

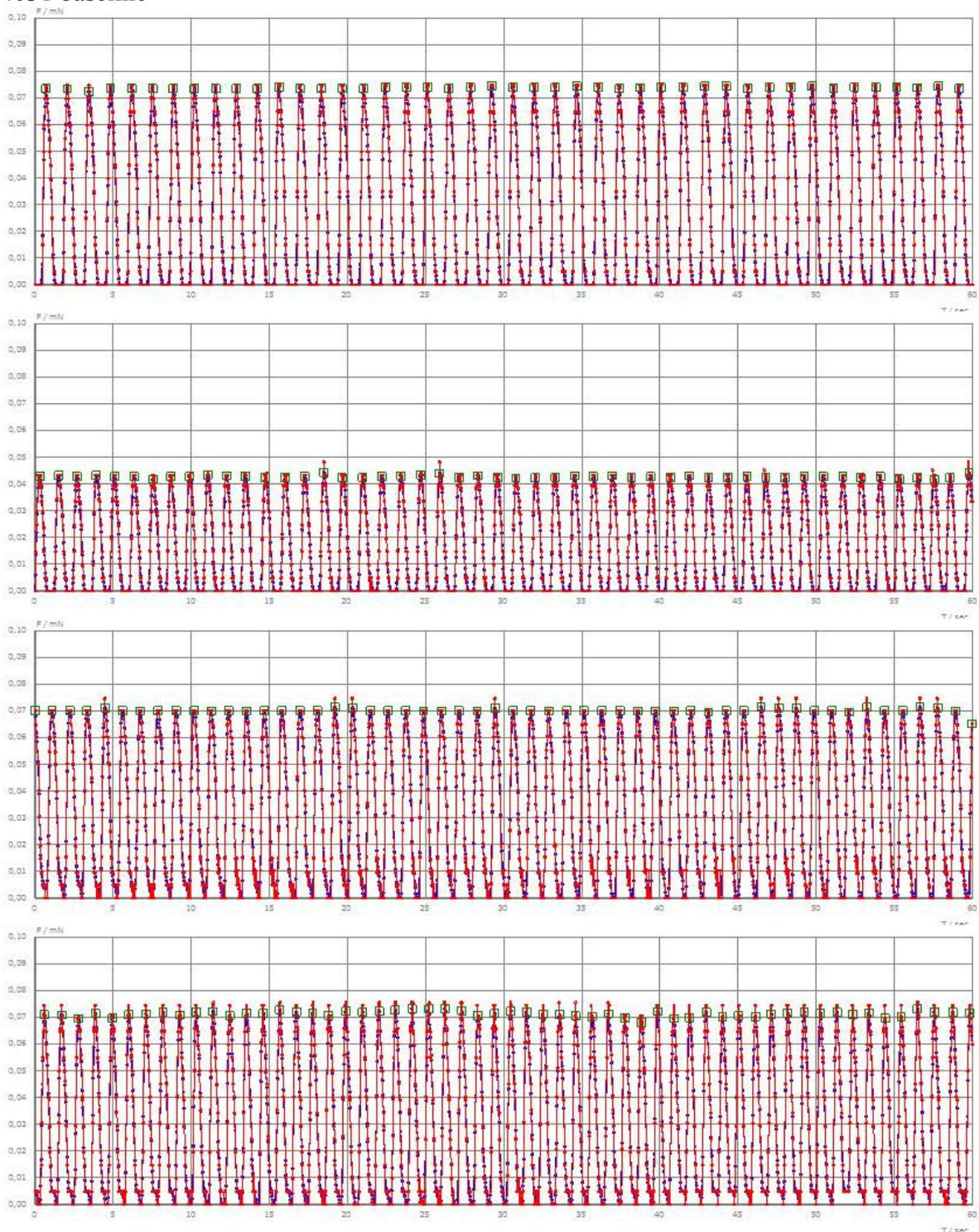
Supporting Information S1

Original recordings and analysis

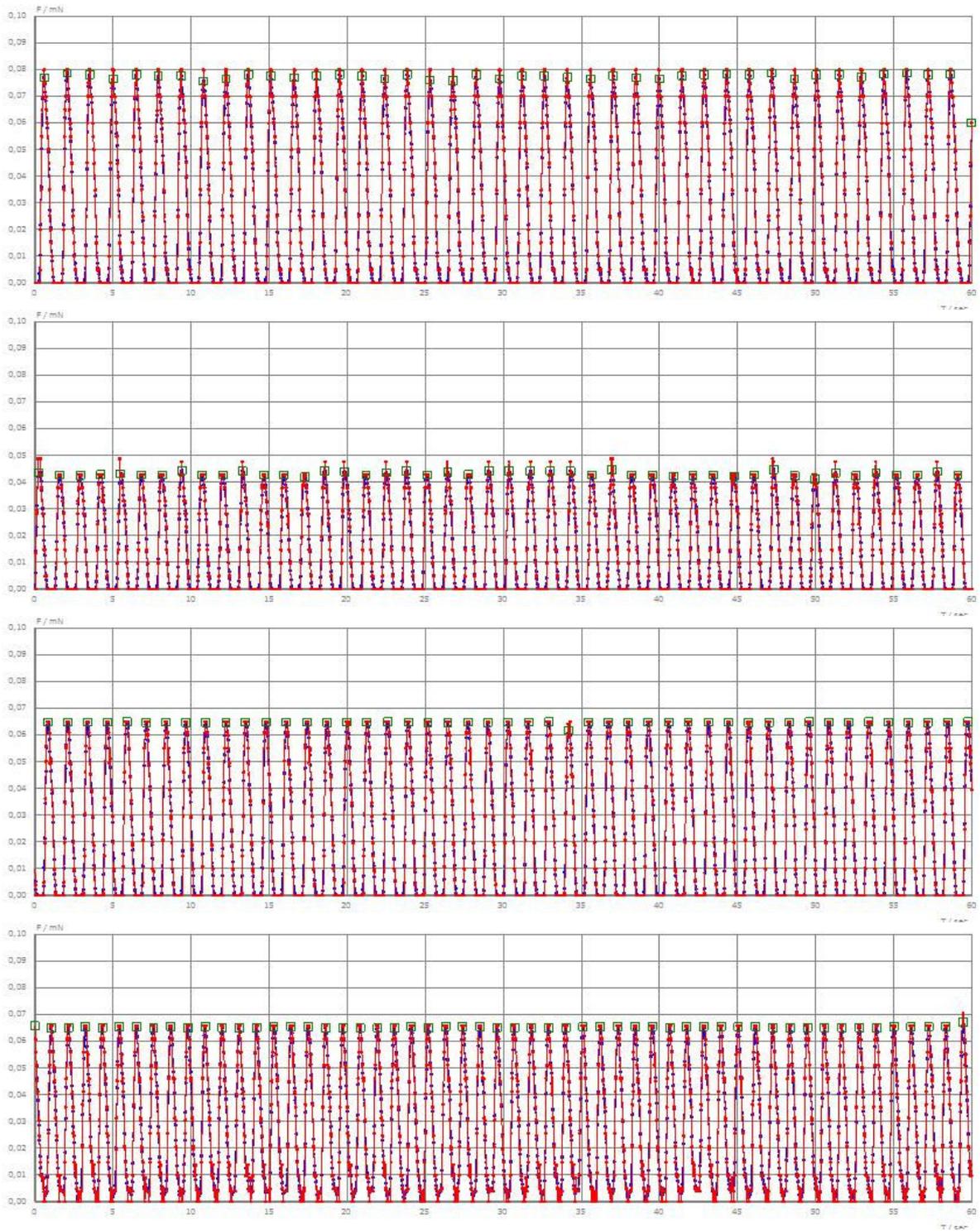
Original recordings of contraction patterns of four EHTs are presented under baseline condition and in the presence of proarrhythmic compounds.

E-4031 original recordings

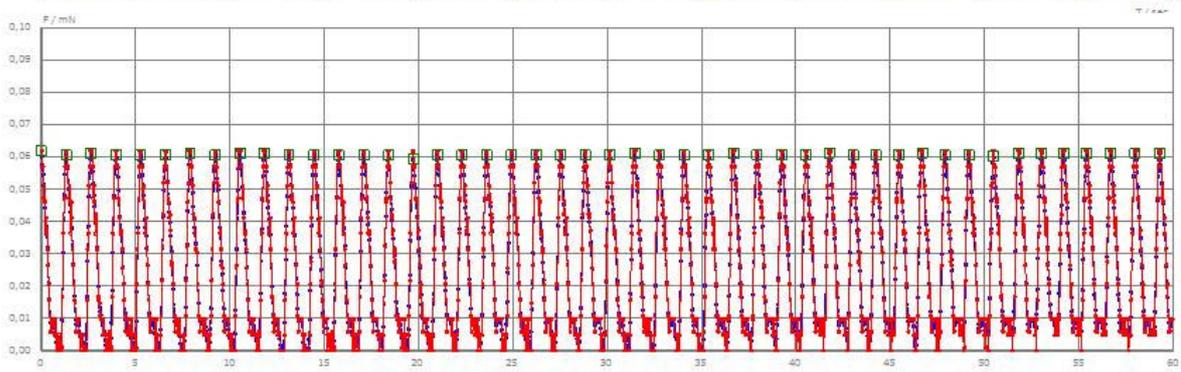
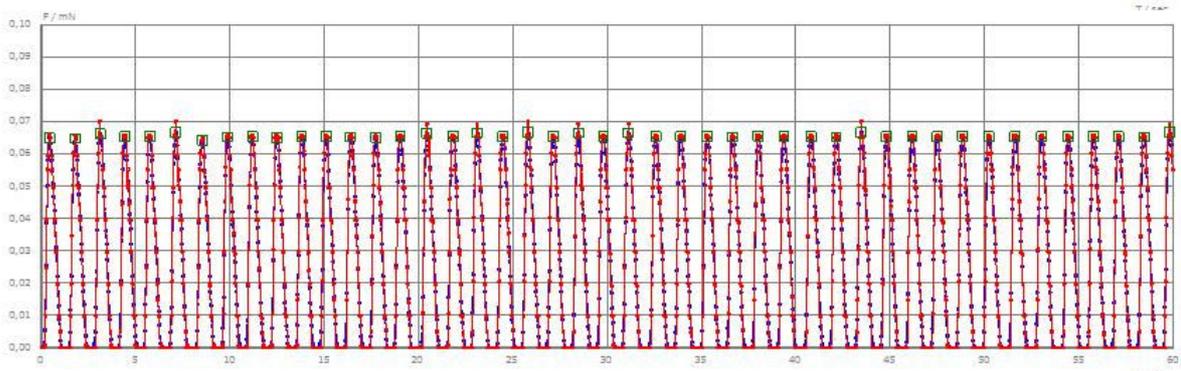
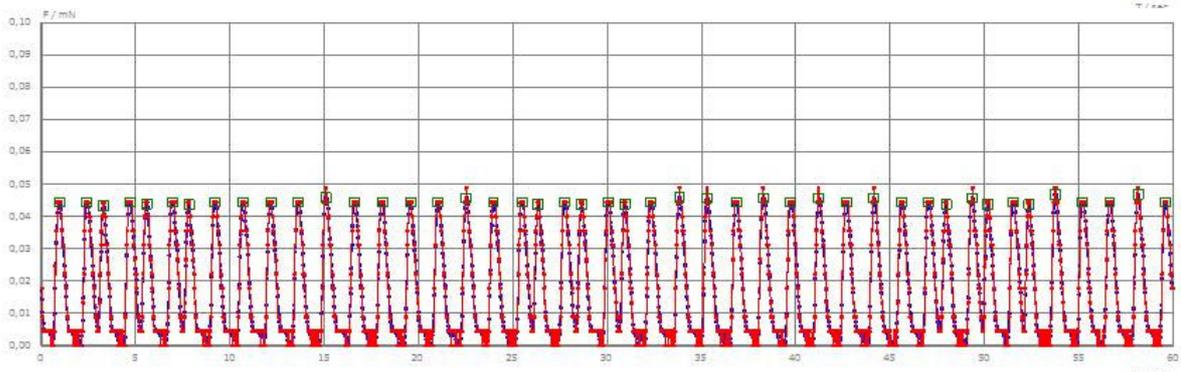
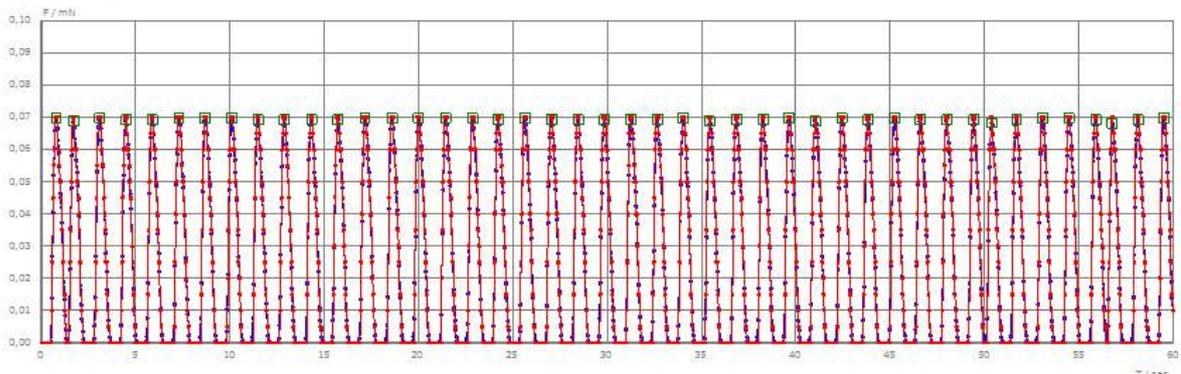
E-4031 baseline



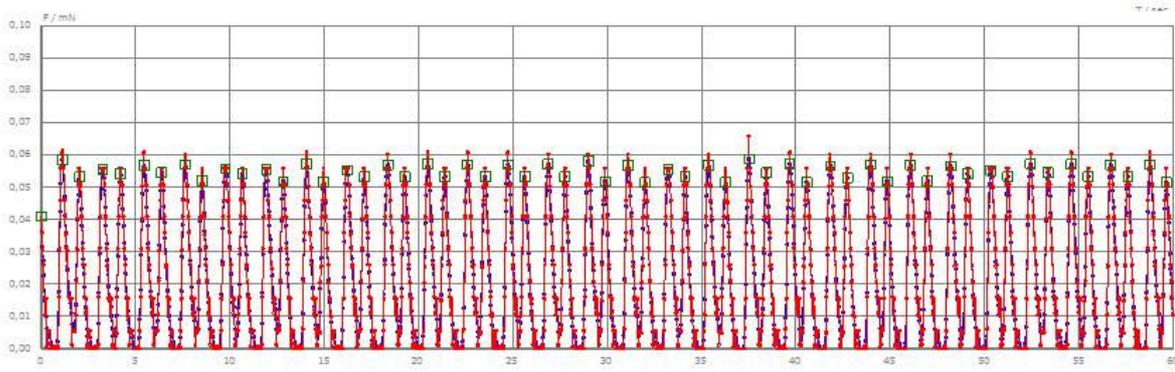
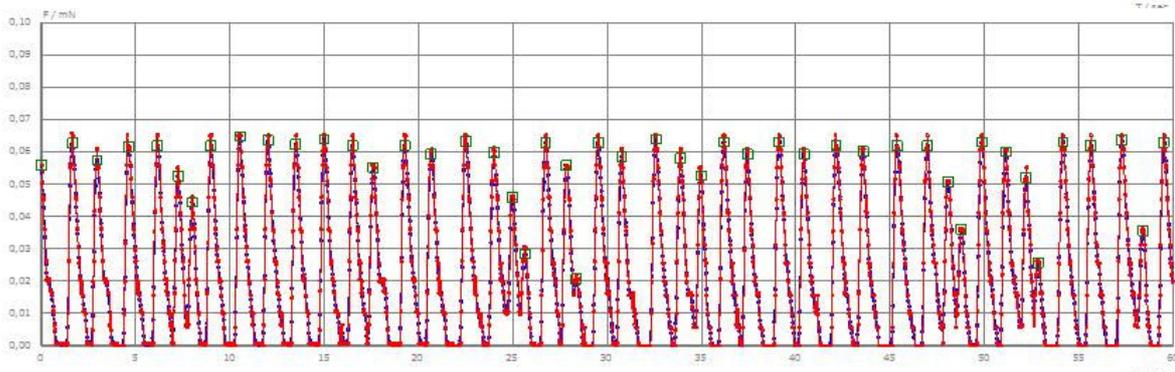
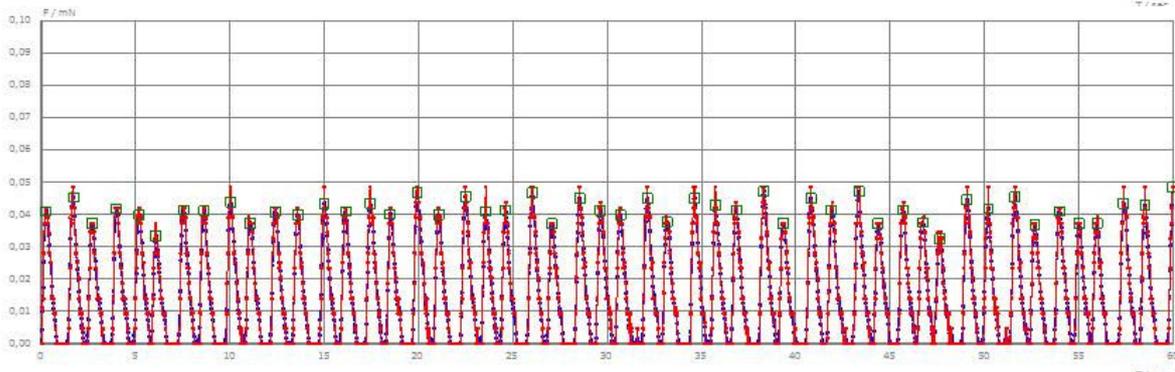
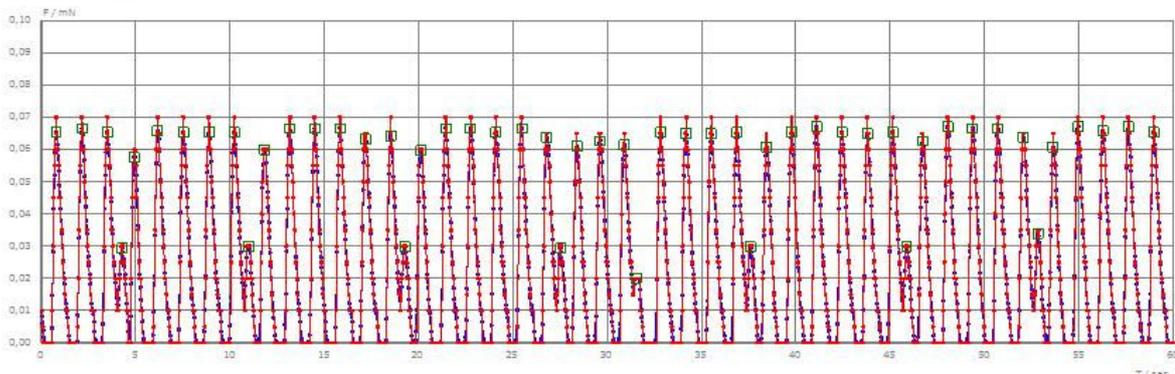
E-4031 1 nM



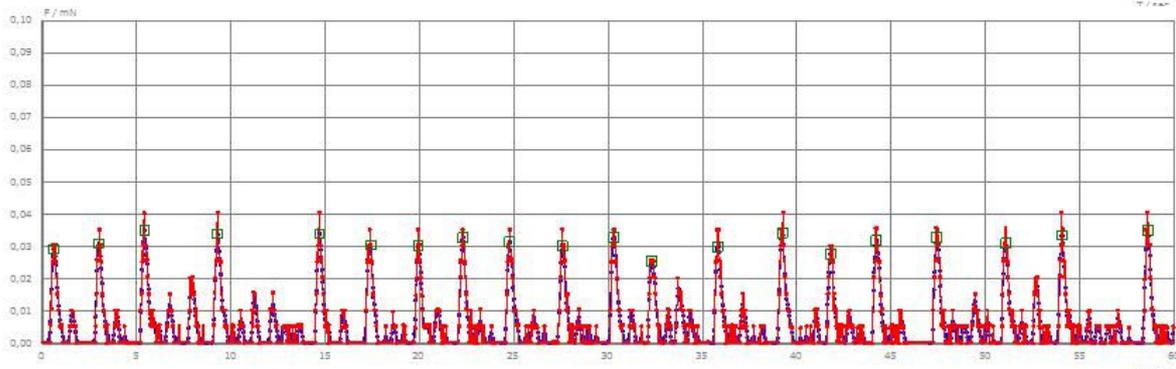
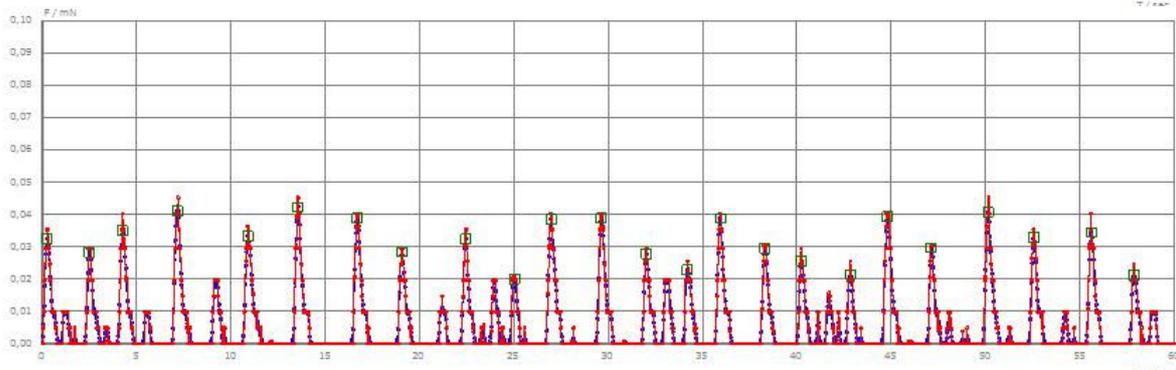
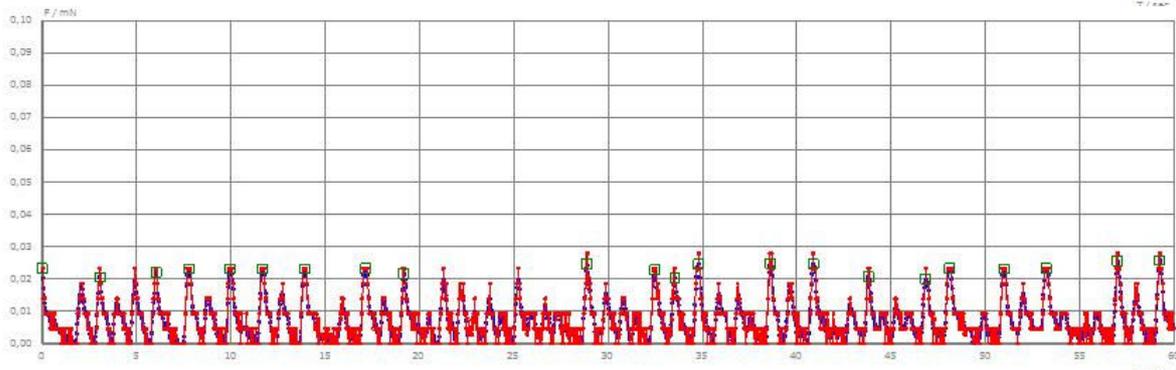
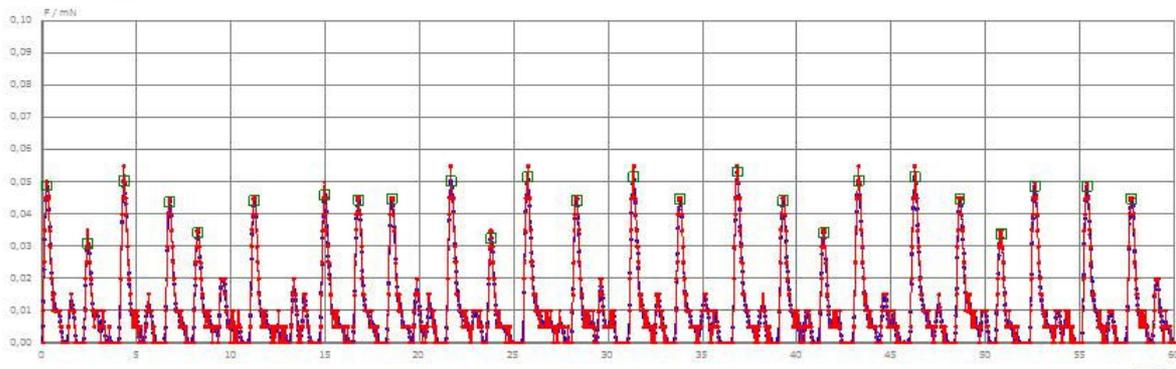
E-4031 3 nM



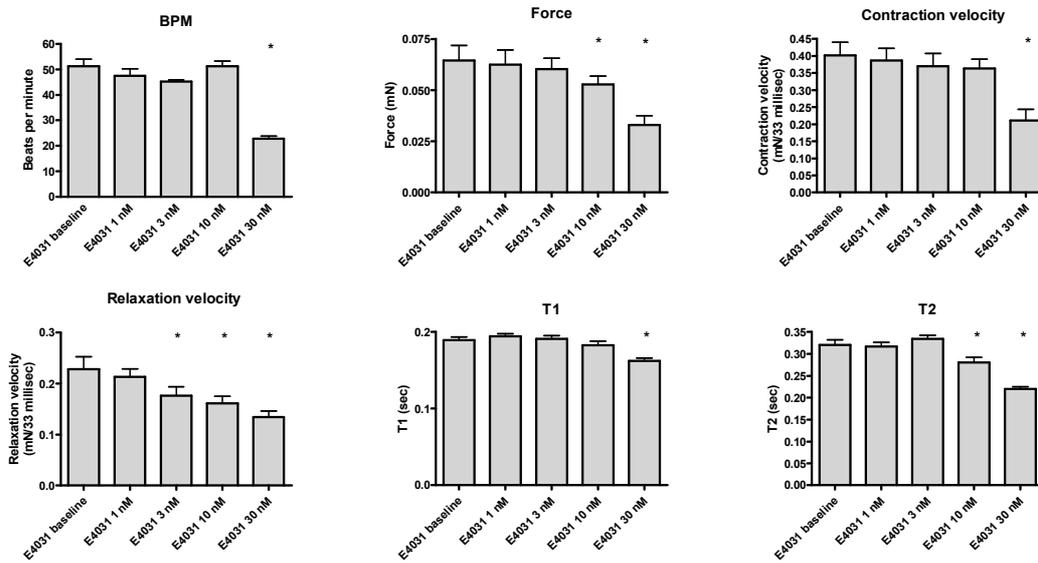
E-4031 10 nM



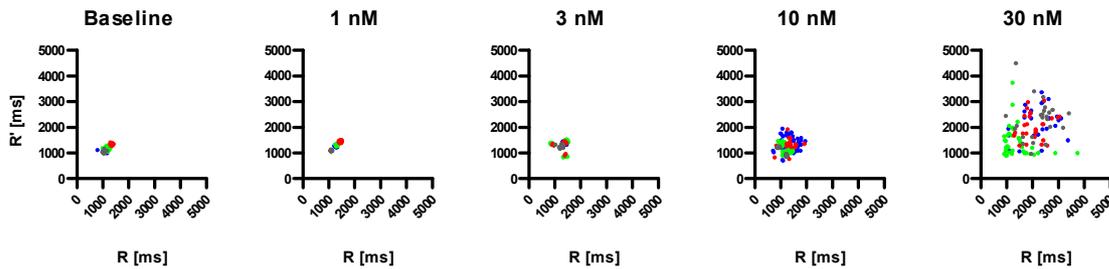
E-4031 30 nM



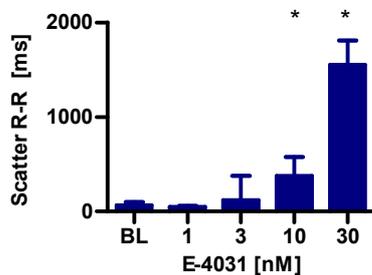
Analysis E-4031



Analysis of contraction. * $P < 0.05$ (Student's t-test), 4 biological replicas, bars show means \pm SD. Beats per minute (BPM), contraction time (T1), relaxation time (T2).



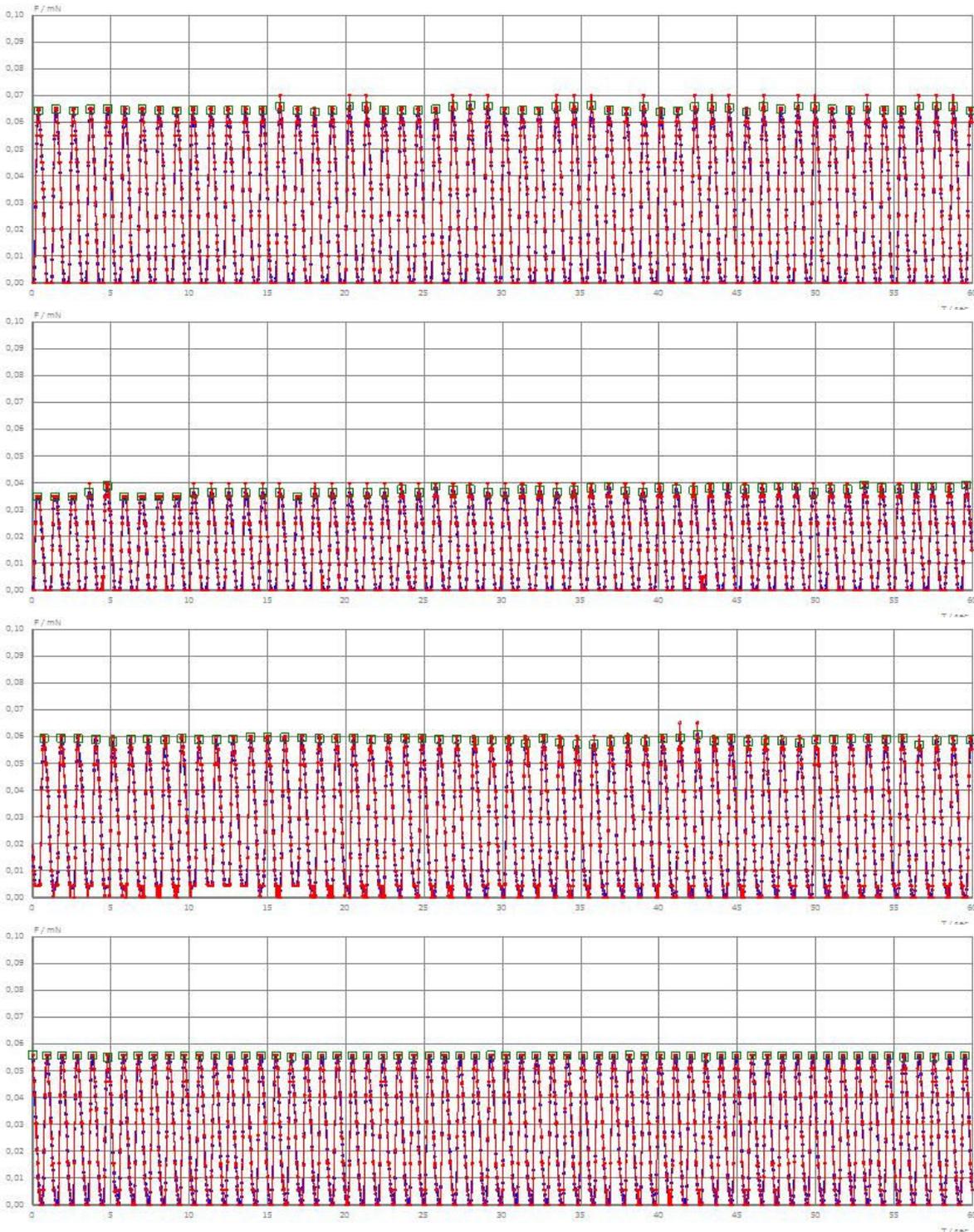
Graphical illustration of irregularity integrating 4 biological replicas. Ordinates indicate the distance from a given twitch to the following, the abscissa the distance to the previous twitch. Biological replicas are discriminated by color code.



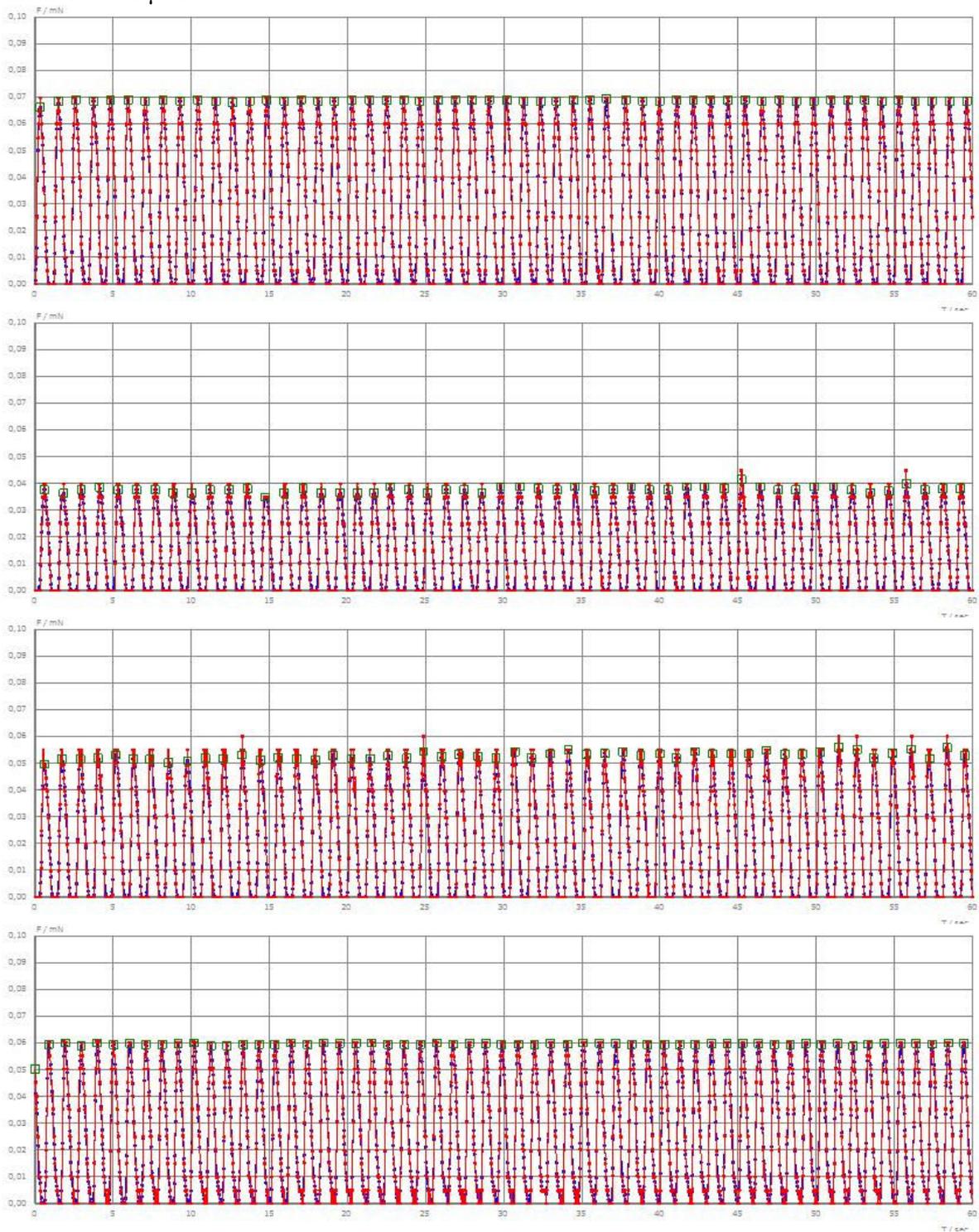
Scatter of beat-to-beat variability in the presence of E-4031, * $P < 0.05$ (Mann-Whitney U test), 4 biological replicas, bars show median \pm interquartile range.

Procainamide - original recordings

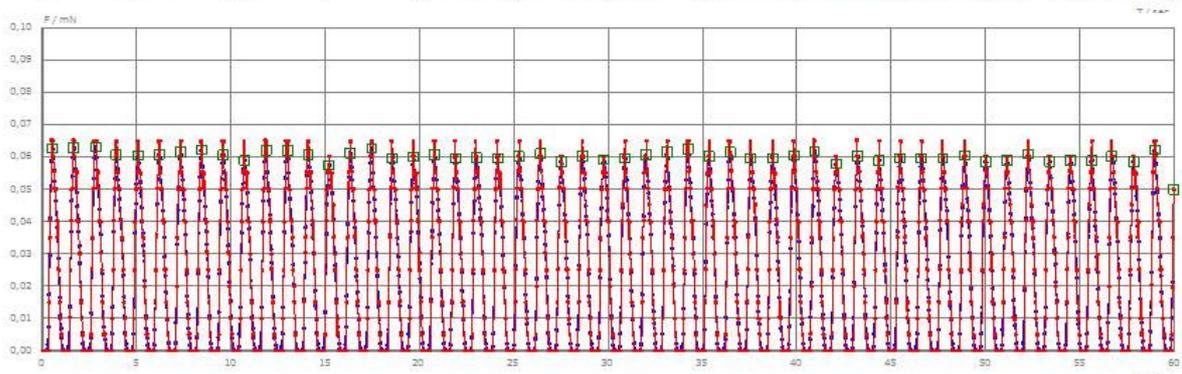
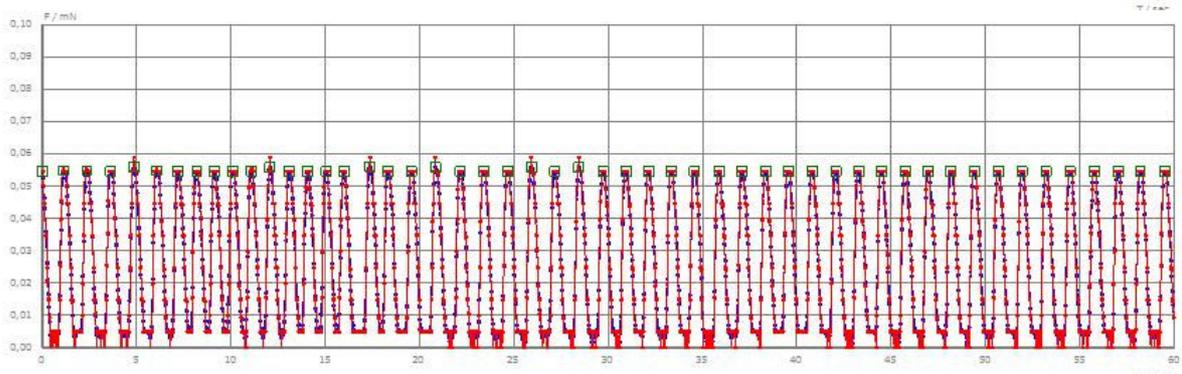
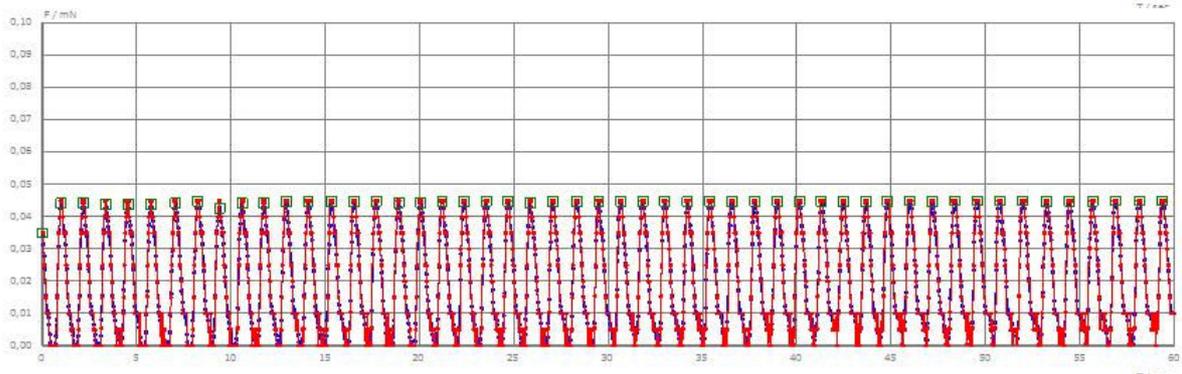
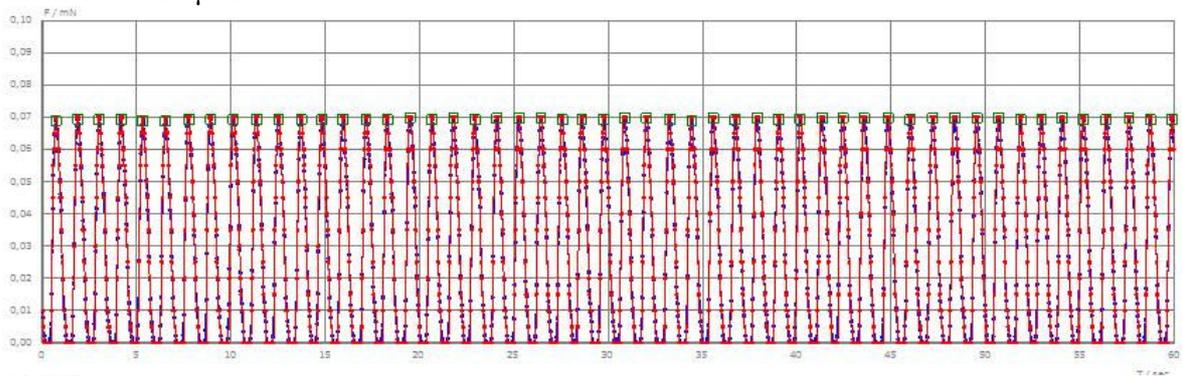
Procainamide baseline



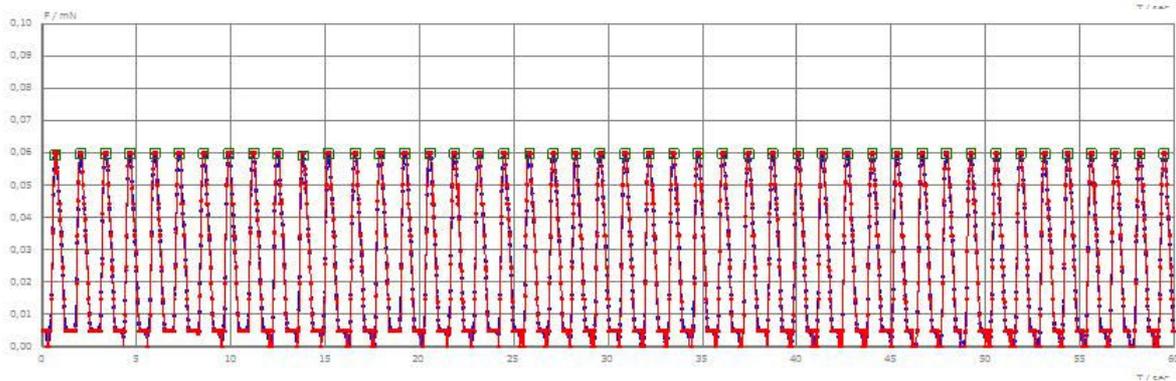
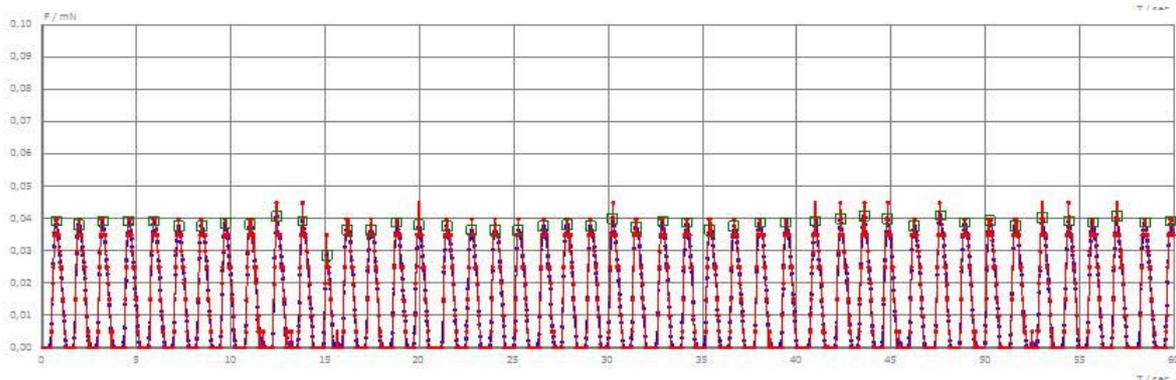
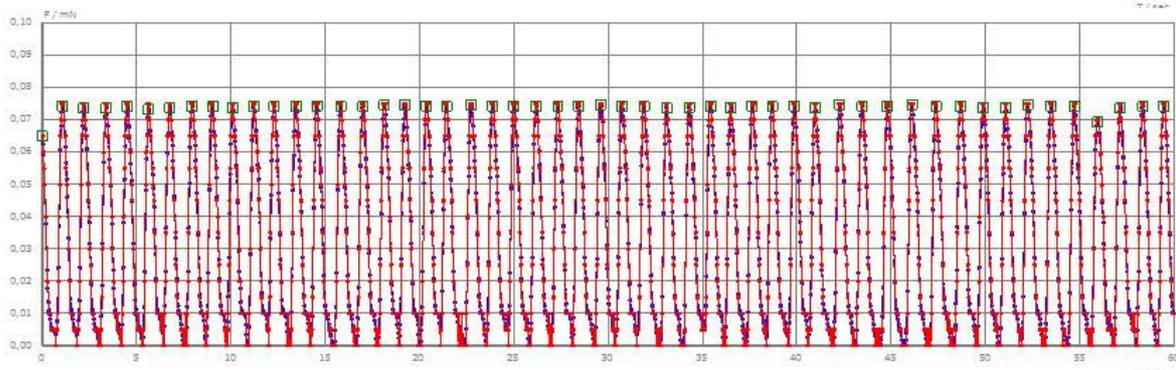
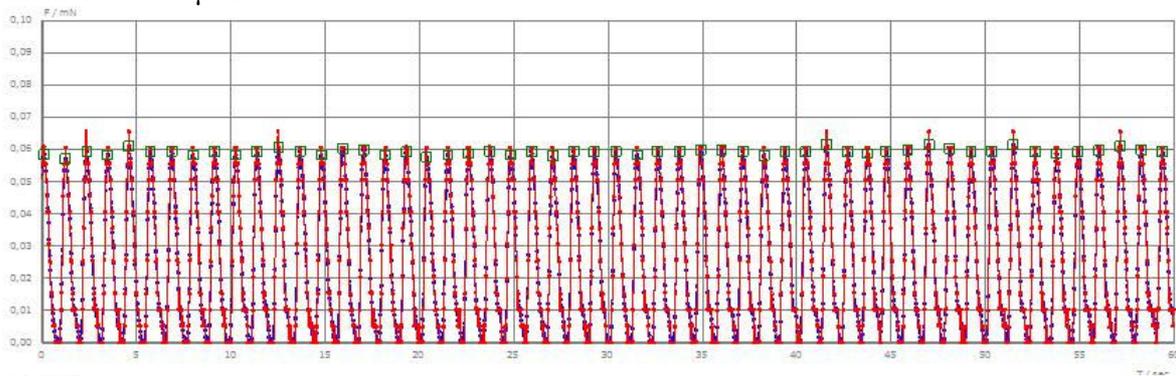
Procainamide 1 μM



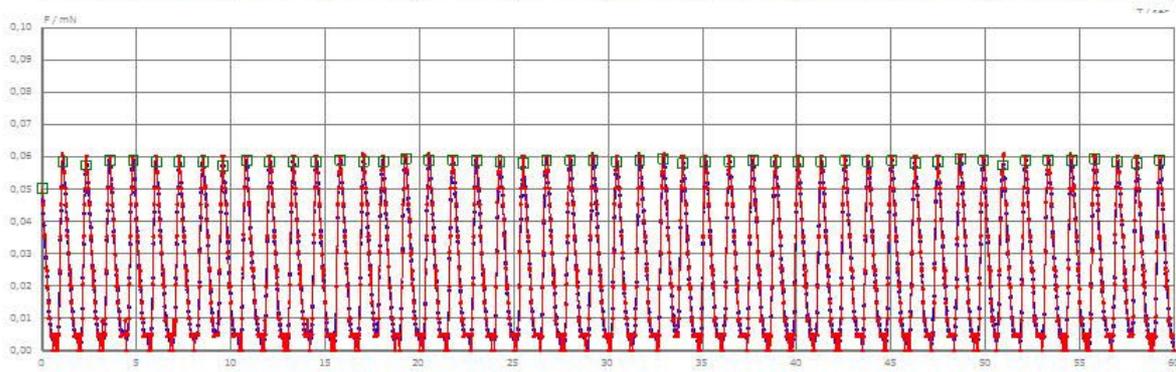
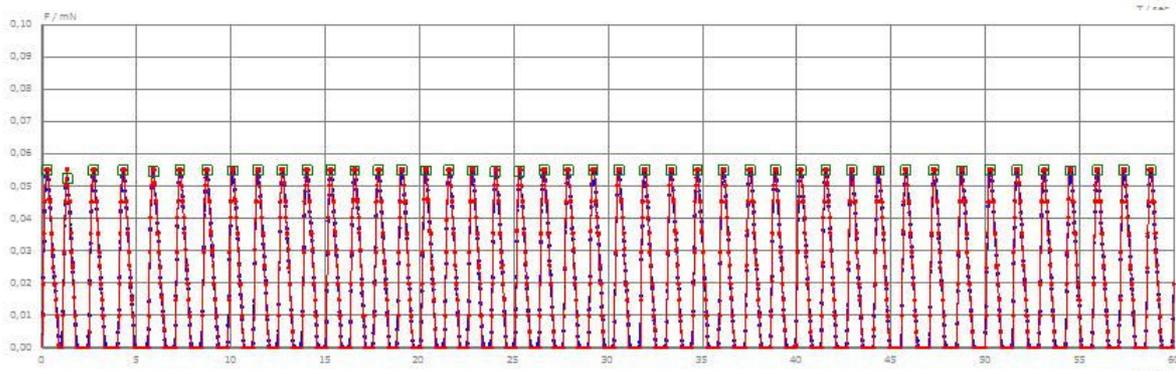
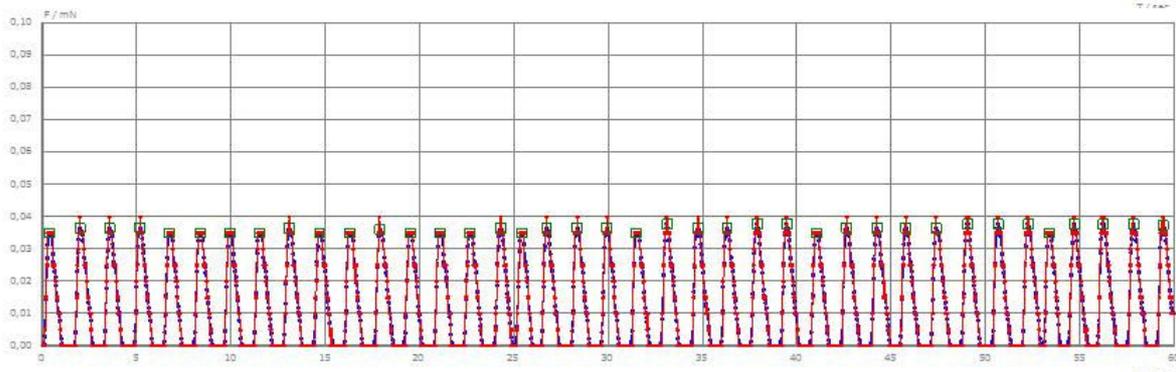
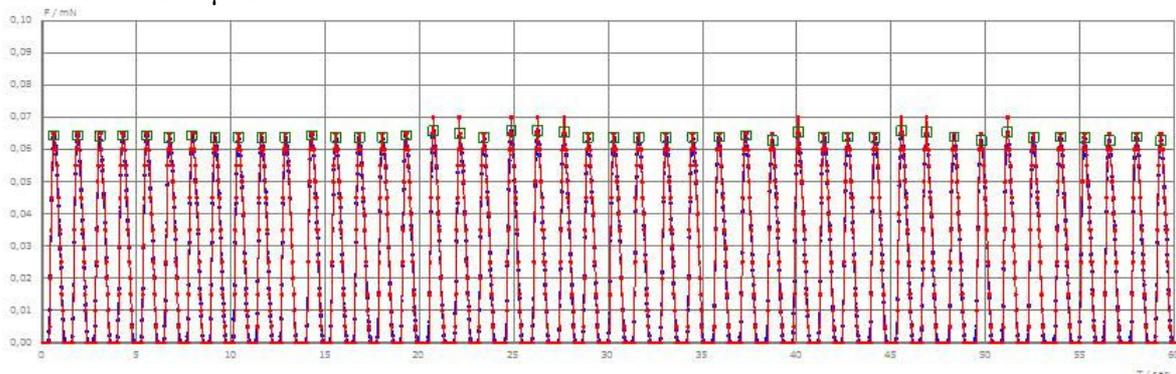
Procainamide 10 μM



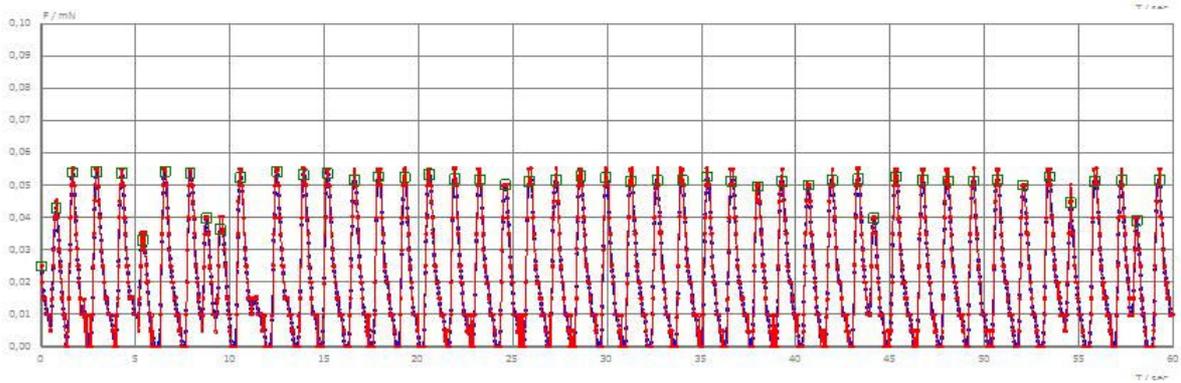
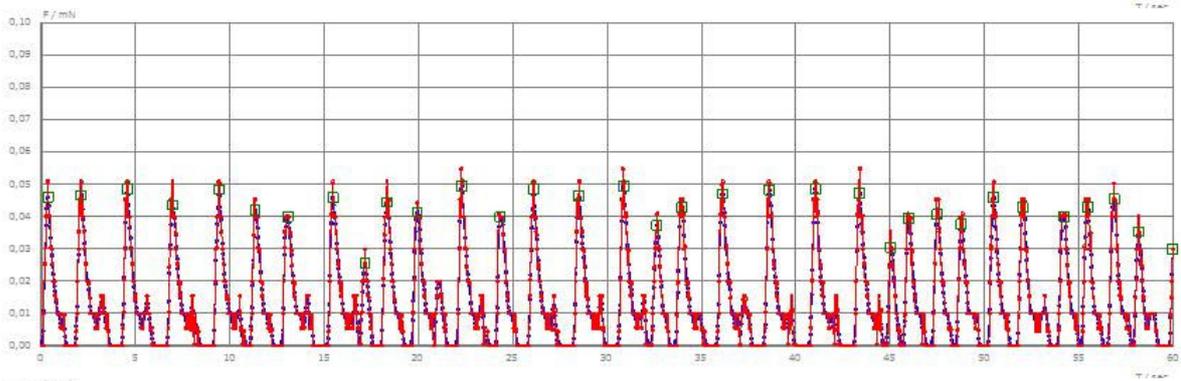
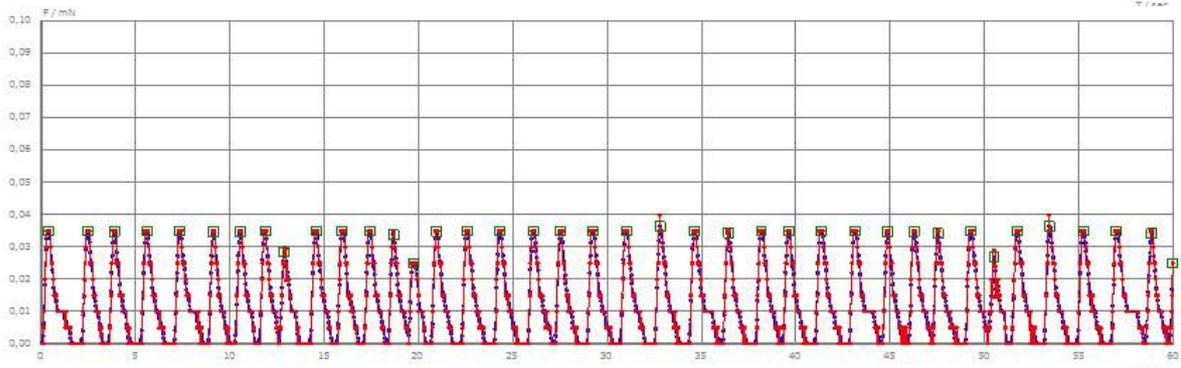
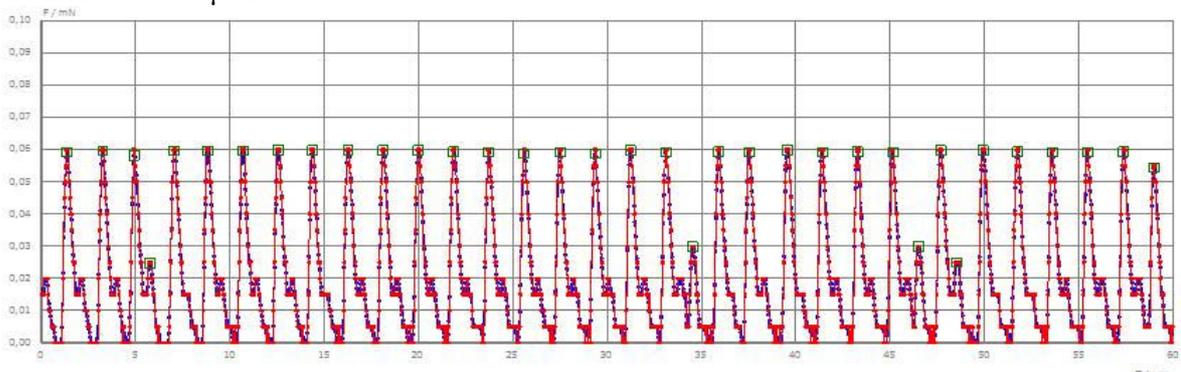
Procainamide 30 μM



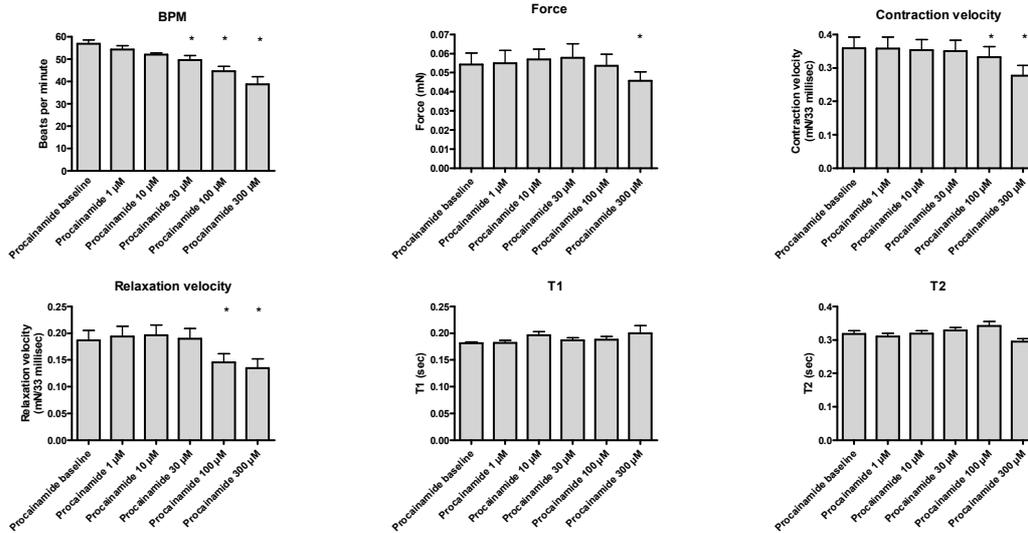
Procainamide 100 μM



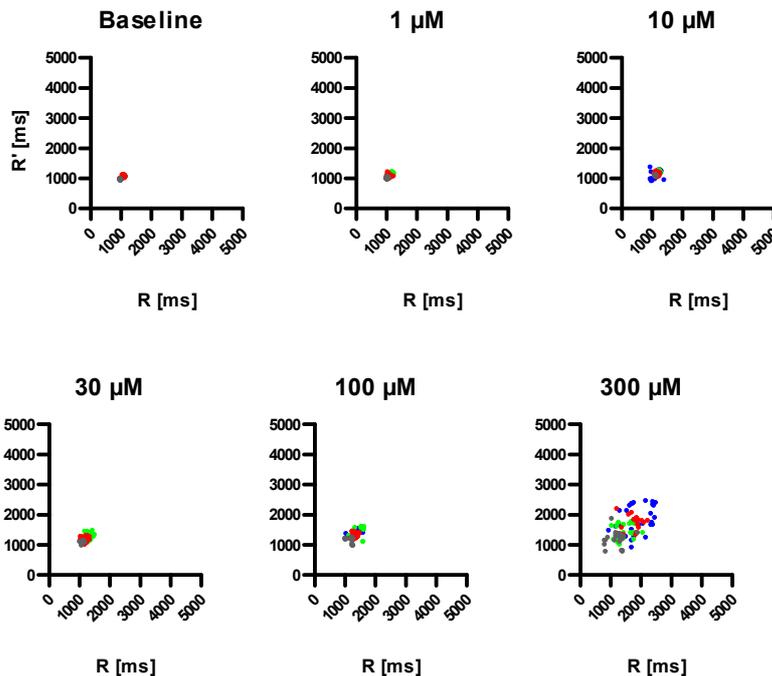
Procainamide 300 μM



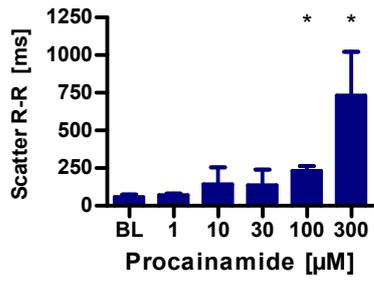
Analysis procainamide



Analysis of contraction. * $P < 0.05$ (Student's t-test), 4 biological replicas, bars show means \pm SD. Beats per minute (BPM), contraction time (T1), relaxation time (T2).



Graphical illustration of irregularity integrating 4 biological replicas. Ordinates indicate the distance from a given twitch to the following, the abscissa the distance to the previous twitch. Biological replicas are discriminated by color code.



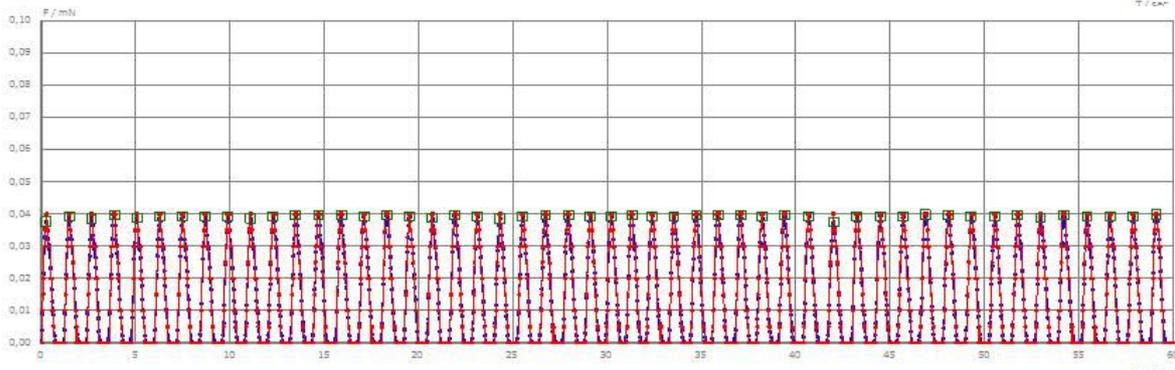
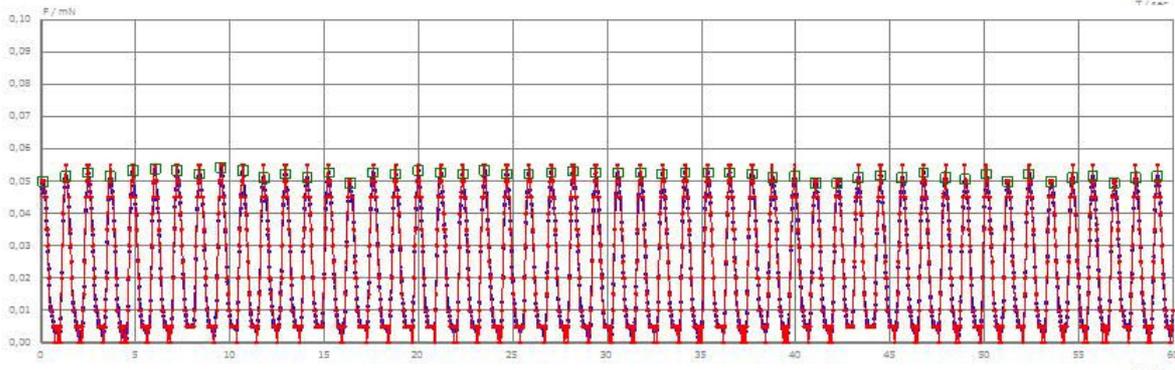
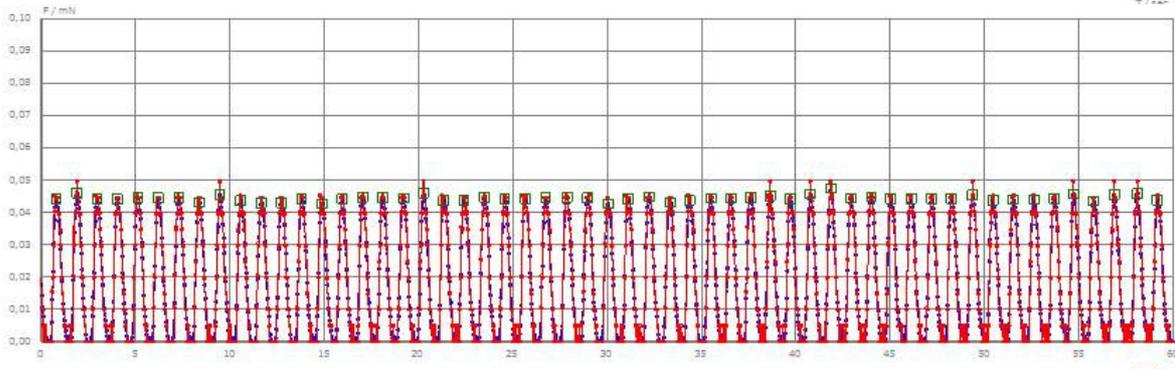
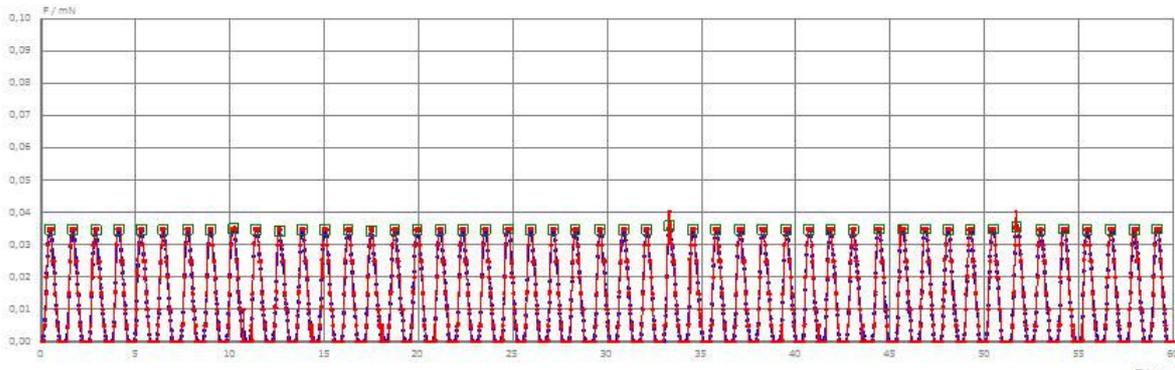
Scatter of beat-to-beat variability in the presence of procainamide, * $P < 0.05$ (Mann-Whitney U test), 4 biological replicas, bars show median \pm interquartile range.

Quinidine - original recordings

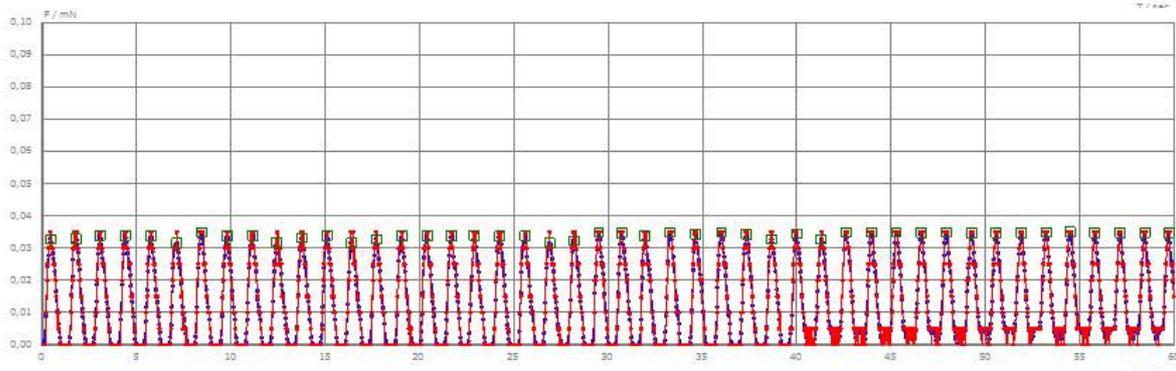
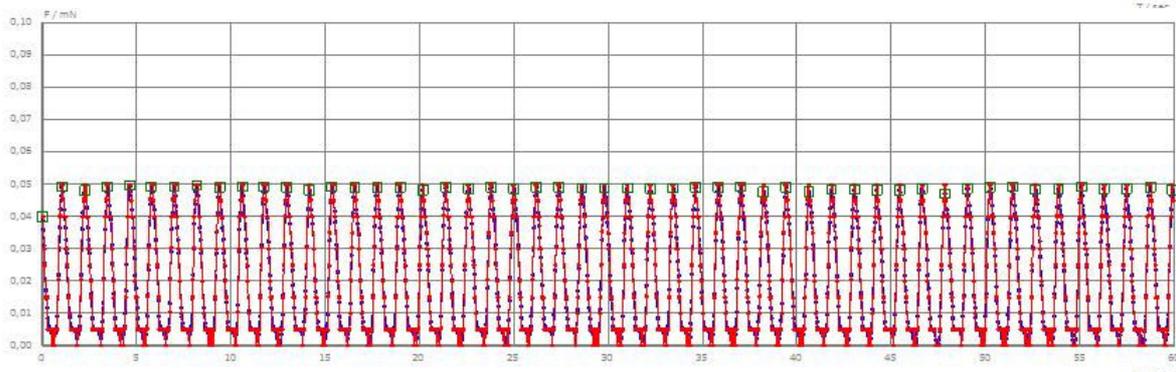
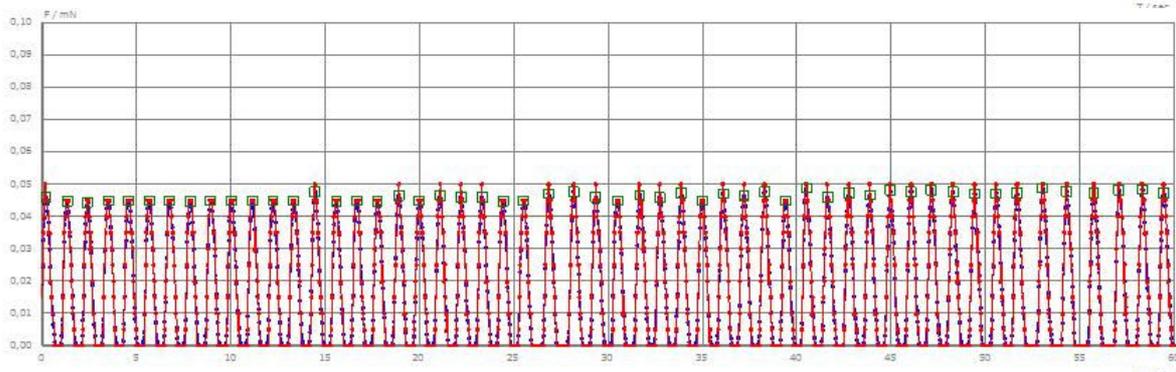
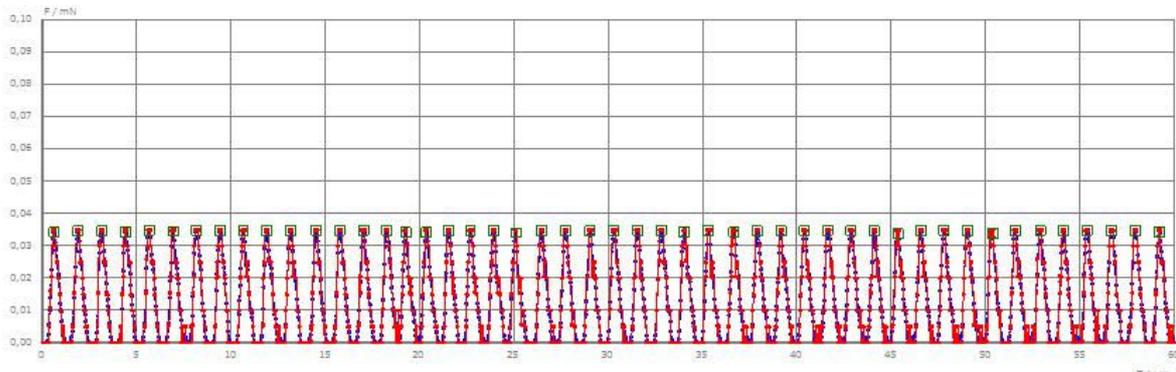
Quinidine baseline



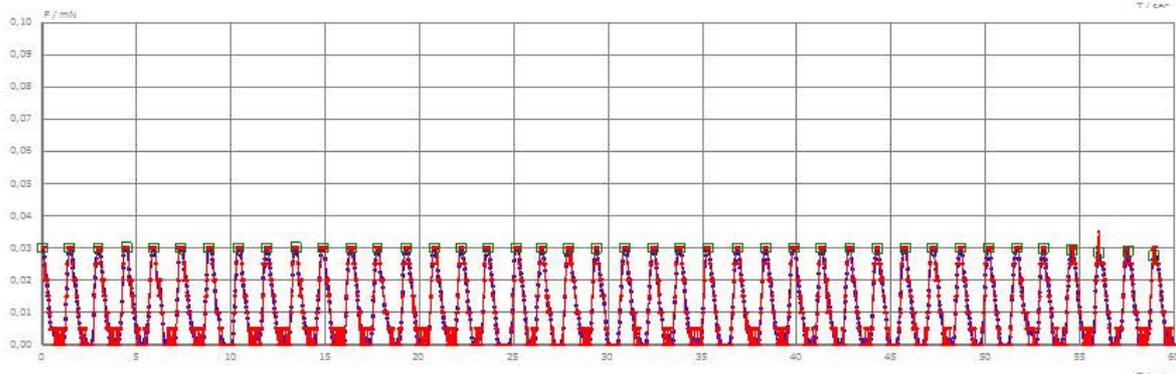
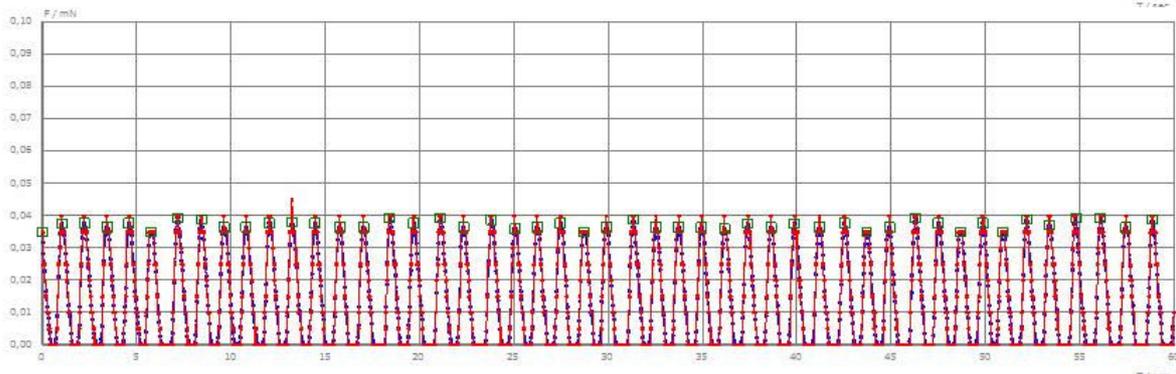
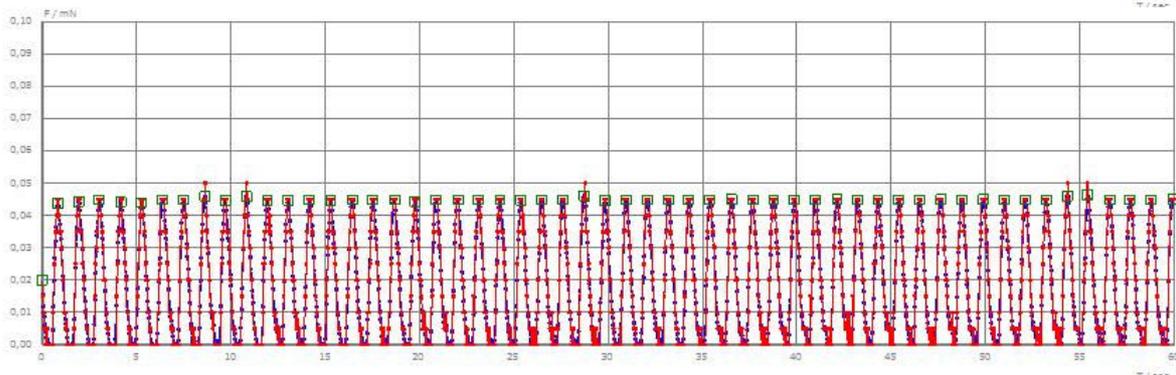
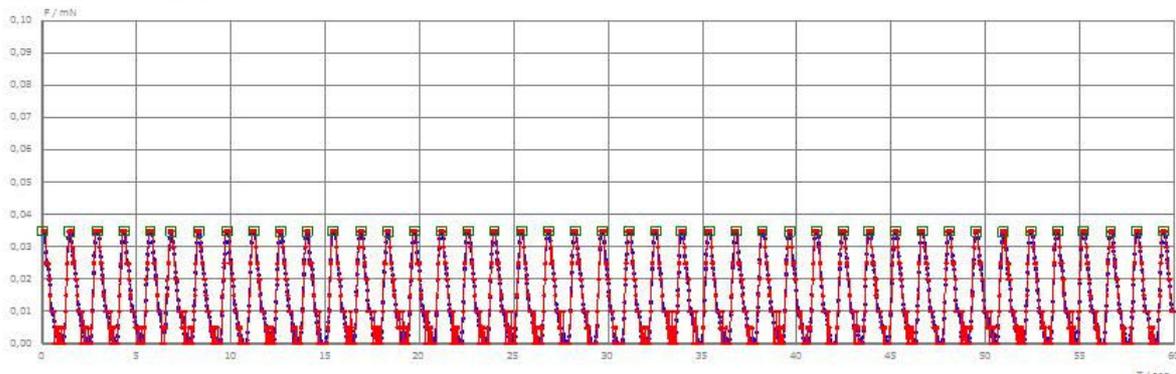
Quinidine 10 nM



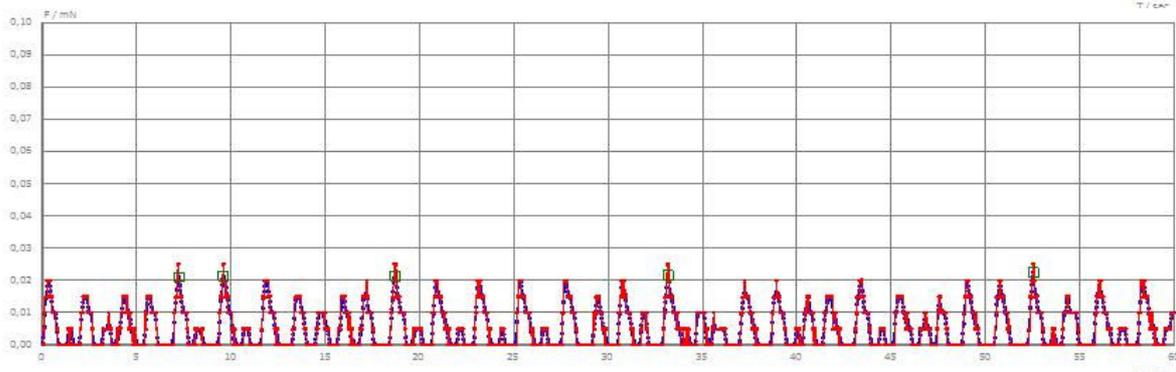
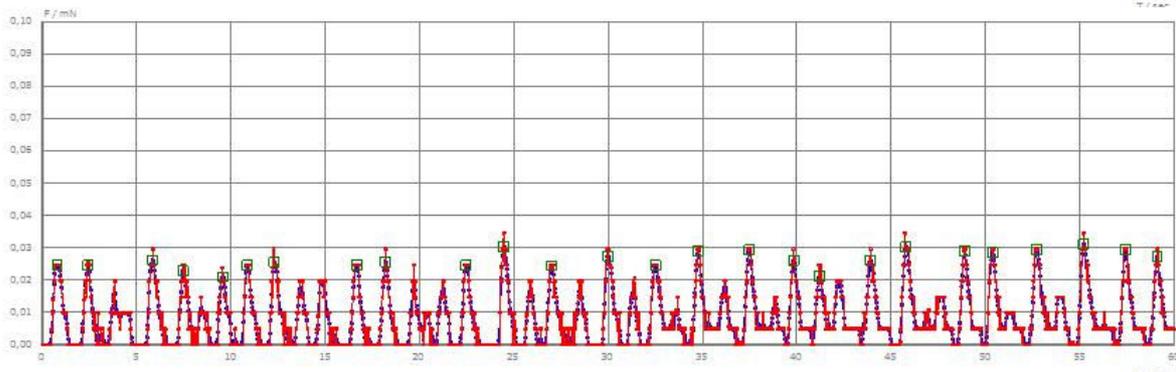
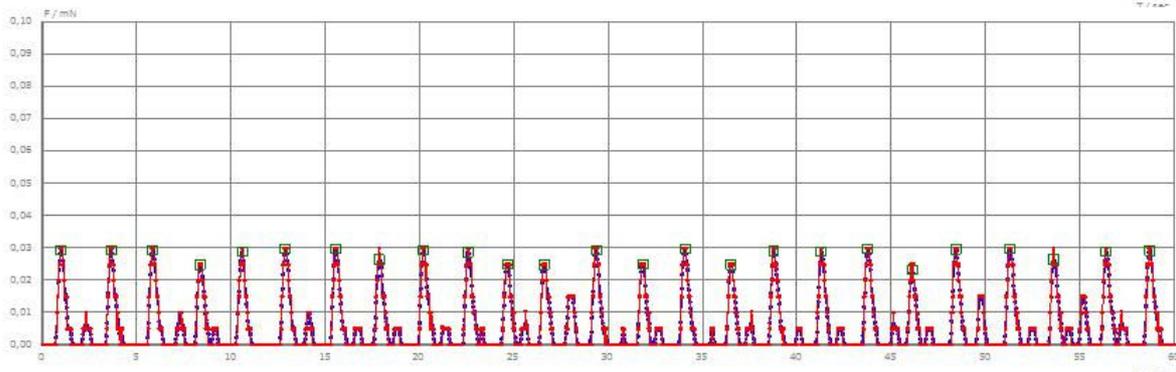
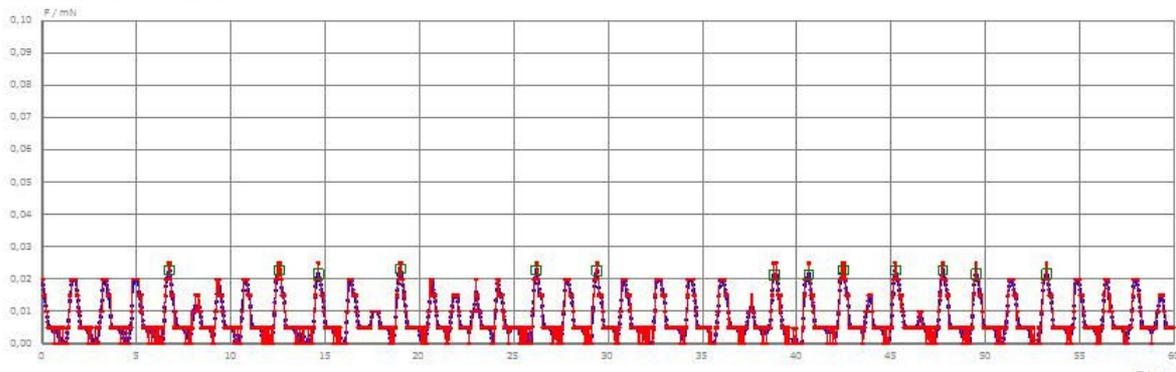
Quinidine 100 nM



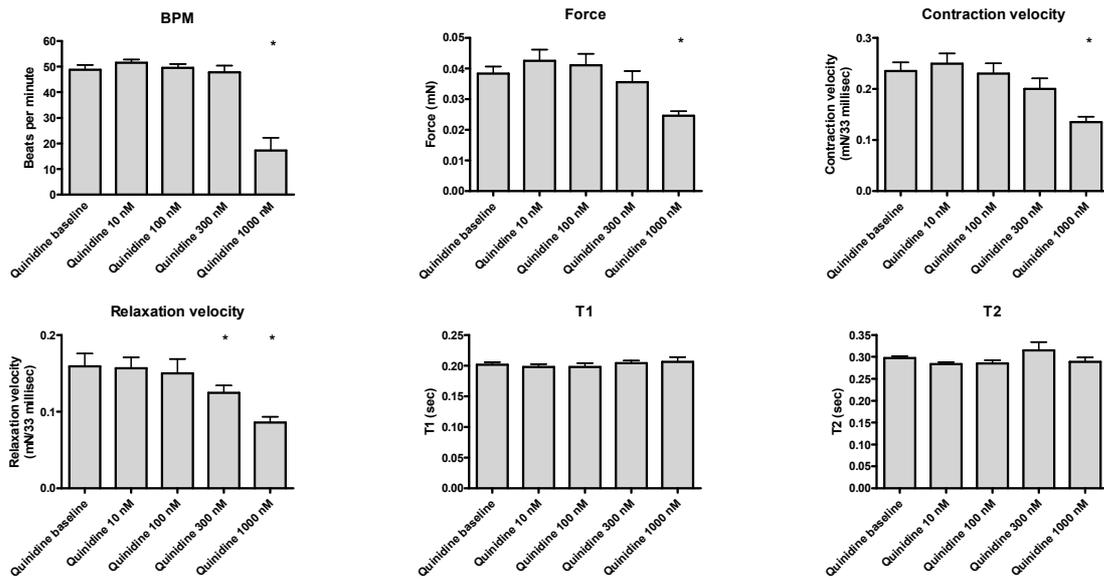
Quinidine 300 nM



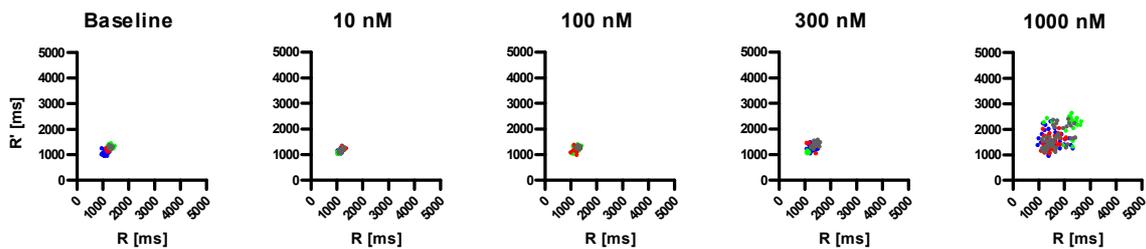
Quinidine 1000 nM



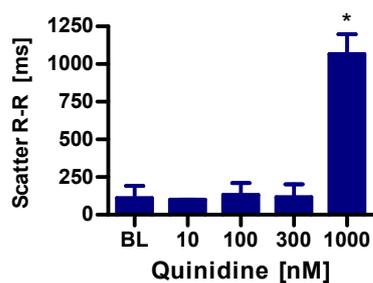
Analysis quinidine



Analysis of contraction. * $P < 0.05$ (Student's t-test), 4 biological replicas, bars show means \pm SD. Beats per minute (BPM), contraction time (T1), relaxation time (T2).



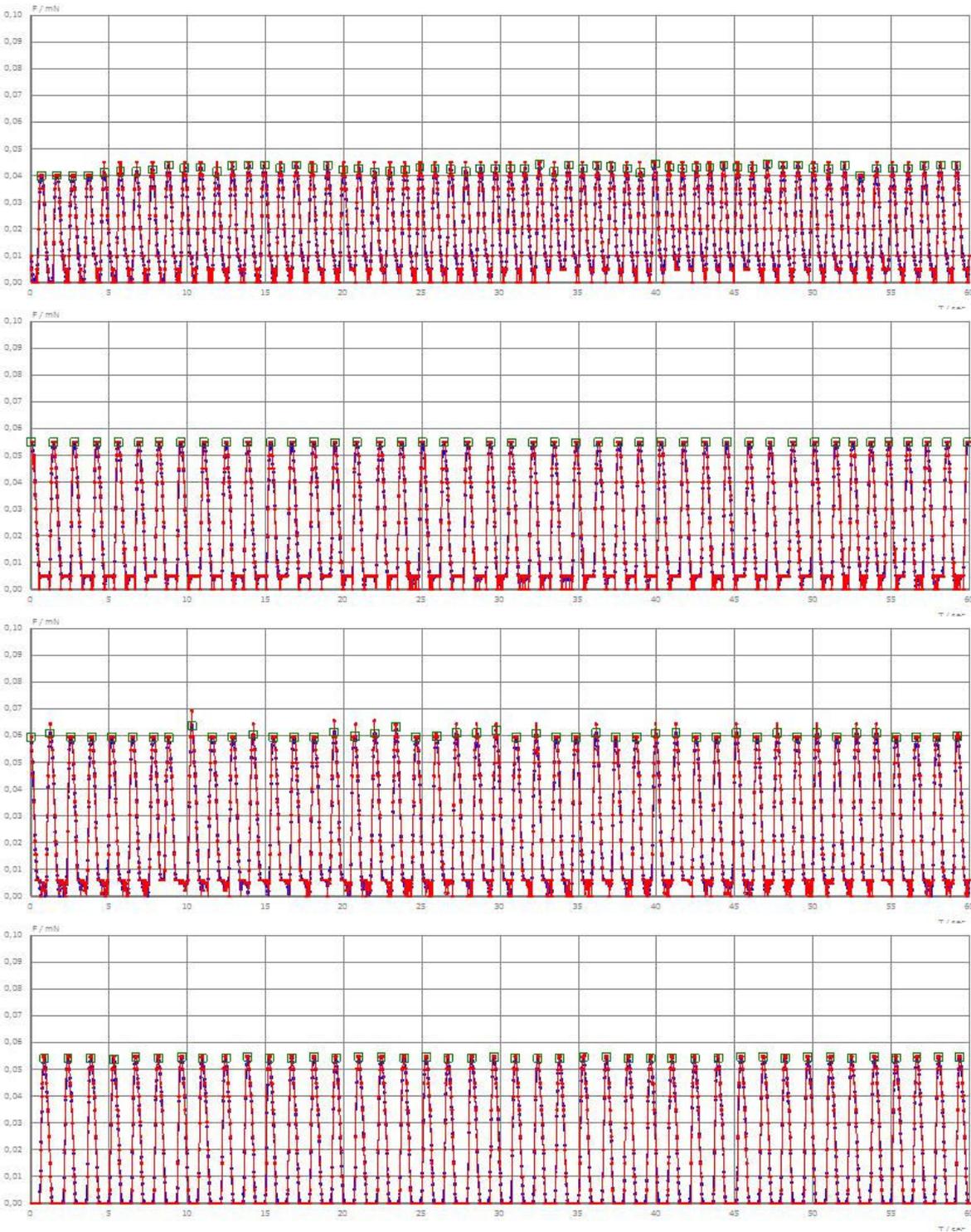
Graphical illustration of irregularity integrating 4 biological replicas. Ordinates indicate the distance from a given twitch to the following, the abscissa the distance to the previous twitch. Biological replicas are discriminated by color code.



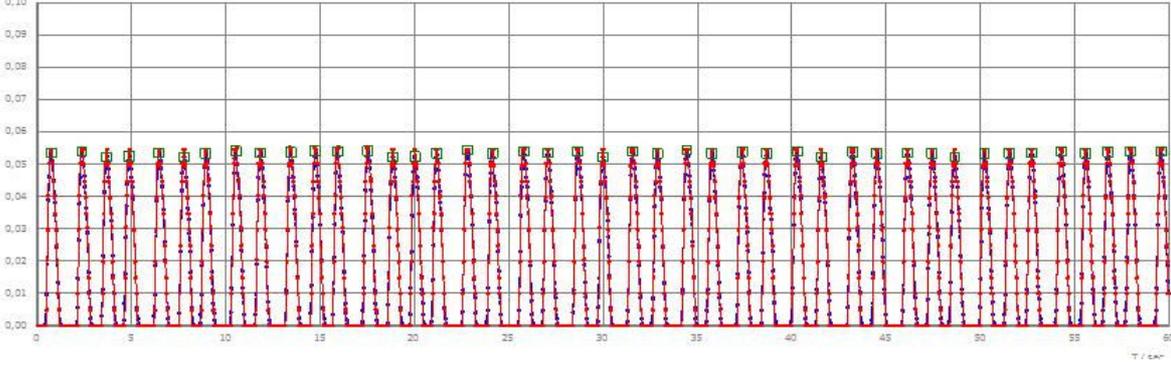
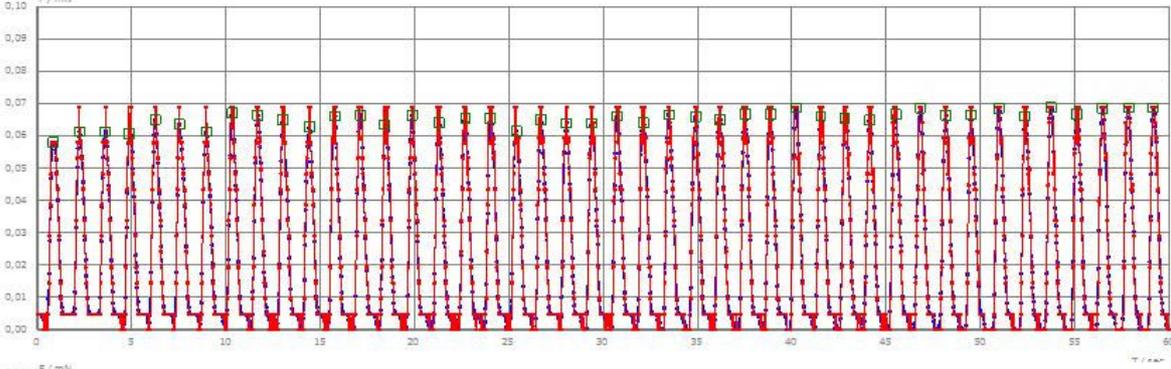
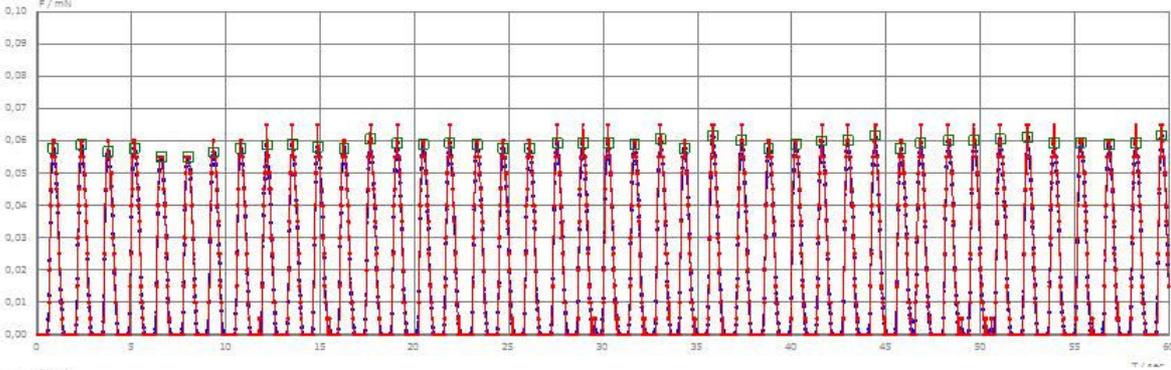
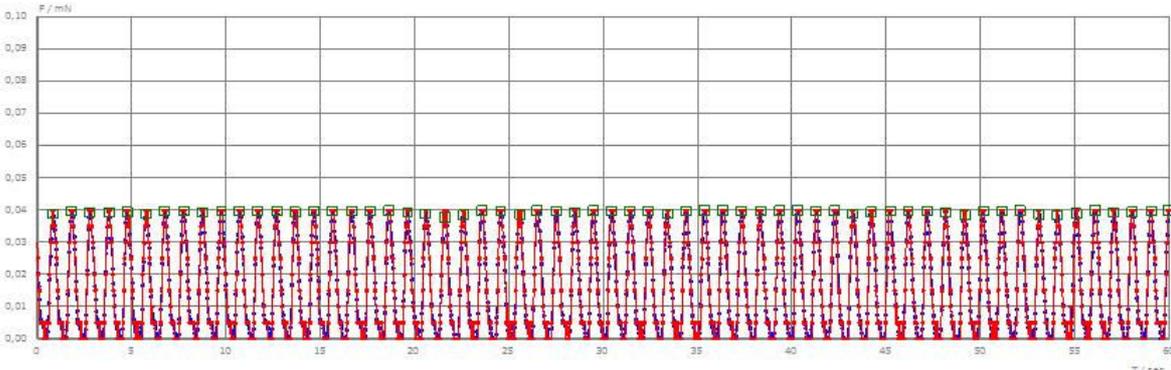
Scatter of beat-to-beat variability in the presence of quinidine, * $P < 0.05$ (Mann-Whitney U test), 4 biological replicas, bars show median \pm interquartile range.

Sertindole - original recordings

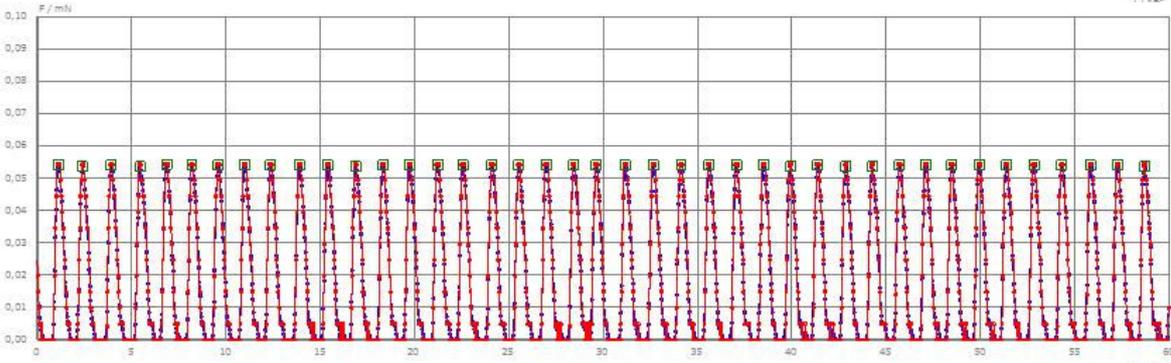
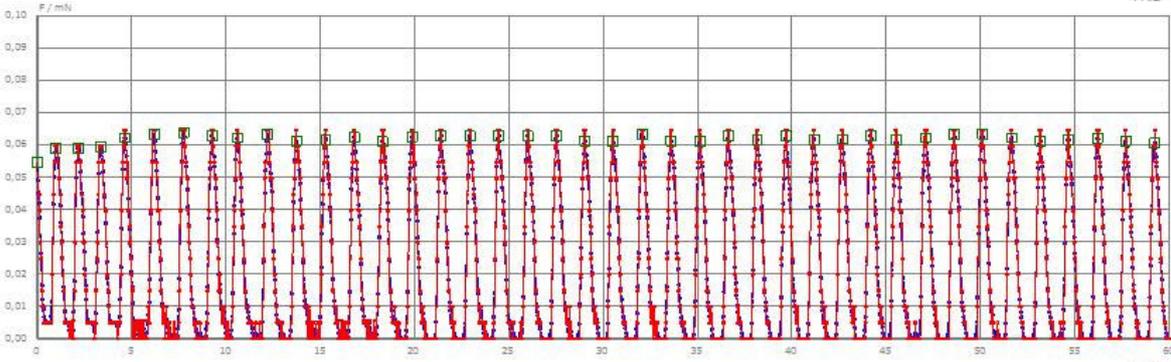
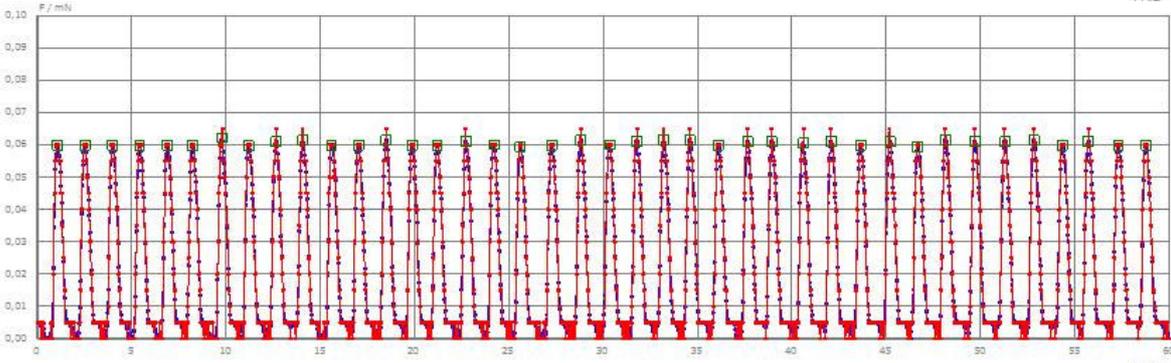
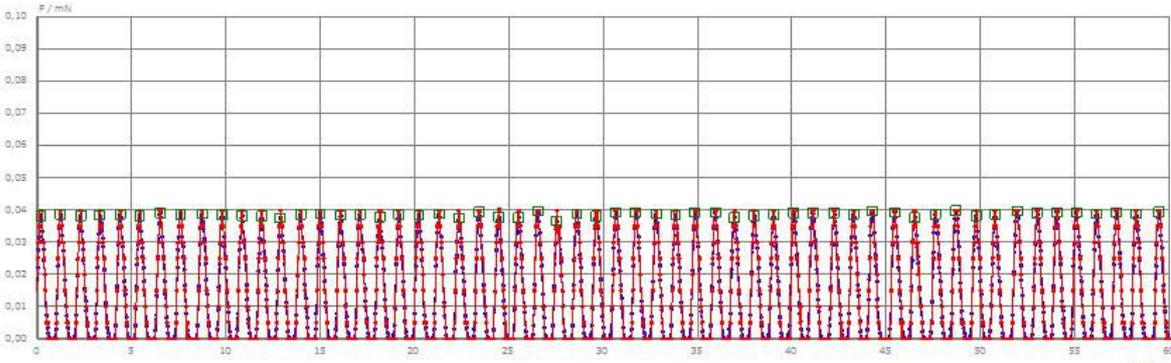
Sertindole baseline



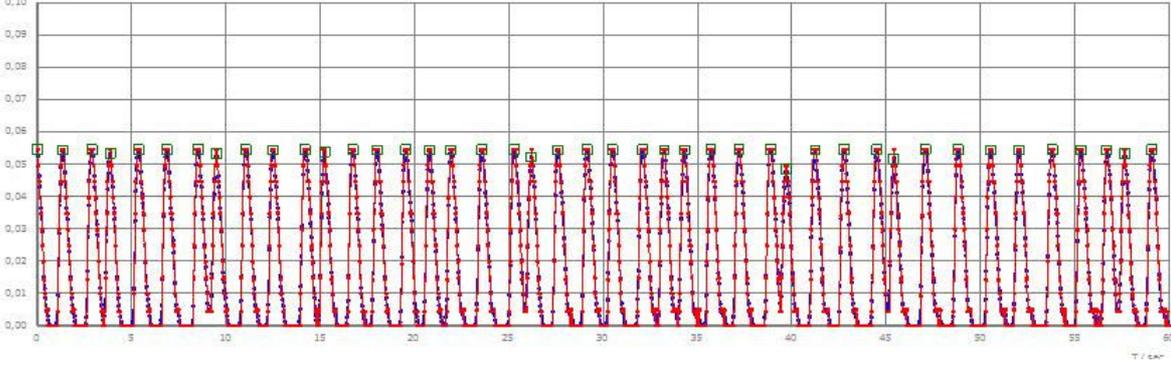
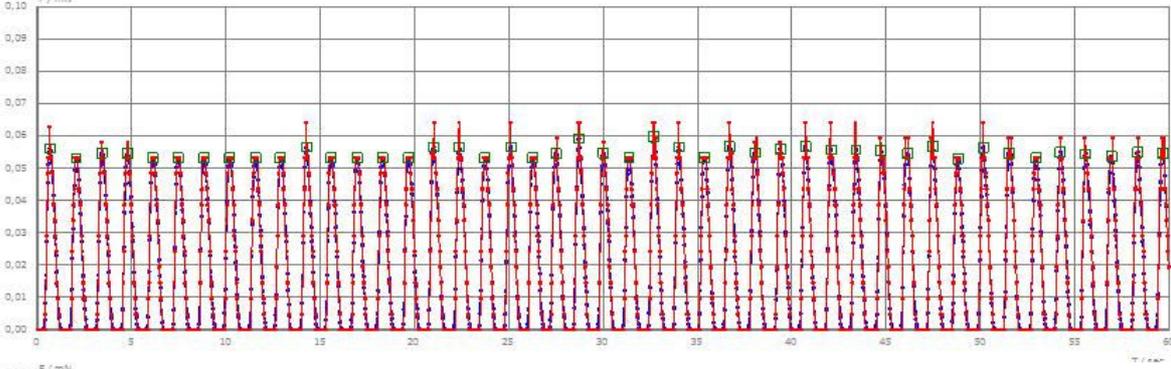
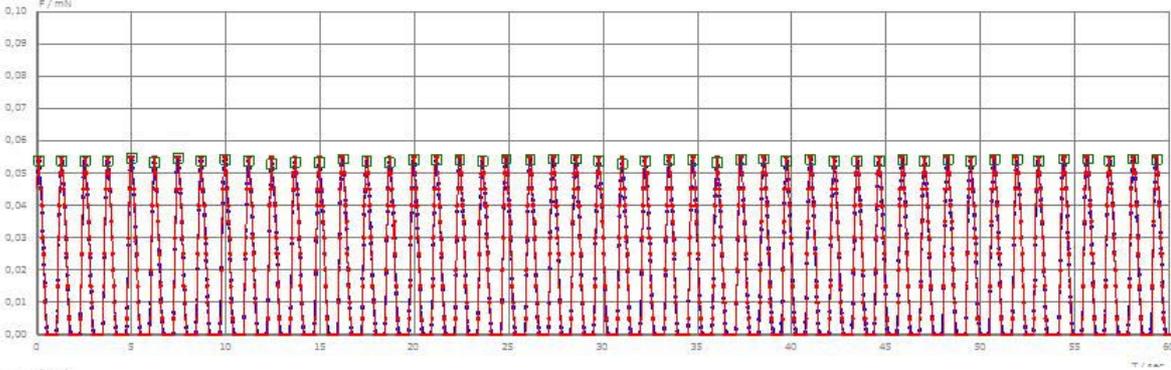
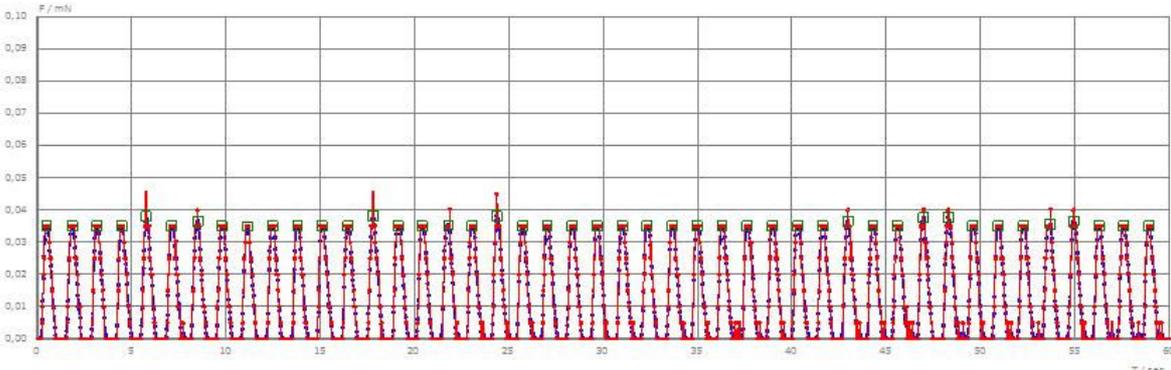
Sertindole 1 nM



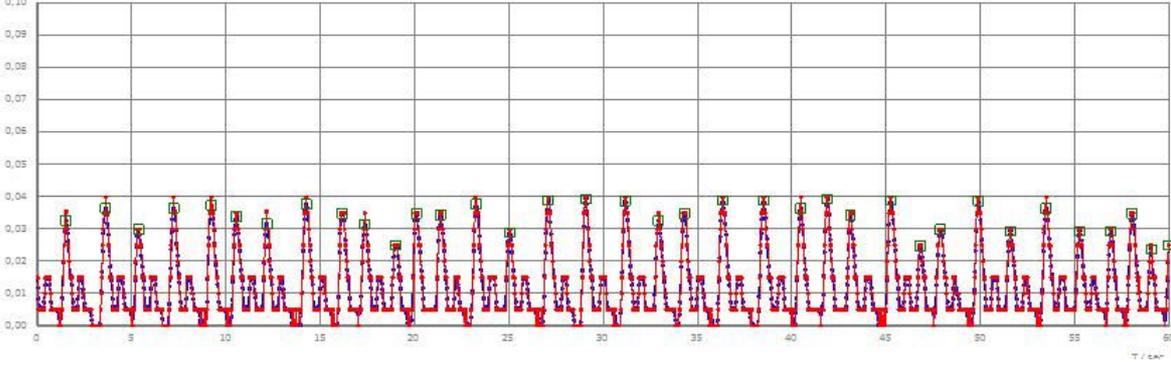
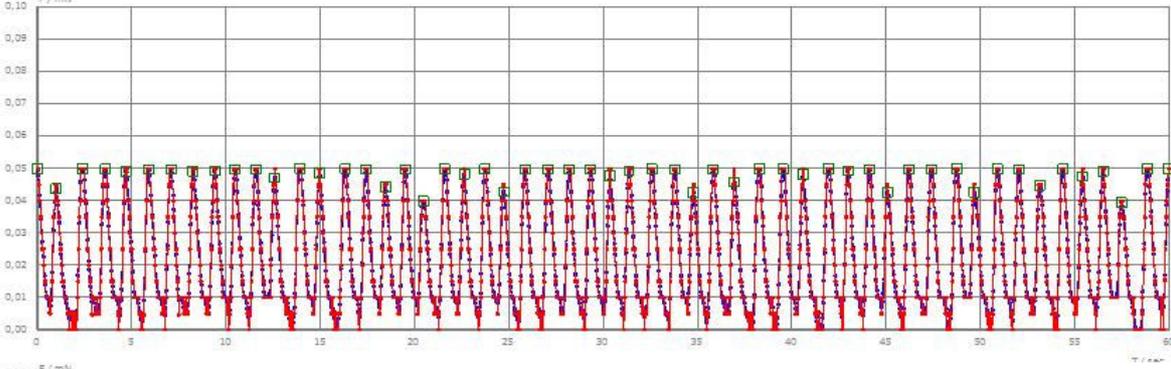
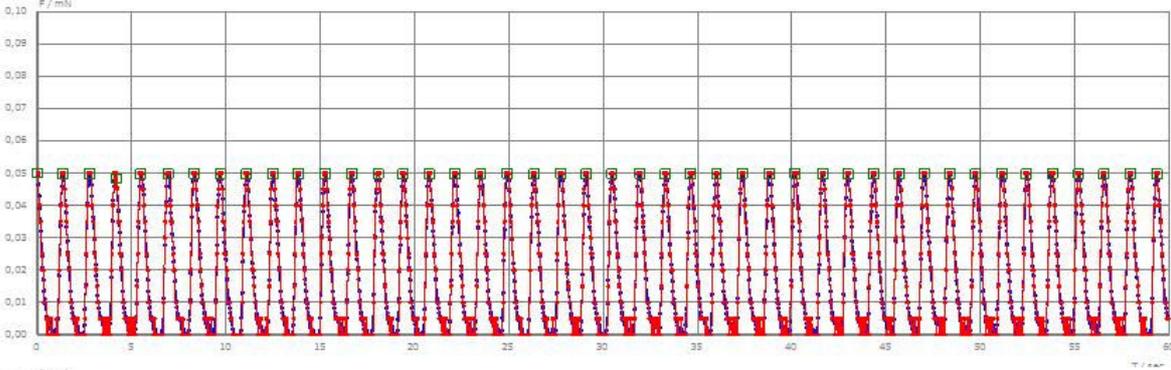
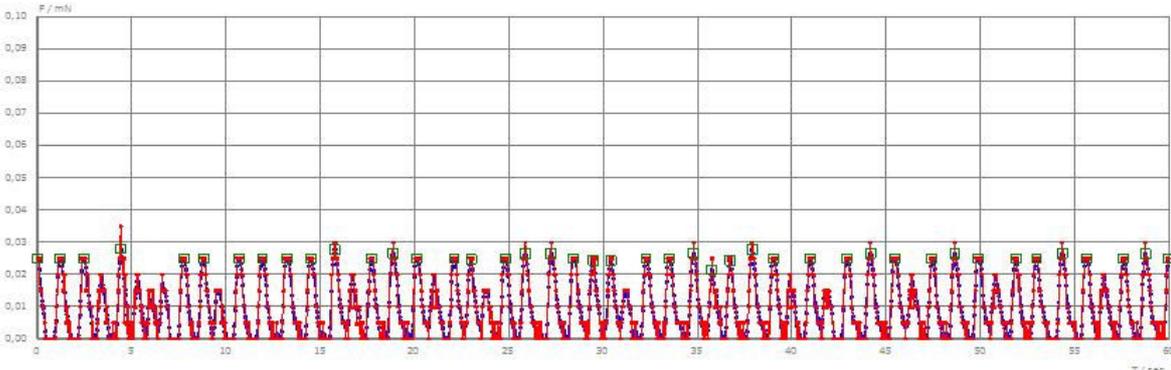
Sertindole 10 nM



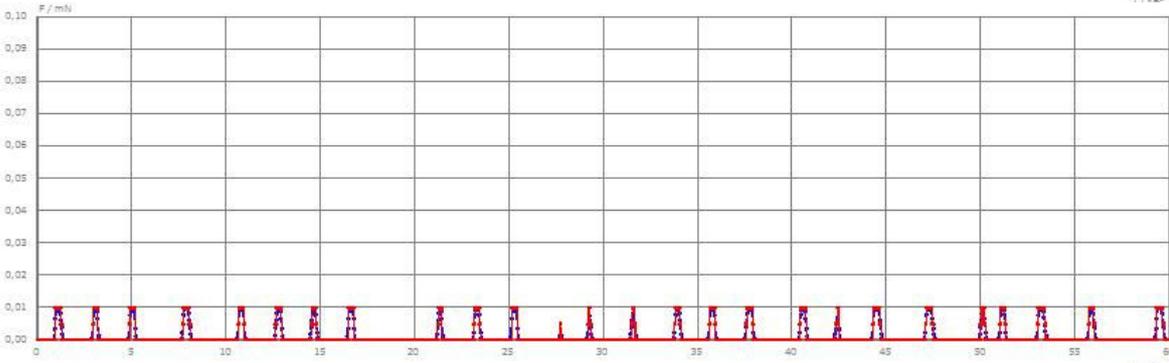
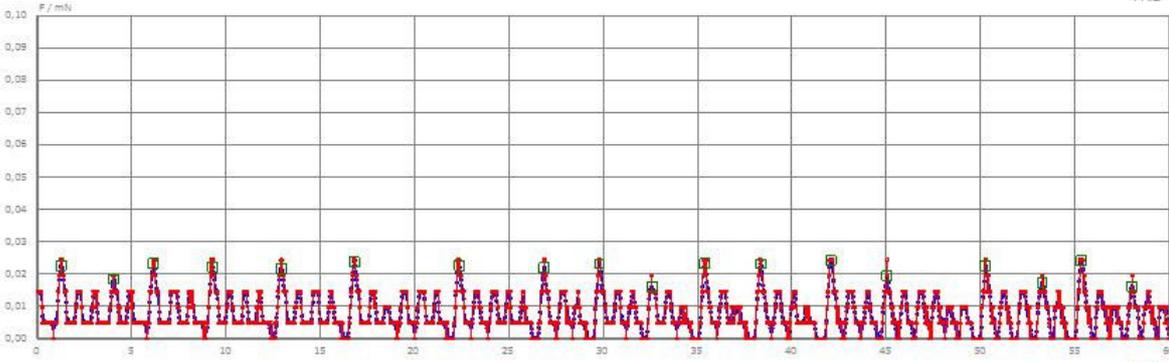
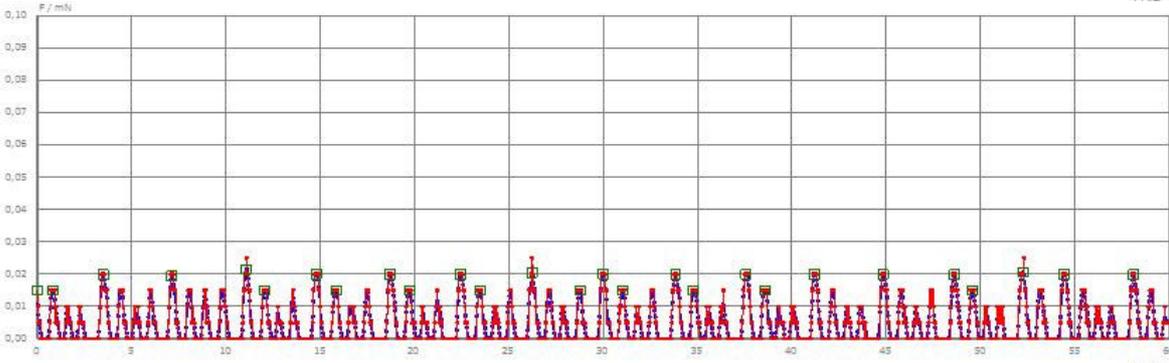
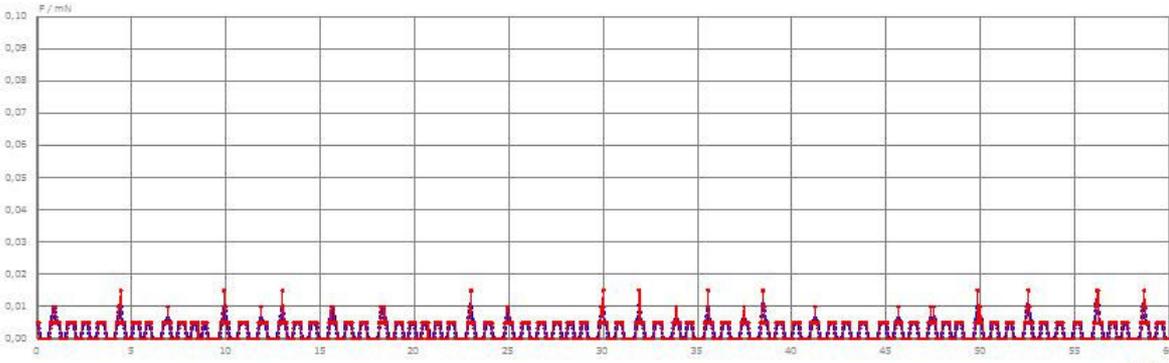
Sertindole 30 nM



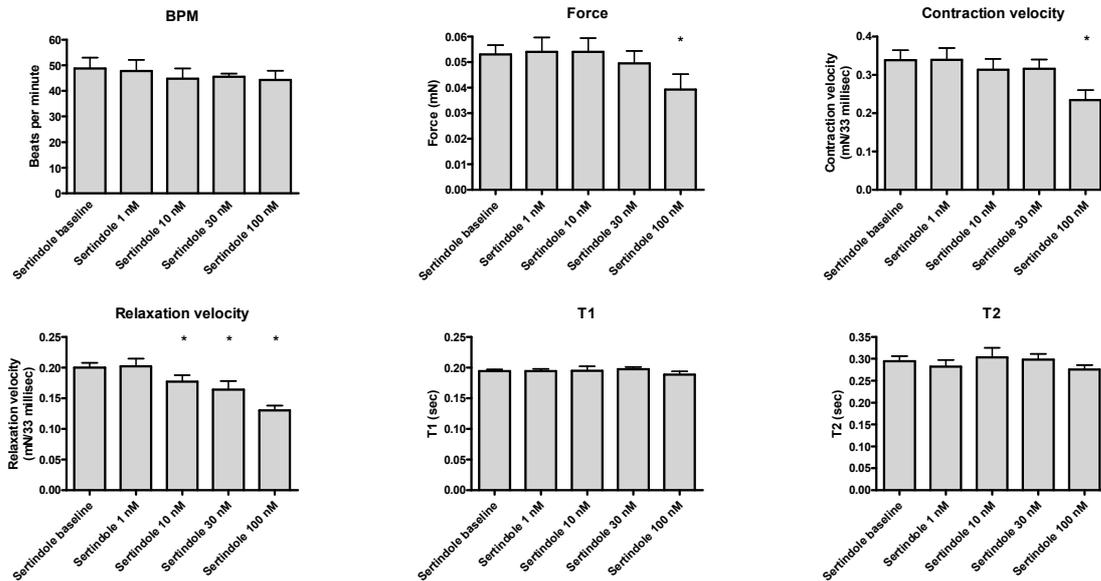
Sertindole 100 nM



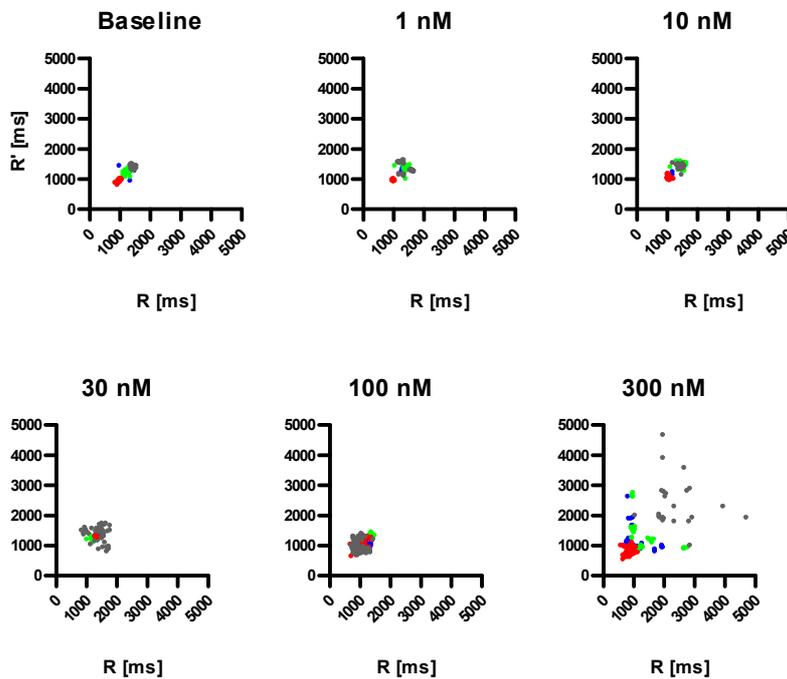
Sertindole 300 nM



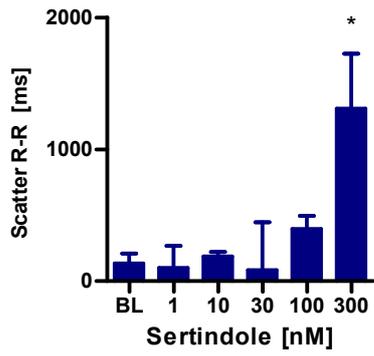
Analysis sertindole



Analysis of contraction. * $P < 0.05$ (Student's t-test), 4 biological replicas, bars show means \pm SD. Beats per minute (BPM), contraction time (T1), relaxation time (T2).



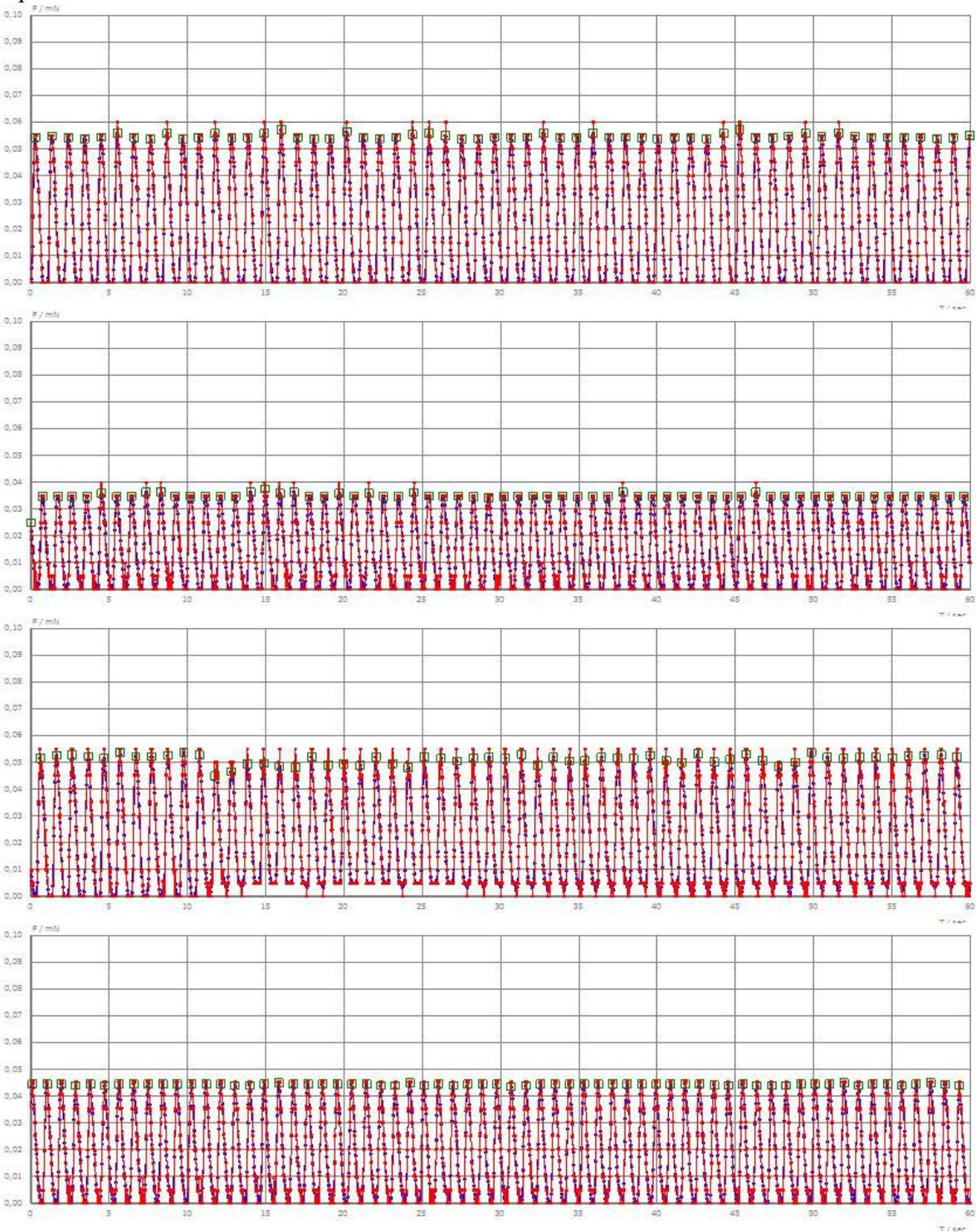
Graphical illustration of irregularity integrating 4 biological replicas. Ordinates indicate the distance from a given twitch to the following, the abscissa the distance to the previous twitch. Biological replicas are discriminated by color code.



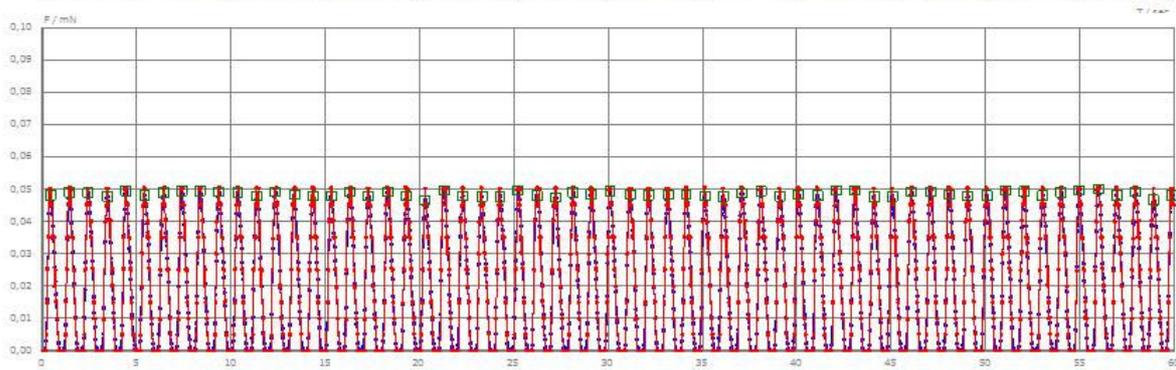
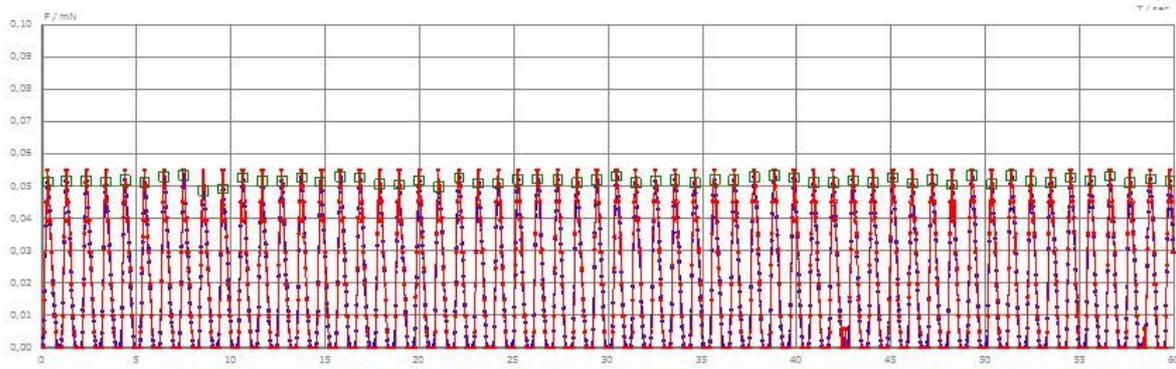
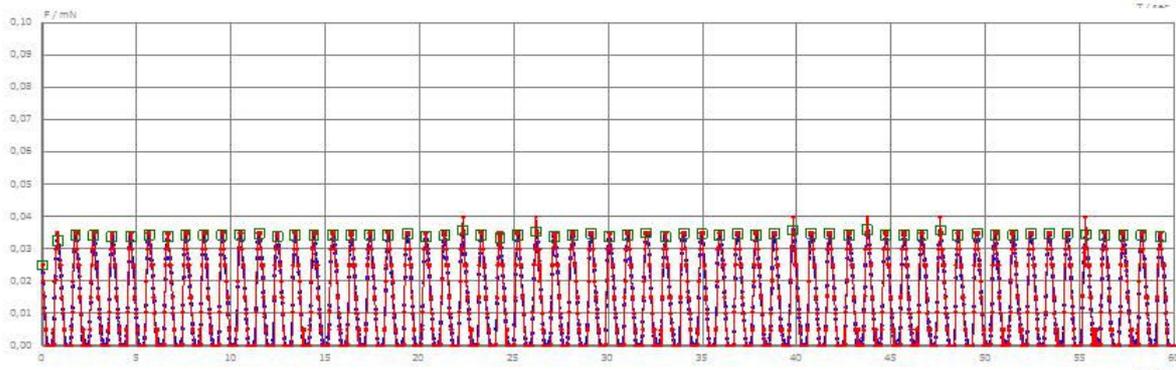
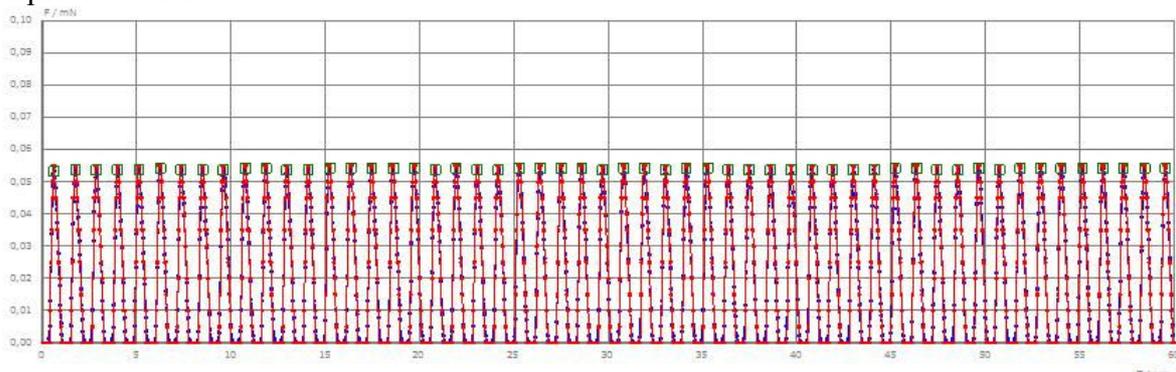
Scatter of beat-to-beat variability in the presence of sertindole, * $P < 0.05$ (Mann-Whitney U test), 4 biological replicas, bars show median \pm interquartile range.

Cisapride - original recordings

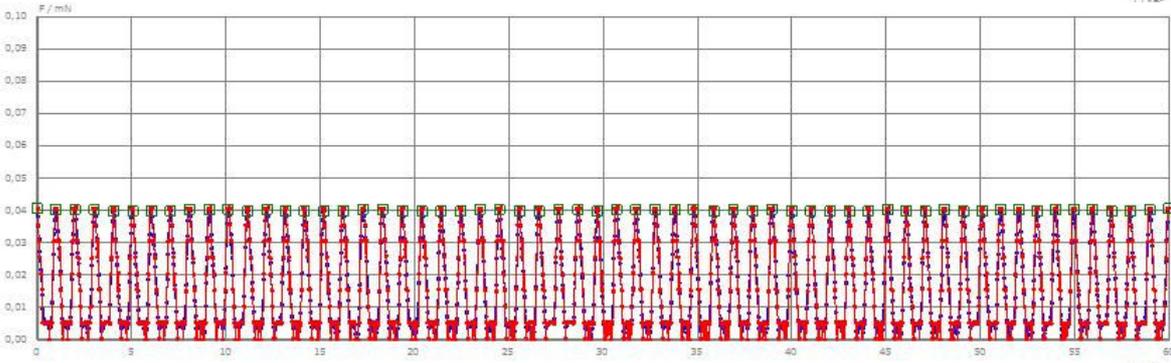
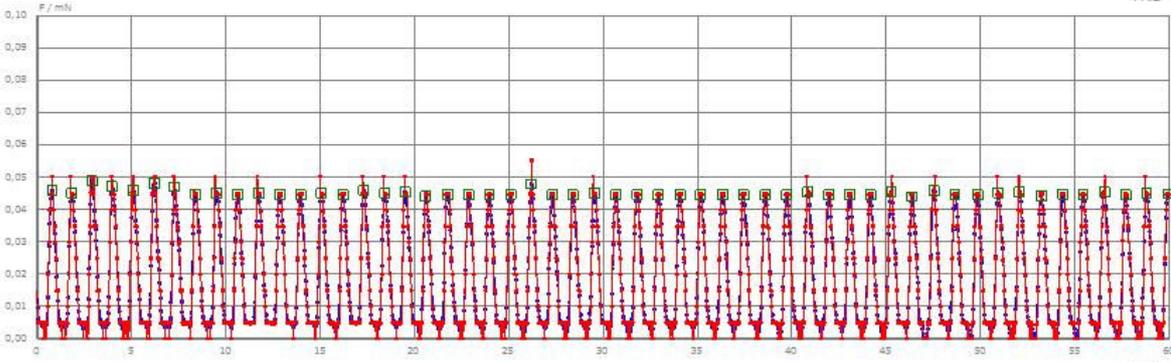
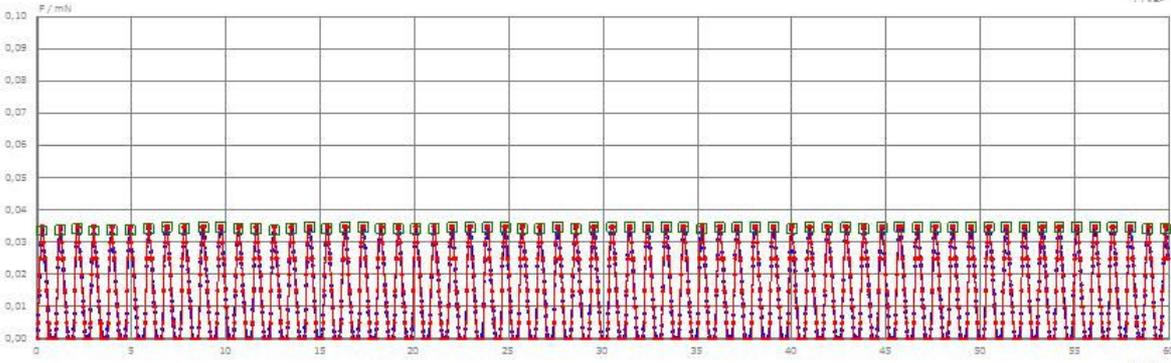
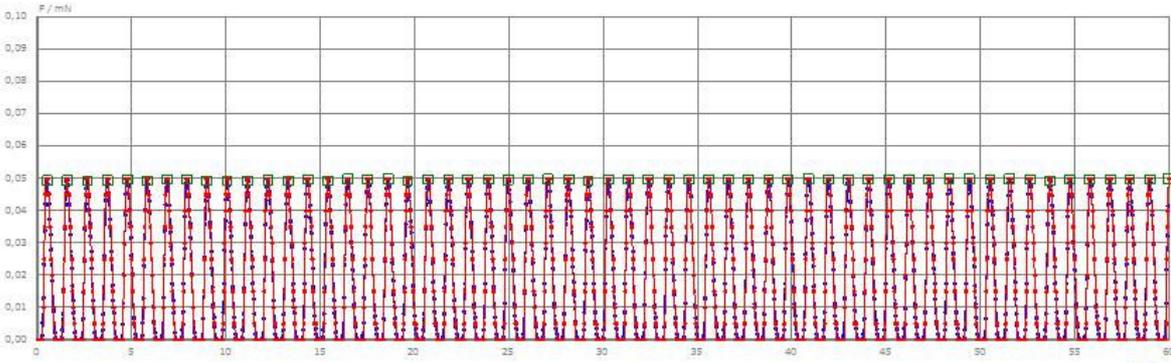
Cisapride baseline



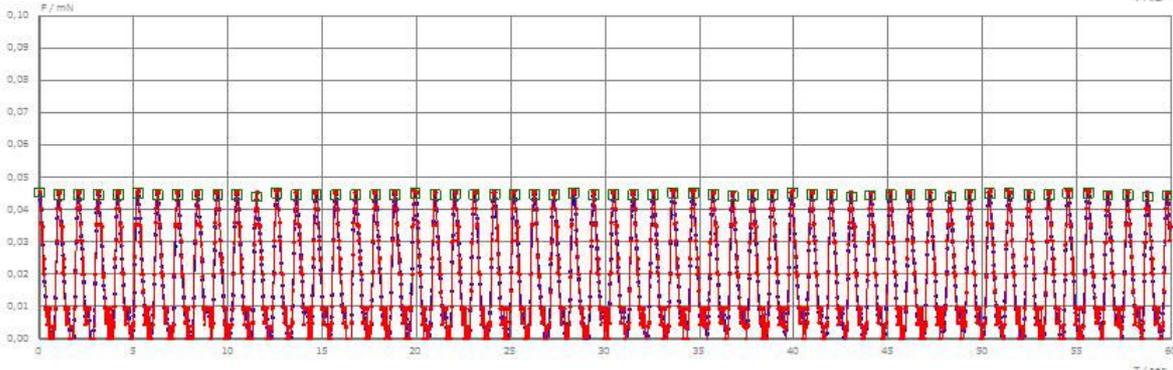
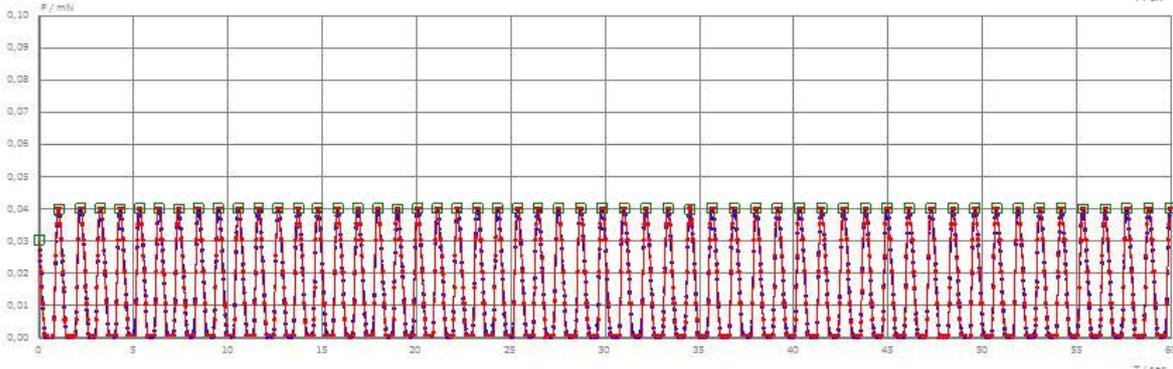
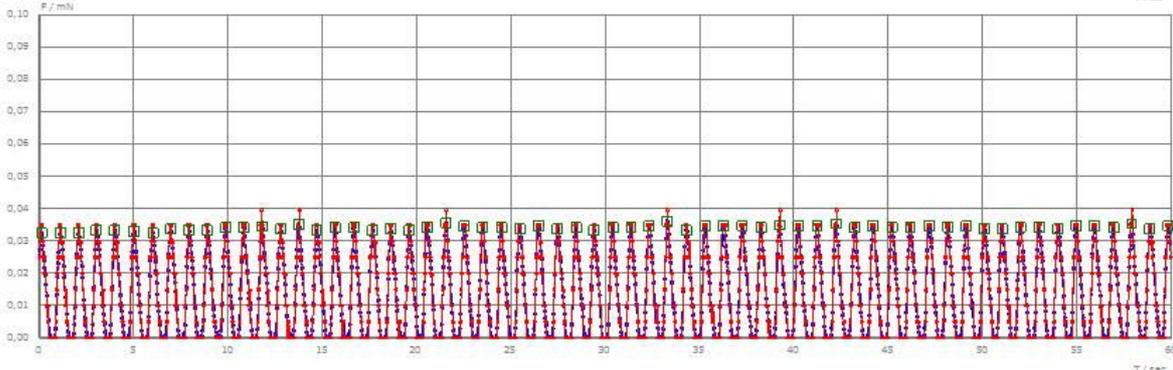
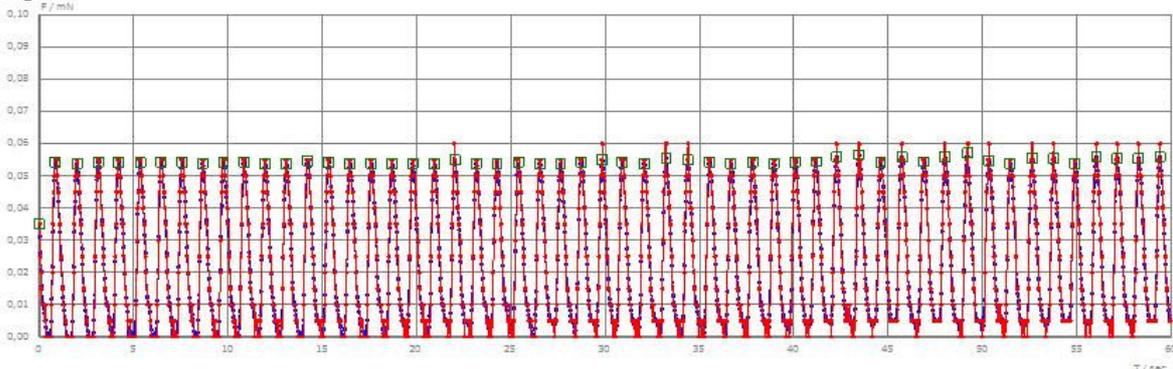
Cisapride 0.3 nM



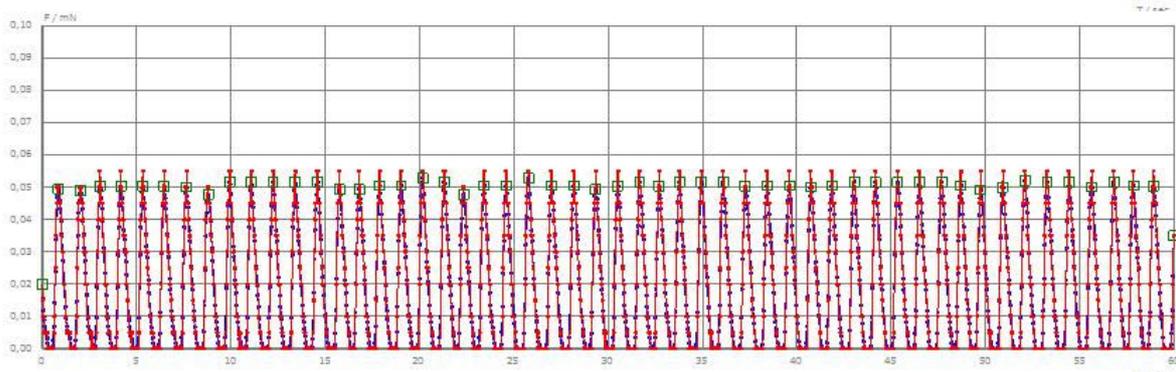
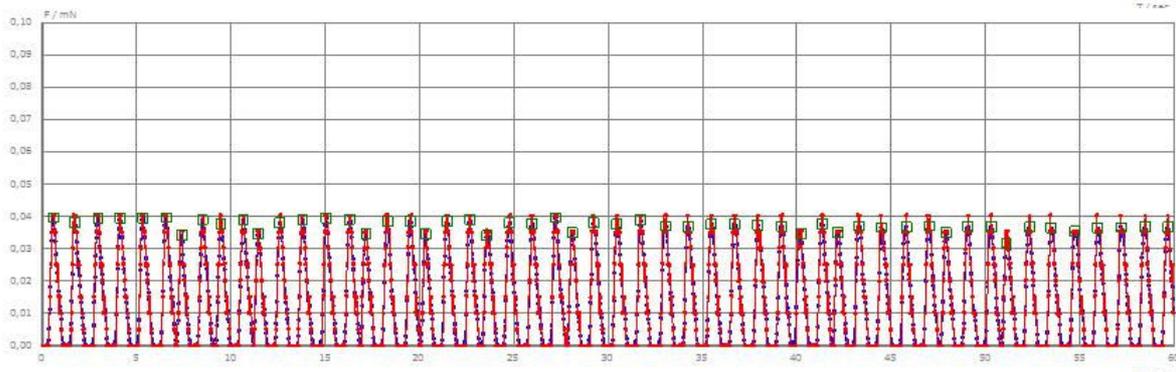
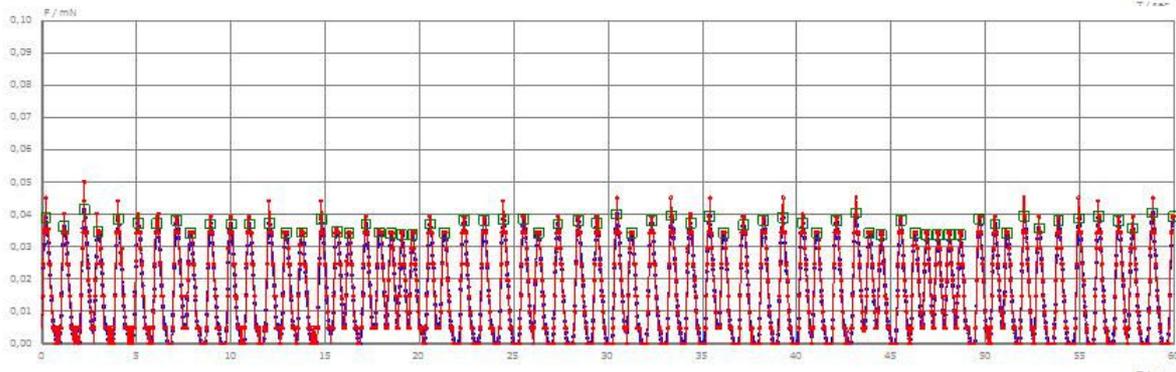
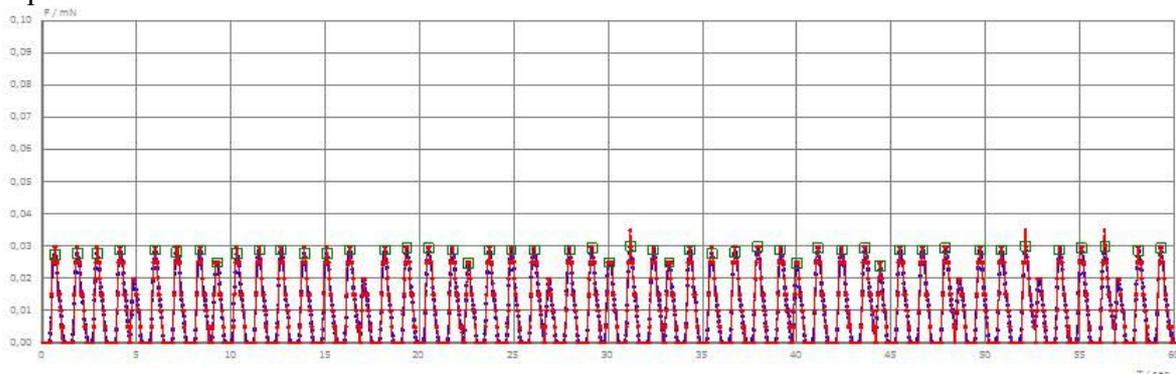
Cisapride 3 nM



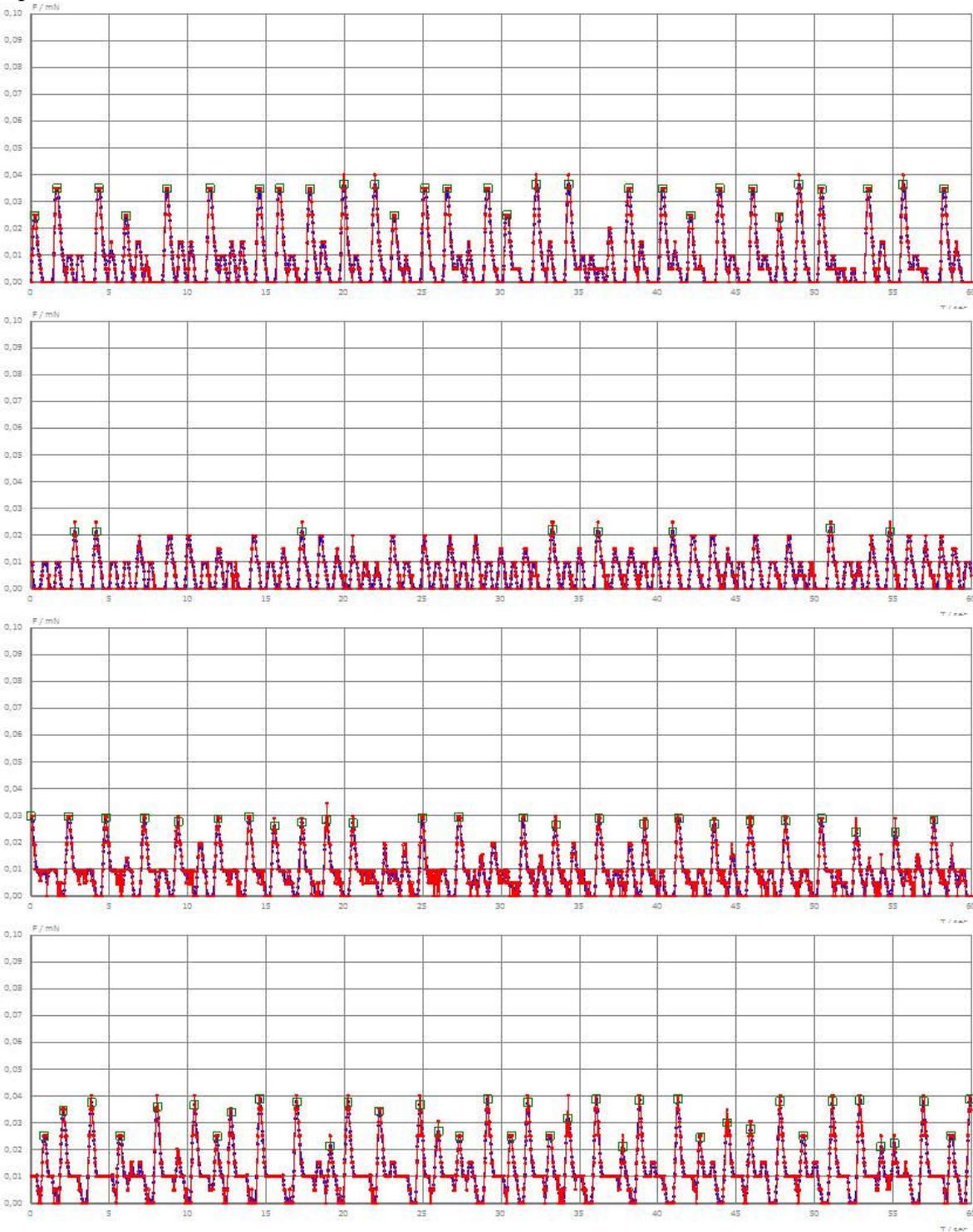
Cisapride 10 nM



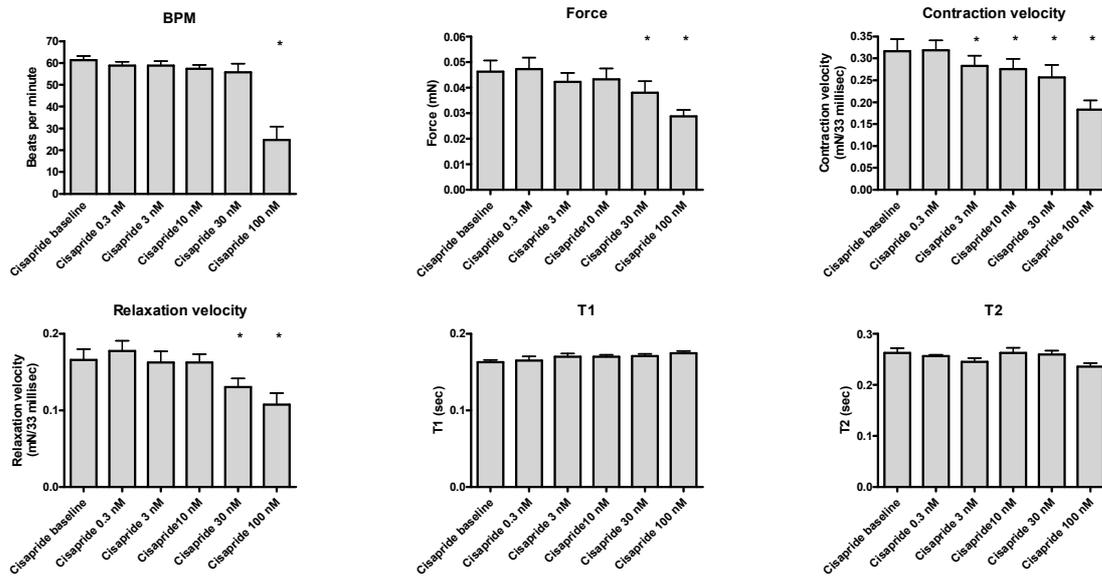
Cisapride 30 nM



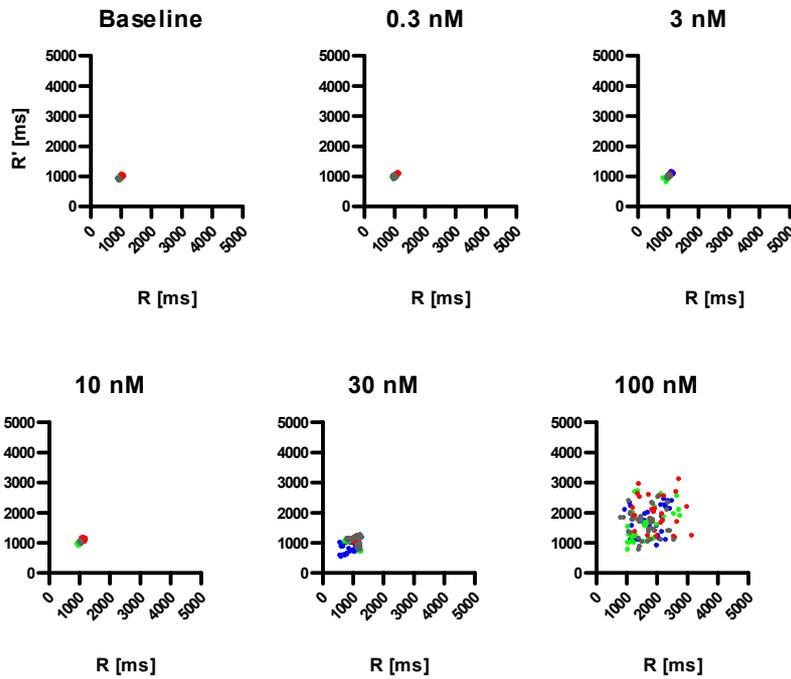
Cisapride 100 nM



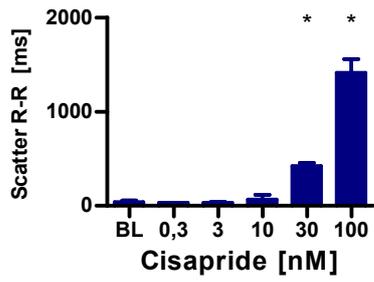
Analysis cisapride



Analysis of contraction. * $P < 0.05$ (Student's t-test), 4 biological replicas, bars show means \pm SD. Beats per minute (BPM), contraction time (T1), relaxation time (T2).



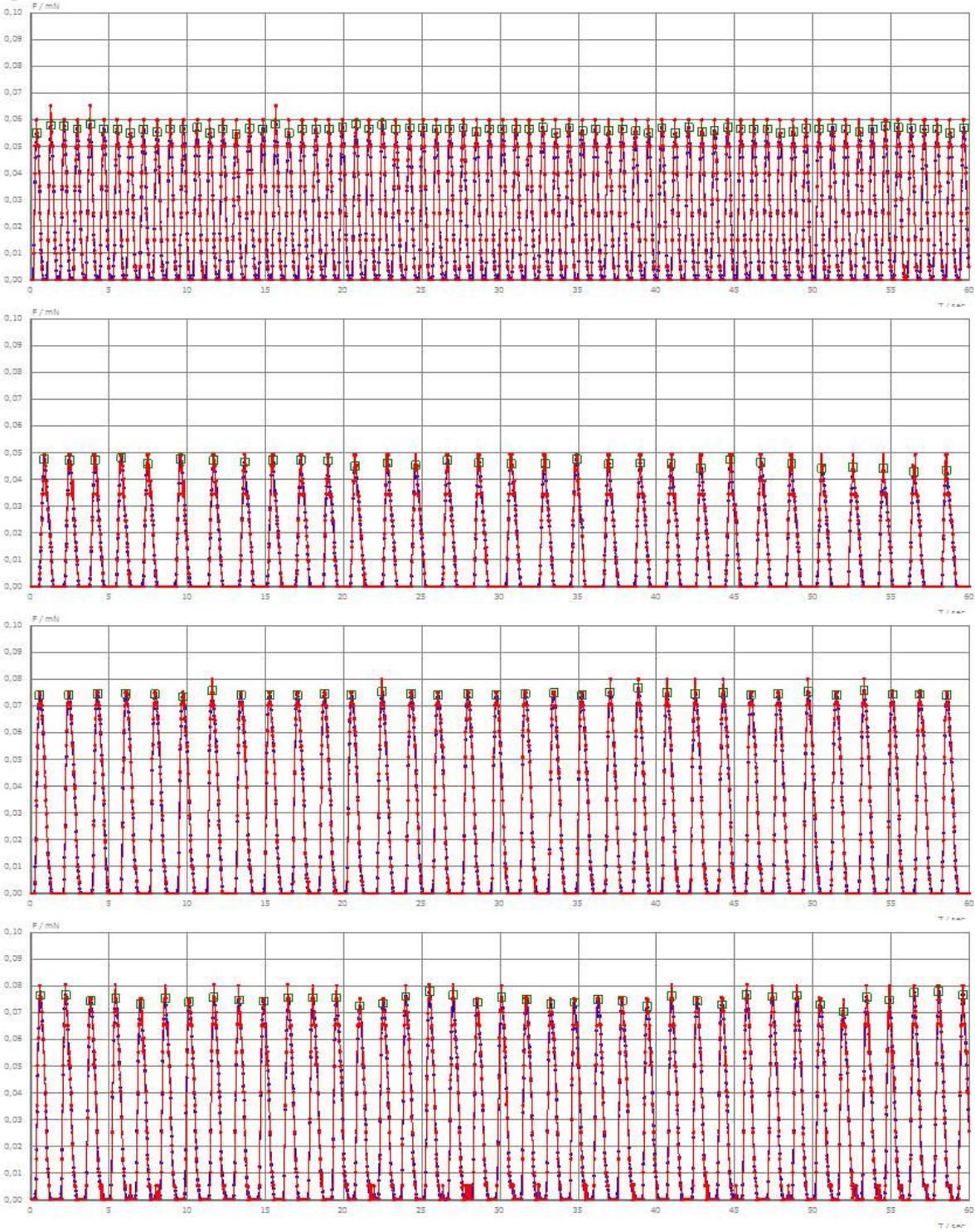
Graphical illustration of irregularity integrating 4 biological replicas. Ordinates indicate the distance from a given twitch to the following, the abscissa the distance to the previous twitch. Biological replicas are discriminated by color code.



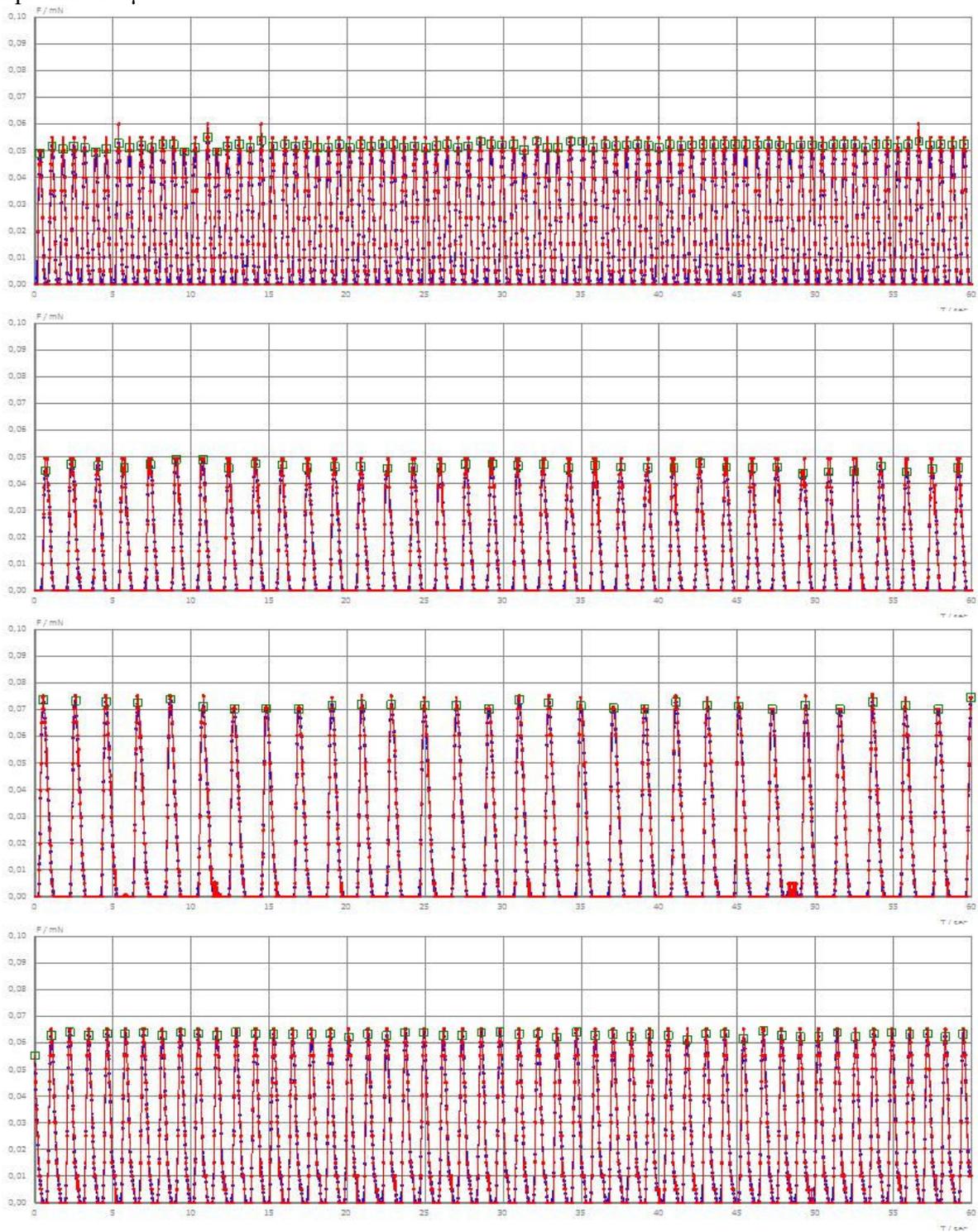
Scatter of beat-to-beat variability in the presence of cisapride, $*P < 0.05$ (Mann-Whitney U test), 4 biological replicas, bars show median \pm interquartile range.

Ampicillin - original recordings

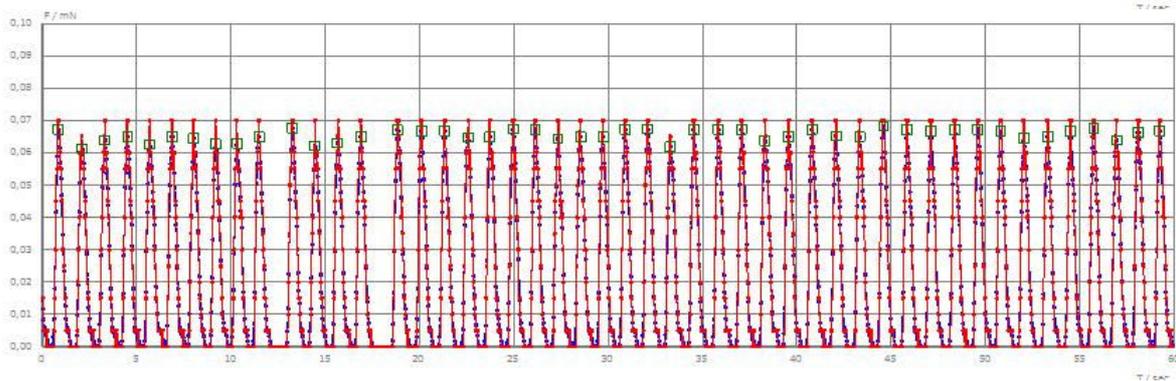
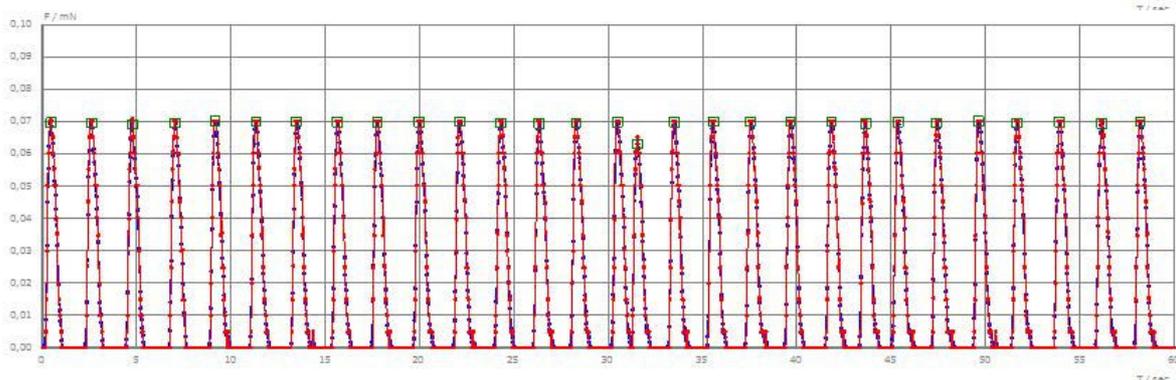
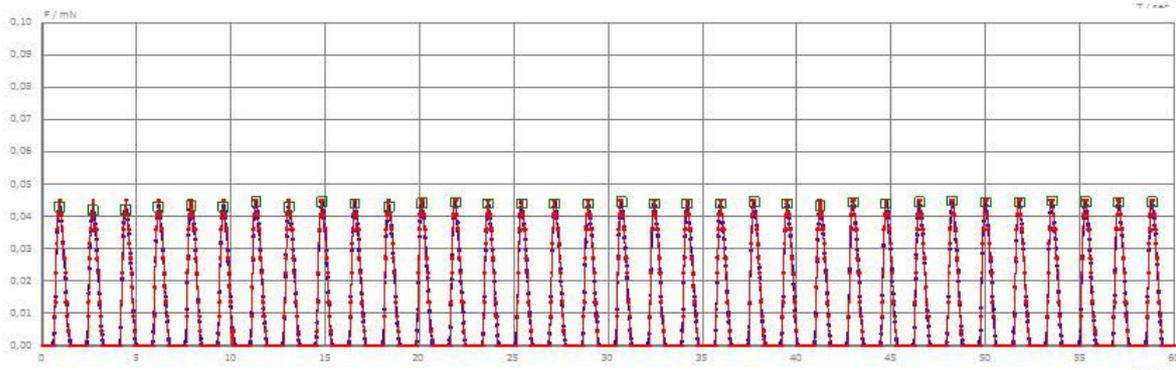
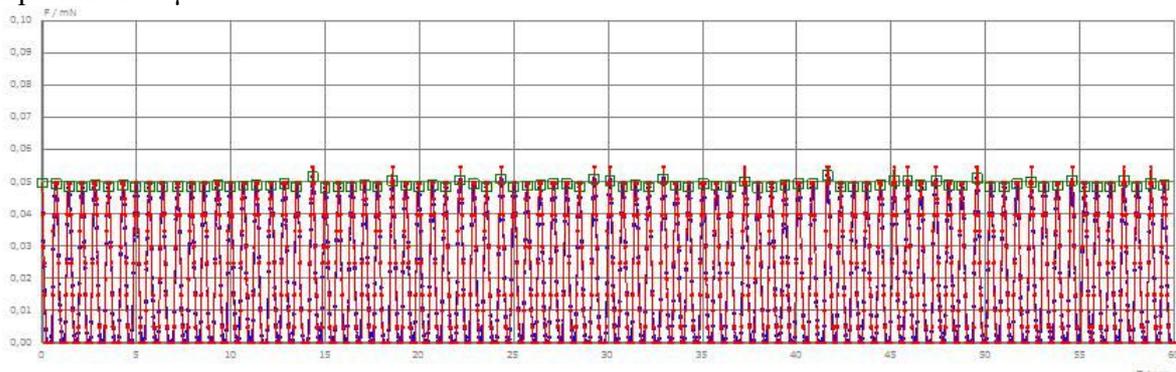
Ampicillin baseline



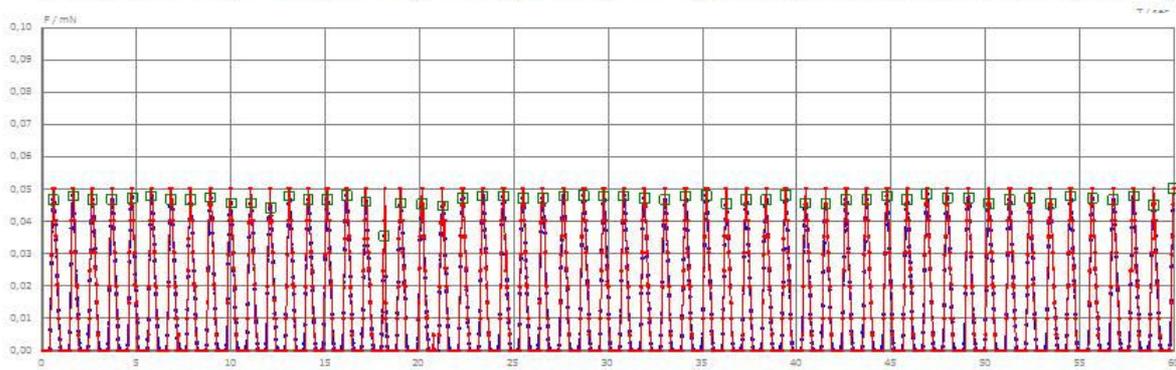
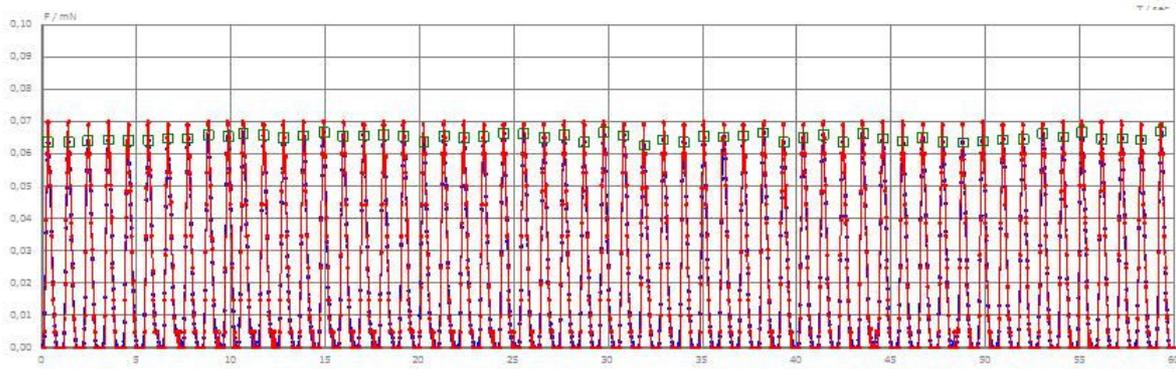
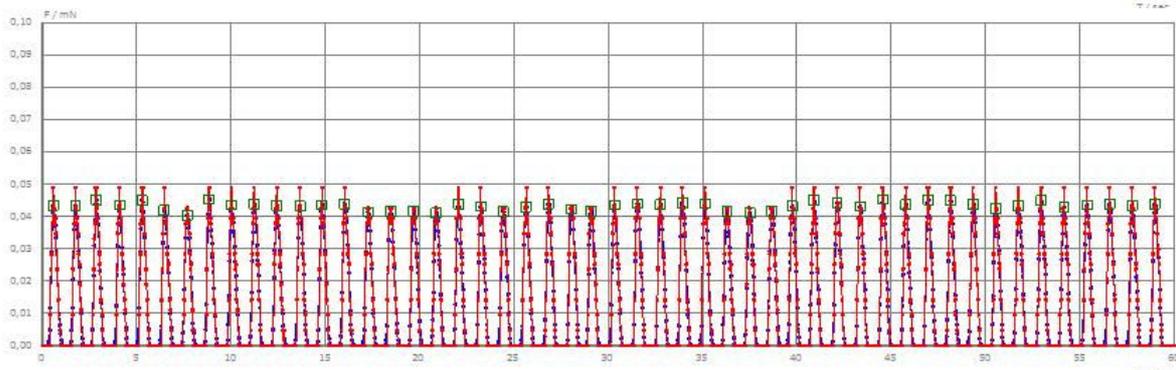
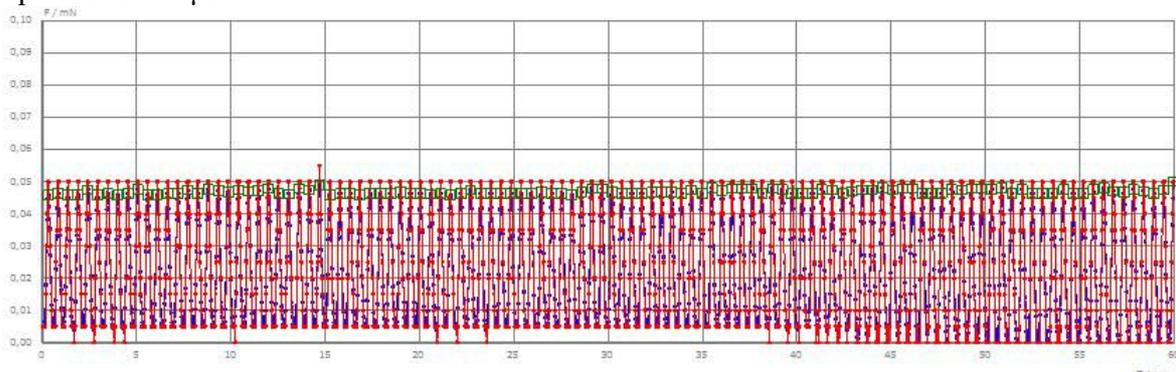
Ampicillin 10 μM



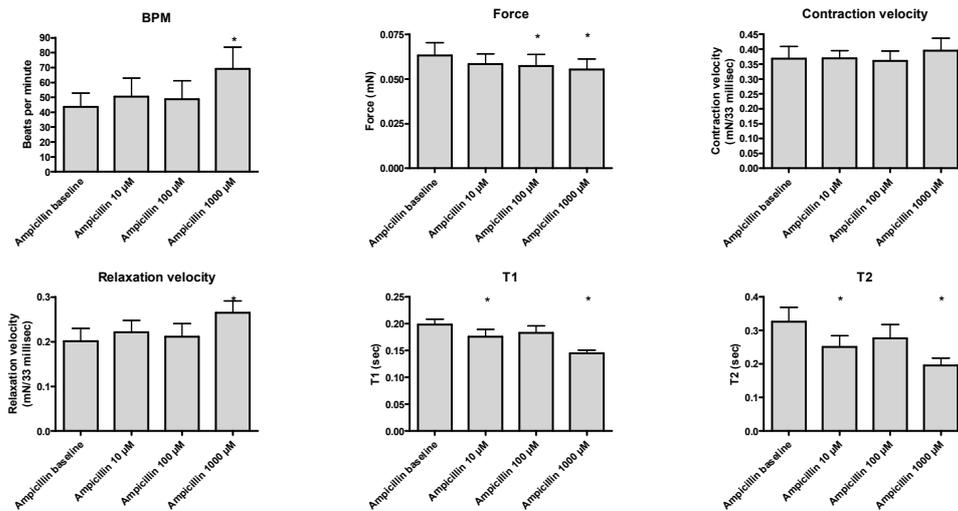
Ampicillin 100 μ M



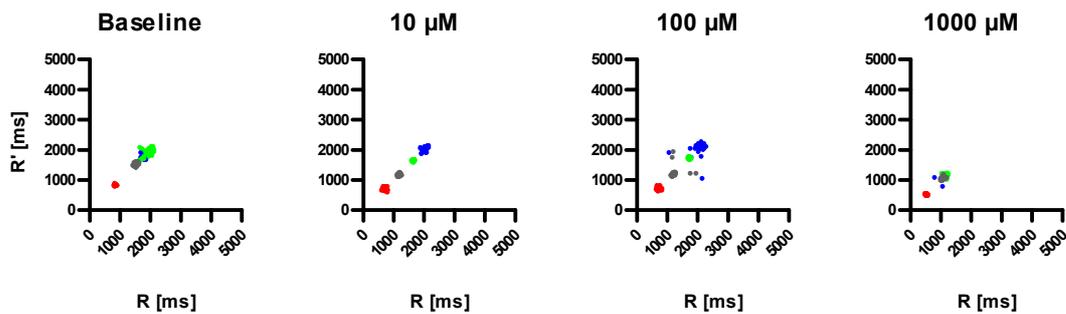
Ampicillin 1000 μM



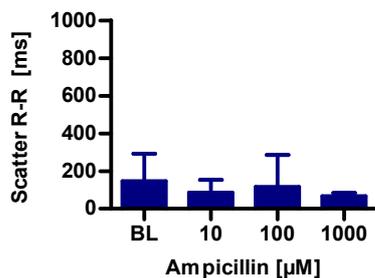
Analysis ampicillin



Analysis of contraction. * $P < 0.05$ (Student's t-test), 4 biological replicas, bars show means \pm SD. Beats per minute (BPM), contraction time (T1), relaxation time (T2).



Graphical illustration of irregularity integrating 4 biological replicas. Ordinates indicate the distance from a given twitch to the following, the abscissa the distance to the previous twitch. Biological replicas are discriminated by color code.



Scatter of beat-to-beat variability in the presence of ampicillin, * $P < 0.05$ (Mann-Whitney U test), 4 biological replicas, bars show median \pm interquartile range.