## Participant Flow



## Baseline Characteristics

| Number of patients | $\mathrm{n}=747$ |  |
| :---: | :---: | :---: |
| Study sites | $\mathrm{n}=260$ |  |
| Patients per study site (mode) | $\mathrm{n}=3$ |  |
| Duration of observation per patient (months, $\mathrm{n}=732$ ) |  |  |
| Mean $\pm$ SD | $3.94 \pm 0.92$ |  |
| Sex (gender, $\mathrm{n}=729$ ) |  |  |
| Female | $\mathrm{n}=274$ | 37.59\% |
| Male | $\mathrm{n}=455$ | 62.41\% |
| Age (years, $\mathrm{n}=744$ ) |  |  |
| Mean $\pm$ SD | $66.39 \pm 10.75$ |  |
| Minimum | 25.00 |  |
| Maximum | 95.00 |  |
| Body mass index (kg/m², $\mathrm{n}=739$ ) |  |  |
| Mean $\pm$ SD | $28.60 \pm 4.77$ |  |
| Minimum | 17.04 |  |
| Maximum | 51.98 |  |
| Duration of coronary artery disease (months, $\mathrm{n}=717$ ) |  |  |
| Mean $\pm$ SD | $69.95 \pm 64.18$ |  |
| Minimum | 0.20 (six days) |  |
| Maximum | 325.28 (27.10 years) |  |
| Anamnesis of coronary artery disease ( $\mathrm{n}=711$ ) |  |  |
| Single vessel disease | $\mathrm{n}=227$ | 31.93\% |
| Double vessel disease | $\mathrm{n}=213$ | 29.96\% |
| Triple vessel disease | $\mathrm{n}=152$ | 21.38\% |
| Coronary sclerosis | $\mathrm{n}=118$ | 16.60\% |
| Vasospastic angina | $\mathrm{n}=1$ | 0.14\% |
| Duration of chronic stable angina pectoris (months, $n=716$ ) |  |  |
| Mean $\pm$ SD | $\mathrm{n}=59.39 \pm 60.43$ |  |
| Minimum | $\mathrm{n}=0.20$ (six days) |  |
| Maximum | $\mathrm{n}=469.80$ (39.15 years) |  |


| Heart rate (bpm; $\mathrm{n}=732$, LOCF) |  |  |
| :---: | :---: | :---: |
| Mean $\pm$ SD | 76.83 | $\pm 11.75$ |
| Blood pressure systolic/diastolic (mmHg; $\mathrm{n}=729$, LOCF) |  |  |
| Mean $\pm$ SD | 137.64/81.81 | $\pm 15.14 / 10.12$ |
| Angina pectoris attacks in the last seven days ( $\mathrm{n}=736$, LOCF) |  |  |
| Mean $\pm$ SD | 1.02 | $\pm 2.52$ |
| Need for short acting nitrates in the last seven days ( $\mathrm{n}=717$, LOCF) |  |  |
| Mean $\pm$ SD | 1.25 | $\pm 3.03$ |
| Angina pectoris classification according to the Canadian Cardiovascular Society (CCS, $n=724$ ) |  |  |
| Class I | $\mathrm{n}=194$ | 26.80\% |
| Class II | $\mathrm{n}=393$ | 54.28\% |
| Class III | $\mathrm{n}=134$ | 18.51\% |
| Class IV | $\mathrm{n}=3$ | 0.41\% |
| Classification of heart failure according to the to the New York Heart Association (NYHA, $n=722$ ) |  |  |
| NYHA-class I | $\mathrm{n}=92$ | 12.74\% |
| NYHA-class II | $\mathrm{n}=243$ | 33.66\% |
| NYHA-class III | $\mathrm{n}=61$ | 8.45\% |
| NYHA-class IV | $\mathrm{n}=5$ | 0.69\% |
| No heart failure | $\mathrm{n}=321$ | 44.46\% |
| Ejection fraction (\%, n=99) |  |  |
| Mean $\pm$ SD | 43.00 | $\pm 10.89$ |
| Myocardial infarction ( $n=744$ ) |  |  |
| Yes | $\mathrm{n}=233$ | 31.32\% |
| No | $\mathrm{n}=511$ | 68.68\% |
| Frequency of a myocardial infarction ( $\mathrm{n}=220$ ) |  |  |
| One infarction | $\mathrm{n}=181$ | 82.27\% |
| Two infarctions | $\mathrm{n}=28$ | 12.73\% |
| Three infarctions | $\mathrm{n}=6$ | 2.73\% |
| Four infarctions | $\mathrm{n}=5$ | 2.27\% |
| Chronic heart failure ( $\mathrm{n}=743$ ) |  |  |


| Yes | $\mathrm{n}=414$ | 55.72\% |
| :---: | :---: | :---: |
| No | $\mathrm{n}=329$ | 44.28\% |
| Left ventricular dysfunction ( $\mathrm{n}=739$ ) |  |  |
| Yes | $\mathrm{n}=242$ | 32.75\% |
| No | $\mathrm{n}=497$ | 67.25\% |
| Cardiac valve disease ( $\mathrm{n}=741$ ) |  |  |
| Yes | $\mathrm{n}=149$ | 20.11\% |
| No | $\mathrm{n}=592$ | 79.89\% |
| Disturbance of conduction ( $\mathrm{n}=730$ ) |  |  |
| Yes | $\mathrm{n}=166$ | 22.74\% |
| No | $\mathrm{n}=564$ | 77.26\% |
| Atrioventricular block ( $\mathrm{n}=747$ ) |  |  |
| Yes | $\mathrm{n}=56$ | 7.50\% |
| No | $\mathrm{n}=691$ | 92.50\% |
| Atrioventricular block degree ( $\mathrm{n}=48$ ) |  |  |
| First degree | $\mathrm{n}=37$ | 77.08\% |
| Second degree | $\mathrm{n}=7$ | 14.58\% |
| Third degree | $\mathrm{n}=4$ | 8.33\% |
| Atrial fibrillation ( $\mathrm{n}=716$ ) |  |  |
| Yes | $\mathrm{n}=109$ | 15.22\% |
| No | $\mathrm{n}=607$ | 84.78\% |
| Non-medicinal treatments ( $\mathrm{n}=747$ ) |  |  |
| Without non-medicinal treatments | $\mathrm{n}=316$ | 42.30\% |
| With non-medicinal treatments | $\mathrm{n}=431$ | 57.70\% |
| - Percutaneous coronary intervention ( PCl ) with implantation of stent | $\mathrm{n}=376$ | 50.33\% |
| - Coronary artery bypass graft (CABG) | $\mathrm{n}=76$ | 10.17\% |
| - Heart pacemaker implantation | $\mathrm{n}=56$ | 7.50\% |
| Cardiovascular risk factors ( $\mathrm{n}=747$ ) |  |  |
| Without risk factors | $\mathrm{n}=5$ | 0.67\% |


| With risk factors | $\mathrm{n}=742$ | 99.33\% |
| :---: | :---: | :---: |
| Concomitant diseases ( $\mathrm{n}=747$ ) |  |  |
| Without concomitant diseases | $\mathrm{n}=126$ | 16.87\% |
| With concomitant diseases | $\mathrm{n}=621$ | 83.13\% |
| Adherence at admission (Patient questionnaire, $\mathrm{n}=747$ ) |  |  |
| Complete compliance (all questions were answered with never) | $\mathrm{n}=241$ | 33.66\% |
| Incomplete compliance | $\mathrm{n}=475$ | 66.34\% |
| Missing | $\mathrm{n}=31$ |  |
| Pre-treatment of stable angina pectoris ( $\mathrm{n}=747$ ) |  |  |
| Beta-blocker | $\mathrm{n}=706$ | 94.51\% |
| Ivabradine (Procoralan ${ }^{\text {® }}$ ) | $\mathrm{n}=665$ | 89.02\% |
| Hospitalization due to deterioration of angina pectoris symptoms within the last three months prior to admission ( $n=747$ ) |  |  |
| Yes | $\mathrm{n}=47$ | 6.30\% |
| No | $\mathrm{n}=699$ | 93.70\% |
| Missing | $\mathrm{n}=1$ |  |

## Outcome Measures

| Heart rate (bpm; n=732, LOCF) |  |  |
| :---: | :---: | :---: |
| Mean $\pm$ SD | 67.23 | $\pm 8.21$ |
| Blood pressure systolic/diastolic (mmHg; $\mathbf{n = 7 2 9 , ~ L O C F ) ~}$ |  |  |
| Mean $\pm$ SD | 129.07/77.45 | $\pm 12.54 / \pm 7.79$ |
| Angina pectoris attacks in the last seven days ( $\mathrm{n}=736$, LOCF) |  |  |
| Mean $\pm$ SD | 0.17 | $\pm 0.85$ |
| Need for short acting nitrates in the last seven days ( $n=717$ ) |  |  |
| Mean $\pm$ SD | 0.24 | $\pm 1.27$ |
| Angina pectoris classification according to the Canadian Cardiovascular Society (CCS, $\mathrm{n}=672$ ) |  |  |
| Class I | $\mathrm{n}=423$ | 62.95\% |
| Class II | $\mathrm{n}=212$ | 31.55\% |
| Class III | $\mathrm{n}=36$ | 5.36\% |
| Class IV | $\mathrm{n}=1$ | 0.15\% |
| Classification of heart failure according to the to the New York Heart Association (NYHA, $n=722$ ) |  |  |
| NYHA-class I | $\mathrm{n}=145$ | 20.08\% |
| NYHA-class II | $\mathrm{n}=164$ | 22.71\% |
| NYHA-class III | $\mathrm{n}=37$ | 5.12\% |
| NYHA-class IV | $\mathrm{n}=2$ | 0.28\% |
| No heart failure | $\mathrm{n}=374$ | 51.80\% |
| Ejection fraction (\%, $\mathrm{n}=99$ ) |  |  |
| Mean $\pm$ SD | 46.73 | $\pm 11.22$ |
| Physicians' global assessment of effectiveness ( $\mathrm{n}=730$ ) |  |  |
| Very good | $\mathrm{n}=515$ | 70.55 \% |
| Good | $\mathrm{n}=196$ | 26.85 \% |
| Moderate | $\mathrm{n}=16$ | 2.19 \% |
| Poor | $\mathrm{n}=3$ | 0.41\% |


| Hospitalization due to deterioration of angina pectoris symptoms (n=747) |  |  |
| :--- | :--- | :--- |
| Yes | $\mathrm{n}=8$ | $1.09 \%$ |
| No | $\mathrm{n}=724$ | $98.91 \%$ |
| Missing | $\mathrm{n}=15$ |  |
| Adherence (n=747) | $\mathrm{n}=415$ | $58.04 \%$ |
| Complete compliance (all <br> questions were answered with <br> never) | $\mathrm{n}=300$ | $41.96 \%$ |
| Incomplete compliance | $\mathrm{n}=32$ |  |
| Missing |  |  |

## Adverse Events

Category and frequency of events occurred and number of patients affected ( $n=747$ )

| Category | Frequency events | Frequency patients |  |
| :--- | :---: | :---: | :---: |
|  |  | $\mathbf{\%}$ |  |
| Adverse events (AE), excluding product use issue | 58 | 146 | 19.54 |
| $-\quad$ Non-serious AE | 54 | 40 | $5.35 \%$ |
| $-\quad$ Serious AE | 4 | 36 | $4.82 \%$ |
| Adverse drug reaction (ADR) | 32 | 4 | $0.54 \%$ |

Adverse events listed according to the preferred term by MedDRA 19.1.

| MedDRA 19.1 / Preferred term | Patients | Percent |
| :--- | :---: | :---: |
| Product use issue | 113 | $15.13 \%$ |
| Blood pressure increased | 6 | $0.80 \%$ |
| Dizziness | 6 | $0.80 \%$ |
| Headache | 5 | $0.67 \%$ |
| Chest discomfort | 3 | $0.40 \%$ |
| Dyspnoea | 2 | $0.40 \%$ |
| Angina pectoris | 2 | $0.27 \%$ |
| Bradycardia | 2 | $0.27 \%$ |
| Cardiac failure | 2 | $0.27 \%$ |
| Chest pain | 2 | $0.27 \%$ |
| Drug ineffective | 2 | $0.27 \%$ |
| Fatigue | 2 | $0.27 \%$ |
| Hypotension | 1 | $0.27 \%$ |
| Percutaneous coronary intervention | 1 | $0.13 \%$ |
| Abdominal pain upper | 1 | $0.13 \%$ |
| Angina unstable | 2 | $0.13 \%$ |
| Asthenia | 2 | $0.13 \%$ |
| Blood pressure decreased | 2 | 0. |
| Blood pressure diastolic increased | 2 | 0 |
|  | 2 | 0. |


| MedDRA 19.1 / Preferred term | Patients | Percent |
| :--- | :---: | :---: |
| Cardiac resynchronisation therapy | 1 | $0.13 \%$ |
| Cerebrovascular accident | 1 | $0.13 \%$ |
| Cough | 1 | $0.13 \%$ |
| Dry mouth | 1 | $0.13 \%$ |
| Heart rate decreased | 1 | $0.13 \%$ |
| Heart rate increased | 1 | $0.13 \%$ |
| Hypertensive crisis | 1 | $0.13 \%$ |
| Hypertensive emergency | 1 | $0.13 \%$ |
| Multiple organ dysfunction syndrome | 1 | $0.13 \%$ |
| Myocardial infarction | 1 | $0.13 \%$ |
| Nausea | 1 | $0.13 \%$ |
| Physical examination | 1 | $0.13 \%$ |
| Skin ulcer | 1 | $0.13 \%$ |
| Treatment noncompliance | 1 | $0.13 \%$ |

