http://bjas.bu.edu.eg

Serum Endocan Level, Mean Platelet Volume and Neutrophil to Lymphocyte Ratio in Patients with Erectile Dysfunction: a narrative review Mohammed.H.Shaath¹, Samy.H.Mohamed¹, Neveen .E.Sorour¹, Asmaa.A.El-Fallah² and Karem.T.Khalil¹

¹Dermatology, Venereology and Andrology Dept., Faculty of Medicine, Benha University ²Clinical and Chemical Pathology Dept., Faculty of Medicine, Benha University **E-mail:** m.h.shaath@gmail.com

ABSTRACT

Background: ED is disease with several potential causes, including high blood pressure and diabetes. Serum endocan, MPV, and NLR are only a few of the biomarkers suggested to measure the severity of this inflammatory response. The goals of this study are to examine the relationship between erectile dysfunction and serum endocan level, mean platelet volume, and neutrophil-to-lymphocyte ratio. Serum endocan Level, Mean Platelet Volume, and Neutrophil to Lymphocyte Ratio in patients with erectile dysfunction were retrieved from the Medline databases (Pub Med and Medscape) until the year 2022. Research Studies were chosen after being objectively evaluated by two reviewers. If they met any of the following criteria, we considered them for inclusion: First, it must be written in English. 2. Appearing in journals with a strict peer review process, Third, explain why serum endocan, MCV, and the neutrophil-to-lymphocyte ratio are important in ED patients. Extraction of Data: Research Studies were not included if they did not meet the inclusion criteria. Ethical permission, eligibility criteria, controls, information, and well-defined evaluation measures were all factors in determining the study's quality. Our concerned research results were captured by independently abstracting data utilising a data collecting form from each qualifying study. We conclude that Endocan, MPV, and NLR are indicators implicated in the pathogenesis of ED in individuals with diabetes and hypertension. Endocan and MPV are markers of erectile dysfunction severity in people with diabetes and hypertension.

Keywords: endocan, erectile dysfunction, mean platelet volume, neutrophil/lymphocyte ratio.

1. Introduction

Erectile penile erection necessary for optimal sexual performance for at least 6 months is a hallmark of erectile dysfunction (ED) (Wang et al., 2019) 1. ED has a complex set of causes, some of which are long-term conditions including hypertension (HT) and diabetes mellitus (DM) (Akdemir et al., 2019) Second, vascular illnesses and other organic diseases are the leading causes of ED in elderly men (Sambel et al., 2018) Third, endocan is a new soluble dermatan sulphate proteoglycan that originates from the endothelium (Bessa et al., 2020) A key player in inflammatory processes, it is also engaged in endothelial cell rearrangement, adhesion, migration, proliferation, and angiogenesis (Nalewajska et al., 2020) Five, endocan can foresee endothelial dysfunction (Onuk et al., 2018) Six, illnesses including diabetes and hypertension have been linked to abnormal endocan levels. Endothelial activation and neovascularization are prominent pathophysiological changes associated with inflammation and tumour growth, and a rise in tissue expression or serum levels of endocan reflects these changes (Wang et al., 2019) 1. Platelet activation and functional changes may be predicted using mean platelet volume. The levels of MPV tend to rise in several disorders, including hypertension, diabetes, and others.

Kalliopi et al. (2019). 7. Problems of ED often manifest two to five years before to the onset of cardiovascular symptoms. The cavernous arteries are only around 1-2 mm in diameter, whereas the coronary arteries are about 3-4 mm. An early sign of cardiovascular disease, vasculogenic ED has been proposed (Crafa et al., 2020) Furthermore, the ratio of neutrophils to lymphocytes might reveal systemic inflammation (González et al., 2018). 9 Atherosclerosis formation and progression are both heavily influenced by chronic arterial wall inflammation (Gencer et al., 2021) Atherosclerotic risk factors, like diabetes and hypertension, may influence the NLR 10. (Moghanjoughi et al., 2022) Eleven. The NLR may prevent ED and cardiovascular disease by blocking many pathophysiological pathways that contribute to these conditions, including moderate systemic inflammation. Multiple inflammatory mediators, Myeloperoxidase (MPO) and reactive oxygen species (ROS), are produced and secreted by neutrophils and may contribute to cardiac and non-myocardial tissue damage (Nishida et al., 2017) 12.

print: ISSN 2356-9751

online: ISSN 2356-976x

2. Materials and methods

Data Sources: By Serum endocan Level, Mean Platelet Volume, and Neutrophil to Lymphocyte Ratio in Patients with Erectile Dysfunction, Searching and Reviewing Medline Databases (Pub Med and Medscape) Available Until 2022.

Choice of Studies: Each study that was considered for inclusion was reviewed separately. If they met any of the following criteria, we considered them for inclusion: First, it must be written in English. 2. Appearing in journals with a strict peer review process. Third, explain why serum endocan, MCV, and the neutrophil-to-lymphocyte ratio are important in ED patients.

Data Mining: Studies were not included if they did not meet the inclusion criteria. Ethical permission, eligibility criteria, controls, information, and well-defined evaluation measures were all factors in determining the study's quality. Our concerned research results were captured by independently abstracting data utilising a data collecting form from each qualifying study.

Third, a literature review:

Male sexual dysfunction in people with diabetes

Endothelial dysfunction, accumulation and increased advanced glycation end products (AGEs), impaired nitric oxide (NO) synthesis, increased endothelin B receptor binding sites, oxidative stress, autonomic neuropathy, the Rho/Rho kinase regulation signalling pathway, impaired cyclic guanosine monophosphate, and impaired cyclic guanosine monophosphate are all involved in the pathophysiology of ED in (cGMP). Alterations in the cavernous bodies' smooth muscle's structure and function, as well as hormonal shifts, also play a role (Ibrahim, 2019) Hypertensive patients with erectile dysfunction

In hypertensive individuals, erectile dysfunction may be caused by an aberrant response to contractile stimuli like AngII and ET-1, or a reduced response to NO and other stimuli that induce relaxation of penile vascular smooth muscle. Reduced reactivity to vasodilator mediators and greater sensitivity to vasoconstrictor molecules are symptoms common to both ED and endothelial dysfunction (Gokhan et al., 2018) 14.

Potential inflammatory indicators in vascular disease include endocan, macrophage polarisation molecule 2, and neutrophil elastase-like receptor, and inflammation is a key player in the development of vascular endothelial dysfunction and atherosclerosis (Pujani et al., 2022)

15. Patients with diabetes and hypertension have been shown to have a negative association between serum endocan and ED severity as measured by IIEF-5, as described by Akarsu et al. (2018)16. Studies

by Balta et al. (2015)19, Onuk et al. (2018)6, and Wang et al. (2015)20 and Elkamshoushi et al. (2018)6 all found similar things.

El Taieb et al. (2019)26, Otunctemur et al. (2015)25, Tangal et al. (2020),21 Bayraktar et al. (2017)22, Sönmez et al. (2017)23, Aldemir et al. (2016)24, and Otunctemur et al. (2015)25 all observed substantially higher MPV levels in vasculogenic ED patients than in controls (arterial, venous, or mixed).

No statistically significant variations in MPV values were found between ED patients and healthy controls, as reported by Sentürk et al. (2018)27.

Several studies have shown a negative relationship between MPV and IIEF score, including one by Tangal et al. (2020), another by El Taieb et al. (2019), and still another by Culha et al. The severity of ED was not linked to MPV by either Ciftci et al. (2013)29 or Sentürk et al. (2018)27.

Numerous studies, including those by Liao et al. (2021)30, Sambel et al. (2018)31, Karabakan et al. (2019)18, Bozkurt et al. (2020), Aslan et al. (201933, and Alcaidinho et al. (2021)34, have shown elevated NLR in ED and drawn the conclusion that NLR is a reliable predictor of ED.

While we found a correlation between NLR and ED, Akbas et al. (2016)35 found no such link.

According to Sambel et al. (2018)31 and Ventimiglia et al. (2018)36, NLR levels are inversely linked with IIEF-5.ED severity.

4. Conclusion

Our findings suggest that serum endocan, MPV, and NLR can be used to evaluate ED severity.

References

- [1] Wang X, Guo L, Xiao ZY, Guan Y, Zhang JY, Li M, et al. Mean platelet volume might be an effective indicator of arterial erectile dysfunction. Asian J Androl 2019; 21(1): 62-66.
- [2] Akdemir A, Karabakan M, Aktas BK, Bozkurt A, Ozgur EG, Akdogan N et al. Visceral adiposity index is useful for evaluating obesity effect on erectile dysfunction. Andrologia 2019; 51(6): e13282.
- [3] Sambel M, Kilic M, Demirbas M, Onen E, Oner S, Erdogan A, et al. Relationship between erectile dysfunction and the neutrophil to lymphocyte and platelet to lymphocyte ratios. Int J Impot Res 2018; 30(1): 27-35.

- [4] Bessa J, Teixeira A, Couto M and Sousa T. Endocan: A novel biomarker for risk stratification, prognosis and therapeutic monitoring in human cardiovascular and renal diseases. Clin Chim Acta 2020; 509: 310-335.
- [5] Nalewajska M. Gurazda Marchelek-Myśliwiec M, Pawlik A and Dziedziejko V. The role of endocan in selected kidney diseases. Int. J. Mol. Sci. 2020; 21(17): 6119.
- [6] Onuk O, Arslan B, Gezmis T, Cetin B, Göv T, Yazıcı G, et al. Is the plasma endocan level a reliable predictor for the severity of erectile dysfunction? Int Urol Nephrol. 2018; 50(9): 1577-1582.
- [7] Kalliopi P, Theano P, Dimitri PM and Nikolaos P. Mean platelet volume and coronary artery disease. Current Opinion in Cardiology 2019; 34(4): 390-398.
- [8] Crafa A, Condorelli R, Mongioì L, Cannarella R, Barbagallo F, Aversa A, et al. Mean platelet volume as a marker of vasculogenic erectile dysfunction and future cardiovascular risk. Journal of Clinical Medicine 2020; 9(8): 2513.
- [9] González Y, Sánchez M, Dorca J and Santos S. Defining the role of neutrophil-to-lymphocyte ratio in COPD. Int J Chron Obstruct Pulmon Dis. 2018;(13): 3651-3662.
- [10] Gencer S, Evans B, Vorst E, Yvonne Döring and Weber C. Inflammatory Chemokines in Atherosclerosis. Cells 2021; 10(2): 226.
- [11] Moghanjoughi P, Neshat S, Rezaei A and Ghahdarijani K. Is the neutrophilto-lymphocyte ratio an exceptional indicator for metabolic syndrome disease and outcomes? Endocrine Practice 2022; 28(3): 342-348.
- [12] Nishida Y, Hosomi S, Yamagami H, Yukawa T, Otani K, Nagami Y, et al. Neutrophil to lymphocyte ratio for predicting loss of response to infliximab in ulcerative colitis. PLoS ONE 2017; 12(1): e0169845.
- [13] Ibrahim AM. Erectile dysfunction in diabetes. Inter Journal of Innovative Studies in Medical Sciences 2019; 3(1): 13-14.
- [14] Gokhan K, Ecem S, Didem O and Serap G. Management of erectile dysfunction: An under- recognition of hypertension. Current Pharmaceutical Design 2018; 24(30): 3506-3519.

- [15] Pujani M, Agarwal C, Chauhan V, Agarwal S, Passi S and Singh K, et al. Platelet parameters, neutrophillymphocyte ratio, platelet lymphocyte ratio, red cell distribution width: can they serve as biomarkers in evaluation of severity of psoriasis? Journal of applied hematology 2022; 13(2): 95-102.
- [16] Akarsu M, Atalay HA, Canat L, Ozcan M, Arman Y, Aydın S, et al. Endocan is markedly overexpressed severe erectile dysfunction. Andrologia 2018; 50(3): e12912.
- [17] El-kamshoushi M. Hassan M. El-Abd M. Hassan Z and Maher A. Serum endocan as a predictive biomarker of cardiovascular risk in erectile dysfunction patients. Andrologia 2018; 50(10): e13113.
- [18] Karabakan M, Bozkurt A, Akdemir S, Gunay M and Keskin E. Significance of serum endothelial cell specific molecule-1 (Endocan) level patients with erectile dysfunction: a pilot study. International Journal of Impotence Research 2017; 29: 175-178.
- [19] Balta S, Mikhaildis DP, Demirkol S, Celik T, Ozturk C and Iyisoy A. Endocan and atherosclerosis. Angiology 2015; 66(5): 490.
- [20] Wang X, Yang W, Luo T, Wang J and Jing Y. Serum endocan levels are correlated with the presence and severity of coronary artery disease in patients with hypertension. Genet Test Biomarkers Mol 2015 19(3):124-127.
- [21] Tangal S, Ozayar A, Ener K, Gokce MI and Haliloglu AH. Does mean platelet volume (MPV) have a role in evaluation of erectile dysfunction and its severity? Rev Int Androl. 2020; 18(1): 1-6.
- [22] Bayraktar Z and Albayrak S. Blood platelet activity in men with vasculogenic erectile dysfunction. Archivio Italiano di Urologia e Andrologia 2017; 89(1): 51-54.
- [23] Sönmez MG, Göğer YE, Sönmez LÖ, Aydın A, Balasar M and Kara C. Can eosinophil count, platelet count, and mean platelet volume be a positive predictive factor in penile arteriogenic erectile dysfunction etiopathogenesis? American Journal of Men's Health 2017; 11(3): 678-683.

- [24] Aldemir M, Akdemir F, Okulu E, Ener K, Ozayar A and Gudeloglu A. Evaluation of blood platelet count and function in patients with erectile dysfunction. Andrologia 2016; 48(2): 189-192.
- [25] Otunctemur A, Bozkurt M, Besiroglu H, Polat EC, Ozcan L and Ozbek E. Erectile dysfunction is positively correlated with mean platelet volume and platelet count, but not with eosinophil count in peripheral blood. Urol J. 2015; 12(5): 2347-52.
- [26] El-Taieb MA, Hegazy EM, Maklad SM and Khairy R. Platelet Indices as a marker for early prediction of erectile dysfunction in diabetic patients. Andrologia 2019; 51(1): e13163.
- [27] Sentürk AB, Yaytokgil M, Yilmaz AH, Ekici M, Aydin C, Demir E and Cakiroglu B. Can platelets be the early biomarkers of erectile dysfunction? J Pak Med Assoc 2018; 68(4): 515-518.
- [28] Culha MG, Atalay HA, Canat HL, Alkan I, Ozbir S, Can O, et al. The relationship between erectile dysfunction severity, mean platelet volume and vitamin D levels. The Aging Male 2020; 23(3): 173-178.
- [29] Ciftci, H, Yeni E, Demir M, Yagmur I, Gümü K, Celik H, Savas M, Gulum M. Can the mean platelet volume be a risk factor for vasculogenic erectile dysfunction? World J Men's Health 2013; 31(3): 215-219.
- [30] Liao Z, Tang Y, Li X and Li D. The relationship between hematologic parameters and erectile dysfunction. Sex Med. 2021; 9(4): 100401.

- [31] Sambel M, Kilic M, Demirbas M, Onen E, Oner S, Erdogan A, et al. Relationship between erectile dysfunction and the neutrophil to lymphocyte and platelet to lymphocyte ratios. Int J Impot Res. 2018; 30(1): 27-35.
- [32] Bozkurt E, Gumus A and Koban Y. Can serum endocan level predict stage of diabetic retinopathy? Retina-Vitreus 2020; 29(4): 318-23.
- [33] Aslan A, Kaya Y, Cirakoglu A, Benli E, Demir EY and Calgin MK. Neutrophil-Lymphocyte Ratio could be a marker for erectile dysfunction. Urol J. 2019; 16(2): 216-220.
- [34] Alcaidinho A, Nguyen D, Delouya G, Zorn KC, Saad F, Taussky D and Liberman D. The interaction between inflammation, urinary symptoms and erectile dysfunction in early-stage prostate cancer treated with brachytherapy. Andrologia 2021; 53(6): e14070.
- [35] Akbas A, Gulpınar MT, Sancak EB, Gunes M, Ucar M, Altok M, et al. The relationship between platelet—lymphocyte ratio and severity of erectile dysfunction. Kaohsiung J Med Sci 2016; 32(2): 91-95.
- [36] Ventimiglia E, Cazzaniga W, Pederzoli F, Frego N, Chierigo F, Capogrosso P, et al. The role of neutrophil-to lymphocyte ratio in men with erectile dysfunction-preliminary findings of a real-life cross-sectional study. Andrology 2018; 6(4): 559-563.