

| Supplementary Table 1. Groups calcium scores                                                                                                             |                          |
|----------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------|
| Groups                                                                                                                                                   | Total calcium score (HU) |
| ATTR with cardiac involvement                                                                                                                            | 87 (6.25–573)            |
| ATTR without cardiac involvement                                                                                                                         | 2.5 (0–8.2)              |
| Values are expressed as median (25 <sup>th</sup> –75 <sup>th</sup> percentiles); <i>P</i> < 0.05. HU, Hounsfield units; ATTR, transthyretin amyloidosis. |                          |

| Supplementary Table 2. Baseline characteristics of the three groups                                                                                                                                                                                                                                                                                                                                                                                                                                 |                          |                             |                     |                |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------|-----------------------------|---------------------|----------------|
|                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     | ATTR with CA<br>(n = 20) | ATTR without CA<br>(n = 20) | Control<br>(n = 20) | <i>P</i> value |
| Clinical data                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       |                          |                             |                     |                |
| Age (years)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         | 69.6 ± 8.2               | 41.6 ± 10.8                 | 51.7 ± 13.3         | <0.001***      |
| Male sex (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        | 16 (80)                  | 11 (55)                     | 11 (55)             |                |
| BMI                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 | 23.85 ± 3                | 25.8 ± 4.7                  | 25.9 ±2.6           | 0.1            |
| Hypertension (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    | 5 (25)                   | 4 (20)                      | 3 (15)              | 0.2            |
| Diabetes mellitus (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                               | 4 (20)                   | 3 (15)                      | 3 (15)              | 0.6            |
| History of carpal tunnel syndrome                                                                                                                                                                                                                                                                                                                                                                                                                                                                   | 13 (65)                  | 3 (15)                      | 0 (0)               | 0.005          |
| Extracardiac involvement                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | 14 (70)                  | 7 (35)                      | 0                   | 0.027          |
| Nervous system                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      | 13 (93)                  | 7 (100)                     | 0                   | -              |
| Lung                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                | 1 (7)                    | 0                           | 0                   | -              |
| 6-minute walk test distance (m)                                                                                                                                                                                                                                                                                                                                                                                                                                                                     | 336.9 ± 83.7             | 420.4 ± 112.8               | 460 ± 56.9          | <0.001*        |
| Functional class (NYHA)                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |                          |                             |                     | 0.001          |
| -NYHA I                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             | 4 (20)                   | 20 (100)                    | -                   | -              |
| -NYHA II                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | 14 (70)                  | 0 (0)                       | -                   | -              |
| -NYHA III                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 2 (10)                   | 0 (0)                       | -                   | -              |
| Laboratory tests                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    |                          |                             |                     |                |
| Troponin (ng/L)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     | 61 (30.7–86.2)           | 3 (2–4.2)                   | 2.7 (2.5–4)         | <0.001*        |
| BNP (pg/mL)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         | 389.5 (210–545)          | 14 (12.5–24)                | 16.5 (7.5–24)       | <0.001*        |
| Echocardiogram (2D)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 |                          |                             |                     |                |
| LVEF (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | 43.1 ± 12.7              | 59.2 ± 2.9                  | 62.6 ± 4.1          | <0.001*        |
| LVGLS (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 9.2 ± 3.3                | 17.8 ± 1.9                  | 19.1 ± 2.5          | <0.001*        |
| Septum (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         | 17.8 ± 3.5               | 8.6 ± 1.3                   | 8 ± 0.9             | <0.001*        |
| Posterior wall (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 | 17.5 ± 3.3               | 8.4 ± 1.3                   | 7.8 ± 0.9           | <0.001*        |
| LVDD (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 44 ± 5.5                 | 47 ± 2.8                    | 48.7 ± 4.8          | <0.01**        |
| LVSD (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 35 ± 6.2                 | 31.2 ± 2.8                  | 31.7 ± 5.2          | 0.06           |
| Index mass                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          | 194 ± 52.9               | 74.158 ± 16.2               | 70.2 ± 11.1         | <0.001*        |
| LA diameter (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    | 49 ± 12.5                | 32.7 ± 4.5                  | 34.6 ± 4            | <0.001*        |
| LA volume (mL)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      | 61 ± 16.6                | 28.4 ± 4                    | 27.3 ± 7.7          | <0.001*        |
| RA volume (mL)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      | 47.5 ± 17.36             | 19.2 ± 4.8                  | 21.4 ± 6.6          | <0.001*        |
| Diastolic function                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  |                          |                             |                     |                |
| -Grade II dysfunction                                                                                                                                                                                                                                                                                                                                                                                                                                                                               | 5 (25)                   | 0 (0)                       | N/A                 | 0.017          |
| -Grade III dysfunction                                                                                                                                                                                                                                                                                                                                                                                                                                                                              | 12 (60)                  | 0 (0)                       | N/A                 | <0.001         |
| -E/e’ ratio                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         | 20.6 ± 10.5              | 8.37 ± 9.2                  | 6.3 ± 2.2           | <0.001*        |
| -TAPSE                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              | 13.2 ± 4.5               | 20.8 ± 3.2                  | 22.5 ± 9            | <0.001*        |
| Pericardial effusion                                                                                                                                                                                                                                                                                                                                                                                                                                                                                |                          |                             |                     | <0.001         |
| -Mild                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               | 10 (50)                  | 0 (0)                       | -                   | -              |
| -Moderate                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 1 (5)                    | 0 (0)                       | -                   | -              |
| 99mTc-PYP scintigraphy                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |                          |                             |                     |                |
| 1-hour uptake                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       | 1.7 ± 0.2                | 1.1 ± 0.2                   | -                   | <0.001         |
| 3-hour uptake                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       | 1.6 ± 0.1                | 1.1 ± 0.1                   | -                   | <0.001         |
| * <i>P</i> < 0.05; ** <i>P</i> < 0.01; *** <i>P</i> < 0.001. ATTR, transthyretin amyloidosis; CA, cardiac amyloidosis; BMI, body mass index; NYHA, New York Heart Association classification; 2D, two dimensional; LVEF, left ventricular ejection fraction; LVGLS, left ventricular global longitudinal strain; LVDD, left ventricular diastolic dysfunction; LVSD, left ventricular systolic dysfunction; LA, xx; RA, xx; TAPSE, tricuspid annular plane systolic excursion; N/A, not applicable. |                          |                             |                     |                |

| Supplementary Table 3. Baseline echocardiographic variables                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         |                       |                          |                |
|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------|--------------------------|----------------|
|                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     | ATTR with CA (n = 20) | ATTR without CA (n = 20) | <i>P</i> value |
| LVEF (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | 43.1 ± 12.7           | 59.2 ± 2.9               | < 0.001        |
| LVGLS (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 9.2 ± 3.3             | 17.8 ± 1.9               | < 0.001        |
| Septum (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         | 17.8 ± 3.5            | 8.6 ± 1.3                | < 0.001        |
| Posterior wall (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 | 17.5 ± 3.3            | 8.4 ± 1.3                | < 0.001        |
| LVDD (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 44 ± 5.5              | 47 ± 2.8                 | < 0.040        |
| LVSD (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 35 ± 6.2              | 31.2 ± 2.8               | 0.019          |
| Mass (indexed)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      | 194 ± 52.9            | 74.158 ± 16.2            | < 0.001        |
| Relative thickness                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  | 0.8 ± 0.21            | 0.36 ± 0.05              | < 0.001        |
| RV diameter (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    | 40.8 ± 7.7            | 34.21 ± 5.2              | < 0.001        |
| Left atrial diameter (mm)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 49 ± 12.5             | 32.7 ± 4.5               | < 0.001        |
| Left atrial volume (mL)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             | 61 ± 16.6             | 28.4 ± 4                 | < 0.001        |
| Right atrium volume (mL)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | 47.5 ± 17.36          | 19.2 ± 4.8               | < 0.001        |
| Apical preservation                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 | 19/20 (95%)           | 2/20 (10%)               |                |
| Diastolic function                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  |                       |                          |                |
| Normal diastole                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     | 0/20 (0%)             | 14/20 (70%)              | < 0.001        |
| Diastolic dysfunction grade I                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       | 1/20 (5%)             | 3/20 (15%)               | 0.292          |
| Diastolic dysfunction grade II                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      | 5/20 (25%)            | 0/20 (0%)                | 0.017          |
| Diastolic dysfunction grade III                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     | 12/20 (60%)           | 0/20 (0%)                | < 0.001        |
| Indeterminate diastole                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              | 2/20 (10%)            | 0/20 (0%)                | 0.147          |
| E/e’ ratio                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          | 20.6 ± 10.5           | 8.37 ± 9.2               | < 0.001        |
| e’ velocity                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         | 4.2 ± 1.2             | 10.2 ± 3.5               | <0.001         |
| E-wave deceleration time                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | 162 ± 39.8            | 206.5 ± 47.7             | 0.001          |
| Right ventricular function                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          |                       |                          |                |
| RVGLS                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               | 16.1 ± 6.1            | 25.8 ± 3.6               | < 0.001        |
| FAC                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 | 33.2 ± 8.8            | 46.2 ± 10.6              | < 0.001        |
| S’ velocity of the RV                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               | 6.5 ± 3.3             | 11.1 ± 2.1               | < 0.001        |
| TAPSE                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               | 13.2 ± 4.5            | 20.8 ± 3.2               | < 0.001        |
| Pericardial effusion                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                |                       |                          |                |
| Mild                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                | 10/20 (50%)           | 0/20 (0%)                | -              |
| Moderate                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | 1/20 (5%)             | 0/20 (0%)                | -              |
| Severe                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              | 0/20 (0%)             | 0/20 (0%)                | -              |
| RVGLS (%)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           | 16.1 ± 6.1            | 25.8 ± 3.6               | < 0.001        |
| Values are expressed as mean ± standard deviation, n/N (%), or median (25 <sup>th</sup> –75 <sup>th</sup> percentiles), as appropriate. ATTR, transthyretin amyloidosis; CA, cardiac amyloidosis; LVEF, left ventricular ejection fraction; LVGLS, left ventricular global longitudinal strain; LVDD, left ventricular diastolic dysfunction; LVSD, left ventricular systolic dysfunction; RVGLS, right ventricular global longitudinal strain; FAC, fractional area change (right ventricular area variation); TAPSE, tricuspid annular plane systolic excursion (systolic excursion of the tricuspid annulus toward the apex); LA, left atrial; RA, right atrial. |                       |                          |                |

| Supplementary Table 4. Baseline ECG                                                                                                                                                                                                                                                                                                                                                  |                       |                          |                |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------|--------------------------|----------------|
| ECG                                                                                                                                                                                                                                                                                                                                                                                  | ATTR with CA (n = 20) | ATTR without CA (n = 20) | <i>P</i> value |
| Rhythms                                                                                                                                                                                                                                                                                                                                                                              |                       |                          |                |
| Sinus rhythm                                                                                                                                                                                                                                                                                                                                                                         | 9/20 (45%)            | 19/20 (95%)              | <0.01          |
| Non-sinus rhythm                                                                                                                                                                                                                                                                                                                                                                     | 11/20 (55%)           | 1/20 (5%)                | <0.01          |
| Atrioventricular and intraventricular conduction                                                                                                                                                                                                                                                                                                                                     |                       |                          |                |
| AVB 1°                                                                                                                                                                                                                                                                                                                                                                               | 6/20 (30%)            | 2/20 (10%)               | 0.114          |
| AVB 2°                                                                                                                                                                                                                                                                                                                                                                               | 0/20 (0%)             | 0/20 (0%)                | -              |
| RBBB                                                                                                                                                                                                                                                                                                                                                                                 | 2/20 (10%)            | 0/20 (0%)                | -              |
| LBBB                                                                                                                                                                                                                                                                                                                                                                                 | 2/20 (10%)            | 0/20 (0%)                | -              |
| LAFB                                                                                                                                                                                                                                                                                                                                                                                 | 3/20 (15%)            | 0/20 (0%)                | -              |
| Overloads                                                                                                                                                                                                                                                                                                                                                                            |                       |                          |                |
| Left ventricle                                                                                                                                                                                                                                                                                                                                                                       | 3/20 (15%)            | 2/20 (10%)               | 0.633          |
| Right ventricle                                                                                                                                                                                                                                                                                                                                                                      | 0/20 (0%)             | 0/20 (0%)                | -              |
| Left atrium                                                                                                                                                                                                                                                                                                                                                                          | 4/20 (20%)            | 0/20 (0%)                | 0.035          |
| Right atrium                                                                                                                                                                                                                                                                                                                                                                         | 1/20 (5%)             | 0/20 (0%)                | 0.311          |
| Low voltage                                                                                                                                                                                                                                                                                                                                                                          | 11/20 (55%)           | 3/20 (15%)               | 0.008          |
| Repolarization changes                                                                                                                                                                                                                                                                                                                                                               | 13/20 (65%)           | 6/20 (30%)               | 0.027          |
| Inactive area                                                                                                                                                                                                                                                                                                                                                                        | 6/20 (30%)            | 0/20 (0%)                | 0.008          |
| Anteroseptal wall                                                                                                                                                                                                                                                                                                                                                                    | 1/6 (16%)             | N/A                      | -              |
| Inferior wall                                                                                                                                                                                                                                                                                                                                                                        | 5/6 (84%)             | N/A                      | -              |
| Values are expressed as mean ± standard deviation, n/N (%), or median (25 <sup>th</sup> –75 <sup>th</sup> percentiles), as appropriate. N/A, not applicable; ATTR, transthyretin amyloidosis; CA, cardiac amyloidosis; ECG, electrocardiography; AVB, atrioventricular block; RBBB, right bundle branch block; LBBB, left bundle branch block; LAFB, left anterior fascicular block. |                       |                          |                |

| Supplementary Table 5. Global CFR clinical correlations                                                                                                                                                            |        |                |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------|----------------|
| Correlation                                                                                                                                                                                                        | r      | <i>P</i> value |
| Age                                                                                                                                                                                                                | −0.701 | <0.001         |
| Weight (kg)                                                                                                                                                                                                        | 0.174  | 0.309          |
| Height (m)                                                                                                                                                                                                         | 0.309  | 0.067          |
| BMI                                                                                                                                                                                                                | 0.049  | 0.785          |
| Heart rate (bpm)                                                                                                                                                                                                   | −0.285 | 0.079          |
| SBP (mmHg)                                                                                                                                                                                                         | 0.149  | 0.367          |
| MBP (mmHg)                                                                                                                                                                                                         | 0.344  | 0.030          |
| Functional class (NYHA)                                                                                                                                                                                            | −0.553 | <0.001         |
| 6-minute walk test distance (m)                                                                                                                                                                                    | 0.696  | <0.001         |
| Troponin (ng/L)                                                                                                                                                                                                    | −0.581 | <0.001         |
| BNP (pg/mL)                                                                                                                                                                                                        | −0.734 | <0.001         |
| CFR, coronary flow reserve; BMI, body mass index; SBP, systolic blood pressure; MBP, mean blood pressure; bpm, beats per minute; BNP, B-type natriuretic peptide; NYHA, New York Heart Association classification. |        |                |