

ORIGINAL ARTICLE

Effectiveness of Beta Blockers against Hypertrophic Cardiomyopathy

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ABSTRACT

Aim: To evaluate the effectiveness of beta blockers against hypertrophic cardiomyopathy.**Study design:** Retrospective study**Place and duration of study:** Department of Medicine, Islam Medical College, Sialkot from 01-10-2020 30-01-09-2021.**Methodology:** One hundred and thirty six confirmed hypertrophic cardiomyopathy cases within the age of 3-75 years were enrolled. Different types of beta blocker used and its efficacy result and side effects were documented. The average efficacy was assessed for each beta blocker upto 5 years' survival rate.**Results:** High doses of beta blocker were indicated for hypertrophic cardiomyopathy patients in majority of the cases. The efficacy of various beta blockers shows that cardio-selective blocker is much better in efficacy. Left ventricle outflow tract (LVOT) with a value greater or equal to 50 mmHg was observed in patients at or greater than 70 years.**Conclusion:** Beta blockers are effective in hypertrophy cardiomyopathy cases depending upon the left ventricle out flow tract for better outcome.**Key words:** Beta blockers, Cardiomyopathy, Efficacy, Therapeutic interventions

INTRODUCTION

Hypertrophic Cardiomyopathy (HCM) is the disease of heart muscles in which heart muscles get thickened abnormally, it makes harder for the heart to pump blood normally. It is the most common genetic cardiomyopathy characterized by heterogeneous morphology, complex pathophysiology and variable clinical manifestations¹⁻⁴. It is also regarded as rare malignant disease. The spectrum of HCM has expanded over the recent years due to its increasing prevalence and clinical profile^{3,5}. Hypertrophic cardiomyopathy is ranging from severe manifestations to absence of morphological expression^{6,7}.

Present pharmacological interventions have not shown enough results to modify or reduce the symptomology of HCM^{8,9} except diltiazem in preventing LV remodeling¹⁰. Only present therapy to reduce long term prognosis of this rare disease is implantable cardiac defibrillator (ICD), which is known as myectomy⁸. However, pharmacological therapies play a subsequent role in reducing the risk disease complications and disease implications. Targeted goals of these therapies is to control of HCM symptoms, treatment of LV dysfunction, reduction of intraventricular gradients, control of atrial fibrillation, heart failure, prevention of cardioembolism and ventricular arrhythmias.

Few hypertrophic cardiomyopathic patients were included in randomized control trial to evaluate the efficacy of different drugs used against this disease⁸. Scientific community is hoping for disease specific/better treatment plans and options^{11,12}.

This study is aimed to design to find the efficacy of beta-blockers in the treatment or reducing symptomology of hypertrophic cardiomyopathy.

MATERIALS AND METHODS

This retrospective study was carried out at from 1st October 2020 to 30th September 2021 after IRB permission. The present study was conducted on 136 hypertrophic cardiomyopathy confirmed cases within the age 3-75 years who visit the indoor outdoor patients were enrolled. The required past data analysis for assessing the efficacy of beta blockers in HCM patients. This research was approved from the ethical institutional committee. Patient's demographic information as well as type of beta blocker used, its efficacy result and side effects was documented as an overall picture of all the enrolled patients. Those patients who were only suffering from hypertension and not HCM were not included in this study. The average efficacy was assessed for each beta blocker upto 5 yrs survival rate. The data was entered in SPSS.

RESULTS

The mean age was 35±10.1 years. High doses of beta blocker were indicated for HCM patients in majority of the cases. Indication of beta blockers were either angina, dyspnoea reduction, ventricular-ectopic beats control, ventricular response control in AF patients or in heart failure cases (Table 1)

Table 1: Indications for beta blockers

Beta blocker	Indications	Initial dose	Maximum dose
Propranolol	Angina, dyspnoea reduction, ventricular-ectopic beats control, ventricular response control in AF patients control of	40 mg BD	80 mg TDS
Atenolol	Like propranolol	25 mg QD	150 mg QD
Nadolol	Like propranolol, in NSVT, SCD incidence prevention	40 mg QD	80 mg BD
Metoprolol	Like propranolol	50 mg QD	100 mg BD
Bisoprolol	Systolic dysfunction treatment and terminal Heart Failure patients	1.25 mg QD	15 mg QD

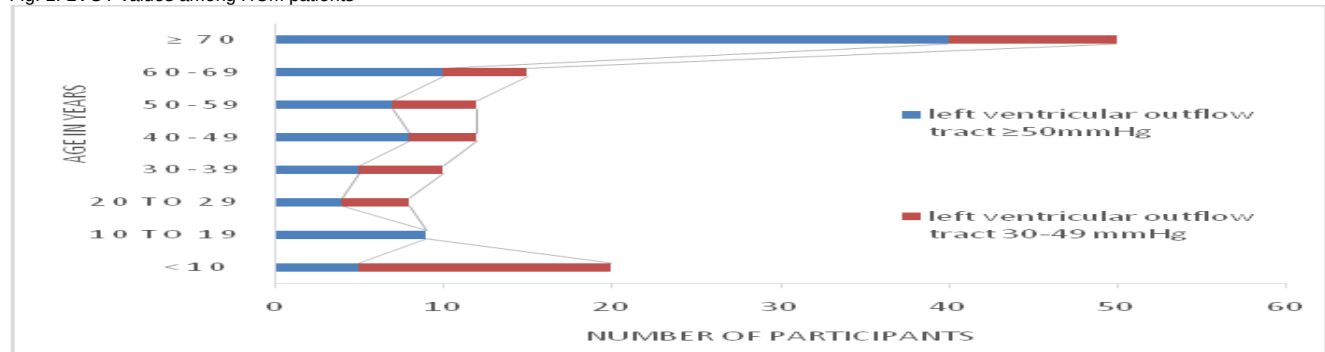
Table 2: Beta blockers and their side effects

Beta Blocker	Efficacy	Features	Side effects
Propranolol	Drug of choice in neonates/infants	Short - half-life	Depression, chronotropic-incompetence AV decrease and asthma formation
Atenolol	Drug of choice in HC with hypertension	-----	Hypotension chronotropic-incompetence asthma
Nadolol	Effective in obstruction control	Increase patient compliance in QD	Chronotropic-incompetence, AV conduction reduction
Metoprolol	Not useful in hypertrophic obstructive cardiomyopathy	Short half-life	Asthma
Bisoprolol	Not useful in hypertrophic obstructive cardiomyopathy	-----	Chronotropic-incompetence asthma

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Fig. 1: LVOT values among HCM patients



Beta blockers have various side effects due to its treatment. These include depression, asthmatic attacks, hypotension, chronotropic incompetence as well as AV conduction reduction. The efficacy of various beta blockers shows that cardioselective blocker is much better in efficacy (Table 2). In patients with higher LVOT rate the beta blockers did not work well and showed poor health prognosis. As an additional information it was observed that the left ventricle outflow tract (LVOT) with a value greater or equal to 50mm Hg was observed in patients at or greater than 70 years while lesser LVOT than 50mm Hg was observed in paediatric cases at highest peak (Fig. 1).

DISCUSSION

Beta adrenergic blockers remain an important pharmacological intervention in the treatment of hypertrophic cardiomyopathy. It can be used for with and without outflow obstructions. Beta blockers shown great pharmacologic properties for the reduction of physiologic outflow obstruction, risk of ventricular arrhythmias, angina and dyspnea. These effects are achieved by reducing heart rate which leads to elevated diastolic filling time, decrease in ventricular stiffness and inotropy.¹³

Different beta blockers are used for the treatment of hypertrophic cardiomyopathy¹³, propranolol was the first introduced BAB in the earliest trials.¹⁴⁻¹⁷ Other trial compared the result of two beta blockers; nadolol and verapamil (calcium channel blocker). Nadolol shows better result in symptomatic relief as compared to verapamil¹⁸, while bisoprolol proved effectiveness against gradient on provocation reduction.¹⁹ It is widely accepted that non vasodilating beta blockers prove great efficacy in avoiding obstruction in HCM patients²⁰.

Long-term consequences of beta blockers are still undefined but it is considered potential therapy and standard care in with and without outflow obstruction patients.¹³ Patient symptomology and stability is took into consideration while deciding treatment and rhythm control strategy by beta blockers. Insufficient data and literature is available to assess patient' outcomes in each strategy. Certain medication should be avoided while using beta blockers including digoxin.²⁰ On the other, hand, few beta blockers showed better efficacy when use in combination with other^{20,21}.

CONCLUSION

Beta blockers are effective in hypertrophy cardiomyopathy cases depending upon the left ventricle out flow tract for better outcome.

Conflict of interest: Nil

REFERENCES

- Alcalai R, Seidman JG, Seidman CE. Genetic basis of hypertrophic cardiomyopathy: From bench to the clinics. *J Cardiovasc Electrophysiol* 2008;19:104-10.
- Mogensen J, Murphy RT, Kubo T, Bahl A, Moon JC, Klausen IC, et al. Frequency and clinical expression of cardiac troponin I mutations in 748 consecutive families with hypertrophic cardiomyopathy. *J Am Coll Cardiol* 2004; 44:2315-25.
- Maron BJ. Hypertrophic cardiomyopathy: a systematic review. *JAMA* 2002;287:1308-20.
- Maron BJ, Casey SA, Hauser RG, Aeppli DM. Clinical course of hypertrophic cardiomyopathy with survival to advanced age. *J Am Coll Cardiol* 2003;42:882-8.
- Semsarian C, Ingles J, Maron MS, Maron BJ. New perspectives on the prevalence of hypertrophic cardiomyopathy. *J Am Coll Cardiol* 2015;65:1249-54.
- Ho CY, Seidman CE. A contemporary approach to hypertrophic cardiomyopathy. *Circulation* 2006;113:e858-62.
- Captur G, Lopes LR, Mohun TJ, Patel V, Li C, Bassett P, et al. Prediction of sarcomere mutations in subclinical hypertrophic cardiomyopathy. *Circ Cardiovasc Imaging* 2014; 7:863-71.
- Spoladore R, Maron MS, D'Amato R, Camici PG, Olivetto I. Pharmacological treatment options for hypertrophic cardiomyopathy: High time for evidence. *Eur Heart J* 2012;33:1724-33.
- Axelsson A, Iversen K, Vejstrup N, Ho C, Norsk J, Langhoff L, et al. Efficacy and safety of the angiotensin II receptor blocker losartan for hypertrophic cardiomyopathy: The INHERIT randomised, double-blind, placebo-controlled trial. *Lancet Diabetes Endocrinol* 2015;3:123-31.
- Ho CY, Lakdawala NK, Cirino AL, Lipshultz SE, Sparks E, Abbasi SA, et al. Diltiazem treatment for pre-clinical hypertrophic cardiomyopathy sarcomere mutation carriers: a pilot randomized trial to modify disease expression. *JACC Heart Fail* 2015;3:180-88.
- Elliott PM, Anastakis A, Borger MA, Borggreve M, Cecchi F, Charron P, et al. 2014 ESC guidelines on diagnosis and management of hypertrophic cardiomyopathy: The task force for the diagnosis and management of hypertrophic cardiomyopathy of the European Society of Cardiology (ESC). *Eur Heart J* 2014;35:2733-79.
- Gersh BJ, Maron BJ, Bonow RO, Dearani JA, Fifer MA, Link MS, et al. 2011 ACCF/AHA guideline for the diagnosis and treatment of hypertrophic cardiomyopathy: a report of the American College of Cardiology Foundation/American Heart Association Task Force on practice guidelines. *Circulation* 2011;124:e783-831.
- Spoladore R, Maron MS, D'Amato R, Camici PG, Olivetto I. Pharmacological treatment options for hypertrophic cardiomyopathy: high time for evidence. *Eur Heart J* 2012;33(14):1724-33.
- Cohen LS, Braunwald E. Amelioration of angina pectoris in idiopathic hypertrophic subaortic stenosis with beta-adrenergic blockade. *Circulation* 1967;35:847-51.
- Flamm MD, Harrison DC, Hancock EW. Muscular subaortic stenosis. Prevention of outflow obstruction with propranolol. *Circulation* 1968;38:846-58.
- Adelman AG, Shah PM, Gramiak R, Wigle ED. Long-term propranolol therapy in muscular subaortic stenosis. *Br Heart J* 1970;32:804-11.
- Thompson DS, Naqvi N, Juul SM, et al. Effects of propranolol on myocardial oxygen consumption, substrate extraction, and haemodynamics in hypertrophic obstructive cardiomyopathy. *Br Heart J* 1980;44:488-98.
- Gilligan DM, Chan WL, Joshi J, et al. A double-blind, placebo-controlled crossover trial of nadolol and verapamil in mild and moderately symptomatic hypertrophic cardiomyopathy. *J Am Coll Cardiol* 1993;21:1672-9.
- Cabrera-Bueno F, García-Pinilla JM, Gómez-Doblas JJ, Montiel-Trujillo A, Rodríguez-Bailón I, de Teresa-Galván E. Beta-blocker therapy for dynamic LV outflow tract obstruction induced by exercise. *Int J Cardiol* 2007;117:222-6.
- Elliott PM, Anastakis A, Borger M, et al. ESC Guidelines on diagnosis and management of hypertrophic cardiomyopathy The Task Force for the Diagnosis and Management of Hypertrophic Cardiomyopathy of the European Society of Cardiology (ESC). *Eur Heart J* 2014;2014;35:2733-79.
- Guttmann OP, Rahman MS, O'Mahony C, Anastakis A, Elliott PM. Atrial fibrillation and thromboembolism in patients with hypertrophic cardiomyopathy: systematic review. *Heart* 2014;100(6):465-72.