

**Supplemental Table 1: Devices Under Investigation for the Treatment of HFpEF and Clinical Trials**

Device category	Physiological target	Study Type	Primary Endpoints	RCT identifier	RCT status	Reference
Atrial shunts	Elevated LA pressure	Interventional, Est. Size: 150	Incidence of and time-to cardiovascular mortality or first non-fatal, ischemic stroke (12 months) -Total rate per patient year of HF admissions (~24 months) -Improvement in baseline KCCQ score (12 months)	NCT04632160	Not yet recruiting	88-97
		Observational, Est. Size: 100	-Device serious adverse events (30 days) -Improvement in KCCQ score, NYHA classification and EQ5D score (12 months)	NCT03191656	Recruiting	
		Interventional, Est. Size: 608	-Incidence of and time-to cardiovascular mortality or first non-fatal, ischemic stroke (12 months) -Total rate per patient year of HF admissions (~24 months) -Improvement in baseline KCCQ score (12 months)	NCT03088033	Recruiting	
		Interventional, Act. Size: 44	-Peri-procedural and post implant serious adverse events such as cardiovascular death or event, embolic stroke, kidney dysfunction (1 month) -Change in PCWP in supine exercise (1 month)	NCT02600234	Active	
		Interventional, Act. Size: 64	-Major adverse cardiac and cerebrovascular events (6 months)	NCT01913613	Completed	
		Interventional, Act. Size: 11	-Device serious adverse events (30 days)	NCT01570517	Completed	
		Interventional, Est. Size: 20	-Device and non-device associated major adverse cardiac and neurological events (6 months) -Percentage of patients successfully implanted with study device (3 months) -Percentage of patients implanted with right to left interatrial flow	NCT03838445	Recruiting	102-106
		Interventional, Est. Size: 500	-Device associated major adverse cardiac and neurological events (30 days) -Hierarchical composite of death, heart transplant or LVAD implantation, HF hospitalizations, and change in 6MWT	NCT03499236	Recruiting	

		Interventional, Act. Size: 16	-Device associated major adverse cardiac and neurological events (3 months)	NCT01965015	Completed	
		Interventional, Est. Size: 230	-Change in 6MWT (12 months)	NCT03751748	Not yet recruiting	111-115
		Observational, Est. Size: 100	- Device serious adverse events (12 months)	NCT04405583	Recruiting	
		Interventional, Est. Size: 30	- Incidence of Serious Adverse Device Effects (~3 months)	NCT03030274	Recruiting	
		Interventional, Est. Size: 30	- Incidence of Serious Adverse Device Effects (~3 months)	NCT03022851	Recruiting	
		Interventional, Est. Size: 40	- Incidence of device associated major adverse cardiac, cerebrovascular, or renal events (30 days)	NCT03523416	Recruiting	116,117
LV expanders	Diminished LV compliance	Interventional, Act. Size: 19	- Number of any adverse cardiac event (36 months)	NCT01347125	Terminated	120,121
		Interventional, Est. Size: 10	- Number of participants with all-cause mortality and serious adverse events (6 months) - Incidence of successful implant surgical procedure (~24 hours)	NCT02499601	Recruiting	122-124
Stimulators	Low baroreflex sensitivity	Observational, Est. Size: 70	-Changes in office cuff systolic blood pressure (6 months)	NCT02876042	Recruiting	130
	Diminished diastolic function	Interventional, Est. Size: 60	- Change in KCCQ score (24 weeks)	NCT03240237	Recruiting	145
	Mechanical desynchrony	Interventional, Est. Size: 10	- Change in diastolic reserve index (after 12 weeks of rate responsive pacing and again after 12 weeks of biventricular pacing) - Change in systolic reserve index (after 12 weeks of rate responsive pacing and again after 12 weeks of biventricular pacing)	NCT03338374	Recruiting	158,160-164,172
		Interventional, Est. Size: 20	- Change in MLHFQ score (~5 months) - Percent change in NTproBNP (~5 months)	NCT04546555	Recruiting	
		Interventional, Est. Size: 30	- Oxygen consumption at ventilatory anaerobic threshold (~4 weeks)	NCT02145351	Recruiting	
		Interventional, Act. Size: 22	-Number of adverse events while the Fusion Pacing download is active vs. inactive (~8 months) -Percentage of time the Fusion Pacing is active throughout a four-month follow-up period	NCT01045291	Completed	

		Interventional	N/R	NCT01618981	Completed	
MCS	Diminished cardiac output	-Clinical studies for the MCS devices mentioned are yet to be conducted” and have that centred across the columns and rows potentially				23 176,177 178-182 87 87

Reference numbers refer to the reference numbers in the main article.

6MWT= 6-minute walk test;

EQ5D = EuroQol five dimensional;

KCCQ= Kansas City Cardiomyopathy Questionnaire;

LVAD= Left ventricular assist device;

MLHFQ= Minnesota Living with Heart Failure Questionnaire;

NTproBNP= N-terminal pro-brain natriuretic peptide;

PCWP= Pulmonary capillary wedge pressure;

N/R= Not Reported