

The Effect of Dapagliflozin in Improving Ejection Fraction

Gurindapalli Tejaswi, Danduprolu Dhana Lakshmi, Kandula Rohith Kumar,

Dr.Shaik Shakeela , Dr.Anurag Polavarapu

Pharm D, Hindu College of Pharmacy

Pharm D, Hindu college of pharmacy;

Pharm D, Hindu college of pharmacy

Pharm D, Assistant professor, Hindu college of pharmacy

MD,DM, Interventional cardiologist, Director-Endo vascular Intervention. Lalitha super specialties Hospital.

Date of Submission: 01-06-2025

Date of Acceptance: 10-06-2025

ABSTRACT

This project is about to evaluate the effect of Dapagliflozin in improving ejection fraction and observe the effect of dapagliflozin in hospitalization of cardiovascular diseases like Heart failure. It was a Prospective observational study conducted from October 2023 to April 2024 at Lalitha Super Specialty Hospital Guntur. A sample size of 60 patients were taken, all the necessary and relevant data was collected from the patient case notes, treatment charts and laboratory reports.

Heart failure is a serious, progressive condition that occurs when heart muscle can't able to pump enough blood to meet body needs for blood and oxygen. Between 1990 and 2000, the number of deaths in India from cardiovascular illnesses increased from 2.26 million to 4.7 million annually. The amount of blood that the left ventricle pumps out with each contraction is measured and represented as the ejection fraction. Guidelines strongly advise using SGLT2 inhibitors to treat individuals with HF who have a decreased ejection fraction.

In our study among 60 patients during and after therapy the considerable changes have been seen in cardiac function through ECHO, ECG and symptomatic relief. The positive aspects towards the EF and symptomatic relief have been observed. The patients who are admitted with heart failure have shown positive response towards SGLT2 inhibitors.

KEYWORDS: Heart failure, Hypertension, SGLT2 inhibitors, symptomatic relief, ejection fraction.

I. INTRODUCTION

Heart failure is a long-term condition that gets worse over time. Although the name sounds like heart has stopped working, heart failure means your heart isn't able to pump blood as well as it should. Heart failure continues to be a prevalent

and relevant clinical problem. The current prevalence of heart failure in the adult population is estimated to be 1-2%. Approximately 50% of these cases can be attributed to heart failure with reduced ejection fraction, referring to a left ventricular ejection fraction of 40% or less⁽¹⁾.

Traditionally, guideline direct medical therapy (GDMT) for HFrEF has been focused on renin angiotensin aldosterone system and sympathetic pathways through agents including: ACE inhibitors, ARBs, ARNI, beta blockers and mineralocorticoids. More recently, sodium glucose cotransporter 2 (SGLT2) inhibitors have shown further reduction in hospitalization of heart failure, cardiovascular events and mortality, especially for HFrEF patients.

EF is a measurement of the percentage of blood leaving the heart each time it squeezes. 2D-Echo, CMR, CT, cardiac catheterization are most commonly used methods for EF estimation. Among these methods, echocardiogram is non-invasive and mostly used of its short-time and bedside properties along with its ability to create 2 and 3 dimensional images, which provide useful information about shape, size and other volumetric information.

• Ejection fraction is calculated as:

$$EF = \frac{SV}{EDV} = \frac{EDV - ESV}{EDV}$$

Where,

EF = ejection fraction.

SV = stroke volume.

EDV = end diastolic volume.

ESV = end systolic volume.

DAPAGLIFLOZIN

Dapagliflozin is a sodium glucose cotransporter 2(SGLT2) inhibitor. The usually recommended daily dose of dapagliflozin is 5mg once daily at afternoon, in addition to recommended therapy. SGLT2 inhibitors are originally developed as glucose lowering agents in patients with type 2 diabetes mellitus. they selectively and reversibly inhibit SGLT2, which is selectively express in PCTs of kidney and responsible for reabsorption of glucose from the glomerular filtrate back into the circulation as shown in the fig no:1

The mechanism underlying the clinical benefit in heart failure is not well known. Different studies suggest that the reduction of hospitalization of heart failure could be explained by structural and functional improvement of the left ventricle.

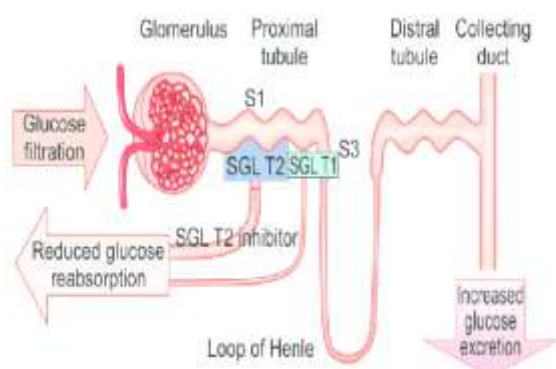


Fig.no.1 MECHANISM OF ACTION OF DAPAGLIFLOZIN

II. MATERIALS AND METHODS

Study design: A prospective observational study was carried out to assess the effect of Dapagliflozin in improving ejection fraction.

Study sample: The sample taken for the study was 60

Study criteria:

Inclusion criteria:

- Both male and female patients of age >18years.
- Patients with left ventricular ejection fraction <40%.
- Patients with heart failure with type-2 diabetes mellitus.

Exclusion criteria:

- Patients with type-1 diabetes mellitus.
- Patients with Chronic kidney Disease.
- Patients with LVEF >40%.
- Both male and female patients of age<18years.

III. SOURCE OF DATACOLLECTION

Data was collected from the patient case files. Additional information was collected from the patient by using an interviewer-administered questionnaire. If the participant was unable to understand the questions, he/she was explained in detail in the local language. The participants were encouraged to provide answers to questions without assistance from the family, to avoid bias

IV. RESULTS

A total of 60 patients were included in the study during the period from October to April. We have conducted an analysis on how dapagliflozin in increasing ejection fraction in cardiac patients who had decreased EF less 40 percentage.

TABLE.5.1 DIFFERENT AGE GROUPS

AGE INTERVAL	NO. OF PATINETS	PERCENTAGE
35-45	8	13%
46-55	15	25%
56-65	17	28%
66-75	18	30%
76-85	2	3%

Observation:From the table it is observed that, there are 8 patients within the age group 35-45, 15 patients within the 46-55, 17 patients within the 56-

65 age group, 18 patients within the 66-75 age group, 2 patients within the 76-85 age group.

GRAPH

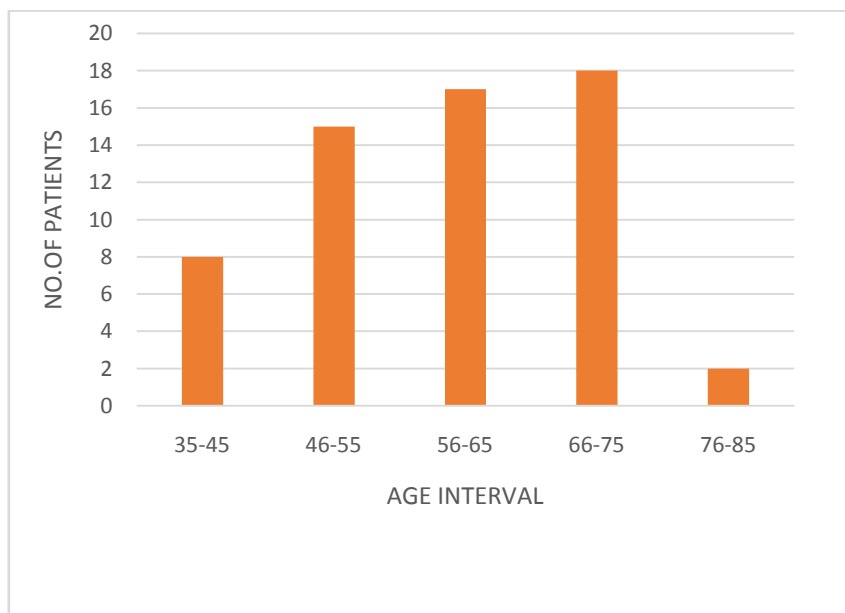


Fig.no 5.1 DIFFERENT AGE GROUP PATIENTS

TABLE.5.2 DISTRIBUTION OF GENDER DIFFERENTIATION

GENDER	NO. OF PATIENTS	PERCENTAGE
Male	39	65%
Female	21	35%

Observation: From the table we can observe that number of male patients are 39 and the female patient are 21 out of 60 patients

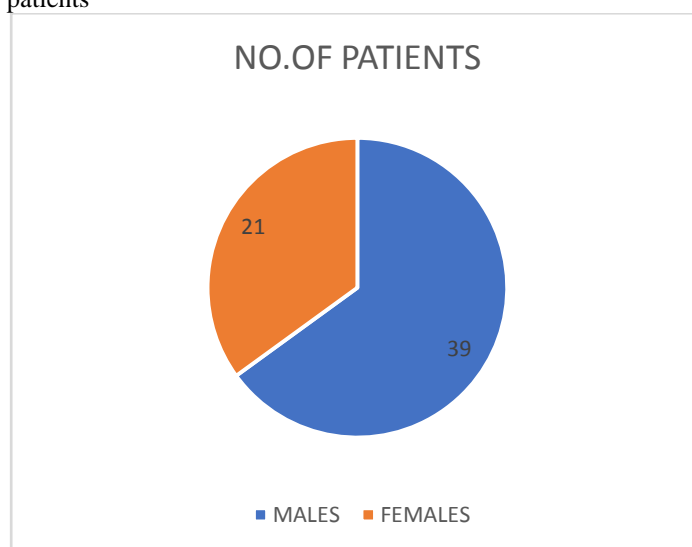


Fig no.5.2 REPRESENTATION OF GENDER

TABLE.5.3 TOTAL NUMBER OF PATINETS RECEIVING DAPAGLIFLOZIN

NO. OF PATIENTS	EF BEFORE TREATMENT	EF AFTER TREATMENT	INCREASE IN EF
1	40	50	10
2	25	48	23
3	35	48	13
4	30	43	13
5	35	56	21
6	35	50	15
7	38	50	12
8	30	48	18
9	32	48	16
10	36	56	20
11	40	50	10
12	34	49	15
13	30	45	15
14	28	40	12
15	34	47	13
16	35	50	15
17	30	50	20
18	35	42	7
19	38	48	10
20	20	45	25
21	32	45	13
22	25	39	14
23	25	35	10
24	28	38	10
25	25	45	20
26	29	40	11
27	20	45	25
28	35	50	15
29	30	45	15
30	32	42	10
31	30	43	13
32	40	55	15
33	32	43	11
34	32	45	13
35	30	45	15
36	30	50	20
37	34	46	12
38	40	50	10
39	35	49	14
40	35	52	17
41	30	46	16
42	38	43	5
43	40	56	16
44	35	52	17
45	35	49	14
46	25	40	15
47	28	42	14
48	25	43	14
49	25	40	18

50	40	50	15
51	38	52	10
52	30	48	14
53	40	52	18
54	30	49	12
55	34	50	19
56	28	47	16
57	35	50	19
58	20	43	15
59	20	45	23
60	33	48	25
Mean	31.31	46.62	15
STDEV	5.688	4.006	

Observation: A total of sixty samples were analysed. The ejection fraction is higher in patients who received dapagliflozin (46.62±4.006)

compared to those who did not receive it (31.31±5.688).

GRAPH

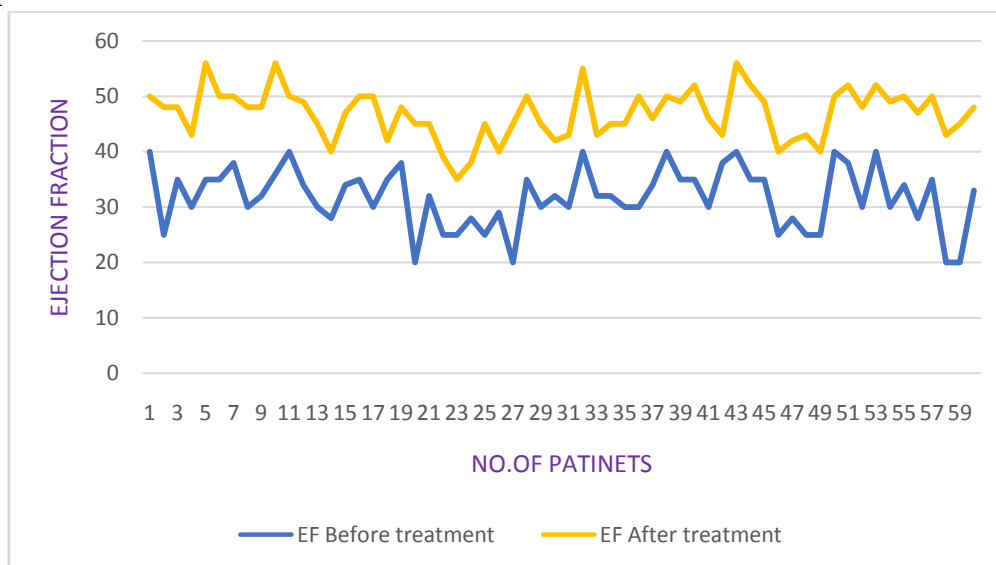


Fig no.5.3 EJECTION FRACTION TOTAL NO. OF PATIENTS

TABLE.5.8 GRBS LEVELS OF PATIENTS ADMINISTERING DAPAGLIFLOZIN

NO. OF PATIENTS	GRBS BEFORE TREATMENT (mg/dl)	GRBS AFTER TREATMENT (mg/dl)
1	136	116
2	220	200
3	156	136
4	165	145
5	148	128
6	246	226
7	178	158
8	134	114
9	234	214
10	112	92
11	223	203



12	130	110
13	200	180
14	157	137
15	167	147
16	200	180
17	240	220
18	234	214
19	143	123
20	156	136
21	226	126
22	146	134
23	164	112
24	154	132
25	174	110
26	130	126
27	146	124
28	144	120
29	140	115
30	135	115
31	162	108
32	128	150
33	170	130
34	160	130
35	180	144
36	164	130
37	150	127
38	147	123
39	143	123
40	135	115
41	158	128
42	164	134
43	172	152
44	126	106
45	134	114
46	142	122
47	132	122
48	120	100
49	146	123
50	164	143
51	172	153
52	175	154
53	175	155
54	175	156
55	160	138
56	140	129
57	130	110
58	146	126
59	148	128
60	160	145
MEAN	160.98	139.50
STDEV	31.767	30.547

OBSERVATION:It was observed that the patients who received dapagliflozin (n=60) are diabetic and there shows a decrease in serum glucose levels and

no patients has hypoglycemia even when other anti-diabetics were being administered.

GRAPH

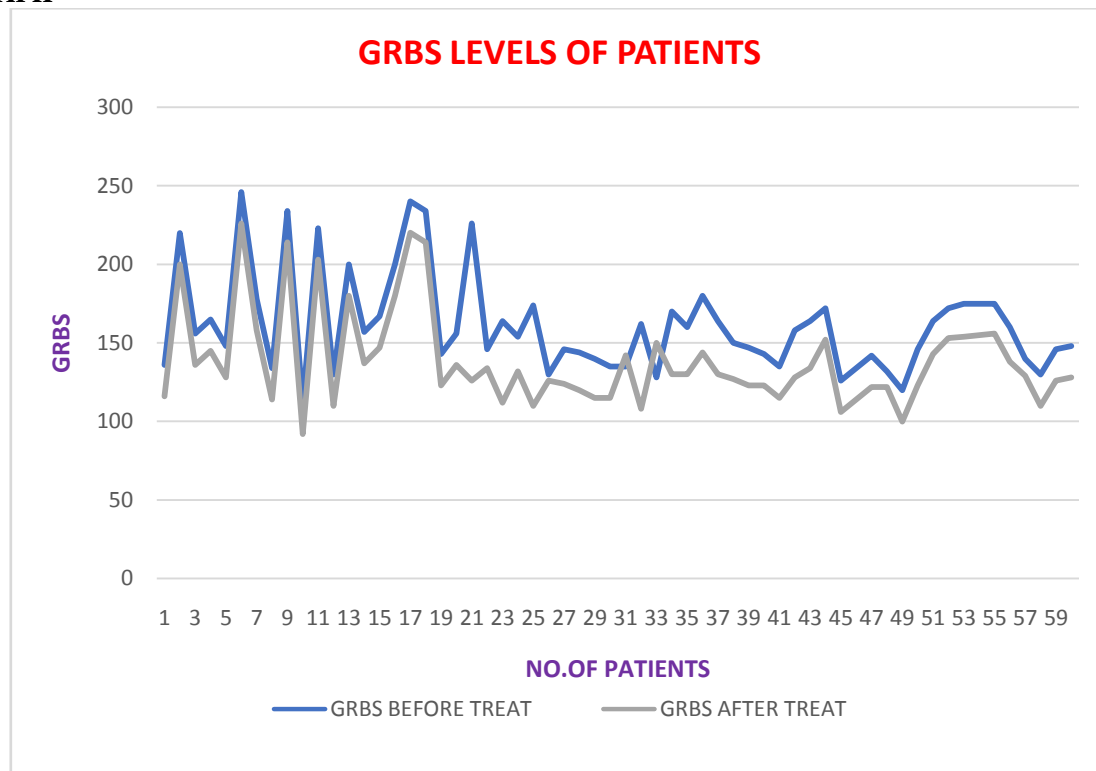


Fig no.5.7 GRBS LEVELS OF TOTAL NO. OF PATIENT

V. CONCLUSION

There are sixty individuals with type 2 diabetes mellitus and HF in our study. Most of the patients had a history of cardiovascular disease that spans approximately two to three years. Patients who are experiencing a reduced ejection fraction respond positively to dapagliflozin.

Improvements in EF and symptom alleviation are seen in HF patients taking dapagliflozin as opposed to those who did not get medication. It was observed that all the patients are diabetic in our study. When they received dapagliflozin there shows a decrease in serum glucose levels and no patients had hypoglycemia even when other anti-diabetics were being administered. Dapagliflozin administration results in higher EF, symptomatic alleviation, and reduced hospitalizations in Heart Failure patients.

REFERENCES:

- [1]. Mann, Zipes, Libby, Bonow 'BRAUNWALDS HEART DISEASE'- A

Text Book of Cardiovascular Medicine, 10th Edition, Chapter 3 page no- 613-615, chapter 38:799-801.

- [2]. Vateria C. Scanlon, Tina Sanders Essentials of Anatomy and Physiology 5th edition page no:219-317.
- [3]. BD Chaurasia Human Anatomy Regional and Applied Dissection and clinical volume-1 chapter 18 page no:239-255.
- [4]. Richard L. Drake A. Wayne VOGI Adam W. M. Mitchell- GRAYS Anatomy for students- 3rd edition chapter-3 page no:183-196.
- [5]. Andrea Fields MHA, RDCS- Mitral valve anatomy: name 5 components. Cardio serv net.
- [6]. Heart Failure(congestive heart failure)- MY Cleveland clinic.org.
- [7]. Marjonie Hecht on December 20, 2019, medically reviewed by Dr.Payalkohli,MD,FACC,types and stages of hypertension.

- [8]. KD Tripathi, Essentials of Medical Pharmacology, section 8, cardio vascular Drugs, anti-anginal and anti-ischemic drugs page no:539-541.
- [9]. Roger Walker and late Whittelsa, clinical pharmacy and therapeutics- 6th edition, section 8, cardiovascular disorder, acute myocardial infarction.
- [10]. Kristian Thygesen, Joseph. S et.al, white and the executive group on behalf of the joint European society of cardiology (ESC), American college of cardiology (ACC)/American heart association (AHA)world-mheart federation (WHF) task force for the universal definition of myocardial infarction(2018), published 24 Aug 2018, circulation, 2018, 138: E618-E651.
- [11]. Shervin Assari, cherlywisseh; Mohsen Bazargan obesity and polypharmacy among African American older adults: gender as the moderator and multi morbidity as the mediator 2019 june20.
- [12]. Access NCBI through the world wide web(www) molecular biology, 2016:3(1).
- [13]. Fulton M, Riley Allen E poly pharmacy in the elderly: Aliterature review journal of the American academy of nurse, practitioners.2005;17(4).
- [14]. paoloFabbietti, Giuseppina Di Stefano, rafellaMorresi et.al, Impact of potentially inappropriate medications and polypharmacy on 3- month readmission among older patients dischared from acute care hospital: A prospective study, 2018 Aug;30(8):977-984.
- [15]. K. Sembulingam, Essentials of Medical Physiology 9th edition, section-8page no:496-497.
- [16]. Cate Whittilesa, Karen Hodson, Clinical Pharmacy and Therapeutics, 6th edition, section-3chapter-45 page no: 784.
- [17]. Petrie, M.C., Verma, S., Docherty, K.F., Inzucchi, S.E., Anand, I. et. Al (2020). Effect of Dapagliflozin on Worsening Heart Failure and Cardiovascular Death in Patients with Heart Failure with and without Diabetes. JAMA, 323(14), 1353-1368.
- [18]. Peikert A, Martinez FA, Vaduganathan M, Claggett BL, Kulac IJ, et al. Efficacy and Safety of Dapagliflozin in Heart Failure With Mildly Reduced or Preserved Ejection Fraction According to Age: 2022 Aug 27. JAMA.
- [19]. Nassif ME, Windsor SL, Borlaug BA, Kitzman DW, Shah SJ, Tang F, Khariton Y. et. al, The SGLT2 inhibitor dapagliflozin in heart failure with preserved ejection fraction:2021 Oct 28.
- [20]. Solomon SD, McMurray JJV, Claggett B, de Boer RA, DeMets D, Hernandez AF, Inzucchi SE, et al., for the DELIVER Trial Committees and Investigators. Dapagliflozin in Heart Failure with Mildly Reduced or Preserved Ejection Fraction. N Engl J Med. 2022 Aug 27;387(12).
- [21]. Solomon SD, de Boer RA, DeMets D, Hernandez AF, Inzucchi SE, et. al, Dapagliflozin in heart failure with preserved and mildly reduced ejection fraction: rationale and design of the DELIVER trial. Eur J Heart Fail. 2021 Jul;23(7):1217-1225.
- [22]. Cunningham JW, Vaduganathan M, Claggett BL, Kulac IJ, et.al, Dapagliflozin in Patients Recently Hospitalized With Heart Failure and Mildly Reduced or Preserved Ejection Fraction 2022 Oct 4;80(14):1302-1310.
- [23]. Avgerinos, I., Liakos, A., Tsapas, A., &Bekiari, E. (2019). Cardiovascular Risk Reduction in Type 2 Diabetes: Therapeutic Potential of Dapagliflozin. Diabetes, Metabolic Syndrome and Obesity: Targets and Therapy, 12, 2549-2557.
- [24]. Alkagiet, S., &Tziomalos, K. (2020). Role of sodium-glucose co-transporter-2 inhibitors in the management of heart failure in patients with diabetes mellitus. World Journal of Diabetes, 11(5), 150-154.
- [25]. Butt, J. H., Docherty, K. F., Petrie, M. C., Schou, et. Al, (2021). Efficacy and Safety of Dapagliflozin in Men and Women With Heart Failure With Reduced Ejection Fraction: A Prespecified Analysis of the Dapagliflozin and Prevention of Adverse Outcomes in Heart Failure Trial. JAMA Cardiology, 6(6), 678-689.
- [26]. Nakagawa, Y., & Kuwahara, K. (2020). Sodium-Glucose Cotransporter-2 inhibitors are potential therapeutic agents for treatment of non-diabetic heart failure patients. Journal of Cardiology, 76(2), 123-131.

- [27]. McMurray, J. J. V., Solomon, S. D., Inzucchi, S. E., Køber, et.al (2019). Dapagliflozin in Patients with Heart Failure and Reduced Ejection Fraction. *New England Journal of Medicine*, 381(21), 1995-2008.
- [28]. Selvaraj, S., Vaduganathan, M., Claggett, B. L., Miao, Z. M., Fang, J. C., et.al (2023). Blood Pressure and Dapagliflozin in Heart Failure With Mildly Reduced or Preserved Ejection Fraction: DELIVER. *JACC: Heart Failure*, 11(1), 76-89.
- [29]. Kondo, T., Wang, X., Yang, M., Jhund, P. S., Claggett, B. L., Vaduganathan et.al (2023). Efficacy of Dapagliflozin According to Geographic Location of Patients With Heart Failure. *Journal of the American College of Cardiology*, 82(10), 1014-1026.
- [30]. Lipeng Mao, Dabei Cai, Boyuchi, Tingting xiao, et.al (2023). Dapagliflozin reduces risk of heart failure rehospitalization in diabetic acute myocardial infarction patients. 79(7).
- [31]. Jawad H Butt, Kieran F Docherty, Brian Claggett, et.al (2023). Dapagliflozin in black and white patients with heart failure across the ejection fraction spectrum. 14(5).
- [32]. Safia c hatur MD, Muthiah vaduganathan MD, Brian L. Claggett et.al (2023). Dapagliflozin in patients with heart failure and deterioration in renal function. 82(19):1854-1863.
- [33]. Smith M, Mytilinaios D. Diagrams, quizzes and worksheets of the heart. Last reviewed: October 30, 2023.
- [34]. Karki G. Human Heart-Gross structure and Anatomy. *Anatomy and Physiology*, Class 12, Transport in Animals, Zoology. March 18, 2017.
- [35]. Carpentier A, Adams DH, Filsoufi F. Carpentier's Reconstructive Valve Surgery: From Valve Analysis to Valve Reconstruction. Saunders Elsevier; 2010.
- [36]. Lorenzini M, Ricci C, Riccomi S, Abate F, Casalgrandi B, Quattrini B, Spagnoli G, Reggiani L, Capelli O. Integrated Care for Heart Failure in Primary Care. Published: May 11, 2016.
- [37]. Canadian Cardiovascular Society, Canadian Heart Failure Society. CCS/CHFS Heart Failure Guidelines Update: Defining a New Pharmacologic Standard of Care for Heart Failure with Reduced Ejection Fraction. *Can J Cardiol*. 2021 Apr;37(4):531-546.
- [38]. Boden, W. E., Marzilli, M., & Crea, F. (2023). Evolving Management Paradigm for Stable Ischemic Heart Disease Patients. *Journal of the American College of Cardiology*, 81(5), 505-514.