## 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 | $\begin{aligned} & \text { Crude IR } \\ & (95 \% \text { CI) } \end{aligned}$ | Agestandardize d IR | Crude IR | $\begin{gathered} \text { Age- } \\ \text { standardized } \\ \text { IR } \end{gathered}$ | Cases | $\begin{aligned} & \text { Crude IR } \\ & \text { (95\% CI) } \end{aligned}$ | Age- <br> standardized <br> IR | Crude IR | Age- <br> standardized <br> IR |
| Lung | 7 | $\begin{gathered} 275.1 \\ (71.3-478.8) \end{gathered}$ | 272.2 | 22.4 | 15.6 | 28 | $\begin{gathered} 447.3 \\ (281.6-613) \end{gathered}$ | 210.3 | 100.5 | 69.7 |
| Colorectal | 1 | $\begin{gathered} 39.3 \\ (1.0-218.9) \end{gathered}$ | - | 63.1 | 35.5 | 17 | $\begin{gathered} 271.6 \\ (142.5-400.7) \end{gathered}$ | 142.8 | 104.7 | 70 |
| Breast | 8 | $\begin{gathered} 314.4 \\ (96.5-532.2) \end{gathered}$ | 260.3 | 125.2 | 94.4 | 1 | $\begin{gathered} 16.0 \\ (0.4-89.0) \end{gathered}$ | - | - | - |
| Prostate | - | - | - | - | - | 23 | $\begin{gathered} 367.4 \\ (217.3-517.6) \end{gathered}$ | 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: https://gco.iarc.fr/overtime/en.
**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: https://gco.iarc.frrovertime/en.

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 | $\mathbf{9 5 \%}$ Confidence Interval | $\mathbf{P}$ 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 <br> malignancy (n=98) | Newly diagnosed <br> malignancy (n=77) | No malignancy <br> $(\mathbf{n}=\mathbf{3 6 1})$ |
| :--- | :---: | :---: | :---: |
| 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 | $22(22.4 \%)$ | $52(67.5 \%)$ | $32(8.9 \%)$ |
| Non cardiovascular | $18(18.4 \%)$ | $4(5.2 \%)$ | $45(12.5 \%)$ |
| Cancer | $11(11.2 \%)$ | $1(1.3 \%)$ | $24(6.6 \%)$ |
| Infection | 0 | 0 | $11(3 \%)$ |
| Other non-cardiovascular causes |  |  |  |
| Not specified |  |  |  |

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 <br> (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 |

