm S.D) or N (%)			
Age, years	61 (\pm 11)	60 (\pm 11)	62 (\pm 11)	0.233
Ethnicity				<0.001
Caucasian	269 (81%)	188 (87%)	81 (70%)	<0.001
Marital status				
Married/ cohabitate	181 (55%)	141 (65%)	40 (35%)	
Single/divorced/ widowed	150 (45%)	75 (35%)	75 (65%)	
Ischemic etiology of HF	179 (54%)	137 (65%)	42 (38%)	<0.001
NYHA class				0.203
I/II	128 (39%)	91 (42%)	37 (32%)	
III	145 (44%)	90 (42%)	55 (48%)	
IV	58 (18%)	35 (16%)	23 (20%)	
Body mass index (kg/m ²)	32 (\pm 7)	31 (\pm 7)	32 (\pm 8)	0.548
Charlson comorbidity index	3.4 (\pm 2.0)	3.4 (\pm 2.0)	3.4 (\pm 2.0)	0.837
ACE I or ARB	278 (84%)	186 (86%)	92 (80%)	0.078
Beta blocker	287 (87%)	185 (86%)	102 (89%)	0.630
Total scores of physical	10 (\pm 6)	10 (\pm 6)	12 (\pm 6)	0.004
Total scores of emotional/	5 (\pm 5)	5 (\pm 4)	5 (\pm 5)	0.453

Table -1, Sample characteristics (N=331)

Patients have been predominately male, Caucasian, married or cohabitating, and overweight. Most of the people of patients had an ischemic HF etiology, and had been in NYHA purposeful lessons III and IV. A extra percentage of ladies have been non-Caucasian (i.e., African-American and Hispanics), and had non-ischemic HF etiology (e.g., idiopathic and hypertension) than guys ($p < 0.05$).

Symptom misery scores from each individual symptom are illustrated in table - 2.

	Male (n=216)	Female (n=115)	<i>p</i> -value
Edema	1.1 (\pm 1.5)	1.4 (\pm 1.6)	0.130
Dyspnea	2.7 (\pm 1.8)	2.9 (\pm 1.7)	0.747
Fatigue/increased need to rest	2.0 (\pm 1.7)	2.8 (\pm 1.8)	0.011
Fatigue/low energy	2.9 (\pm 1.7)	3.3 (\pm 1.6)	0.150
Sleep disturbances	2.0 (\pm 1.8)	2.5 (\pm 1.9)	0.010
Worrying	1.7 (\pm 1.7)	1.7 (\pm 1.9)	0.305
Feeling depressed	1.4 (\pm 1.6)	1.6 (\pm 1.9)	0.021
Cognitive problems	1.6 (\pm 1.7)	1.8 (\pm 1.9)	0.213

Table – 2, Symptom distress scores between men and women (N=331)

All patients stated that among bodily symptoms fatigue/low power turned into the most distressful and edema the least (imply \pm S.D three zero \pm 1.7 vs. 1.2 \pm 1.5), even as among emotional/cognitive symptoms, disturbing become the maximum distressful and feeling depressed the least (imply \pm S.D 1.7 \pm 1.Eight vs. 1.Five \pm 1.Five). Ladies mentioned substantially better levels of distress from fatigue/expanded need to rest, sleep disturbances, and feeling depressed than guys, while all different symptoms had been rated in addition with the aid of women and men.

Symptom Clusters

Three equal clusters had been recognized in men and women. The first cluster classified the bodily symptom cluster blanketed dyspnea, fatigue/increased need to relaxation, fatigue/low power, and sleep disturbances. The second one cluster categorized the emotional/cognitive symptom cluster, blanketed traumatic, feeling depressed, and cognitive issues. Edema formed a third, single symptom cluster. The dendrograms and the pseudo-F and pseudo-T squared statistics indicated that 3 clusters were the most useful answer, which changed into additionally showed through the main factor analysis

(Figures 1 and a pair of). Because the definition of a symptom cluster is two or greater symptoms that arise simultaneously, edema changed into excluded from further analyses. Additionally, due to the fact gender variations in symptom clusters have been now not determined, the entire pattern become used for in addition analyses.

Traits of Symptom Cluster companies

Patients have been divided into four groups based totally at the median break up of overall symptom misery scores of the bodily and emotional/cognitive symptom clusters. The “low distress” organization included patients with low misery in both physical and emotional/cognitive symptom clusters. The “bodily misery” institution included sufferers with high distress scores in the bodily symptom cluster and coffee misery scores within the emotional/cognitive symptom cluster. The “emotional/cognitive distress” institution protected patients with high distress ratings inside the emotional/cognitive symptom cluster and occasional misery rankings inside the physical symptom cluster. The “high distress” organization blanketed patients with excessive distress ratings in both bodily and emotional/cognitive symptom clusters.

Sufferers within the bodily misery and excessive distress agencies consisted commonly of girls and people in NYHA functional elegance III and IV. Patients in the emotional/cognitive distress and excessive distress organizations were more youthful than sufferers inside the low misery and physical distress groups. Sufferers within the excessive misery organization had a greater comorbidity burden than the ones in the different three organizations. Fewer beta blockers, angiotensin-converting enzyme inhibitors (ACE I), or angiotensin-receptor blockers (ARB) were taken with the aid of patients within the physical distress group compared to the opposite 3 agencies

Prediction of Cardiac occasion-free Survival

Throughout a mean comply with-up period of 361 days, there had been eighty-two cardiac activities: dying 2% (2/82); hospitalization eighty-one% (66/82); and ED go to 17% (14/82). The cardiac event costs of the four organizations have been 17.5% (22/126) within the low misery institution, 16.7% (6/36) inside the bodily misery organization, 32.6% (15/46) in the emotional/cognitive distress

organization, and 31.7% (39/123) in the excessive misery organization. The consequences of the multivariate hierarchical Cox regression analysis for symptom clusters are summarized in table 5. Because the whole symptom misery ratings in the bodily and emotional/cognitive symptom clusters had been considerably correlated ($r=0.644$, $p<0.01$), the interplay impact of the two symptom clusters became blanketed in the analysis. The overall symptom distress rating in the emotional/cognitive symptom cluster, however not the bodily symptom cluster, turned into an unbiased predictor of cardiac occasion-loose survival after adjusting for age, gender, total comorbidity scores, BMI, and NYHA useful elegance ($p=0.007$). Every one-unit increment in misery ratings in the emotional/cognitive symptom cluster was associated with an 18% growth within the threat for a cardiac event.

The groups have been entered as categorical variables into the multivariate hierarchical Cox regression to decide which organization had a higher threat for a cardiac event. Figure three depicts the survival curves of the four businesses after controlling for age, gender, general comorbidity rating, BMI, and NYHA purposeful class. Sufferers within the emotional/cognitive distress (threat ratio [HR]: 2.4, ninety five% self-assurance c programming language [CI] 1.31-4.41) and excessive distress companies (HR: 2.02, ninety five% CI 1.03-three.Ninety five) had a higher chance for a cardiac event in comparison to the ones inside the low misery organization. However, there have been no variations in time to first cardiac occasion between sufferers inside the bodily distress and coffee misery corporations.

Discussion

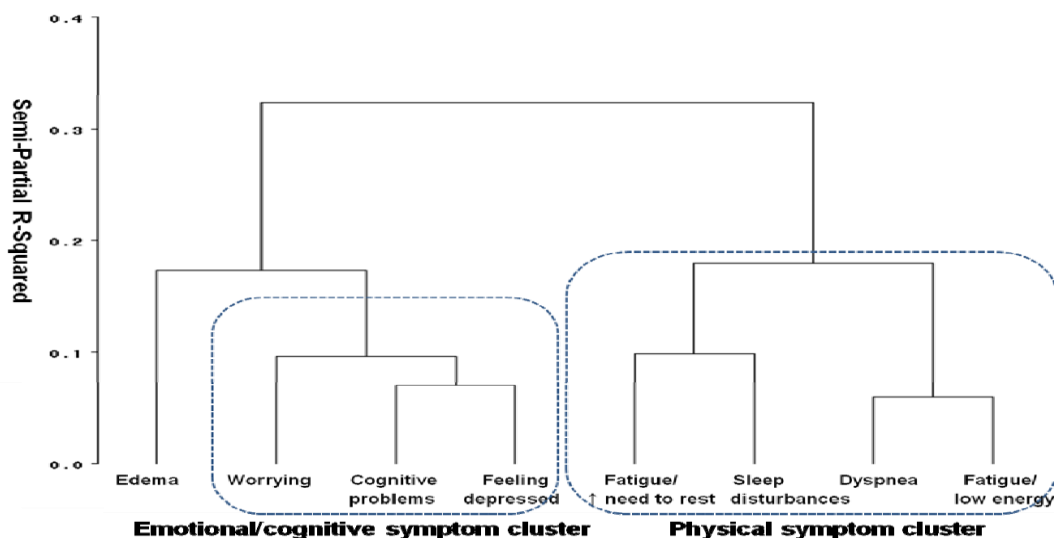
This becomes the primary examine to examine symptom clusters among genders in patients with HF and to determine whether symptom clusters expected cardiac occasion-free survival. Opposite to what we hypothesized based totally on previous studies, gender variations in HF symptom clusters were no longer determined. Between the two symptom clusters, simplest the emotional/cognitive symptom cluster expected higher threat for a cardiac occasion.

The relationships among person symptoms and outcomes in patients with HF were suggested in previous research. For instance, in the study of Ekman and colleague's dyspnea severity become a

predictor of improved demise and all-cause rehospitalization and fatigue severity become a predictor of rehospitalization due to HF exacerbation. Fifty-five coronary heart failure patients with depressive signs and symptoms have also been suggested to have up to a 2.25-fold extra hazard for dying (HR=1.08-2.25). Whilst those results are informative, they do not offer a full understanding of the connection between symptoms and consequences of sufferers with HF given that patients commonly revel in more than one signs simultaneously.

There are statistics suggesting that the presence of co-occurring signs may also convey a higher chance for bad outcomes that is probably handiest identifiable whilst signs are taken into consideration collectively. The coexistence of tension and depressive signs and symptoms in patients with coronary heart ailment became associated with a higher risk for mortality whilst compared with the presence of both symptoms alone. Superior most cancers sufferers with concurrent 4 symptoms were nine times much more likely to die in comparison to people with one symptom. For that reason, exploring symptom clusters seems to be critical for accurate risk evaluation of patients with HF.

Figure -1. Symptom cluster dendrogram



We verified a extra danger for shorter cardiac occasion-unfastened survival time in patients with better distress scores from the emotional/cognitive symptom cluster than the physical symptom cluster. A ability purpose for this locating is that healthcare providers and sufferers frequently focus extra on bodily than mental signs.

Because the number one motives for looking for fitness care are typically associated with bodily manifestations of HF, healthcare companies have a tendency to treat physical signs and no longer explore mental elements that is probably an underlying cause of HF events. As an instance, the ACC/AHA tenet for analysis and management of HF cope with bodily signs and symptoms, however now not patient emotional or cognitive signs and symptoms. Therefore, the assessment of emotional or cognitive symptom fame may not frequently occur and subsequently now not be controlled correctly.

The dearth of an impact of bodily symptom cluster on cardiac event-unfastened survival may also be explained via the affiliation between self-care and signs skilled through sufferers with HF. Coronary heart failure patients whose symptoms are extreme enough to impair each day sports have a better expertise of the significance of self-care. Excessive bodily signs and symptoms might also motivate sufferers to carry out higher self-care to prevent worsening symptoms. Alternatively, the negative effect of man or woman emotional/cognitive signs and symptoms on accomplishing self-care changed into said in preceding research: patients who are depressed, aggravating, or cognitively impaired have a tendency to revel in extra problem with and less benefits to the overall performance of self-care (e.g., taking medicinal drugs as directed and monitoring signs and symptoms regularly), difficulty remembering complex pointers on medication, food regimen, or symptom tracking, and feature much less ability to as it should be interpret the changes in symptom fame.

There is evidence that patients with depressive symptom often revel in cognitive impairment simultaneously. Mind structural modifications that worried in emotional and cognitive features (e.g., hippocampus and caudate nuclei) had been located in sufferers with HF, which can also provide an explanation for why they arise as a cluster. Different evidence to aid this end consists of the look at via Alves and associates in which depressive symptom ratings expected typical cognitive characteristic in sufferers with HF. Cognitive feature become advanced through eight-week antidepressant remedy in HF patients with principal depressive disease. Our data endorse that coexistence of both signs would possibly magnify the detrimental effect of each symptom and get worse results extra than the presence of a single symptom. For that reason, it's miles essential to reveal and manipulate signs in cluster in sufferers with HF.

Within the cluster analysis, edema became now not included in both symptom cluster for either

gender. This will be associated with decrease misery scores for edema compared to the opposite signs in this take a look at. Given that clusters were constructed by way of minimizing the heterogeneity within clusters, the decrease distress score for edema led to extra dissimilarity with other signs. Patients who experience edema commonly delay contacting their healthcare issuer earlier than acute cardiac decompensation; this may occur because they're less possibly to revel in distress from edema.

Statistics about affected person characteristics may be of value for healthcare vendors to broaden and supply efficacious techniques, along with chance assessment, which can help to save you capacity unfavorable effects. Accordingly, we tested characteristics of patients in line with symptom cluster organizations. There were more girls within the “bodily distress” and “high distress” organizations in which patients skilled extra distress from bodily signs. Comparable results had been suggested in previous research in which ladies experienced greater bodily impairment than guys. Sufferers were generally more youthful inside the “high distress” and “emotional/cognitive misery” organizations, which shows that younger sufferers with HF revel in greater misery from the emotional/cognitive symptom cluster irrespective of the misery associated with the physical symptom cluster. This finding is in line with preceding research reporting higher bodily and worse emotional reputation in younger patients with HF. Given that greater distress from the emotional/cognitive symptom cluster turned into related to earlier cardiac events on this take a look at, healthcare providers need to attention on younger patients who've terrible mental status irrespective of their physical popularity.

There had been numerous limitations of this look at. First, the outcomes of this study might not be generalizable to all sufferers with HF, specifically older patients and ethnic businesses apart from Caucasian. Second, the MLHF become used as the degree of symptom misery. This device is frequently a measure of health-related best of lifestyles. But, given the proven reliability and validity of the MLHF, the usage of symptom information from this device become a scientifically sound approach to fulfill the objectives of this observe. Even though sufferers with HF enjoy a huge variety of signs and symptoms, we simplest evaluated seven symptoms. The limited quantity of symptoms covered may not fully seize affected person symptom experiences. However, the signs covered on this look at our said maximum frequently by way of sufferers with HF. 0.33, we assessed best symptom misery levels.

Extra symptom dimensions consisting of frequency and intensity may additionally want to be

taken into consideration. In most cancers' patients, symptom misery become incredibly continual over time compared to intensity⁹⁰ and baseline symptom distress expected misery degrees three and six months later. As a result, symptom misery is probably the great predictor of lengthy-term outcomes.

Conclusions

On this take a look at we proven that symptoms occur in clusters rather than in isolation. Those findings provide a new angle on symptom evaluation and management in sufferers with HF by way of highlighting the importance of signs clusters. Identifying symptom clusters may additionally guide and help the improvement of more comprehensive interventions. Teaching sufferers approximately symptom clusters can also enhance symptom popularity by using promoting more patient self-focus. If sufferers realize that signs and symptoms arise in clusters, awareness of one symptom may additionally cause self-assessment for presence of additional symptoms, which would possibly facilitate fitness care-in search of behaviors for changes in symptom fame in a timely manner. As a consequence, specializing in symptom clusters can also result in better affected person outcomes.

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