Supple. 1. Effects of tramadol, duloxetine, fluvoxamine, sertraline, morphine or reboxetine on the combination of collagen and CXCL12-indued platelet aggregation.

| agent | $\begin{aligned} & \text { dose } \\ & (\mu \mathrm{M}) \end{aligned}$ | Transmittance (\%) |  | Size of aggregation |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  |  | Large (\%) |  | Medium (\%) |  | Small (\%) |  |
| tramadol | 0 | 73.7 | $\pm 4.3$ | 56.3 | $\pm 4.1$ | 17.7 | $\pm 1.5$ | 26.0 | $\pm 2.9$ |
|  | 100 | 53.3 | $\pm 5.2^{*}$ | 32.0 | $\pm 6.7^{*}$ | 24.3 | $\pm 0.9 *$ | 44.0 | $\pm 6.1^{*}$ |
|  | 200 | 34.0 | $\pm 1.5^{*}$ | 12.3 | $\pm 3.4 *$ | 20.7 | $\pm 2.3$ | 67.0 | $\pm 5.7^{*}$ |
|  | 300 | 26.3 | $\pm 1.2 *$ | 6.0 | $\pm 1.0^{*}$ | 15.7 | $\pm 0.3$ | 78.3 | $\pm 1.2^{*}$ |
| duloxetine | 0 | 74.7 | $\pm 0.9$ | 48.7 | $\pm 3.8$ | 19.0 | $\pm 0.6$ | 32.0 | $\pm 3.8$ |
|  | 10 | 61.0 | $\pm 1.5^{*}$ | 43.3 | $\pm 3.2$ | 21.7 | $\pm 0.6$ | 37.0 | $\pm 0.6$ |
|  | 20 | 40.3 | $\pm 3.8 *$ | 16.0 | $\pm 3.8 *$ | 21.7 | $\pm 1.9$ | 62.3 | $\pm 5.5^{*}$ |
|  | 30 | 20.7 | $\pm 3.8 *$ | 7.0 | $\pm 1.5^{*}$ | 11.3 | $\pm 2.6 *$ | 81.3 | $\pm 4.3 *$ |
| fluvoxamine | 0 | 80.3 | $\pm 3.5$ | 58.3 | $\pm 3.8$ | 16.0 | $\pm 0.6$ | 25.7 | $\pm 3.3$ |
|  | 10 | 75.3 | $\pm 5.2$ | 45.3 | $\pm 1.8{ }^{*}$ | 19.7 | $\pm 1.2 *$ | 35.0 | $\pm 1.0^{*}$ |
|  | 20 | 56.3 | $\pm 4.7 *$ | 25.3 | $\pm 2.3 *$ | 24.0 | $\pm 0.0$ * | 50.7 | $\pm 2.0^{*}$ |
|  | 30 | 26.3 | $\pm 6.6^{*}$ | 6.7 | $\pm 3.2 *$ | 12.0 | $\pm 2.0$ | 81.3 | $\pm 5.1^{*}$ |
| sertraline | 0 | 81.7 | $\pm 5.5$ | 47.0 | $\pm 8.4$ | 18.7 | $\pm 1.8$ | 34.0 | $\pm 6.9$ |
|  | 10 | 54.3 | $\pm 2.3 *$ | 23.3 | $\pm 3.4 *$ | 22.3 | $\pm 1.8$ | 54.7 | $\pm 4.8$ |
|  | 20 | 45.7 | $\pm 9.0^{*}$ | 18.3 | $\pm 9.6$ | 17.7 | $\pm 4.1$ | 64.0 | $\pm 13.4$ |
|  | 30 | 26.3 | $\pm 1.2 *$ | 6.0 | $\pm 1.0^{*}$ | 15.7 | $\pm 0.3$ | 78.3 | $\pm 1.2^{*}$ |
| morphine | 0 | 80.0 | $\pm 2.1$ | 52.3 | $\pm 1.8$ | 17.7 | $\pm 0.3$ | 30.3 | $\pm 1.9$ |
|  | 100 | 81.7 | $\pm 5.5$ | 48.7 | $\pm 2.2$ | 19.0 | $\pm 0.6$ | 32.0 | $\pm 1.5$ |
|  | 200 | 83.0 | $\pm 5.6$ | 45.0 | $\pm 0.6$ | 18.7 | $\pm 1.3$ | 35.0 | $\pm 0.6$ |
|  | 300 | 85.3 | $\pm 5.2$ | 51.0 | $\pm 3.6$ | 17.3 | $\pm 0.3$ | 32.3 | $\pm 3.4$ |
| reboxetine | 0 | 80.7 | $\pm 6.8$ | 49.3 | $\pm 3.0$ | 18.3 | $\pm 0.9$ | 32.0 | $\pm 2.5$ |
|  | 30 | 76.7 | $\pm 6.4$ | 48.0 | $\pm 5.0$ | 19.0 | $\pm 1.0$ | 33.3 | $\pm 3.7$ |
|  | 70 | 77.3 | $\pm 4.7$ | 48.0 | $\pm 3.6$ | 20.7 | $\pm 0.7$ | 31.0 | $\pm 3.0$ |
|  | 100 | 72.0 | $\pm 6.4$ | 48.7 | $\pm 7.6$ | 17.0 | $\pm 1.5$ | 34.7 | $\pm 6.4$ |

PRP was pretreated with various doses of tramadol, duloxetine, fluvoxamine, sertraline, morphine or reboxetine for 3 min , and then simultaneously stimulated by collagen ( $0.075-0.45 \mu \mathrm{~g} / \mathrm{ml}$ ) and $10 \mathrm{ng} / \mathrm{ml}$ of CXCL12 for 5 min . The results analyzed with the aggregometer for the transmittance and the ratio of the platelet aggregates size in large, medium and small, are summarized. Each value represents the mean $\pm \mathrm{SEM}$ of 3 healthy donors. $* p<0.05$, compared to the value of control in each.

