Supplemental Table 1.

Crude and age-standardized incidence rates of the most frequent cancer sites (excluded non-melanoma skin cancer), both in the study population and in the general Spanish population. Incidence rates are expressed in episodes per 100,000 patients-years. IR, Incidence Rate.

	Women				Men					
	Study population			General Spanish population*		Study population			General Spanish population**	
	Cases	Crude IR (95% CI)	Age- standardize d IR	Crude IR	Age- standardized IR	Cases	Crude IR (95% CI)	Age- standardized IR	Crude IR	Age- standardized IR
Lung	7	275.1 (71.3–478.8)	272.2	22.4	15.6	28	447.3 (281.6–613)	210.3	100.5	69.7
Colorectal	1	39.3 (1.0–218.9)	_	63.1	35.5	17	271.6 (142.5–400.7)	142.8	104.7	70
Breast	8	314.4 (96.5–532.2)	260.3	125.2	94.4	1	16.0 (0.4–89.0)	Ι	_	_
Prostate	_	_	_	_	_	23	367.4 (217.3–517.6)	135.6	144	101.7

*Crude and age-standardized incidence rates of selected cancer sites (lung, colorrectum, breast) expected for the general Spanish population of women aged \geq 15 years in the year 2010, according to the Global Cancer Observatory. Source: <u>https://gco.iarc.fr/overtime/en</u>.

**Crude and age-standardized incidence rates of selected cancer sites (lung, colorrectum, prostate) expected for the general Spanish population of men aged \geq 15 years in the year 2010, according to the Global Cancer Observatory. Source: <u>https://gco.iarc.fr/overtime/en</u>.

Age-standardized rates were calculated by the direct method, taking the modified world standard population proposed by Doll et al as a reference. Doll R, Payne P, Waterhouse J. Cancer incidence in five countries: a technical report. Berlin: Springer-Verlag Berlin Heidelberg; 1966. Available in *https://www.springer.com/gp/book/97835400347*

Supplemental Table 2. Univariate coefficients of all baseline variables explored as potential risk factors for malignancy in the study cohort: Fine-Gray competing risk regression analyses.

Variable	Univariate Hazard Ratio	95% Confidence Interval	P value*
Age (years)	1.03	1.02–1.04	<0.01
Female gender	0.68	0.46-1.01	0.05
Previous history of malignancy	1.29	0.83–1.99	0.258
History of alcohol abuse	1.21	0.86–1.70	0.27
History of smoking	1.45	1.04-2.03	<0.01
Body mass index (kg/m2)	1.00	0.98–1.03	0.88
Hypertension	1.16	0.84–1.59	0.37
Diabetes mellitus	1.09	0.78–1.52	0.61
Dyslipidaemia	1.19	0.87–1.65	0.26
Coronary artery disease	1.31	0.95–1.79	0.09
Chronic pulmonary obstructive disease	1.71	1.12–2.61	0.01
Chronic renal dysfunction	1.13	0.81-1.59	0.47
Anaemia	1.16	0.82–1.64	0.39
New York Heart Association class III or IV	0.79	0.56–1.12	0.19
Serum NTproBNP (ng/ml)	1.00	0.99–1.00	0.36
Left ventricular ejection fraction (%)	1.00	0.99–1.01	0.67
Diuretic use	0.98	0.66–1.65	0.92
Angiotensin converter enzyme inhibitor use	1.40	1.00-1.95	0.05
Angiotensin II receptor blocker use	0.85	0.54–1.35	0.49
Sacubitril-valsartan use	0.74	0.34–1.56	0.43
Beta-blocker use	1.13	0.67–1.91	0.64
Mineralocorticoid receptor antagonists	0.80	0.58–1.10	0.17
Digoxin use	1.08	0.63–1.85	0.77
Ivabradine use	0.94	0.46-1.92	0.86

*Variables that showed a univariate p-value <0.10 were selected for entering the first step of multivariable backward stepwise analysis.

Supplemental Table 3. Causes of death in patients with heart failure and a history of pre-existing malignancy, newly diagnosed malignancy during follow-up or no malignancy.

Causes of death (n=536)	Pre-existing malignancy (n=98)	Newly diagnosed malignancy (n=77)	No malignancy (n=361)	
Cardiovascular				
Sudden death	14 (14.3%)	9 (11.7%)	109 (30.2%)	
Heart failure	22 (22.4%)	10 (13%)	140 (38.8%)	
Other cardiovascular causes	11 (11.2%)	1 (1.3%)	32 (8.9%)	
Non cardiovascular				
Cancer	22 (22.4%)	52 (67.5%)	0	
Infection	18 (18.4%)	4 (5.2%)	45 (12.5%)	
Other non-cardiovascular causes	11 (11.2%)	1 (1.3%)	24 (6.6%)	
Not specified	0	0	11 (3%)	

Supplemental Table 4. Source of data about cancer diagnoses in previous studies that addressed the incidence of cancer in patients with heart

failure.

Authors	Reference	Region/country	Source of data	Description
Hasin et al.	J Am Coll Cardiol 2013; 62: 881-86.	Oldmest County (Minnesota, United States)	Rochester Epidemiology Project	Medical records linkage system
Hasin et al.	J Am Coll Cardiol 2016; 68: 265-71.	Oldmest County (Minnesota, United States)	Rochester Epidemiology Project	Medical records linkage system
Banke et al.	Eur J Heart Fail 2016; 18: 260-66	Denmark	Danish National Patient Registries	Administrative database
Kwak S et al.	J Cardiol 2021; 77: 231-38.	South Korea	Korean National Health Insurance Database	Administrative database
Roderburg C et al.	ESC Heart Fail 2021; 8: 3628-33	Germany	Disease Analyzer Database	Administrative database
Bertero E et al.	JACC CardioOncol 2022; 4: 98-109	Puglia (Italy)	Various administrative databases	Administrative databases
Scwartz B et al.	Int J Cardiol 2020; 316:209-213.	Denmark	Danish Nationwide Administrative Database	Administrative database