

The Impact Of EECp On Improving Exercise Capacity In Patients With Stable Ischemic Heart Disease Who Treated Medically Or Percutaneously Over One Year

Ammar Jabbar Majeed¹, Yasseen Abdulruda Yasseen², Ali Yahya Abdullah Alsallami³, Khalid Ibrahim Amber⁴

¹M.B.Ch.B; F.I.C.M.S (Medicine), DM (cardiology); Assistant professor, Department of Interventional Cardiology, Medical College, University of Kufa, Iraq

²M.B.Ch.B; F.I.C.M.S (Medicine), DM (cardiology); Assistant professor, Department of Interventional Cardiology, Medical College, University of Kufa, Iraq

³M.B.Ch.B; F.I.C.M.S (Medicine), DM (cardiology); Assistant professor, Department of Interventional Cardiology, Medical College, University of Kufa, Iraq

⁴M.B.Ch.B; F.I.C.M.S (Medicine), Consultant cardiology, Head of Al Najaf cardiac center, Iraq

*Corresponding Author:

Ammar Jabbar Majeed,

Email ID: ammarmajeed48@gmail.com

Cite this paper as: Ammar Jabbar Majeed, Yasseen Abdulruda Yasseen, Ali Yahya Abdullah Alsallami, Khalid Ibrahim Amber, (2025) The Impact Of EECp On Improving Exercise Capacity In Patients With Stable Ischemic Heart Disease Who Treated Medically Or Percutaneously Over One Year. *Journal of Neonatal Surgery*, 14 (11s), 20-26.

ABSTRACT

Objective: Enhanced External Counterpulsation (EECP) is a non-invasive therapy involving sequential inflation of external cuffs placed on the lower limbs in sync with the cardiac cycle. This method is primarily used to treat refractory angina (persistent for at least three months) in patients who do not respond to medical therapy, surgical intervention, or percutaneous coronary procedures.

Methods: A total of 91 patients with refractory angina who were either unresponsive to medical therapy, ineligible for surgical intervention, or had undergone PCI without symptom relief were enrolled in the study. Patients were selected from the Al-Najaf Cardiac Center and private clinics between January 2018 and December 2019. All participants underwent coronary angiography, and those with significant three-vessel disease (>70% stenosis in each vessel) were included.

Results: Over 12 months, 91 patients (32 women and 59 men) aged 45 to 80 years (mean 61 ± 8.2) were analyzed. Most patients completed an average of 26 EECp sessions ($SD \pm 7$), with a response rate of 88.7%, as measured by symptom improvement based on Canadian Cardiovascular Society (CCS) classification. No significant difference in response to EECp was observed between the PCI group and those treated with medical therapy alone ($p = 0.87$, $p = 0.47$). Additionally, sex ($p = 0.185$), smoking history ($p = 0.67$), hypertension ($p = 0.4$), diabetes ($p = 0.12$), and age ($p = 0.26$) did not significantly impact the response to EECp.

Conclusion: EECp proved to be a safe and effective treatment for select patients with refractory angina who were unresponsive to conventional medical or surgical interventions.

Keywords: Lipid nanoparticles, curcumin, antifungal therapy, targeted drug delivery, fungal infections, nanotechnology.

1. INTRODUCTION

Refractory angina is defined as chronic stable ischemic heart disease lasting three months or more, unmanageable through medical therapy, surgical intervention, or PCI. EECp, a non-invasive technique, involves placing compressive cuffs on the lower limbs that inflate sequentially from distal to proximal during the cardiac cycle, mimicking the effects of an intra-aortic balloon pump. This therapy enhances venous return, improves coronary perfusion, and reduces vascular resistance, thereby alleviating cardiac workload and increasing systemic circulation.

EECP has been shown to improve endothelial function, reduce vascular stiffness, and enhance collateral blood flow. The International EECp Patient Registry has reported long-term benefits in heart failure patients, including decreased angina

symptoms, reduced nitroglycerin dependence, and improved quality of life. These benefits have been sustained for up to three years post-treatment.

While EECp has demonstrated efficacy in managing stable angina, its application is restricted by contraindications, including coagulation disorders (INR >2.5), severe COPD, cardiac arrhythmias, and significant peripheral vascular disease. Additionally, studies suggest that EECp reduces hospitalization costs, making it a viable option for future widespread adoption.

Study Objectives: To evaluate the efficacy of EECp in treating refractory angina in patients undergoing PCI versus those managed medically, with a focus on short-term outcomes.

Methods: A total of 518 patients with refractory angina who were unresponsive to medical therapy or ineligible for surgical intervention were initially considered. All underwent coronary angiography, and those with significant three-vessel disease were divided into two groups:

1. Patients deemed unfit for intervention or surgery.
2. Patients eligible for PCI on one or two vessels but experiencing persistent ischemic symptoms post-PCI.

After six weeks, only those with ongoing symptoms were included in the study, resulting in 91 enrolled patients (48 in the medical group and 43 in the PCI group). All were referred for EECp treatment eight weeks post-angiography.

Patients were treated according to standard guidelines, with comprehensive baseline data collected, including age, sex, blood pressure, glucose levels, renal function, and CBC results. Inclusion criteria required symptomatic angina despite medical therapy or intervention. Exclusion criteria included severe peripheral vascular disease, aortic aneurysm, severe aortic regurgitation, uncontrolled hypertension, and coagulopathies.

Each EECp session lasted one hour, with most patients completing 30 sessions. Treatment response was assessed using CCS grading before and after EECp, along with echocardiographic evaluations at six months and one year.

2. RESULTS

Of the 91 patients, 59 were men and 32 were women. Most completed an average of 26 EECp sessions. After treatment, 88.7% showed symptomatic improvement.

No significant difference was observed between the PCI and non-PCI groups in response to EECp ($p = 0.87$) or echocardiographic parameters ($p = 0.47$). Similarly, age ($p = 0.26$), sex ($p = 0.185$), smoking status ($p = 0.67$), hypertension ($p = 0.4$), and diabetes ($p = 0.12$) did not significantly impact EECp outcomes.

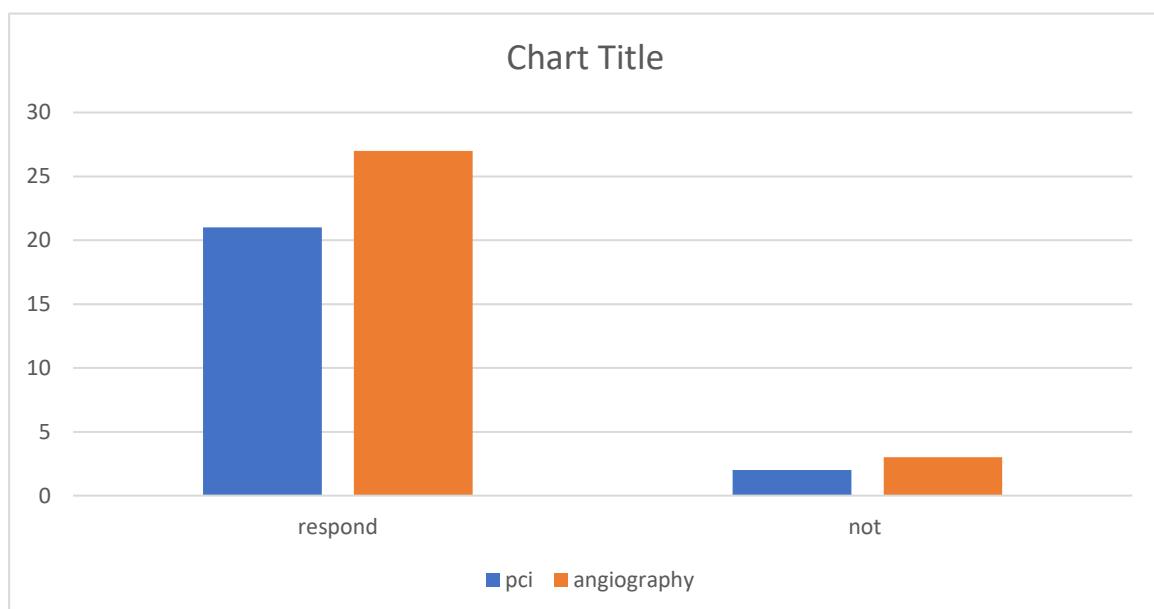


Figure 1: Relation of PCI group and angiography only group in response to EECp,

P-value = 0.87.

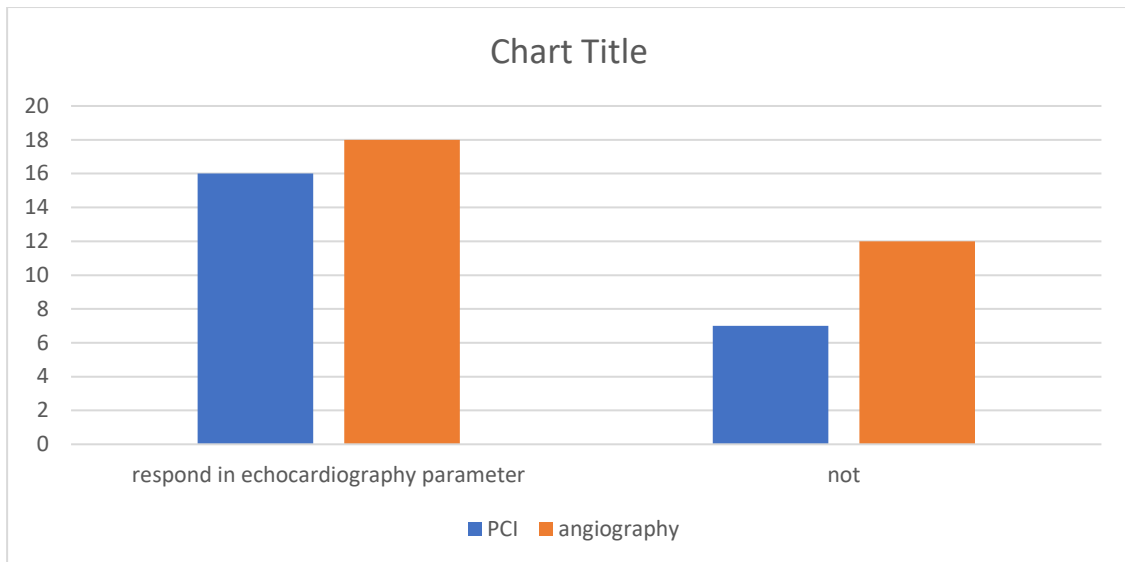


Figure 2: Relation of PCI group and angiography only group in response to echocardiography parameter, P-value = 0.4.

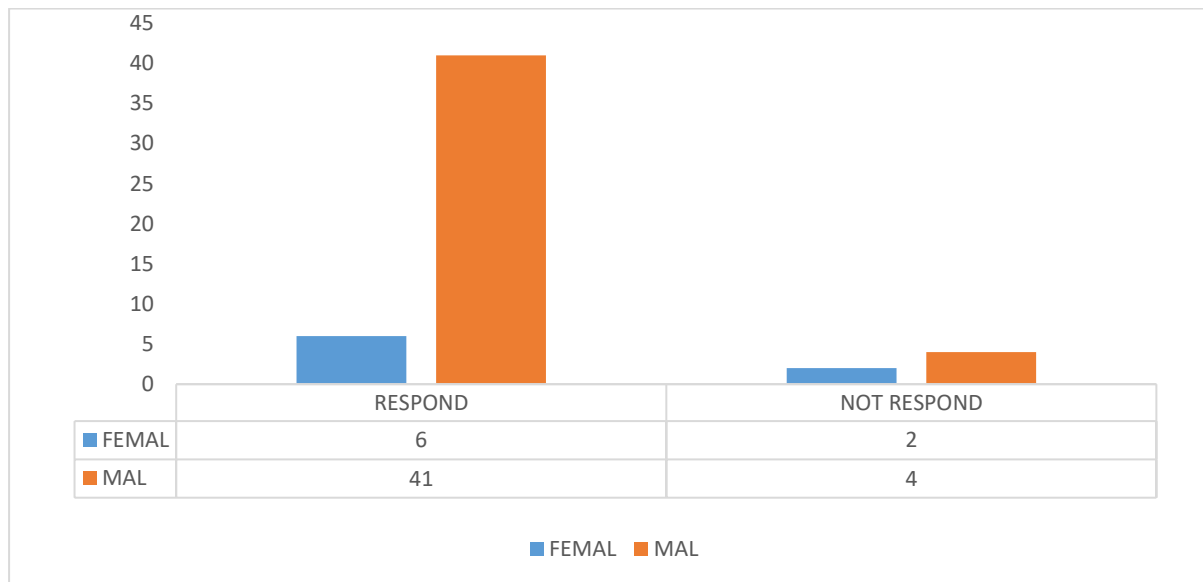


Figure 3: Sex and response to EECP, P-value = 0.185.

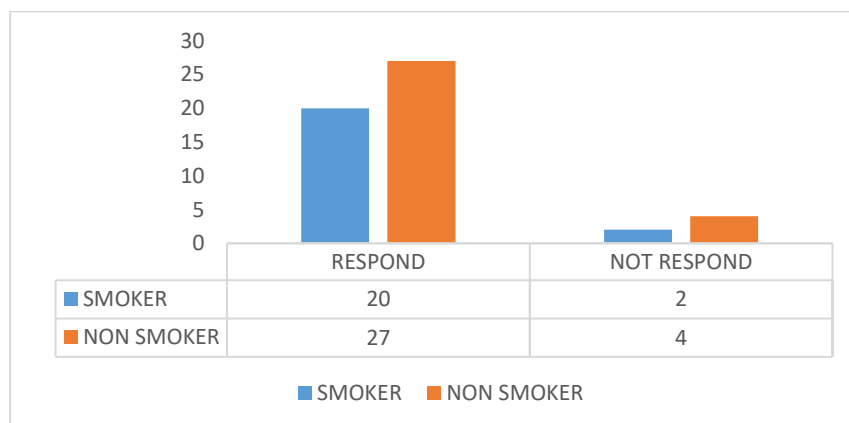


Figure 4: Smoker response to EECP, P-value = 0.67.

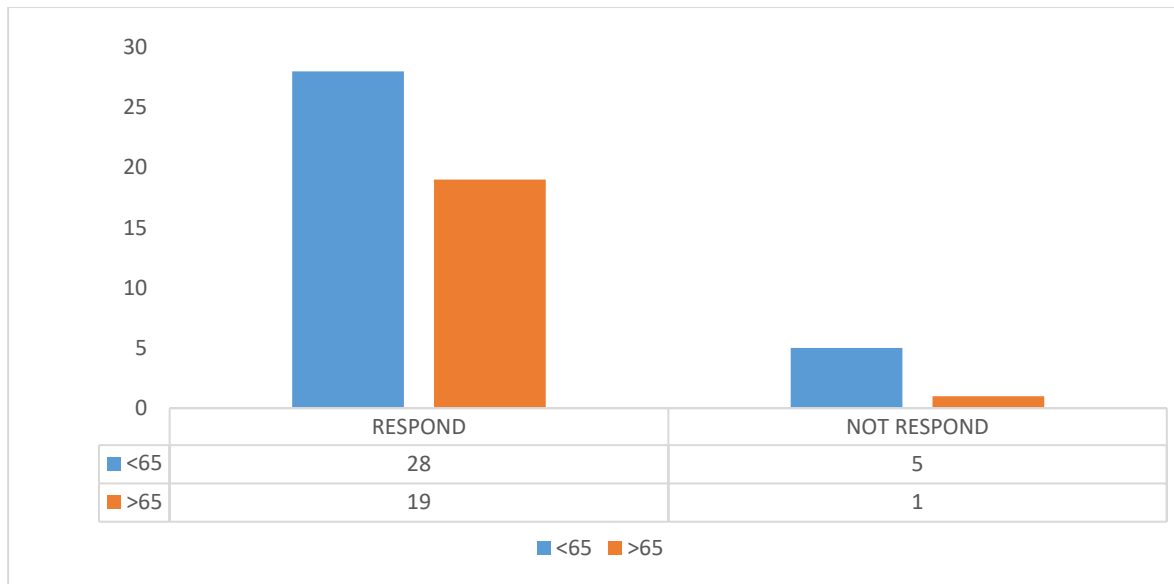


Figure 5: Age and response to EECP, P-value < 0.26.

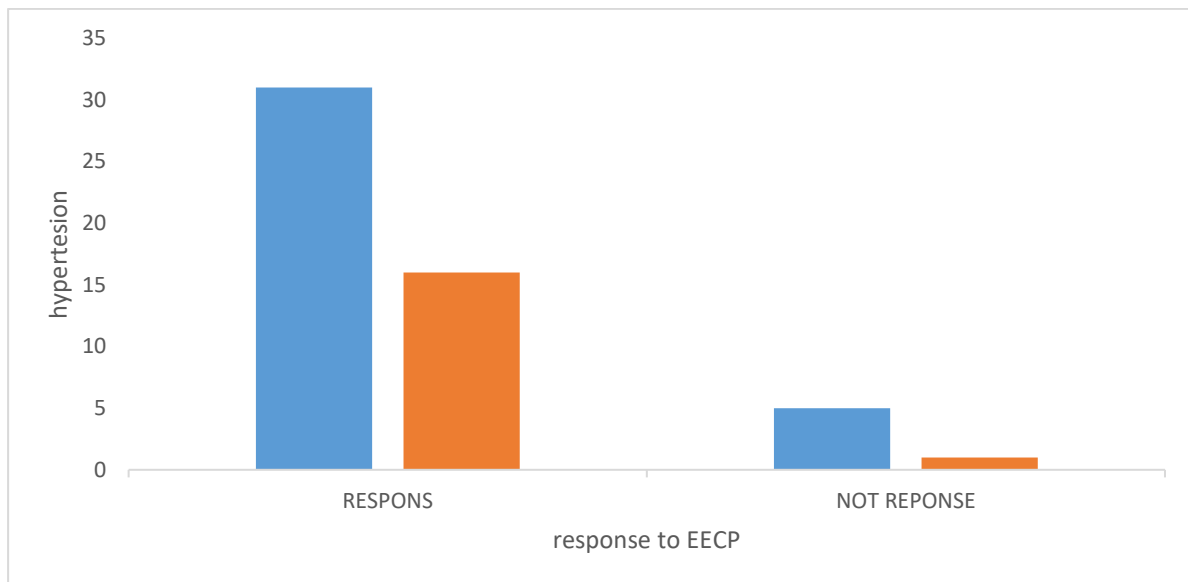


Figure 6: Hypertension and response to EECP, P-value < 0.4

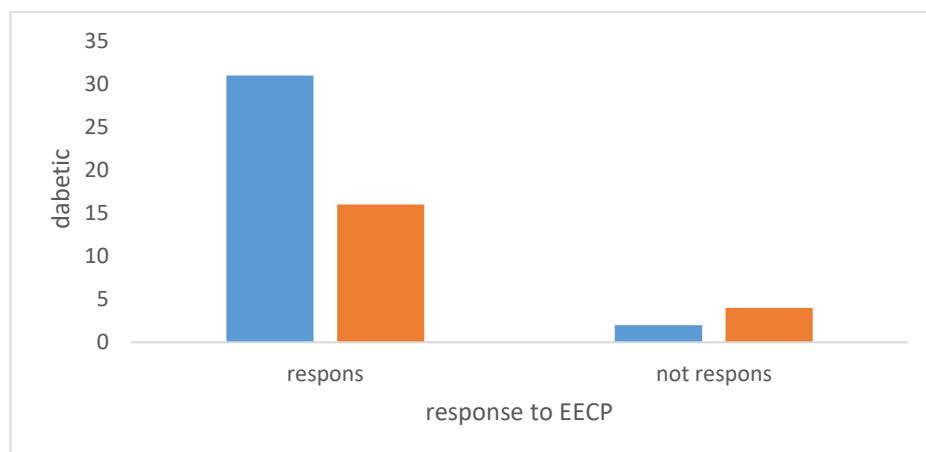


Figure 7: Diabetic and response to EECP, P-value < 0.12.

3. DISCUSSION

Most study participants had diffuse coronary artery disease confirmed via angiography. Some underwent PCI, while others were ineligible for the procedure. Given their persistent symptoms, they were referred for EECP therapy, which resulted in significant symptomatic relief, particularly in those classified as CCS3 and CCS4.

After completing EECP sessions, most patients improved to CCS1 or CCS2, with sustained benefits observed at one-year follow-up, consistent with previous studies. The improvements noted may not solely result from increased myocardial perfusion but also from secondary effects similar to those seen in physical rehabilitation.

Long-term benefits of EECP include enhanced endothelial function, angiogenesis, neurohormonal modulation, and exercise training effects. The International EECP Patient Registry has reported sustained symptom relief for up to five years post-treatment.

This study found no added benefit of PCI for three-vessel disease patients in terms of EECP response or echocardiographic improvement. Furthermore, both diabetic and non-diabetic patients exhibited significant symptomatic relief and improved quality of life post-EECP.

Conclusion:

1. PCI for single-vessel disease in three-vessel patients does not improve EECP response or echocardiographic outcomes.
2. EECP is a safe and effective treatment for select patients with refractory angina unresponsive to medical therapy or ineligible for intervention.
3. EECP provides long-lasting symptomatic relief, with benefits persisting for at least one year.

REFERENCES

- [1] Mannheimer C, Camici P, Chester MR, et al. The problem of chronic refractory angina; report from the 4e ESC Joint Study Group on the Treatment of Refractory Angina. *Eur Heart J* 2002;23:355–70. <https://doi.org/10.1053/euhj.2001.2706>; PMID: 11846493.
- [2] Povsic TJ, Broderick S, Anstrom KJ, et al. Predictors of longterm clinical endpoints in patients with refractory angina. *J Am Heart Assoc* 2015;e001287. Lawson WE, Hui JKC, Zheng ZS, Oster Z, Katz JP, Diggs P, Burger L, Cohn CD, Soroff HS, Cohn PE. Three-year sustained benefit from enhanced external counterpulsation in chronic angina pectoris. *Am J Cardiol* 1995;75:840-841.
- [3] Henry TD, Satran D, Hodges JS, et al. Long-term survival in patients with refractory angina. *Eur Heart J* 2013;34:2683–8. <https://doi.org/10.1093/eurheartj/ehf165>; PMID: 23671156.
- [4] Beck CS, Leighninger DS. Scientific basis for the surgical treatment of coronary artery disease. *J Am Med Assoc* 1955;159:1264–71. <https://doi.org/10.1001/jama.1955.02960300008003>; PMID: 13271060.
- [5] Werner GS, Hochadel M, Zeymer U, et al. Contemporary success and complication rates of percutaneous coronary intervention for chronic total coronary occlusions: results from the ALKK quality control registry of 2006. *Euro Intervention* 2010;6:361–6. <https://doi.org/10.4244/EIJV6I3A60>; PMID: 0884415.
- [6] Manchanda A, Soran O. Enhanced external counterpulsation and future directions: step beyond medical management for patients with angina and heart failure. *J Am Coll Cardiol* 2007;50:1523–31. [tps://doi.org/10.1016/j.jacc.2007.07.024](https://doi.org/10.1016/j.jacc.2007.07.024); PMID: 17936150.
- [7] Bonetti PO, Barsness GW, Keelan PC, et al. Enhanced external counterpulsation improves endothelial function in patients with symptomatic coronary artery disease. *J Am Coll Cardiol* 2003;41:1761–8. [https://doi.org/10.1016/S0735-1097\(03\)00329-2](https://doi.org/10.1016/S0735-1097(03)00329-2); PMID: 12767662.
- [8] Nichols WW, Estrada JC, Braith RW, et al. Enhanced external counterpulsation treatment improves arterial wall properties and wave reflection characteristics in patients with refractory angina. *J Am Coll Cardiol* 2006;48:1208–14. <https://doi.org/10.1016/j.jacc.2006.04.094>; PMID: 16979007 Lawson WE, Hui JC, Cohn PF. Long-term prognosis of patients with angina treated with enhanced external counter pulsation: fiveyears follow-up study. *Clin Cardiol* 2000;23:254-258.
- [9] Levenson J, Pernollet MG, Iliou MC, et al. Cyclic GMP release by acute enhanced external counterpulsation. *Am J Hypertens* 2006;19:867–72. <https://doi.org/10.1016/j.amjhyper.2006.01.003>; PMID: 16876689.
- [10] Buschmann EE, Utz W, Pagonas N, et al. Improvement of fractional flow reserve & collateral flow by treatment with external counterpulsation (Art.Net.-2 Trial). *Eur J Clin Invest* 2009;39:866–75. <https://doi.org/10.1111/j.1365-2362> PMID: 19572918.
- [11] Gloekler S, Meier P, de Marchi SF, et al. Coronary collateral growth by external pulsation: a randomised

- controlled trial. *Heart BrCardSoc*2010;96:202–7. <https://doi.org/10.1136/hrt.2009.184507>; PMID: 19897461.
- [12] Soran O, Kennard ED, Kfoury AG, Kelsey SF. IEPR Investigators. Two-year clinical outcomes after enhanced external counterpulsation (EECP) therapy in patients with refractory angina pectoris and left ventricular dysfunction (report from The International EECP Patient Registry). *Am J Cardiol*2006;97:17–20. <https://doi.org/10.1016/j.amjcard.2005.07.122>;
- [13] Loh PH, Cleland JGF, Louis AA, et al. Enhanced external counterpulsation in the treatment of chronic refractory angina: a long-term follow-up outcome from the International Enhanced External Counterpulsation Patient Registry. *Clin Cardiol*2008;31:159–64. <https://doi.org/10.1002/clc.20117>;
- [14] Lawson WE, Hui JCK, Kennard ED, Linnemeier G. IEPR-II Investigators. Enhanced external counterpulsation is costeffective in reducing hospital costs in refractory angina patients. *Clin Cardiol*2015;38:344–9. <https://doi.org/10.1002/clc.22395>; PMID: 25962616.
- [15] McGillion M, Arthur HM, Cook A, et al. Management of patients with refractory angina: Canadian Cardiovascular Society/Canadian Pain Society Joint Guidelines. *Can J Cardiol*. England: 2012 Canadian Cardiovascular Society. Published by Elsevier Inc;2012:S20-S41.
- [16] Henry TD, Satran D, Jolicœur EM. Treatment of refractory angina in patients not suitable for revascularization. *Nat Rev Cardiol*. 2014;11:78–95.
- [17] Manchanda A, Aggarwal A, Aggarwal N, et al. Management of refractory angina pectoris. *Cardiol J*. 2011;18:343–351.
- [18] Montalescot G, Sechtem U, Achenbach S, et al. 2013 esc guidelines on the management of stable coronary artery disease: the task force on the management of stable coronary artery disease of the European society of cardiology. *Eur Heart J*. 2013;34:2949–3003.
- [19] Soran O. Treatment options for refractory angina pectoris: enhanced external counterpulsation therapy. *Curr Treat Options Cardiovasc Med*. 2009;11:54–60.
- [20] Michaels AD, Linnemeier G, Soran O, et al. Two-year outcomes after enhanced external counterpulsation for stable angina pectoris (from the international eecp patient registry iepr). *Am J Cardiol*. 2004;93:461–464.
- [21] Yavari M, Montazeri HR. Effects of enhanced external counterpulsation on anginal symptoms and improvements in objective measures of myocardial ischaemia. *Cardiovasc J Afr*. 2007;18:154–156.
- [22] Kumar A, Aronow WS, Vadnerkar A, et al. Effect of enhanced external counterpulsation on clinical symptoms, quality of life, 6-minute walking distance, and echocardiographic measurements of left ventricular systolic and diastolic function after 35 days of treatment and at 1-year follow up in 47 patients with chronic refractory angina pectoris. *Am J Ther*. 2009;16:116–118.
- [23] Urano H, Ikeda H, Ueno T, et al. Enhanced external counterpulsation improves exercise tolerance, reduces exercise-induced myocardial ischemia and improves left ventricular diastolic filling in patients with coronary artery disease. *J Am Coll Cardiol*. 2001;93–99 United States.
- [24] Masuda D, Nohara R, Hirai T, et al. Enhanced external counterpulsation improved myocardial perfusion and coronary flow reserve in patients with chronic stable angina-evaluation by n-13-ammonia positron emission tomography. *Eur Heart J*. 2001;22:1451–1458.
- [25] Arora RR, Chou TM, Jain D, et al. The multicenter study of enhanced external counterpulsation (must-eeep): effect of eecp on exercise-induced myocardial ischemia and anginal episodes. *J Am Coll Cardiol*. 1999;33:1833–1840.
- [26] Akhtar M, Wu G-F, Du Z-M, et al. Effect of external counterpulsation on plasma nitric oxide and endothelin-1 levels. *Am J Cardiol*. 2006;98:28–30.
- [27] Sessa WC, Pritchard K, Seyedi N, et al. Chronic exercise in dogs increases coronary vascular nitric oxide production and endothelial cell nitric oxide synthase gene expression. *Circ Res*. 1994;74:349–353.
- [28] Flynn MS, Kern MJ, Donohue TJ, et al. Alterations of coronary collateral blood flow velocity during intraaortic balloon pumping. *Am J Cardiol*. 1993;71:1451–1455.
- [29] Kern MJ, Aguirre FV, Tatineni S, et al. Enhanced coronary blood flow velocity during intraaortic balloon counterpulsation in critically ill patients. *J Am Coll Cardiol*. 1993;21:359–368.
- [30] 30. Casey DP, Beck DT, Nichols WW, Conti CR, Choi CY, Khuddus MA, Braith RW. Effects of enhanced external counterpulsation on arterial stiffness and myocardial oxygen demand in patients with chronic angina pectoris. *The American journal of cardiology*. 2011; 07(10):1466–72. [PubMed: 21420062]
- [31] 31. Arora RR, Chou TM, Jain D, Fleishman B, Crawford L, Mckiernan T, Nesto RW. The multicenter study of enhanced external counter pulsation (MUST- EECP): effect of enhanced external counter pulsation on

exercise-induced myocardial ischemia and angina episodes. *J Am Coll Cardiol*1999;33:1833-1840.

- [32] Aroru RR, Chou TM, Jain D, Fleishman B, Crawford L, Mckiernan T, Nesto R, Ferrans CE, Keller S. Effects of enhanced external counter pulsation on health-related quality of life continue 12 months after treatment: a substudy of the multicenter study of enhanced external counter pulsation. *J Invest Med* 2002;50:25-32.
 - [33] Lawson WE, Hui JKC, Zheng ZS, Oster Z, Katz JP, Diggs P, Burger L, Cohn CD, Soroff HS, Cohn PE. Three-year sustained benefit from enhanced external counterpulsation in chronic angina pectoris. *Am J Cardiol*1995;75:840-841.
 - [34] Lawson WE, Hui JC, Cohn PF. Long-term prognosis of patients with angina treated with enhanced external counter pulsation: fiveyears follow-up study. *Clin Cardiol*2000;23:254-258.
 - [35] Pettersson T, Bondesson S, Cojocaru D, Ohlsson O, Wackenfors A, Edvinsson L. One-year follow-up of patients with refractory angina pectoris treated with enhanced external counter pulsation. *BMC Cardiovasc Disord*2006;6:28.
 - [36] Michaels AD, Raisinghani A, Soran O, De lame PA, Lemaire ML, Kligfield P, Watson DD, Conti CR, Beller G. The effects of enhanced external counter pulsation on myocardial perfusion in patients with stable angina: a multicenter radionuclide study. *Am Heart J* 2005;150:1066-1073.
 - [37] Masuda D, Nohara R, Hirai T, Kataoka K, Chen LG, Hosokawa R, Inubushi M, Tadamura E, Fujita M, Sasayama S. Enhanced external counterpulsation improved myocardial perfusion and coronary flow reserve in patients with chronic stable angina; evaluation by(13)N-ammonia positron emission tomography. *Eur Heart J* 2001;22:1451-1458.
 - [38] Bonetti PO, Holmes DR, Lerman A, Barsness GW, Jr. Enhanced external counterpulsation for ischemic heart disease: what's behind the curtain? *J Am Coll Cardiol*2003;41:1918-1925.
 - [39] Michael AD, Linnemeier G, SoranO, Kelsey SF, Kennard ED. Two-years out comes after enhanced external counter pulsation for stable angina pectoris (from the international EECp patient Registry (IEPR)). *Am J Cardiol*2004;93:461-464.
 - [40] Lawson WE, Hui JC, Cohn PF. Long-term prognosis of patients with angina treated with enhanced external counter pulsation: fiveyears follow-up study. *Clin Cardiol*2000;23:254-258.
-