

## SUPPLEMENTARY DATA

### Supplementary methods

All supplementary analyses were performed in the study cohort ( $n = 77$  with available LGE data). Genotype was classified as genotype-positive (pathogenic/likely pathogenic variant) or genotype-negative and, among genotype-positive individuals, as desmosomal (*DSP*, *DSG2*, and *DSC2*) vs nondesmosomal. The discriminative capacity of univariable models for predicting LGE was compared using DeLong's test for 2 correlated ROC curves. To account for the modest sample size and limited number of events, the primary multivariable model ( $\text{LGE} \sim \text{EI-RBBB-VE} + \text{indexed CMR LVEDD [z-score]}$ ) was additionally estimated using Firth's penalized logistic regression with profile-likelihood confidence intervals. EI-RBBB-VE was characterized by precordial transition, QRS duration, fragmentation, and coupling interval. An epicardial proxy was defined as QRS duration  $\geq 140$  ms combined with a late precordial transition ( $\geq V_4$ ). Categorical variables were compared using the Fisher exact test, whereas continuous variables were compared using either the Student  $t$ -test or the Mann-Whitney U test, as appropriate.

**Table S1.** Morphological characteristics of RBBB ectopy by LGE status (48 morphologies in 37 patients)

| Feature (RBBB morphologies, n = 48) | LGE(+)        | LGE(-)        | <i>P</i> |
|-------------------------------------|---------------|---------------|----------|
| QRS duration, ms                    | 130 [120-145] | 130 [125-145] | .851     |
| Precordial transition lead          | 3 [2-4]       | 4 [3-4]       | .703     |
| Coupling interval, ms               | 415 [380-478] | 360 [330-425] | .082     |
| Fragmented QRS                      | 7/35 (20)     | 2/13 (15)     | .999     |

IQR, interquartile range; LGE, late gadolinium enhancement; RBBB, right bundle branch block.

The data are presented as n/N (%) or median [interquartile range].

**Table S2.** Resting Holter ectopic burden by LGE status

| Variable (n = 69)              | LGE(+)          | LGE(-)        | <i>P</i> |
|--------------------------------|-----------------|---------------|----------|
| PVC count / 24 h, median [IQR] | 1167 [250-2360] | 292 [26-1486] | .065     |
| PVC %, median [IQR]            | 1 [0-3]         | 0 [0-1]       | .058     |
| NSVT on Holter, n (%)          | 8/38 (21)       | 5/31 (16)     | .760     |

IQR, interquartile range; LGE, late gadolinium enhancement; NSVT, nonsustained ventricular tachycardia; PVC, premature ventricular complex.

The data are presented as n/N (%) or median [interquartile range].

**Table S3. Genotype vs EI-RBBB-VE as predictors of LGE**

| Predictor/model                                       | Estimate                 | 95%CI      | <i>P</i> |
|-------------------------------------------------------|--------------------------|------------|----------|
| Desmosomal genotype and LGE (Fisher)                  | 11/12 (92) vs 30/65 (46) | —          | .004     |
| AUC—EI-RBBB-VE (univariable)                          | 0.638                    | —          | —        |
| AUC—Genotype-positive (univariable)                   | 0.606                    | —          | —        |
| DeLong (EI-RBBB-VE vs genotype)                       | ΔAUC                     | —          | .656     |
| Genotype-positive (univariable)                       | 3.60                     | 1.12-11.53 | .031     |
| Genotype-positive — adjusted for EI-RBBB-VE + LVEDD-z | 2.88                     | 0.79-10.56 | 0.111    |

AUC, area under the ROC curve; CI, confidence interval; CMR, cardiac magnetic resonance; EI-RBBB-VE, exercise-induced right bundle branch block-pattern ventricular ectopy; LGE, late gadolinium enhancement; LVEDD, left ventricular end-diastolic diameter.

The data are presented as n/N (%) or odds ratios.

Univariable analyses and AUC comparison performed in the full cohort (n = 77); the multivariable model adjusted for CMR-derived LVEDD was restricted to patients with available CMR measurements (n = 66).



**Table S4.** Association between EI-RBBB-VE and LGE according to genetic subgroup

| Subgroup                       | n  | LGE(+) among EI-RBBB-VE(+) | LGE(+) among EI-RBBB-VE(-) | <i>P</i> |
|--------------------------------|----|----------------------------|----------------------------|----------|
| <i>Desmosomal</i>              | 12 | 9/9 (100)                  | 2/3 (67)                   | .250     |
| <i>Nondesmosomal</i>           | 65 | 16/28 (57)                 | 14/37 (38)                 | .140     |
| Nondesmosomal P/LP             | 48 | 13/21 (62)                 | 12/27 (44)                 | .259     |
| Genotype-negative/inconclusive | 17 | 3/7 (43)                   | 2/10 (20)                  | .593     |

EI-RBBB-VE, exercise-induced right bundle branch block-pattern ventricular ectopy; LGE, late gadolinium enhancement; P/LP, pathogenic/likely pathogenic.

**Table S5.** Comparison of standard and Firth penalized logistic regression models for the association between EI-RBBB-VE and myocardial fibrosis

| Model                                 | EI-RBBB-VE OR (95%CI) | <i>P</i> | CMR LVEDD-z OR (95%CI) | <i>P</i> |
|---------------------------------------|-----------------------|----------|------------------------|----------|
| Standard logistic regression (n = 66) | 3.92 (1.28-12.00)     | .017     | 0.53 (0.28-1.04)       | .064     |
| Firth penalized regression            | 3.59 (1.26-11.18)     | .016     | 0.59 (0.28-0.99)       | .047     |

CI, confidence interval; CMR, cardiac magnetic resonance; EI-RBBB-VE, exercise-induced right bundle branch block-pattern ventricular ectopy; LVEDD, left ventricular end-diastolic diameter; OR, odds ratio.

Both models included EI-RBBB-VE and CMR-derived LVEDD z-score as independent variables.