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## *Supplementary Materials*

# Utility of the JT Peak Interval and the JT Area in Determining the Proarrhythmic Potential of QT-Shortening Agents

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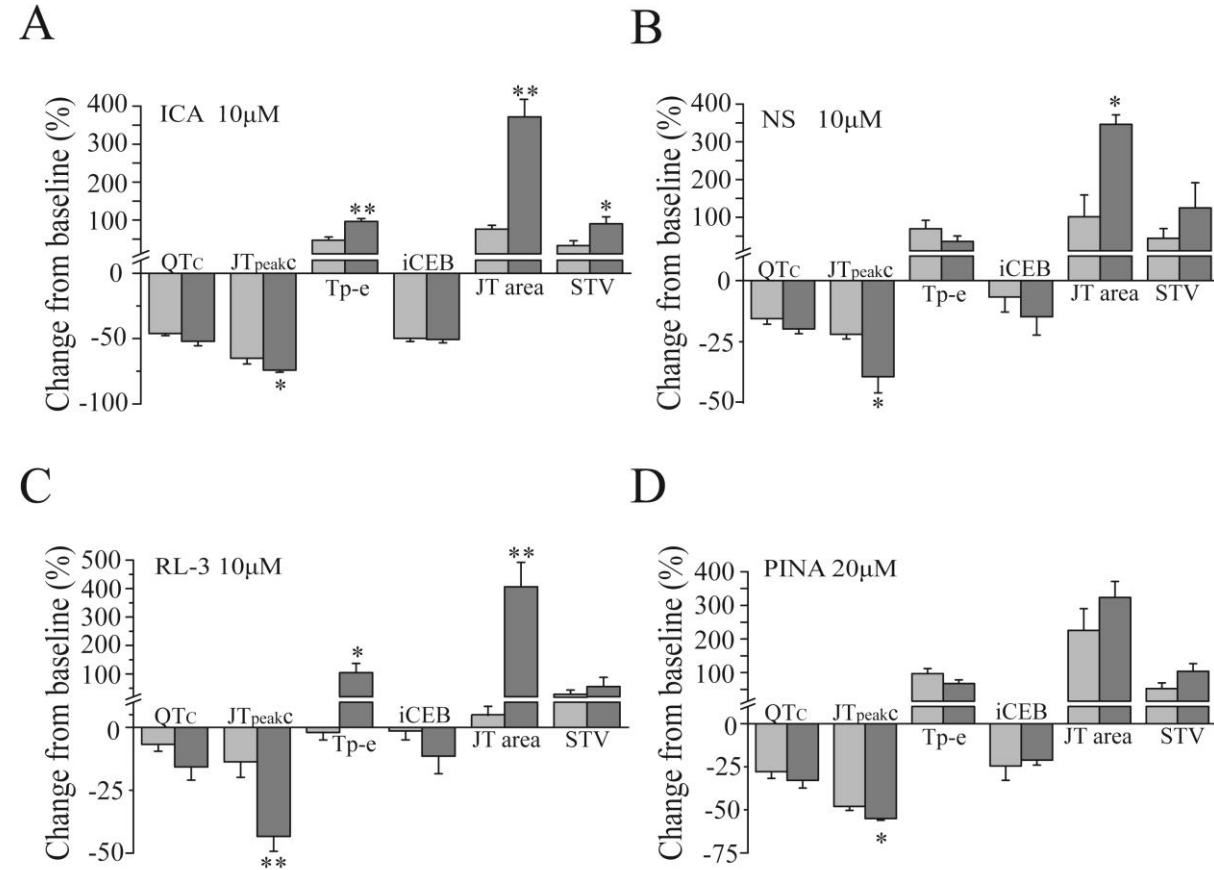
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## Supplementary Figures and tables

Supplementary Figure.1



**Supplementary Figure 1.** Effects of potassium channel openers on ECG parameters in isolated rabbit (light-colored) and guinea pig hearts (dark-colored) at high concentrations.

**Supplement Table 1. Effects of different potassium channel openers on the ECG parameters in the Langendorff-perfused guinea-pigs hearts.**

|     |                 | QTc (ms)            | JT <sub>peakc</sub> (ms) | Tp-e (ms)          | iCEB                  | JTarea (mV•ms)     | STV (ms)              |
|-----|-----------------|---------------------|--------------------------|--------------------|-----------------------|--------------------|-----------------------|
| n=8 | Control         | 159±2               | 120±1                    | 25±1               | 9±0.1                 | 8±0.5              | 1.59±0.1              |
|     | ICA-105574 1μM  | 138±3 <sup>**</sup> | 107±3 <sup>**</sup>      | 36±1 <sup>**</sup> | 7±0.2 <sup>**</sup>   | 16±1 <sup>**</sup> | 2.1±0.2 <sup>*</sup>  |
|     | ICA-105574 5μM  | 106±3 <sup>**</sup> | 60±5 <sup>**</sup>       | 48±1 <sup>**</sup> | 6±0.2 <sup>**</sup>   | 27±2 <sup>**</sup> | 1.9±0.2               |
|     | ICA-105574 10μM | 77±2 <sup>**</sup>  | 33±1 <sup>**</sup>       | 47±1 <sup>**</sup> | 4±0.1 <sup>**</sup>   | 38±3 <sup>**</sup> | 3.2±0.3 <sup>**</sup> |
| n=5 | Control         | 159±3               | 113±5                    | 28±2               | 5.6±0.6               | 8±1                | 1.6±0.4               |
|     | NS-1643 1μM     | 146±3 <sup>**</sup> | 96±5 <sup>**</sup>       | 33±1               | 5.1±0.4               | 18±1 <sup>**</sup> | 1.9±0.8               |
|     | NS-1643 5μM     | 134±3 <sup>**</sup> | 85±6 <sup>**</sup>       | 33±1               | 5.3±0.5               | 29±4 <sup>**</sup> | 2.8±0.5               |
|     | NS-1643 10μM    | 127±2 <sup>**</sup> | 69±5 <sup>**</sup>       | 34±1               | 4.9±0.6               | 37±3 <sup>**</sup> | 3.7±1.1 <sup>*</sup>  |
| n=5 | Control         | 161±2               | 113±5                    | 22±4               | 8.2±0.7               | 9±1                | 1.7±0.1               |
|     | L-3 1μM         | 154±2 <sup>*</sup>  | 95±5 <sup>**</sup>       | 27±3               | 7.5±0.7               | 16±2 <sup>*</sup>  | 2.1±0.3               |
|     | L-3 5μM         | 145±3 <sup>*</sup>  | 78±4 <sup>**</sup>       | 42±4               | 7.6±0.7               | 33±2 <sup>*</sup>  | 2.3±0.1               |
|     | L-3 10μM        | 136±4 <sup>**</sup> | 66±5 <sup>**</sup>       | 44±4               | 7.4±0.8               | 47±4 <sup>**</sup> | 2.5±0.4               |
| n=7 | Control         | 158±2               | 116±4                    | 22±1               | 7.6±0.1               | 8.3±1              | 1.8±0.1               |
|     | Pinacidil 5μM   | 143±3 <sup>**</sup> | 84±4 <sup>**</sup>       | 35±2 <sup>*</sup>  | 7.4±0.1               | 23±3 <sup>*</sup>  | 2.2±0.2               |
|     | Pinacidil 10μM  | 126±3 <sup>**</sup> | 68±3 <sup>**</sup>       | 37±2 <sup>**</sup> | 6.5±0.1 <sup>**</sup> | 28±3 <sup>*</sup>  | 2.9±0.3 <sup>*</sup>  |
|     | Pinacidil 20μM  | 106±4 <sup>**</sup> | 50±2 <sup>**</sup>       | 36±2 <sup>*</sup>  | 5.9±0.1 <sup>**</sup> | 37±4 <sup>*</sup>  | 3.6±0.5 <sup>*</sup>  |

**Supplement Table 2. Effects of different potassium channel openers on the parameters of PR (in ms), QRS (in ms) and heart rate (HR) in Langendorff-perfused guinea-pig hearts.**

|                 | HR                  | PR                 | QRS  |
|-----------------|---------------------|--------------------|------|
| Control         | 199±4               | 65±1               | 18±1 |
| ICA-105574 1µM  | 193±5               | 61±1               | 18±1 |
| ICA-105574 5µM  | 197±5               | 68±3               | 17±1 |
| ICA-105574 10µM | 186±8               | 70±1               | 18±1 |
| Control         | 187±4               | 68±3               | 30±3 |
| NS-1643 1µM     | 183±4               | 69±2               | 30±3 |
| NS-1643 5µM     | 176±4 <sup>**</sup> | 69±2               | 27±3 |
| NS-1643 10µM    | 175±4 <sup>**</sup> | 95±6 <sup>**</sup> | 31±3 |
| Control         | 186±6               | 63±1               | 26±3 |
| L-3 1µM         | 191±7               | 65±1               | 28±3 |
| L-3 5µM         | 174±6 <sup>*</sup>  | 66±1               | 27±3 |
| L-3 10µM        | 168±5 <sup>**</sup> | 68±2               | 27±3 |
| Control         | 187±3               | 62±2               | 23±1 |
| Pinacidil 5µM   | 175±5               | 59±2               | 22±2 |
| Pinacidil 10µM  | 161±5 <sup>**</sup> | 60±1               | 24±1 |
| Pinacidil 20µM  | 155±3 <sup>**</sup> | 64±2               | 23±2 |