

**Supplementary Table 1: Overlap of events in cases considered in the analysis**

<b>Patients with MI</b>	
No other event	30 (34.5%)
One additional event	
With SCD	31 (35.6%)
With HFH	14 (16.1%)
Two additional events	
With HFH and SCD	8 (9.2%)
With stroke and SCD	3 (3.5%)
Three additional events	
With stroke, HFH and SCD	1 (1.1%)
<b>Patients with stroke</b>	
No other event	27 (60.0%)
One additional event	
With SCD	14 (31.1%)
With HFH	1 (2.2%)
With MI	1 (2.2%)
Two additional events	
With HFH and SCD	1 (2.2%)
With MI and SCD	1 (2.2%)
<b>Patients with SCD</b>	
No other event	152 (99.4%)
One additional event	
With stroke	1 (0.6%)
<b>Patients with HFH</b>	
No other event	128 (64.0%)
One additional event	
With SCD	62 (31.0%)
With stroke	4 (2.0%)
With MI	1 (0.5%)
Two additional events	
With MI and SCD	3 (1.5%)
With stroke and SCD	2 (1.0%)

**Supplementary Table 2: Patient characteristics according to MI/stroke, SCD, HF rehospitalization overall in the sex and age-matched COMMANDER HF population**

Characteristic	MI/ Stroke				SCD				HF Rehospitalization			
	Controls (n=127)	Cases (n=132)	SMD	P	Controls (n=149)	Cases (n=153)	SMD	P	Controls (n=179)	Cases (n=200)	SMD	P
Age (yrs)	68.1 ± 10.4	67.7 ± 10.6	4.40%	0.72	66.6 ± 10.1	66.7 ± 10.0	1.10%	0.92	67.6 ± 9.7	67.5 ± 10.2	1.40%	0.89
Study drug	60 (47.2%)	56 (42.1%)	10.40%	0.45	68 (45.6%)	73 (47.7%)	4.20%	0.73	82 (45.8%)	101 (50.5%)	9.40%	0.41
Female sex	38 (29.9%)	41 (30.8%)	2.00%	0.89	29 (19.5%)	31 (20.3%)	2.00%	0.89	43 (24.0%)	48 (24.0%)	0.10%	1
Race			24.90%	0.54			16.90%	0.43			15.80%	0.66
White	114 (89.8%)	120 (90.2%)			133 (89.3%)	143 (93.5%)			158 (88.3%)	170 (85.0%)		
Black	0 (0.0%)	2 (1.5%)			2 (1.3%)	1 (0.7%)			0 (0.0%)	2 (1.0%)		
Asian	12 (9.4%)	11 (8.3%)			12 (8.1%)	6 (3.9%)			18 (10.1%)	24 (12.0%)		
Other	1 (0.8%)	0 (0.0%)			2 (1.3%)	3 (2.0%)			3 (1.7%)	4 (2.0%)		
Region			17.30%	0.68			34.00%	0.058			33.60%	0.04
Eastern Europe	85 (66.9%)	88 (66.2%)			102 (68.5%)	123 (80.4%)			128 (71.5%)	121 (60.5%)		
North America	2 (1.6%)	1 (0.8%)			3 (2.0%)	2 (1.3%)			0 (0.0%)	6 (3.0%)		
Asia Pacific	12 (9.4%)	11 (8.3%)			12 (8.1%)	6 (3.9%)			18 (10.1%)	24 (12.0%)		
Latin America	13 (10.2%)	10 (7.5%)			22 (14.8%)	10 (6.5%)			15 (8.4%)	18 (9.0%)		
Western Europe	15 (11.8%)	23 (17.3%)			10 (6.7%)	12 (7.8%)			18 (10.1%)	31 (15.5%)		
BMI (kg/m <sup>2</sup> )	27.4 ± 5.1	27.9 ± 5.2	9.00%	0.47	27.6 ± 4.8	27.6 ± 5.0	0.40%	0.97	27.7 ± 5.1	28.2 ± 5.2	9.20%	0.37
eGFR (ml/min/1.73 m <sup>2</sup> )	<b>66.8 ± 21.7</b>	<b>65.0 ± 24.1</b>	<b>7.60%</b>	<b>0.54</b>	<b>72.7 ± 23.3</b>	<b>66.3 ± 22.3</b>	<b>28.10%</b>	<b>0.016</b>	<b>70.4 ± 23.2</b>	<b>63.2 ± 22.7</b>	<b>31.40%</b>	<b>0</b>
eGFR (ml/min/1.73 m <sup>2</sup> )			<b>18.40%</b>	<b>0.67</b>			<b>32.00%</b>	<b>0.036</b>			<b>27.40%</b>	<b>0.07</b>
<30 ml/min/1.73 m <sup>2</sup>	4 (3.1%)	8 (6.0%)			2 (1.3%)	3 (2.0%)			5 (2.8%)	14 (7.0%)		
30 to <60 ml/min/1.73 m <sup>2</sup>	55 (43.3%)	56 (42.1%)			41 (27.5%)	65 (42.5%)			61 (34.1%)	78 (39.0%)		
60 to <90 ml/min/1.73 m <sup>2</sup>	50 (39.4%)	54 (40.6%)			73 (49.0%)	61 (39.9%)			81 (45.3%)	86 (43.0%)		
>=90 ml/min/1.73 m <sup>2</sup>	18 (14.2%)	15 (11.3%)			33 (22.1%)	24 (15.7%)			32 (17.9%)	22 (11.0%)		
BNP level (pg/ml)	924.7 ± 839.0	860.8 ± 557.3	8.70%	0.79	856.9 ± 720.3	1045.6 ± 861.6	24.10%	0.38	859.4 ± 831.9	929.3 ± 714.1	9.00%	0.7
Log2 BNP (pg/ml)												
BNP rank (pg/ml)	19.2 ± 12.8	21.2 ± 9.3	18.00%	0.57	26.1 ± 16.3	30.6 ± 15.6	28.00%	0.3	33.5 ± 21.4	38.4 ± 19.9	23.90%	0.31
NT-proBNP (pg/ml)	5966 ± 4396 ± 4464	557.3	17.80%	0.31	5209 ± 7171	6835 ±	18.10%	0.25	6312 ± 7619	5242 ± 5229	16.40%	0.22

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Log2 NT-proBNP (pg/ml)												
NT-proBNP rank	74.6 ± 41.2	72.3 ± 43.8	5.50%	0.74	8.0 ± 1.0	8.2 ± 1.0	22.20%	0.14	109.4 ± 70.0	114.5 ± 58.9	7.90%	0.55
D-dimer level (mg/L)	545.6 ± 614.7	801.6 ± 1456.0	22.70%	0.093	634.7 ± 1002.2	698.5 ± 742.3	7.30%	0.53	704.4 ± 1225.3	712.2 ± 860.7	0.70%	0.94
Log2 D-dimer	5.9 ± 0.8	6.2 ± 0.9	30.90%	0.015	6.0 ± 0.9	6.2 ± 0.8	20.60%	0.075				
D-dimer rank									172.2 ± 116.6	205.9 ± 100.4	31.10%	0.00 3
<b>Ejection fraction (%)</b>	<b>33.5 ± 6.5</b>	<b>32.6 ± 6.5</b>	<b>14.10%</b>	<b>0.26</b>	<b>33.2 ± 6.7</b>	<b>30.7 ± 7.2</b>	<b>36.10%</b>	<b>0.002</b>	<b>32.6 ± 6.7</b>	<b>30.4 ± 7.7</b>	<b>31.10%</b>	<b>0</b>
<b>Ejection fraction rank</b>	<b>137.5 ± 75.8</b>	<b>123.8 ± 73.8</b>	<b>18.20%</b>	<b>0.14</b>	<b>168.2 ± 85.8</b>	<b>135.3 ± 85.6</b>	<b>38.40%</b>	<b>0.001</b>	<b>206.8 ± 106.4</b>	<b>174.9 ± 109.9</b>	<b>29.50%</b>	<b>0.01</b>
NYHA			20.30%	0.99			29.00%	0.068			18.10%	0.18
I	2 (1.6%)	3 (2.3%)			5 (3.4%)	2 (1.3%)			6 (3.4%)	8 (4.0%)		
II	63 (49.6%)	67 (50.4%)			72 (48.3%)	58 (37.9%)			90 (50.3%)	88 (44.0%)		
III	61 (48.0%)	61 (45.9%)			71 (47.7%)	88 (57.5%)			82 (45.8%)	97 (48.5%)		
IV	1 (0.8%)	2 (1.5%)			1 (0.7%)	5 (3.3%)			1 (0.6%)	7 (3.5%)		
<b>Myocardial infarction</b>	<b>95 (74.8%)</b>	<b>108 (81.2%)</b>	<b>15.50%</b>	<b>0.23</b>	<b>107 (71.8%)</b>	<b>123 (80.4%)</b>	<b>20.20%</b>	<b>0.1</b>	<b>129 (72.1%)</b>	<b>154 (77.0%)</b>	<b>11.30%</b>	<b>0.29</b>
Stroke	17 (13.4%)	14 (10.5%)	8.80%	0.57	9 (6.0%)	16 (10.5%)	16.10%	0.21	18 (10.1%)	28 (14.0%)	12.10%	0.27
<b>Diabetes</b>	<b>43 (33.9%)</b>	<b>69 (51.9%)</b>	<b>37.00%</b>	<b>0.004</b>	<b>57 (38.3%)</b>	<b>67 (43.8%)</b>	<b>11.30%</b>	<b>0.35</b>	<b>63 (35.2%)</b>	<b>87 (43.5%)</b>	<b>17.10%</b>	<b>0.11</b>
Hypertension	96 (75.6%)	106 (79.7%)	9.90%	0.46	112 (75.2%)	117 (76.5%)	3.00%	0.89	131 (73.2%)	152 (76.0%)	6.50%	0.56
ACEI or ARB	118 (92.9%)	127 (95.5%)	11.00%	0.43	144 (96.6%)	145 (94.8%)	9.30%	0.57	169 (94.4%)	180 (90.0%)	16.50%	0.13
Beta blockers	119 (93.7%)	121 (91.0%)	10.30%	0.49	139 (93.3%)	140 (91.5%)	6.70%	0.67	166 (92.7%)	184 (92.0%)	2.80%	0.85
<b>MRA</b>	<b>92 (72.4%)</b>	<b>101 (75.9%)</b>	<b>8.00%</b>	<b>0.57</b>	<b>104 (69.8%)</b>	<b>131 (85.6%)</b>	<b>38.70%</b>	<b>0.001</b>	<b>131 (73.2%)</b>	<b>151 (75.5%)</b>	<b>5.30%</b>	<b>0.64</b>
<b>Digoxin</b>	<b>5 (3.9%)</b>	<b>10 (7.5%)</b>	<b>15.50%</b>	<b>0.29</b>	<b>9 (6.0%)</b>	<b>19 (12.4%)</b>	<b>22.20%</b>	<b>0.073</b>	<b>6 (3.4%)</b>	<b>14 (7.0%)</b>	<b>16.50%</b>	<b>0.17</b>
Aspirin	115 (90.6%)	124 (93.2%)	9.80%	0.5	143 (96.0%)	147 (96.1%)	0.50%	1	168 (93.9%)	179 (89.5%)	15.80%	0.14

**Supplementary Table 3: sIPTW analysis of significant associations of protein biomarkers with CV events (as listed in Table 2).**

Biomarker	sIPTW logistic model		sIPTW logistic model adjusted for NT-proBNP
	OR (95% CI)	FDR	OR (95% CI)
NT-proBNP	<b>1.302 (1.205 - 1.406)</b>	<.0001	
BNP	<b>1.258 (1.178 - 1.343)</b>	<.0001	
TIMD4	<b>1.719 (1.378 - 2.144)</b>	0.0001	<b>1.502 (1.196 - 1.886)</b>
FGF-23	<b>1.229 (1.121 - 1.348)</b>	0.0006	1.068 (0.964 - 1.185)
GDF-15	<b>1.391 (1.186 - 1.631)</b>	0.0015	1.086 (0.905 - 1.303)
PSP-D	<b>1.421 (1.219 - 1.655)</b>	0.0004	1.247 (1.062 - 1.464)
SPON1	<b>1.895 (1.435 - 2.502)</b>	0.0004	1.156 (0.833 - 1.605)
VEGFD	<b>1.743 (1.338 - 2.271)</b>	0.0015	1.157 (0.877 - 1.527)
TNC	<b>1.465 (1.220 - 1.759)</b>	0.0015	1.194 (0.980 - 1.456)
AOC3	<b>1.656 (1.267 - 2.165)</b>	0.0056	1.165 (0.870 - 1.561)
IGFBP-7	<b>1.465 (1.186 - 1.810)</b>	0.0078	0.999 (0.782 - 1.277)
IL-1RT1	<b>1.715 (1.291 - 2.280)</b>	0.0056	1.205 (0.883 - 1.645)
TFF3	<b>1.350 (1.122 - 1.623)</b>	0.0202	1.023 (0.833 - 1.258)
TIMP1	<b>1.576 (1.233 - 2.013)</b>	0.0064	1.090 (0.828 - 1.436)
U-PAR	<b>1.456 (1.170 - 1.811)</b>	0.0121	1.045 (0.818 - 1.335)
ST2	<b>1.360 (1.151 - 1.607)</b>	0.0064	1.117 (0.932 - 1.339)
TR	<b>1.340 (1.135 - 1.582)</b>	0.0101	1.111 (0.930 - 1.328)
OPG	<b>1.591 (1.211 - 2.090)</b>	0.0129	1.086 (0.803 - 1.469)
TFPI	<b>1.663 (1.244 - 2.225)</b>	0.0104	1.457 (1.079 - 1.968)
FCGR2A	<b>1.400 (1.147 - 1.708)</b>	0.0135	1.257 (1.025 - 1.542)
COL18A1	<b>1.398 (1.109 - 1.763)</b>	0.0372	1.027 (0.798 - 1.323)
CXCL1	<b>1.189 (1.063 - 1.331)</b>	0.0308	1.152 (1.026 - 1.293)
IGLC2	<b>1.359 (1.101 - 1.678)</b>	0.0355	1.117 (0.894 - 1.396)
IGFBP-2	<b>1.260 (1.076 - 1.475)</b>	0.0355	0.869 (0.713 - 1.060)
IL6	<b>1.175 (1.063 - 1.299)</b>	0.0211	1.096 (0.991 - 1.212)
TGM2	<b>1.207 (1.066 - 1.368)</b>	0.0322	1.138 (1.001 - 1.293)
ACE2	<b>1.281 (1.083 - 1.515)</b>	0.0355	1.039 (0.870 - 1.242)
CCL18	<b>1.283 (1.092 - 1.507)</b>	0.0303	1.232 (1.044 - 1.454)
CD93	<b>1.471 (1.115 - 1.940)</b>	0.043	0.867 (0.626 - 1.201)
IL1RL2	<b>0.699 (0.541 - 0.904)</b>	0.043	0.715 (0.549 - 0.930)
MMP-2	<b>1.445 (1.128 - 1.851)</b>	0.0355	0.758 (0.550 - 1.044)
PGLYRP1	<b>1.303 (1.085 - 1.566)</b>	0.0372	1.147 (0.948 - 1.388)
Notch 3	<b>1.433 (1.125 - 1.825)</b>	0.0355	0.855 (0.638 - 1.145)
OSMR	<b>2.203 (1.312 - 3.699)</b>	0.031	1.181 (0.672 - 2.076)
CCL24	<b>1.222 (1.072 - 1.393)</b>	0.031	1.200 (1.048 - 1.374)
IGFBP-1	<b>1.143 (1.040 - 1.256)</b>	0.042	0.990 (0.890 - 1.101)
CD163	<b>1.425 (1.119 - 1.815)</b>	0.0355	1.177 (0.914 - 1.516)
MMP-9	<b>1.153 (1.042 - 1.276)</b>	0.043	1.146 (1.033 - 1.272)
ICAM1	<b>1.526 (1.144 - 2.037)</b>	0.0355	1.274 (0.944 - 1.719)
OPN	<b>1.257 (1.062 - 1.487)</b>	0.0493	0.942 (0.777 - 1.142)
SERPINA5	<b>0.721 (0.570 - 0.912)</b>	0.043	0.932 (0.725 - 1.199)
LTBP2	<b>1.464 (1.121 - 1.911)</b>	0.039	0.835 (0.605 - 1.152)
CRTAC1	<b>0.768 (0.633 - 0.932)</b>	0.0488	0.782 (0.641 - 0.955)

**Supplementary Table 4: Associations of protein biomarkers (significantly associated with overall CV events) with specific clinical events (MI/stroke, SCD, HF rehospitalization).**

Biomarker	MI/Stroke		SCD		Rehospitalization for HF		Interaction FDR
	OR (95% CI)	FDR	OR (95% CI)	FDR	OR (95% CI)	FDR	
NT-proBNP	<b>1.343 (1.140 - 1.582)</b>	<b>0.0293</b>	<b>1.504 (1.284 - 1.761)</b>	<0.0001	<b>1.319 (1.162 - 1.497)</b>	<b>0.0051</b>	0.7671
BNP	<b>1.262 (1.100 - 1.447)</b>	<b>0.0411</b>	<b>1.414 (1.241 - 1.612)</b>	<0.0001	1.233 (1.110 - 1.370)	0.0135	0.7668
TIMD4	<b>2.832 (1.701 - 4.713)</b>	<b>0.0086</b>	1.853 (1.240 - 2.768)	0.0769	1.304 (0.928 - 1.831)	0.8652	0.6075
FGF-23	1.244 (1.014 - 1.527)	0.213	1.289 (1.093 - 1.520)	0.0769	1.258 (1.075 - 1.472)	0.2889	0.9766
GDF-15	1.425 (1.046 - 1.940)	0.1786	<b>1.736 (1.265 - 2.380)</b>	<b>0.0345</b>	1.431 (1.076 - 1.904)	0.5591	0.8528
PSP-D	<b>2.111 (1.507 - 2.958)</b>	<b>0.0039</b>	1.451 (1.094 - 1.923)	0.1198	1.125 (0.878 - 1.442)	0.8652	0.596
SPON1	2.407 (1.394 - 4.155)	0.0637	1.868 (1.109 - 3.147)	0.1492	1.665 (1.065 - 2.605)	0.6396	0.8528
VEGFD	2.163 (1.214 - 3.855)	0.1475	2.031 (1.206 - 3.422)	0.1198	1.537 (1.012 - 2.335)	0.8107	0.8528
TNC	1.525 (1.073 - 2.167)	0.1668	1.935 (1.339 - 2.796)	0.0304	1.185 (0.902 - 1.557)	0.8652	0.6912
AOC3	1.870 (1.076 - 3.252)	0.1876	1.423 (0.861 - 2.352)	0.3692	1.841 (1.189 - 2.853)	0.3453	0.8864
IGFBP-7	1.667 (1.069 - 2.598)	0.1786	1.798 (1.187 - 2.724)	0.1071	1.308 (0.921 - 1.858)	0.8652	0.8365
IL-1RT1	1.926 (1.115 - 3.325)	0.1668	1.694 (0.968 - 2.965)	0.2345	1.761 (1.074 - 2.887)	0.6396	0.9705
TFF3	1.539 (1.062 - 2.230)	0.1786	<b>1.933 (1.311 - 2.851)</b>	<b>0.0407</b>	1.275 (0.925 - 1.757)	0.8652	0.7668
TIMP1	2.302 (1.309 - 4.045)	0.1299	1.752 (1.121 - 2.738)	0.1358	1.255 (0.855 - 1.844)	0.8652	0.74
U-PAR	1.796 (1.158 - 2.786)	0.1475	1.702 (1.133 - 2.558)	0.1202	1.250 (0.862 - 1.813)	0.8652	0.7671
ST2	1.599 (1.087 - 2.352)	0.1663	1.442 (1.087 - 1.913)	0.1224	1.185 (0.911 - 1.542)	0.8652	0.7671
TR	1.289 (0.931 - 1.786)	0.3107	1.356 (0.990 - 1.858)	0.2185	1.388 (1.056 - 1.825)	0.6396	0.9705
OPG	1.662 (0.989 - 2.794)	0.2501	1.760 (1.031 - 3.006)	0.1999	1.579 (0.994 - 2.507)	0.8107	0.9757
TFPI	2.130 (1.231 - 3.687)	0.1475	1.790 (1.010 - 3.172)	0.2122	1.272 (0.781 - 2.069)	0.8652	0.7671
FCGR2A	1.718 (1.130 - 2.611)	0.1475	1.269 (0.888 - 1.812)	0.3957	1.340 (0.968 - 1.853)	0.8606	0.8492
COL18A1	1.410 (0.875 - 2.270)	0.3482	<b>1.777 (1.137 - 2.776)</b>	0.1229	1.442 (0.971 - 2.143)	0.8606	0.8867
CXCL1	1.322 (1.065 - 1.641)	0.1475	1.272 (1.032 - 1.570)	0.1541	1.080 (0.891 - 1.309)	0.8656	0.7671
IGLC2	1.851 (1.190 - 2.879)	0.1475	1.366 (0.933 - 1.999)	0.3046	1.249 (0.875 - 1.782)	0.8652	0.7671
IGFBP-2	1.289 (0.942 - 1.765)	0.3107	1.576 (1.170 - 2.124)	0.0769	1.132 (0.858 - 1.493)	0.8652	0.7668
IL6	1.201 (0.971 - 1.485)	0.2916	1.220 (1.034 - 1.440)	0.1492	1.106 (0.929 - 1.315)	0.8652	0.8867
TGM2	1.236 (0.971 - 1.574)	0.2883	1.301 (1.025 - 1.652)	0.1724	1.156 (0.931 - 1.435)	0.8652	0.9103
ACE2	1.498 (1.043 - 2.150)	0.1955	1.299 (0.942 - 1.790)	0.3046	1.218 (0.919 - 1.616)	0.8652	0.8786
CCL18	1.225 (0.890 - 1.687)	0.4115	1.195 (0.914 - 1.564)	0.3957	1.432 (1.075 - 1.907)	0.5591	0.8564
CD93	1.834 (1.037 - 3.245)	0.213	1.342 (0.785 - 2.296)	0.4935	1.592 (1.002 - 2.531)	0.8107	0.8867
IL1RL2	0.659 (0.391 - 1.111)	0.3107	0.591 (0.354 - 0.986)	0.2102	0.721 (0.486 - 1.069)	0.8652	0.9337
MMP-2	1.333 (0.814 - 2.183)	0.4604	1.560 (0.951 - 2.557)	0.2525	1.519 (1.011 - 2.282)	0.8107	0.9475
PGLYRP1	1.267 (0.873 - 1.840)	0.4115	1.463 (1.052 - 2.035)	0.1541	1.286 (0.942 - 1.755)	0.8652	0.9171
CCL14	1.592 (0.975 - 2.602)	0.2584	1.645 (1.069 - 2.530)	0.1541	1.241 (0.824 - 1.871)	0.8652	0.8528
Notch 3	1.274 (0.815 - 1.990)	0.4841	1.730 (1.074 - 2.784)	0.1541	1.442 (0.952 - 2.184)	0.8606	0.8598
OSMR	2.901 (0.946 - 8.891)	0.2584	2.848 (1.106 - 7.334)	0.1724	1.523 (0.665 - 3.491)	0.8652	0.8492
CCL24	1.338 (1.024 - 1.748)	0.2047	1.414 (1.105 - 1.808)	0.1071	1.017 (0.819 - 1.264)	0.9655	0.6912
TIMP4	1.400 (0.916 - 2.140)	0.3107	1.555 (1.020 - 2.370)	0.2036	1.327 (0.915 - 1.923)	0.8652	0.9394
TRAIL-R2	1.833 (1.154 - 2.912)	0.1475	1.856 (1.227 - 2.810)	0.0788	1.055 (0.783 - 1.420)	0.9412	0.596
UMOD	0.399 (0.173 - 0.922)	0.2027	0.740 (0.342 - 1.603)	0.6409	0.482 (0.236 - 0.986)	0.8107	0.8492
CD163	1.592 (0.979 - 2.589)	0.2584	1.596 (0.991 - 2.572)	0.2175	1.218 (0.831 - 1.786)	0.8652	0.8528
ICAM1	<b>3.099 (1.622 - 5.924)</b>	<b>0.0342</b>	1.456 (0.863 - 2.456)	0.3661	1.074 (0.677 - 1.704)	0.9554	0.596
IL-27	2.050 (1.172 - 3.588)	0.1475	<b>2.641 (1.556 - 4.483)</b>	<b>0.0297</b>	0.897 (0.616 - 1.306)	0.886	0.4303
OPN	1.558 (1.126 - 2.155)	0.1475	1.311 (0.930 - 1.846)	0.323	1.113 (0.851 - 1.454)	0.8656	0.7668
RARRES2	2.584 (1.307 - 5.109)	0.1475	1.048 (0.557 - 1.972)	0.9359	1.799 (0.973 - 3.329)	0.8459	0.6912
SERPINA5	0.856 (0.535 - 1.370)	0.6695	0.494 (0.315 - 0.775)	0.0769	0.820 (0.554 - 1.212)	0.8652	0.7173
LTBP2	1.276 (0.721 - 2.259)	0.5749	2.076 (1.210 - 3.563)	0.1198	1.290 (0.867 - 1.919)	0.8652	0.7671
CRTAC1	0.863 (0.582 - 1.279)	0.6193	0.562 (0.384 - 0.824)	0.0782	0.854 (0.627 - 1.164)	0.8652	0.7195
GH	1.014 (0.887 - 1.158)	0.896	1.195 (1.050 - 1.360)	0.1198	1.100 (0.981 - 1.234)	0.8652	0.7668
IGFBP-1	1.015 (0.844 - 1.221)	0.9186	1.258 (1.058 - 1.495)	0.1198	1.161 (0.984 - 1.371)	0.8606	0.7668

Logistic model adjusted for sex, age, study drug and significant factors from Table 1

**Supplementary Table 5: Associations of protein biomarkers (significantly associated with overall CV events) with specific clinical events (MI/stroke, SCD, HF rehospitalization) using sIPTW.**

Biomarker	MI/Stroke		SCD		Rehospitalization		Interaction FDR	
	OR (95% CI)	FDR	OR (95% CI)	FDR	for HF			
					OR (95% CI)	FDR		
NT-proBNP	1.255 (1.081 - 1.457)	0.10	<b>1.426 (1.235 - 1.647)</b>	<b>0.0002</b>	<b>1.252 (1.113 - 1.408)</b>	<b>0.0491</b>	0.7251	
BNP	1.236 (1.086 - 1.407)	0.0735	<b>1.375 (1.217 - 1.553)</b>	<b>&lt;.0001</b>	1.197 (1.084 - 1.322)	0.0505	0.6653	
TIMD4	<b>2.898 (1.750 - 4.799)</b>	<b>0.0049</b>	1.858 (1.273 - 2.713)	0.0608	1.278 (0.918 - 1.779)	0.8726	0.4328	
FGF-23	1.196 (0.989 - 1.446)	0.2806	1.276 (1.091 - 1.491)	0.0723	1.213 (1.050 - 1.402)	0.4932	0.9241	
GDF-15	1.336 (1.003 - 1.780)	0.2613	1.584 (1.191 - 2.105)	0.0608	1.290 (0.993 - 1.676)	0.8726	0.7922	
PSP-D	2.043 (1.492 - 2.797)	0.0023	1.434 (1.099 - 1.872)	0.0883	1.110 (0.875 - 1.409)	0.8726	0.3869	
SPON1	2.292 (1.361 - 3.860)	0.0835	1.862 (1.145 - 3.028)	0.0886	1.654 (1.060 - 2.580)	0.7373	0.8231	
VEGFD	2.230 (1.290 - 3.857)	0.1258	1.980 (1.224 - 3.202)	0.0798	1.414 (0.974 - 2.051)	0.8726	0.7251	
TNC	1.568 (1.109 - 2.218)	0.164	<b>1.955 (1.373 - 2.786)</b>	<b>0.0141</b>	1.184 (0.900 - 1.558)	0.8726	0.54	
AOC3	1.876 (1.105 - 3.186)	0.1665	1.346 (0.848 - 2.135)	0.3798	1.802 (1.182 - 2.745)	0.4247	0.7949	
IGFBP-7	1.544 (1.030 - 2.315)	0.2216	1.757 (1.200 - 2.572)	0.0757	1.225 (0.878 - 1.709)	0.8726	0.7308	
IL-1RT1	1.869 (1.124 - 3.107)	0.1665	1.683 (1.002 - 2.828)	0.183	1.636 (1.031 - 2.597)	0.8433	0.957	
TFF3	1.284 (0.924 - 1.785)	0.3568	1.780 (1.253 - 2.527)	0.0608	1.130 (0.840 - 1.519)	0.8726	0.6367	
TIMP1	2.278 (1.329 - 3.906)	0.10	1.735 (1.141 - 2.638)	0.0883	1.211 (0.834 - 1.758)	0.8726	0.6367	
U-PAR	1.610 (1.078 - 2.405)	0.1665	1.694 (1.159 - 2.476)	0.0817	1.166 (0.814 - 1.672)	0.8726	0.7251	
ST2	1.609 (1.116 - 2.322)	0.164	1.479 (1.122 - 1.950)	0.0798	1.173 (0.904 - 1.521)	0.8726	0.7251	
TR	1.279 (0.934 - 1.753)	0.3403	1.369 (1.016 - 1.845)	0.1624	1.362 (1.050 - 1.766)	0.6838	0.9673	
OPG	1.566 (0.976 - 2.515)	0.2806	1.749 (1.056 - 2.895)	0.1392	1.497 (0.960 - 2.336)	0.8726	0.9434	
TFPI	2.017 (1.202 - 3.384)	0.164	1.886 (1.083 - 3.284)	0.1325	1.288 (0.809 - 2.051)	0.8726	0.7447	
FCGR2A	1.749 (1.161 - 2.634)	0.164	1.286 (0.913 - 1.811)	0.3209	1.295 (0.949 - 1.768)	0.8726	0.757	
COL18A1	1.232 (0.799 - 1.899)	0.5519	1.693 (1.124 - 2.550)	0.0883	1.300 (0.896 - 1.886)	0.8726	0.7831	
CXCL1	1.280 (1.040 - 1.575)	0.1665	1.269 (1.038 - 1.552)	0.1257	1.072 (0.894 - 1.286)	0.8726	0.7308	
IGLC2	1.662 (1.106 - 2.497)	0.1665	1.350 (0.942 - 1.935)	0.2729	1.177 (0.839 - 1.652)	0.8726	0.757	
IGFBP-2	1.213 (0.904 - 1.628)	0.4346	1.514 (1.151 - 1.990)	0.0723	1.086 (0.834 - 1.413)	0.8832	0.6653	
IL6	1.189 (0.968 - 1.460)	0.3079	1.247 (1.057 - 1.472)	0.0883	1.111 (0.940 - 1.313)	0.8726	0.8231	
TGM2	1.235 (0.986 - 1.547)	0.2806	1.295 (1.031 - 1.627)	0.1347	1.130 (0.926 - 1.379)	0.8726	0.8253	
ACE2	1.564 (1.111 - 2.201)	0.164	1.295 (0.958 - 1.751)	0.2576	1.145 (0.890 - 1.473)	0.8726	0.7308	
CCL18	1.220 (0.901 - 1.652)	0.4346	1.215 (0.941 - 1.569)	0.3037	1.428 (1.079 - 1.888)	0.5792	0.8253	
CD93	1.631 (0.967 - 2.752)	0.2806	1.347 (0.818 - 2.219)	0.423	1.455 (0.945 - 2.240)	0.8726	0.9328	
IL1RL2	0.672 (0.406 - 1.113)	0.3403	0.610 (0.370 - 1.003)	0.183	0.759 (0.525 - 1.098)	0.8726	0.8951	
MMP-2	1.333 (0.839 - 2.118)	0.4539	1.582 (0.998 - 2.508)	0.183	1.438 (0.980 - 2.108)	0.8726	0.9328	
PGLYRP1	1.176 (0.831 - 1.664)	0.5631	1.496 (1.095 - 2.044)	0.0883	1.242 (0.916 - 1.685)	0.8726	0.7926	
Notch 3	1.224 (0.812 - 1.846)	0.5465	1.751 (1.120 - 2.735)	0.096	1.414 (0.944 - 2.118)	0.8726	0.7789	
OSMR	3.312 (1.115 - 9.840)	0.2183	3.003 (1.231 - 7.328)	0.1057	1.376 (0.623 - 3.039)	0.8726	0.7251	
CCL24	1.323 (1.029 - 1.701)	0.2104	1.441 (1.134 - 1.832)	0.0723	1.028 (0.837 - 1.262)	0.9457	0.54	
IGFBP-1	1.006 (0.842 - 1.202)	0.9518	1.249 (1.065 - 1.465)	0.0817	1.158 (0.987 - 1.358)	0.8726	0.6585	
CD163	1.529 (0.977 - 2.393)	0.2806	1.673 (1.058 - 2.646)	0.1358	1.225 (0.845 - 1.777)	0.8726	0.7922	

MMP-9	1.087 (0.897 - 1.316)	0.5912	1.255 (1.059 - 1.485)	0.0883	1.115 (0.938 - 1.324)	0.8726	0.7678
<b>ICAM1</b>	<b>3.350 (1.790 - 6.269)</b>	<b>0.0144</b>	1.482 (0.899 - 2.443)	0.2827	1.030 (0.661 - 1.606)	0.9551	0.3869
OPN	1.485 (1.096 - 2.012)	0.164	1.300 (0.937 - 1.804)	0.2827	1.078 (0.832 - 1.398)	0.8985	0.7216
SERPINA5	0.859 (0.550 - 1.341)	0.6674	0.517 (0.337 - 0.794)	0.0723	0.816 (0.563 - 1.184)	0.8726	0.6585
LTBP2	1.278 (0.737 - 2.215)	0.5788	2.084 (1.262 - 3.440)	0.0757	1.255 (0.853 - 1.848)	0.8726	0.7043
CRTAC1	0.860 (0.592 - 1.250)	0.632	0.600 (0.421 - 0.854)	0.0793	0.857 (0.637 - 1.153)	0.8726	0.7043

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