

Original Article

Assessment of Ventricular Dysfunction by BNP in Correlation with Echocardiography

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Abstract

Objectives: The objective of the study was to assess ventricular dysfunction by BNP in correlation with echocardiography

Methods: We enrolled 91 consecutive patients admitted with decompensated CHF due to CAD, RHD, COPD or non ischemic cardiomyopathy. Blood samples for BNP assay were taken and echocardiographic examination done. Data analysis was done using student's t-test and coefficient of simple correlation using SPSS-10 software.

Results: Out of 91 CCF patients 54 were male and 37 female and their mean age was 53.95 ± 17.585 years. The presentation of patients was as; 51(56%) had CAD, 17(18.7%) RHD, 05(5.5%) COPD and 18(19.8 %) had non ischemic cardiomyopathy. Over all mean BNP level was 2019.88 ± 1353.798 pg/ml, mean BNP level for patients with CAD was 2080.94 ± 1433.733 pg/ml, for patients with RHD it was 1811.24 ± 1137.062 pg/ml, for patients with COPD it was 2355.00 ± 1595.115 pg/ml, and for patients with non-ischemic cardiomyopathy it was 1950.83 ± 1322.386 pg/ml. There was significant correlation between blood BNP levels and left ventricular end systolic diameter ($r=0.208$, $p=0.048$), left ventricular fractional shortening ($r=-0.327$, $p=0.002$), left ventricular end systolic volume ($r=0.225$, $p=0.032$), left ventricular ejection fraction ($r=-0.321$, $p=0.002$) and right ventricular end diastolic diameter ($r=0.221$, $p=0.036$). There was no significant correlation between BNP levels and left ventricular end diastolic diameter ($r=0.086$, $p=0.420$), left atrial size ($r=0.023$, $p=0.831$). There was significant difference in the BNP levels in the NYHA III and IV (884 ± 685 pg/ml vs. 2666 ± 1207 pg/ml, $p=0.000$).

Conclusions: BNP levels significantly correlate with the left ventricular size and function in systole on echocardiography but not with the left atrial size.

Key Words: Natriuretic peptide, heart failure, echocardiography.

Introduction

The gratifying reduction in the mortality due to acute myocardial infarction and arrhythmias has resulted in an increase in the elderly population with congestive heart failure (CHF) due to coronary artery disease (CAD).¹ The tremendous burden of CHF on the resources¹ necessitates finding the ways to prevent its occurrence, to halt its progression and to minimize the sufferings from it.² In the diagnosis of CHF brain natriuretic peptide (BNP) has been shown to supercede the symptoms and physical signs including phonocardiographic S3 and S4.^{3,4} Moreover the diagnostic and prognostic role of BNP has been well validated.^{5,6} Plasma BNP level is a sensitive indicator of

ventricular dysfunction both in symptomatic and asymptomatic patients and its plasma concentration increases with volume and pressure overload in patients with heart failure.^{7,8} In addition, the left ventricular (LV) systolic dysfunction, plasma BNP levels have been suggested to be significantly associated with diastolic stage (including newer echocardiographic parameters as tissue Doppler imaging and color M-mode propagation velocity) and right ventricular (RV) functions as well.^{9,10} Though the relationship between BNP and left and right ventricular functions have been elucidated in patients with systolic heart failure¹¹, the diagnostic value of BNP in prediction of right and left ventricular systolic/diastolic functions in patients with acute heart failure is not well established and literature data are controversial.¹² The rationale behind the study was to know the correlation between BNP and various echocardiographic parameters of heart failure in patients admitted with

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Objectives: The objective of the study was to assess ventricular dysfunction by BNP in correlation with echocardiography decompensated CHF in NYHA class III or IV dyspnea.

Material and Methods

This was a cross-sectional study carried out at Cardiology department Post Graduate Medical Institute (PGMI), Lady Reading Hospital (LRH), Peshawar, Pakistan from 15th December 2005 to 7th June 2006. A total of 91 patients (both male and female) presenting with different symptoms were included after informed consent.

Variables Definition:

1. Decompensated CHF: Patient admitted with NYHA class III or IV dyspnea, raised JVP, edema and S₃.
2. NYHA class III dyspnea: Dyspnea on less than ordinary activity i.e. on getting out of bed to the bedside toilet.
3. NYHA class IV dyspnea: Dyspnea at rest i.e. in the bed with head end elevated.
4. Coronary artery disease (CAD): Diagnosed on history or evidence of ischemia on ECG, exercise test, coronary angiography or cardiac imaging.
5. Rheumatic heart disease (RHD): Having deformed regurgitant or stenotic valves on echocardiography¹.
6. Chronic obstructive pulmonary disease (COPD): Right ventricular hypertrophy and dilatation caused by diseases of lung parenchyma and/or pulmonary vasculature unrelated to left side of the heart².
7. Cardiomyopathies: Causes of congestive heart failure with systolic or diastolic dysfunction other than CAD, RHD and COPD.

INCLUSION CRITERIA:

1. Patients with decompensated CHF in NYHA class III or IV dyspnea for which hospitalization was mandatory for stabilization.

EXCLUSION CRITERIA:

1. Congenital heart disease
2. Acquired ventricular septal defect
3. Pericardial diseases
4. Patients with renal failure having serum creatinin more than 2 mg/dl

Blood BNP levels were determined by using AXSYM (Abbott) BNP assay. Dimensions and fractional shortening of LV were determined using M-Mode. LV

Volumes and ejection fraction were determined using modified Simpson's rule.¹⁴

DATA ANALYSIS: The data were analyzed using SPSS version-10 software. The mean and standard deviation of blood levels of BNP in decompensated CHF were determined for each etiological group. Scatter diagrams and curvilinear regression plots were plotted for plasma BNP levels against LV dimensions, LV fractional shortening, LV ejection fraction, left atrial diameter and RV end diastolic diameter and coefficient of simple correlation were determined. Independent sample t-test and box plot was used to find the difference between the BNP levels in patients with NYHA class III and IV.

Results

Out of 91 CCF patients 54 were male and 37 female and their mean age was 53.95±17.585 years. These heart failure patients had presented with different pathological causes given in Table 2. Sixty four (70.3%) patients were in sinus rhythm while 27(29.7) had atrial fibrillation. Thirty three (36.3%) patients had NYHA class III dyspnea while 58 (63.7%) had NYHA class IV dyspnea. Over all mean BNP level was 2019.88±1353.798 pg/ml, mean BNP level for patients with CAD was 2080.94 ±1433.733 pg/ml, for patients with RHD it was 1811.24±1137.062 pg/ml, for patients with COPD it was 2355.00±1595.115 pg/ml, and for patients with nonischemic cardiomyopathy it was 1950.83±1322.386 pg/ml. There was no significant difference in the blood BNP levels in those with and without atrial fibrillation (2058±1330 vs. 1928±1428 p=0.678).

There was significant correlation between blood BNP levels and left ventricular end systolic volume (r=0.225,p=0.032, Fig 1) left ventricular end systolic diameter (r=0.208, p=0.048, Fig 2), left ventricular fractional shortening (r=-0.327, p=0.002), left ventricular ejection fraction (r=-0.321, p=0.002), and right ventricular end diastolic diameter (r=0.218, p=0.038) [Table 2] . Significant correlation was also observed between blood BNP levels and right ventricular end diastolic diameter (r=0.218, p=0.038). There was no significant association between BNP levels and left atrial size (r=0.023, p=0.831). Likewise; no significant correlation between BNP levels Left ventricular end diastolic volume (r=0.136, p=0.200), and left atrial size (r=0.023, p=0.831) [Table2]

Discussion

In our study the most significant correlation of the BNP levels, amongst the echocardiographic parameters, was with the LV ejection fraction followed by LV fractional shortening, right ventricular diameter, LV

end systolic volume and left ventricular end systolic dimension (LVESD). Similar finding were found in a study by Gackowski A et al.¹⁵ The correlation of BNP levels with coronary angiographic ejection fraction was more significant ($r = -0.690, p < 0.001$) in a study by Palumbo B et al than the echocardiographic correlation in our study ($r = -0.321, p = 0.002$).¹⁶ Moreover both the blood BNP levels and LV end systolic volume are determinants of mortality and prognosis in patients with CHF.^{17,18} In our study the blood BNP levels in patients having NYHA class III was very significantly different from those having NYHA IV dyspnea. Our results tally with the results of Wieczorek SJ et al where the BNP levels increased with the severity of CHF based on NYHA class.¹⁹ BNP can thus be used as an objective measure for dyspnea and an alternative to six minute walk test to assess the severity of heart failure.²⁰ In a study Lee SC et al found that BNP levels correlated best with the NYHA functional class both at baseline and follow up so much so that they concluded with remarks that plasma BNP is a useful objective biomarker in monitoring human CHF in the outpatient setting.²¹ Yasue H et al by comparing the blood BNP levels in the blood samples from aortic root, anterior interventricular and coronary sinus found that the predominant source of BNP release was the left ventricle not the atrium. Our study showing no correlation between the left atrial size and BNP levels and a significant correlation with left ventricular end diastolic volume complement the results of Yasue H et al.²²

Thus our null hypothesis was partially rejected i.e. some of the echocardiographic parameters correlated well with plasma BNP levels. The rather by product of our study which came out very strongly was the very significant difference in BNP levels based on the NYHA functional class. This fact, if validated by larger randomized studies will go a long way towards the objective assessment of functional status in CHF.

Study limitations:

1. Small sample size
2. Uneven etiologic groups
3. Difference based on NYHA class was not included in our objectives.

Conclusion

Blood BNP levels significantly correlate with the left ventricular size and function in systole on echocardiography but not with the left atrial size.

Table 1: Distribution according to etiology n=91

Patient's presentation	n (%)
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CAD	51(56%)
RHD	17(18.7%)
COPD	05(5.5%)
Cardiomyopathy	18(19.8 %)
Total	91(100%)

Table 2: Correlations of Echocardiographic parameters with blood BNP levels n=91

Parameters	r	P value
LVESD& BNP	0.086	0.420
LVESD& BNP	0.208	0.048
LVFS& BNP	-.327	0.002
LVEDV& BNP	0.136	0.200
LVESV& BNP	0.225	0.032
LVEF& BNP	-.321	0.002
LA DIAMETER& BNP	0.023	0.831
RVEDD& BNP	0.221	0.036

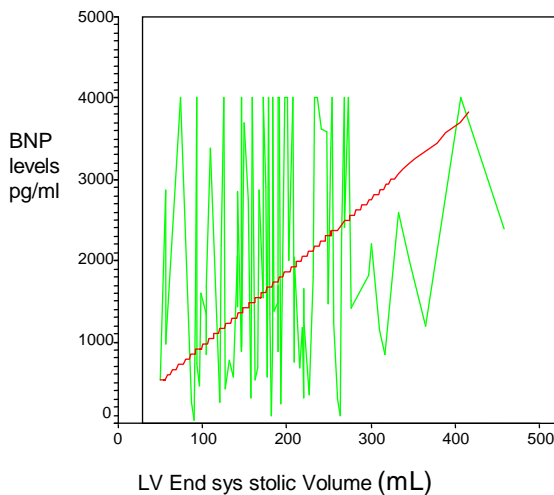


Fig 1: Correlation between BNP and LVESV

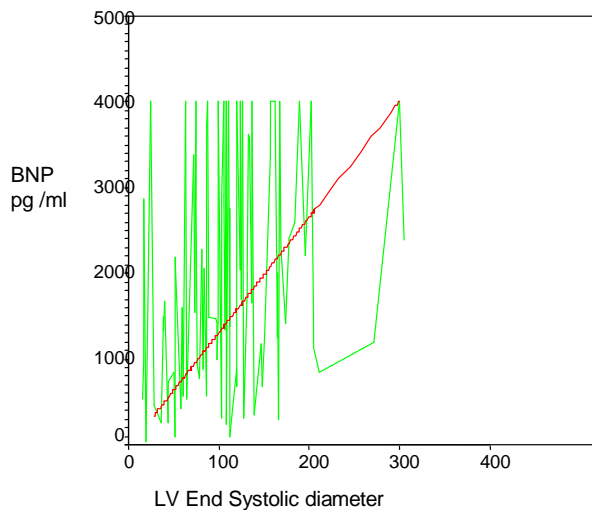


Fig 1: Correlation between BNP and LVES diameter n=91

Brain Natriuretic Peptide (pg/ml)			
Diagnosis	Mean	N	Std. Deviation
CAD	2080.94	51	1433.733
RHD	1811.24	17	1137.062
COPD	2355.00	5	1595.115
CARDIOMYOPATIE S	1950.83	18	1322.386
Total	2019.88	91	1353.798

References

- McCullough PA, Omland T, Maisel AS. B-Type Natriuretic Peptides: A Diagnostic Breakthrough for Clinicians. *Rev Cardiovasc Med* 2003;4:72-80
- Givertz MM, Collucci WS, Braunwald E. Clinical aspects of heart failure. In: Eugene Braunwald, Zepis DP, Libby P, Bonow RO, editors. *Heart Disease 7th ed*. New York: W.B. Saunders company; 2005. p 539.
- Maisel A, Hollander JE, Guss D, McCullough P, Nowak R, Green G et al. Primary results of the Rapid Emergency Department Heart Failure Outpatient Trial (REDHOT). A multicenter study of B-type natriuretic peptide levels, emergency department decision making, and outcomes in patients presenting with shortness of breath. *J Am Coll Cardiol*. 2004; 44:1328-33
- Marcus GM, Gerber IL, McKeown BH, Vessey JC, Jordan MV, Huddleston M et al. Association between phonocardiographic third and fourth heart sounds and objective measures of left ventricular function. *JAMA* 2005; 293:2238-2244.
- Smith H, Pickering RM, Struthers A, Mant D. Biochemical diagnosis of left ventricular dysfunction in elderly

- patients in general practice—an observational study. *BMJ* 2000; 320:906-908.
- Suttner SW, Boldt J. Natriuretic peptide system: Physiology and clinical utility. *Curr Opin Crit Care* 2004;10:336-341.
- Cowie MR, Mendez GF. Brain natriuretic peptide and congestive heart failure. *Prog Cardiovasc Dis* 2002; 44: 293-21.
- Meune C, Fulla Y, Martins E, Bergmann JF, Devaux JY. B-type natriuretic peptide for the diagnostic and prognostic assessment in cardiology. Its interest and perspectives of application. *Presse Med* 2003; 32: 181-5
- Yamaguchi H, Yoshida J, Yamamoto K, Sakata Y, Mano T, Akehi N, et al. Elevation of brain natriuretic peptide is a hallmark of diastolic heart failure independent of ventricular hypertrophy. *J Am Coll Cardiol* 2004; 43: 55-60.
- Troughton RW, Prior DL, Pereira JJ, Martin M, Fogarty A, Morehead A, et al. Plasma B-type natriuretic peptide levels in systolic heart failure: importance of left ventricular diastolic function and right ventricular systolic function. *J Am Coll Cardiol* 2004; 43: 416-22.
- Weber M, Hamm C. Role of B-type natriuretic peptide and NT-proBNP in clinical routine. *Heart* 2006; 92: 843-9.
- Ordonez-Lianos J, Merce-Muntanola J, Santalo-Bel M. Natriuretic peptide testing in emergency settings. *Clin Chem Lab Med* 2008; 46: 1543-9.
- Wu AHB, Smith A, Wiczorek S, Mather JF, Duncan B, White CM et al. Biological variation for N-Terminal Pro- and B-type natriuretic peptides and implications for therapeutic monitoring of patients with congestive heart failure *Am J Cardiol* 2003;92: 628-631.
- William F, Armstrong, Harvey Feigenbalm. Echocardiography. In: Eugene Braunwald, Douglas P. Zepis, Peter Libby, Robert O, Bonow, editors. *Heart Disease 7th ed*. New York: W.B. Saunders company; 2005. p 197.
- Gackowski A, Isnard R, Golmard JL, Pousset F, Carayon A, Montalescot G, et al. Comparison of echocardiography and plasma B-type natriuretic peptide for monitoring the response to treatment in acute heart failure. *Eur Heart J* 2004;25(20):1788-96
- Palumbo B, Siepi D, Lupattelli G, Sinzinger H, Fiorucci G, Annibolletti PF et al. Usefulness of brain natriuretic peptide levels to discriminate patients with stable angina pectoris without and with electrocardiographic myocardial ischemia and patients with healed myocardial infarction. *Am J Cardiol*. 2004 ;94:780-3.
- Doust JA, Pietrzak E, Dobson A, Glasziou P. How well does B-type natriuretic peptide predict death and cardiac events in patients with heart failure: systematic review. *BMJ* 2005;330(7492):625.
- Vanderheyden M, Goethals M, Verstreken S, De Bruyne B, Muller K, Van Schuerbeeck E, et al. Wall stress modulates brain natriuretic peptide production in pressure overload cardiomyopathy. *J Am Coll Cardiol* 2004; 44:2349-54.

19. Wiecezorek SJ, Wu AH, Christenson R, Krishnaswamy P, Gottlieb S, Rosano T et al. A rapid B-type natriuretic peptide assay accurately diagnoses left ventricular dysfunction and heart failure: a multicenter evaluation. *Am Heart J* 2002; 144:834-839.
 20. Wiecezorek SJ, Hager D, Barry MB, Kearney L, Ferrier A, Wu AH. Correlation of B-type natriuretic peptide level to 6-min walk test performance in patients with left ventricular systolic dysfunction. *Clin Chim Acta* 2003; 328:87-90.
 21. Lee SC, Stevens TL, Sandberg SM, Heublein DM, Nelson SM, Jougasaki M et al. The potential of brain natriuretic peptide as a biomarker for New York Heart Association class during the outpatient treatment of heart failure. *J Card Fail* 2002; 8:149-54.
 22. Yasue H, Yoshimura M, Sumida H, Kikuta K, Kugiyama K, Jougasaki M et al. Localization and mechanism of secretion of B-type natriuretic peptide in comparison with those of A-type natriuretic peptide in normal subjects and patients with heart failure. *Circulation* 1994; 90:195-203.
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