

Table 1. Weight and sex of the rats included in the study. The code of the rat indicate that is an anaesthetised rat (A), the first two digits show the month of the experiment, the following two the day, and the last ones the number of the rat in this day.

Rat	Sex	Weight (g)	Drug
A0316-02	Male	297	Bupivacaine
A0311-02	Female	325	Bupivacaine
A0309-02	Male	360	Bupivacaine
A0304-01	Female	260	Bupivacaine
A0302-01	Male	320	Bupivacaine
A0216-02	Female	260	Bupivacaine
A0218-01	Male	279	Bupivacaine
A0219-01	Female	251	Bupivacaine
Mean		274	
SD		34.26	
A0316-01	Male	300	IQB-9302
A0311-01	Female	285	IQB-9302
A0309-01	Male	370	IQB-9302
A0304-02	Female	280	IQB-9302
A0302-02	Male	335	IQB-9302
A0216-03	Female	270	IQB-9302
A0218-02	Male	370	IQB-9302
A0219-02	Female	234	IQB-9302
Mean		267.25	
SD		23.03	
A0318-01	Female	280	Control
A0318-02	Female	235	Control

Table 2a. Rectal temperature before the administration of each dose in the **control** group.

Rat	Drug	Dose (mg/kg)	Temperature (°C)
A0318-02	Control	0.1	36.74
	Control	0.3	36.64
	Control	1	36.13
	Control	3	36.17
	Control	10	
A0318-01	Control	0.1	37.85
	Control	0.3	38.28
	Control	1	37.88
	Control	3	
	Control	10	38.41

Table 2b. Rectal temperature before the administration of each dose in the **bupivacaine** group.

Rat	Drug	Dose (mg/kg)	Temperature (°C)
A0316-02	Bupivacaine	0.1	36.28
	Bupivacaine	0.3	38.03
	Bupivacaine	1	37.71
	Bupivacaine	3	37.42
	Bupivacaine	10	37.52
A0311-02	Bupivacaine	0.1	36.99
	Bupivacaine	0.3	37.12
	Bupivacaine	1	37.39
	Bupivacaine	3	
	Bupivacaine	10	
A0309-02	Bupivacaine	0.1	37.53
	Bupivacaine	0.3	37.48
	Bupivacaine	1	37.49
	Bupivacaine	3	36.39
	Bupivacaine	10	36.28
A0304-01	Bupivacaine	0.1	36.29
	Bupivacaine	0.3	37.03
	Bupivacaine	1	37.52
	Bupivacaine	3	37.86
	Bupivacaine	10	37.84
A0302-01	Bupivacaine	0.1	37.93
	Bupivacaine	0.3	37.5
	Bupivacaine	1	37.12
	Bupivacaine	3	36.96
	Bupivacaine	10	36.72
A0219-01	Bupivacaine	0.1	36.83
	Bupivacaine	0.3	36.72
	Bupivacaine	1	36.88
	Bupivacaine	3	36.97
	Bupivacaine	10	37.35
A0218-01	Bupivacaine	0.1	36.19
	Bupivacaine	0.3	36.39
	Bupivacaine	1	37.34
	Bupivacaine	3	37.78
	Bupivacaine	10	37.78
A0216-02	Bupivacaine	0.1	34.04
	Bupivacaine	0.3	34.57
	Bupivacaine	1	35.59
	Bupivacaine	3	36.25
	Bupivacaine	10	

Table 2c. Rectal temperature before the administration of each dose in the **IQB-9302** group.

Rat	Drug	Dose (mg/kg)	Temperature (°C)
A0316-01	IQB-9302	0.1	36.57
	IQB-9302	0.3	36.48
	IQB-9302	1	36.56
	IQB-9302	3	36.66
	IQB-9302	10	36.47
A0311-01	IQB-9302	0.1	36.49
	IQB-9302	0.3	37.1
	IQB-9302	1	37.74
	IQB-9302	3	37.24
	IQB-9302	10	37.34
A0309-01	IQB-9302	0.1	36.92
	IQB-9302	0.3	37.53
	IQB-9302	1	37.27
	IQB-9302	3	37.17
	IQB-9302	10	37.25
A0304-02	IQB-9302	0.1	37.24
	IQB-9302	0.3	37.24
	IQB-9302	1	36.68
	IQB-9302	3	36.32
	IQB-9302	10	36.06
A0302-02	IQB-9302	0.1	37.32
	IQB-9302	0.3	36.82
	IQB-9302	1	36.78
	IQB-9302	3	37.14
	IQB-9302	10	37.2
A0219-02	IQB-9302	0.1	36.42
	IQB-9302	0.3	37.15
	IQB-9302	1	37.32
	IQB-9302	3	36.92
	IQB-9302	10	36.92
A0218-02	IQB-9302	0.1	36.66
	IQB-9302	0.3	36.67
	IQB-9302	1	
	IQB-9302	3	36.78
	IQB-9302	10	
A0216-03	IQB-9302	0.1	38.17
	IQB-9302	0.3	37.5
	IQB-9302	1	36.42
	IQB-9302	3	36.24
	IQB-9302	10	

Table 3a. Heart rate (bpm) recorded at different times (min) after the administration of **0.1 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	450.3	450.7	445.1	441.6	445.4	446.7	449.6	450.2	452.3	458.9	464.3	460	462.6	454	449.4	385.7	412.8	431.7	368.5	448.6	425.6	430.5
A0311-02	337.3	335.7	335.9	333.7	335.7	335.3	337.8	338.7	342.5	341.6	345.3	345.1	342.6	332.4	336.3	336.5	333.9	335.8	341.6	328.6	326.3	337.5
A0309-02	419.7	416.8	401	404.3	391.3	406.1	408.7	409.9	411.4	411.2	414	418.4	419.8	418.3	419.8	418	419.4	421.2	421.7	428.4	429.8	427.1
A0304-01	386.6	385.6	378.9	379.8	381.1	381.7	383.4	383.1	380.8	379.6	381.7	381.7	384.3	383.3	387.1	385.2	388.6	391.1	396.4	400.3	401	411.1
A0302-01	422.6	421.5	421.4	412.7	417.7	417.3	423.8	430.8	427.4	428.5	439.8	437.8	440.2	425.9	423.5	430	422.4	425.7	425.4	426.4	401.3	436.7
A0216-02	306.4	308.8	301.3	301.2	301.2	301.9	299.4	300.7	300	296.7	295.1	303.6	293	293.6	296.3	292.1	287.8	286.1	286.7	287.8	304.8	315.8
A0218-01	256.2	257	263.6	277.1	274.5	264.5	263.2	260.4	258.6	258.6	251.5	249.1	248.4	254.5	258.6	255.1	248.7	254.4	250.8	251.5	308	331.7
A0219-01	381.7	379.6	377.7	375.1	373.4	376.9	375	379.3	381.8	379.4	377.7	378.4	381.2	383.6	383.5	381.1	385.1	387.6	386.8	386.3	386.4	383.3
Mean	370.1	369.46	365.61	365.69	365	366.3	367.6	369.1	369.4	369.3	371.2	371.8	371.5	368.2	369.3	360.5	362.3	366.7	359.74	369.74	372.9	384.2
SD	65.642	64.784	61.448	57.048	57.8	61.49	63.6	65.21	65.58	67.39	71.86	70.58	73.45	69.12	66.65	61.14	65.35	67.37	63.059	72.176	51.844	49.4

Table 3b. Heart rate (bpm) recorded at different times (min) after the administration of **0.3 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	430.8	413.6	416.3	420	422.9	430.4	434.4	433.9	432.2	434	435	435.5	437.5	442.6	440.8	446.7	443.1	442.5	444.3	451.2	454.2	454.5
A0311-02	337.5	335.8	334.3	329.2	330.7	329.5	330.3	331.2	334.4	334.8	333.9	339	336.5	340.3	339	340.9	346.8	347.9	346.8	342.3	341.7	303.2
A0309-02	432.8	431.1	408.5	412.5	406.6	408.6	408	379.9	411.4	409.9	419.5	413.2	413.7	414.1	416.5	414	413.6	415.9	420.4	426.6	421	400.5
A0304-01	411.1	411.5	395.2	392.4	396	398.3	397.9	401	400.6	400.3	404	405.7	401.1	404.9	405.3	407.1	406.5	411.9	414.3	417.4	423.8	424.6
A0302-01	459.1	454.2	445.9	423.5	424.7	431.6	435.5	438	441.2	444.8	456	463.3	470.4	472	475.7	474.5	472	470.4	460.8	460.8	447.9	448.1
A0216-02	315.8	315.1	305.2	307.6	308.5	310.1	311.3	311.4	311.3	313.9	317.6	320.5	320.8	324.6	325.3	327.3	332.6	332.6	335.2	338.9	351.6	335.9
A0218-01	331.7	332.7	333.9	325.8	327.4	322.3	321.3	324.5	323.9	324.4	329.3	328.3	331.3	332.7	336.9	337.9	337.2	334.2	336.8	343.9	362.6	349.4
A0219-01	383.3	384.8	370.9	363	369.9	371.4	371.5	373.8	374.3	376.4	377.6	381	384.5	387.4	386.3	385.4	384.1	385.4	380.6	384.8	382.8	381.6
Mean	387.76	384.85	376.28	371.75	373.3	375.3	376.3	374.2	378.7	379.8	384.1	385.8	387	389.8	390.7	391.7	392	392.6	392.4	395.74	398.2	387.2
SD	53.96	51.375	48.527	46.609	46.07	49.3	50.34	48.77	50.49	50.62	52.64	52.65	54.06	53.89	54.09	53.82	51.16	51.43	49.595	50.177	44.158	54.72

Table 3c. Heart rate (bpm) recorded at different times (min) after the administration of **1 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	409.1	403.2	413.5	418.9	424.3	429.4	435.2	436	436.6	438.2	440.9	447.6	447.6	444.8	449.3	451.7	449.2	461.1	444.8	439.4	443.1	440.9
A0311-02	303.5	304.1	291.8	289.5	300.1	303.9	310.4	317.1	323.5	331	330.7	329.8	329.9	324.2	322	319.1	313.9	315.1	316	313.1	317.4	335.8
A0309-02	400.5	400.1	357.3	346.9	349.3	357	365.8	369.1	372.7	375.6	381.7	388.8	389.9	387.3	391.8	390	389.8	358.4	361.5	376.7	339.2	343.4
A0304-01	424.6	426.6	404.3	373.8	375.6	370.3	380.9	385.3	383.9	386.2	390.9	392.8	395.2	395.6	393.9	394.2	399.4	402.1	410.7	414	417.6	427.6
A0302-01	448.2	447	433	373.3	368.3	372	379.4	386.7	392.4	397.2	406.7	419	423.4	423.4	426.6	429	420.9	426	422.3	422.9	404.2	370.1
A0216-02	336.5	332.5	273.7	285.7	292.4	301.8	308.9	313.9	312.1	309.3	316.6	334.3	326.8	203	29.7	11	323.8	1.8	0.8	0.8	336.5	335.9
A0218-01	349.4	348.4	343.1	328	321.5	323.5	323	323.2	324.7	327.3	321.1	325.6	330	335.9	335.9	344.5	342.4	350	354.8	351.7	357.3	374.6
A0219-01	375.4	377.1	354.3	325.3	337.3	343.8	349.5	352.5	350.3	354.2	360.5	363	363.8	372.6	372.3	371.5	377.6	379.1	380.2	380.9	379.8	374.4
Mean	380.9	379.88	358.88	342.68	346.1	350.2	356.6	360.5	362	364.9	368.6	375.1	375.8	360.9	340.2	338.9	377.1	336.7	336.39	337.44	374.39	375.3
SD	48.702	48.736	56.78	45.239	43.38	42.15	43.03	42.4	42.4	42.68	44.42	44.72	45.9	75.49	132.4	139.1	47.53	142.8	141.71	141.98	44.272	39.99

Table 3d. Heart rate (bpm) recorded at different times (min) after the administration of **3 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	433.5	440.5	442	440.5	434.1	438.9	440.5	442.2	443.4	437.1	440.2	440.1	438.7	438.1	435.5	434.8	418.9	426.9	445.6	450.3	451	447.4
A0311-02	335.8	333.1	277.9	298.8	243.9	172.7	69.4	3.3														
A0309-02	310.7	308.5	315.7	285.4	277.4	274	275.5	278.4	287.8	291.6	292.4	292	292.6	291.1	289.8	290.7	291.6	289.6	290.5	288.5	287.2	292.3
A0304-01	427.6	426.2	362	314.2	327.7	343.4	346.9	351	356	357.8	354.2	357.2	361.8	364.7	369.3	375	377.8	381.5	383.6	394.6	396.9	395.7
A0302-01	392.1	336.8	367.9	330.6	315.3	311.6	313.6	318	324.4	330	335.1	338.2	340.7	345.5	348.3	350.3	357.9	360.1	352.6	333.4	327.4	358.5
A0216-02	335.9	328.4	244.5	267.9	284.9	284.9	290.7	294.5	297.3	300	300.6	304.2	300.8	302.4	303	308.9	311.3	312.6	318.6	319	333.2	340.1
A0218-01	374.6	376.1	347.7	5.5	2	339.1	328.2	320.2	235.7	327.3	331	335.4	334.3	337.3	332.3	326.3	327.1	324.4	332.7	346.8		
A0219-01	374.4	377.8	313.6	283.6	295.4	312.3	317.8	323.4	325.4	327.4	330.1	336	334.3	338.9	341.6	344.1	349.2	353.1	359.8	367.9	377.4	382.8
Mean	373.08	365.93	333.91	278.31	272.6	309.6	297.8	291.4	324.3	338.7	340.5	343.3	343.3	345.4	345.7	347.2	347.7	349.7	354.77	357.21	362.18	369.5
SD	44.162	47.914	60.584	122.68	122.9	74.99	104.9	126.5	64.68	48.47	48.75	48.07	48.29	48.08	47.91	47.54	42.78	46.04	50.062	53.283	58.305	52.68

Table 3e. Heart rate (bpm) recorded at different times (min) after the administration of **10 mg/kg bupivacaine**.

Rat*	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2
A0316-02	177.6	111.5	170.2	104.2	77.8					
A0309-02	292.3	293.6	23.3	135.6	94.4	76.8				
A0304-01	395.7	394.6	166.4	113.2	104	103.6	104			
A0302-01	358.5	355.6	214.3	114.3	106.4	101.1	57.6			
A0216-02	340.1	338.3	16.2	16.2						
A0218-01	325.5	326.3	216.2	144.6	116.9	65	7.2			
A0219-01	388.8	387.6	282.2	105.8	133.3	128.2	84.9			
Mean	325.5	315.36	155.54	104.84	105.5	94.94	63.43			
SD	74.402	96.432	100.34	41.922	18.95	24.72	42.04			

* There are only 7 rats because rat A0311-02 died 1.5 min after the 3 mg/kg dose.
 All rats were dead at 1.5 min after the administration of the 10 mg/kg dose.

Table 4a. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **0.1 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	114.6	115.82	114.95	118.85	118.4	118.4	119.2	119.4	116.3	119.1	119.1	117	118.5	110.2	111.1	83.64	104.8	112.1	80.76	123.7	115.89	116.3
A0311-02	102.28	102.45	107.65	106.55	108.6	107.9	108.8	108.1	107.7	108.5	109.1	106.2	103.7	81.84	86.05	97.06	99.91	106.6	106.97	88.66	90.64	100.8
A0309-02	126.24	125.89	126.75	126.81	121	126.8	130.6	128.2	127.2	125.9	127.2	128.5	129.1	126.6	128.8	127.9	131.2	133	131.49	133.01	130.26	133.7
A0304-01	122.21	121.49	122.79	125.24	126.2	125.4	126.3	124.7	121	121.4	122.6	121.1	121.9	123.1	125.9	124.2	126.7	125.9	127.36	126.33	121.2	127
A0302-01	101.68	103.66	107.58	108.06	112.3	111.4	112.2	115.7	108.4	112.3	118.1	113.4	114.1	102.4	102.8	109.5	103.6	109.1	111.26	104.11	101.47	117
A0216-02	113.06	115.47	114.49	116.63	113.6	113.8	112	111.7	119.3	115.8	94.38	85.99	83.25	87.39	85.58	90.8	93.74	95.3	95.3	100.84	112.39	112.5
A0218-01	69.85	71.2	75.02	85.52	86.35	80.15	76.93	75.71	73.3	70.49	65.36	62.73	64.14	67.7	66.93	65.12	63.26	67.07	68.44	69.81	104.35	110.7
A0219-01	100.56	97.62	103.34	103.51	105	104.1	97.2	104.8	103.7	104.6	99.2	100.7	100.3	101.5	101	97.3	101.9	100.8	100.73	101.16	101.71	97.67
Mean	106.31	106.7	109.07	111.4	111.4	111	110.4	111	109.6	109.8	106.9	104.4	104.4	100.1	101	99.44	103.2	106.2	102.79	105.95	109.74	114.5
SD	17.584	17.391	15.862	13.515	12.2	14.78	17.07	16.33	16.64	17.3	20.29	21.35	21.7	20.3	21.1	20.82	20.84	20.1	21.521	21.112	12.672	12.09

Table 4b. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **0.3 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	118.68	121.07	123.13	123.74	125.2	126.6	126.3	124.3	120.2	119	117.8	118.5	118.5	120.8	123.4	125.3	118.9	118.8	124.65	126.46	123.91	126.5
A0311-02	100.83	102.07	106.64	106.41	107.1	101.5	101.8	101.9	102.4	99.25	97.9	103.6	100.5	104.1	104	99.2	106.2	106.8	105.27	100.13	93.82	71.33
A0309-02	133	136.59	129.22	135.78	128.8	128.5	130.7	100.3	114.5	107.6	117.4	128.5	132.6	134.2	132.6	134.5	130.3	128.5	122.97	127.62	126.2	105.6
A0304-01	126.99	130	130.18	128.28	131.3	130.5	130.4	130.4	130.6	130.1	130.2	130.1	126.4	126.7	124.2	123.1	123.9	124	125.71	127.29	128.69	129.3
A0302-01	105.81	95.25	78.11	103.66	106.8	108.7	110.6	111	110.7	111.6	113.1	115.4	115.3	114.6	115.5	111.8	111.2	108.6	105.34	105.34	108.45	111.4
A0216-02	112.45	111.9	116.91	118.09	117.7	118.2	116.5	114.7	112.9	114.5	115.2	114.9	116	117.5	116.4	117.4	116.8	115.7	116.93	116.72	122.93	112.5
A0218-01	110.7	113.14	115.25	115.53	114.3	115	115.1	115.6	115.7	114.2	115.1	108.9	112.1	113.7	116.7	118.2	118.5	116.9	114.07	118.93	122.83	121
A0219-01	97.67	98.42	99.77	103.27	103.9	100.4	101	101.6	99.88	101.3	99.93	103.2	102.6	102.8	101.9	98.8	97.81	97.55	97.93	99.45	95.34	96.9
Mean	113.27	113.56	112.4	116.85	116.9	116.2	116.5	112.5	113.4	112.2	113.3	115.4	115.5	116.8	116.8	116	115.4	114.6	114.11	115.24	115.27	109.3
SD	12.351	14.912	17.366	11.995	10.64	11.91	11.85	11.07	9.705	9.882	10.32	10.19	10.84	10.61	10.25	12.41	10.21	9.942	10.344	12.046	14.111	18.73

Table 4c. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **1 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	126.54	130.61	131.33	132.74	130.6	129.8	132.9	134.4	128.5	127.4	129.8	132.3	126.5	126.4	127.2	126.9	127.1	128.4	121.26	119.31	124.21	121.8
A0311-02	75.58	77.96	92.42	82.67	88.66	83.39	79.51	83.24	82.79	86.42	91	95.43	99.13	96.89	96.6	93.54	81.35	76.96	80.62	81.57	86.92	105.6
A0309-02	105.64	108.92	106.33	105.75	108.1	110.5	113.2	116	115.8	116.8	120.1	123.9	123.6	114.4	113.8	114.8	111.6	86.35	97.81	114.57	91.74	84.84
A0304-01	129.32	132.22	130.37	125.66	133	126	130.4	129.3	126.9	127.2	128.3	126.7	124.9	127.9	124.7	124.2	122.8	122.6	124.47	125.53	119.52	125.3
A0302-01	110.98	111.09	114.91	99.56	105.1	108.1	108.2	110.4	111.6	111.4	114.1	114.1	109.5	109.5	110.5	111.6	99.39	105.9	101.69	105.71	92.92	84.01
A0216-02	112.54	113.45	108.8	110.91	112.9	113.9	116.9	118.2	115.8	111.9	116.8	118.8	120.6	121.8	121.3	121.3	117.1	98.26	104.32	104.32	104.62	105.4
A0218-01	120.97	121.29	120.75	120.76	121	119.7	117.3	118	118.6	119.7	114.2	116.3	121	123.7	123.7	128.5	124.7	121.3	126.89	123.85	125.87	131.1
A0219-01	91.88	95.63	100.81	101.77	105.3	104.1	103.4	102.4	102.8	102.2	102.3	100.1	102.1	102	101	100.2	100.3	98.8	98.83	98.36	95.64	97.01
Mean	109.18	111.4	113.22	109.98	113.1	111.9	112.7	114	112.9	112.9	114.6	115.9	115.9	115.3	114.8	115.1	110.5	104.8	106.99	109.15	105.18	106.9
SD	18.132	18.038	13.812	16.127	14.69	14.5	16.77	15.95	14.63	13.59	12.88	12.7	10.77	11.61	11.42	12.75	15.8	18.29	15.976	14.78	15.818	17.96

Table 4d. Mean arterial pressure recorded (mmHg) at different times (min) after the administration of **3 mg/kg bupivacaine**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-02	119.85	124.73	125.68	125.26	120.9	121.8	124	123	123.6	116.8	120.3	119.7	121.3	121.7	122.3	123	109.9	122.4	123.97	123.21	123.62	122
A0311-02	105.64	105.51	102.1	53.3	22.88	22.19	11.21	11.21														
A0309-02	63.83	63.58	86.03	102.69	110.1	102.4	101.6	99.51	99.88	101.1	93.14	87.74	82.46	77.71	71.84	68.81	65.07	61.61	61.31	57.92	59.02	58.1
A0304-01	125.26	127.97	118.6	129.65	134.4	136.4	131.4	132	131.6	128.9	123.3	122.8	119.9	120.2	118.7	119.4	118.4	119.6	118.86	120.06	117.29	116.6
A0302-01	112.86	102.58	83.16	104.12	108	108.5	109.3	109.7	109.1	108.3	104.3	102.3	103.1	103.5	104.6	100.8	105.6	101	86.31	72.27	77.13	107.7
A0216-02	105.43	101.91	112.63	112.6	113.1	112.1	112	110.3	110.1	109.7	109.4	109.9	106.5	104.6	103.9	102.6	102.6	102.3	102.33	102.35	109.32	107.3
A0218-01	131.14	132.02	145.25	152.04	152	152.1	151.2	149.3	143.4	137.3	128.8	125.7	123	123.6	121	120.1	116.3	111.1	109.44	102.13	111.87	110.1
A0219-01	97.01	97.33	98.62	106.19	107.4	107.4	104	102.7	103.5	101.5	99.14	99.61	98.13	97.71	97.45	97.94	98.18	99.49	100.09	100.77	98.48	96.94
Mean	107.63	106.95	109.01	110.73	108.6	107.9	105.6	104.7	117.3	114.8	111.2	109.7	107.8	107	105.7	104.7	102.3	102.5	100.33	96.959	99.533	102.7
SD	20.977	22.06	20.868	28.551	37.94	38.45	41.52	41.23	16	13.76	13.29	13.93	14.83	16.48	17.78	18.89	17.93	20.19	21.246	23.898	23.409	21.17

Table 4e. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **10 mg/kg bupivacaine**.

Rat*	-0.25	0	0.25	0.5	0.75	1	1.25
A0316-02	96.12	55.36	41.65	27.85	0		
A0309-02	58.1	58.92	22.46	43.48	25.25	16.11	
A0304-01	116.6	118.09	79.97	56.46	44.33	31.32	14.95
A0302-01	107.68	103.81	58.99	56.94	36.8	23.69	23.31
A0216-02	107.26	109.52	49.42	49.42			
A0218-01	104.22	105.82	124.33	86.12	53.26	25.29	9.9
A0219-01	101.86	99.9	90.82	42.48	43.56	27.53	10.5
Mean	98.834	93.06	66.806	51.821	33.87	24.79	14.67
SDt	19.016	25.197	34.209	18.08	19.02	5.634	6.188

* There are only 7 rats because rat A0311-02 died 1.5 min after the 3 mg/kg dose.
All rats were dead at 1.5 min after the administration of the 10 mg/kg dose.

Table 5a. Heart rate (bpm) recorded at different times (min) after the administration of **0.1 mg/kg IQB-9302**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	428.1	427.9	426	422.4	416.8	418.2	420.2	423.5	426.2	421.1	434.2	436.9	443.2	442	442.7	442.8	448.1	439.2	437.9	440.9	438	419.8
A0311-01	415.1	417.7	415.9	400.4	407.4	407.2	408.4	412.1	412	413	414.8	418.9	423.9	422.5	422.7	427	430	403.3	384.9	362.2	399.5	419
A0309-01	406.9	406.4	405.8	399.7	408.8	412.7	407.5	415.7	418.9	417.2	407.1	410.1	406.7	410.6	410.4	405.9	399.7	392.6	380.2	386.5	387.3	378.6
A0304-02	416.7	417.8	399.2	410.7	383.1	406.6	405.4	408.9	411.9	414	416.5	416.4	417	422	362	418.2	382.8	374	372.8	319.4	368.9	394.9
A0302-02	349.9	343.5	355.8	363.7	362.9	362.9	366.7	376.2	372	372.1	373.3	364.2	362.2	364.9	359.4	349	347.6	348.7	351.2	349.4	335.1	334.9
A0216-03	393.3	393.8	392.8	393.4	393	377.9	388.4	387.5	390.7	393.4	393.3	393.5	390.3	389.4	390.6	389.6	386.3	388.8	388.5	389.7	385.2	382.2
A0218-02	369.5	368	350.4	354.3	362.4	361.7	362.5	367.7	368.3	370.5	378.5	381.6	384.1	386.6	377.9	375.3	351.8	339.2	343.9	343.7	369.1	367.8
A0219-02	351.2	356.6	370.1	372.2	370.8	377.8	379.9	386.4	386.5	391.2	396.6	396.7	395.6	385.7	371.1	367.1	362.4	349.3	349.4	343.5	372.2	367.9
Mean	391.34	391.46	389.5	389.6	388.2	390.6	392.4	397.3	398.3	399.1	401.8	402.3	402.9	403	392.1	396.9	388.6	379.4	376.1	366.91	381.91	383.1
SD	30.73	31.633	27.879	23.826	21.61	23.02	21.22	20.41	21.92	20.24	20.41	23.06	25.36	25.42	30.36	32.18	36.13	33.6	30.35	37.892	29.554	28.24

Table 5b. Heart rate (bpm) recorded at different times (min) after the administration of **0.3 mg/kg IQB-9302..**

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	416.2	418.4	406	403.6	407.2	410	414.1	418.1	420.5	396.5	417.7	423.2	430.7	437.3	422.5	421.5	426.6	432.9	426.3	434.5	429.2	415.7
A0311-01	413.9	417.1	407.6	398.9	410.2	407.5	408.7	409.3	410.2	410.5	416.1	418.8	412	399.4	421.3	415.2	422.6	426.2	423.7	433.8	430.2	419.7
A0309-01	376.3	375.8	368.4	371.3	374.3	375.8	385.3	372.9	373.8	383.4	380.8	384.1	389.7	384.6	392.5	396.5	384.1	383.1	377.3	392.8	388.3	385.6
A0304-02	394.9	395.9	379.1	369.9	371.2	376.2	381.9	384.1	388	388.8	392.1	394.8	397.7	388.7	403.3	408	415.2	423.8	427.9	428.4	427.4	417.8
A0302-02	351.8	354.7	347.1	336.7	342.9	337.5	359.1	353.9	356.7	362.9	354.1	347.4	349.7	352.8	350	352.7	351.8	348.6	357.8	317.2	305.6	320
A0216-03	385.2	382.3	383.3	370.6	368.6	370.8	374.4	376.6	376.3	377.4	378.2	374	376.6	377.7	377.9	376.5	374.5	374.5	373.3	370.2	364.7	358.6
A0218-02	357.4	355.7	333.9	331	333.3	345.7	352.9	351.2	352.4	359.7	354.1	359.7	350.4	344.5	341.7	341.5	346.2	360.7	341.7	330.3	330.9	384.9
A0219-02	356.1	356.5	357.6	359.7	358.8	363.1	361.5	365.4	366.9	371	378.2	386.5	395.9	399.2	403	407.2	414	424.5	385.3	423.1	390.5	380.9
Mean	381.48	382.05	372.88	367.71	370.8	373.3	379.7	378.9	380.6	381.3	383.9	386.1	387.8	385.5	389	389.9	391.9	396.8	389.16	391.29	383.35	385.4
SD	25.585	26.392	26.423	25.802	27.32	25.87	22.56	24.21	24.32	17.2	24.19	26.37	28.21	29	30.38	29.8	32.16	33.75	33.201	47.388	46.991	34.14

Table 5c. Heart rate (bpm) recorded at different times (min) after the administration of **1 mg/kg IQB-9302**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	415.7	418.3	399.6	389.7	388.3	391.1	398.1	402.2	404.4	412.2	412.2	413.1	415.8	416.4	413.7	416.1	414.8	419.1	407.2	411.3	393.7	381.4
A0311-01	419.7	422.6	383.6	356.6	353	358.6	360.8	365.3	368.3	375.1	378.2	385	390	394.5	398.2	401.2	404.5	401.3	407.3	408.9	408.5	405.1
A0309-01	385.6	386.6	364.5	336	336.8	345.5	360.6	367.2	371.8	381	387.1	388.2	385.8	388.5	389.3	385.2	383.1	388.5	391.8	400.1	410.5	377.1
A0304-02	417.8	378.7	378.1	348.2	346.4	353	357.4	359.6	364.4	368.7	369	375.7	377.6	380.8	386.1	387.9	391.1	352.3	403	401	403.2	393.1
A0302-02	320	318.7	306.6	289.9	302.6	310.3	314.7	312.7	315.4	314.7	319.4	326	327.6	329.5	330.6	337.4	336.1	337.2	339.3	336	348.9	368.4
A0216-03	357.6	358.7	357.2	337.3	326.3	332.7	338.4	339.9	341	342	345.7	347.6	349.5	350.1	349.1	350.3	354.6	354.5	355.2	357.8	358.4	357
A0218-02	380.6	376.1	352.8	340.5	328.4	330.8	344.9	349.7	354.8	356.5	363.1	366.3	371	369.4	372.8	373.6	351	346.7	334.1	346.9	344.9	368.8
A0219-02	371.1	371.3	357.6	348.7	336.7	337.8	342.4	348.5	348.1	357.2	363.6	370.2	378	386	392.4	394	397.4	393.8	399.8	391.2	362.9	354.8
Mean	383.51	378.88	362.5	343.36	339.8	345	352.2	355.6	358.5	363.4	367.3	371.5	374.4	376.9	379	380.7	379.1	374.2	379.71	381.65	378.88	375.7
SD	34.627	32.991	27.62	27.598	24.74	23.85	23.98	25.6	25.91	28.72	27.57	26.4	26.54	27.06	27.21	26.11	28.42	30.08	31.449	29.971	27.817	17.27

Table 5d. Heart rate (bpm) recorded at different times (min) after the administration of **3 mg/kg IQB-9302**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	392	395.1	221.8	260.7	304.2	308.4	133.9	65.1	263.7	329.2	360.3	359.5	360	363.7	367.9	371.7	378.4	382.5	389.1	391.9	405	385.1
A0311-01	405.1	407.1	372	301.8	294.6	292.6	294.6	302.8	313.1	313.1	322.9	331.3	334.4	340	339.7	343.3	357	358.8	365.6	366.6	373.7	369.7
A0309-01	437.1	438.2	345.7	328.7	349.5	380.2	389.4	394.6	400	401.9	406	406.2	401.1	402.4	405.1	406.3	416.5	424.1	433	445.4	439.5	446.8
A0304-02	393.1	393.1	356.2	290.3	302.3	307.6	315.5	325.4	326.9	328.7	333.1	338.3	340.8	343.8	342.7	345.8	354.4	359.1	367.9	376.8	393.2	397.8
A0302-02	373.3	372.6	362.1	348.3	290.7	292.8	294.7	309.3	308.5	319	322.6	328.5	330.7	335.9	335.4	335.6	341.3	342.2	345.1	347.2	349.6	329.1
A0216-03	356.1	357.2	328.9	294.2	291.5	336.6	340.3	322.3	321.5	325.2	328.6	331.6	334.1	336.6	336	338.7	340.8	341.8	339.2	340.6	333.3	325
A0218-02	368.8	367.2	351.2	315.7	281.4	269.4	287.2	298.3	302.6	301.7	301.4	304.7	301.8	306.6	310.2	305.9	307.1	306.7	308.1	335.1	356.2	371.1
A0219-02	354.8	352.3	346.3	351.7	352.6	356.9	359.8	366.4	370.3	369.4	374.9	383.8	389.5	389.6	393.1	396.8	398.1	383.1	380	392.8	359.6	349.6
Mean	385.04	385.35	335.53	311.43	308.4	318.1	301.9	298	325.8	336	343.7	348	349.1	352.3	353.8	355.5	361.7	362.3	366	374.55	376.26	371.8
SD	27.722	28.78	47.67	30.954	27.3	36.98	76.73	99.82	42.02	33.09	34.1	33.16	32.82	31.3	32.19	33.66	34.88	35.05	37.282	36.212	34.548	39.62

Table 5e. Heart rate (bpm) recorded at different times (min) after the administration of **10 mg/kg IQB-9302**.

Rat*	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75
A0316-01	385.1	384.4	105.1	123.9	123	89.4	77.4	32.7	
A0311-01	369.7	369.9	222.2	135.4	130.9	125.1	27.6		
A0309-01	447.8	448.8	189.8	76	121.6	116.3	82.8	77.6	
A0302-02	328.3	343.8	143	167.7	136.1	108	96.1	51.7	3.6
A0216-03	347.2	350.5	323.6	139.1	186	135.8	137.8	27.5	
A0218-02	371.1	363.3	341.3	185.3	115.3	109.5	134.4	112.8	104.8
Mean	374.87	376.78	220.83	137.9	135.5	114	92.68	60.46	54.2
SD	40.985	38.085	95.384	37.868	25.8	15.91	40.88	35.24	71.56

* There are only 6 rats because by mistake rat A0304-02 received two consecutive doses of 3 mg/kg and did not receive the 10 mg/kg dose, and in rat A0219-02 we lost the record of the effects of the administration of 10 mg/kg.rat. All rats were dead at 2 min after the administration of the 10 mg/kg dose.

Table 6a. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **0.1 mg/kg IQB-9302**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	111.38	111.79	115.73	117.49	112.4	107	111.7	116.5	117	104	121.7	119.4	120	119.7	118.4	122	120.6	116	116.14	117.44	118.04	100
A0311-01	110.83	111.58	110.97	113.52	114.7	113.8	113.9	114.1	113.5	112.3	113.2	114.6	115.1	113.9	114.8	114.6	115.2	108.6	102.08	92.94	107.53	114.6
A0309-01	120.22	121.83	126.76	128.16	131.4	131.9	125.8	131.5	132.9	127.8	122.5	122	122.5	123.5	126.7	118.3	119.2	116.2	118.24	115.75	115.02	114.5
A0304-02	117.54	117.05	117.15	118.68	109.5	107.8	110.4	116.3	120.9	120.9	121.2	120	119.3	120.2	118.3	107.7	68.86	66.12	68.85	99.49	94.42	117.8
A0302-02	98.25	94.4	110.41	115.93	111.5	120.7	122.5	121.9	118.3	116.8	115.5	105	104.3	106	99.97	97.2	95.7	98.61	101.33	105.59	100.78	103.8
A0216-03	106.62	107.71	111.74	111.4	111.2	104.6	110.7	109.8	111.5	113.3	111.4	111.3	106.5	107.3	108.6	106.5	107.1	108.7	106.19	110.85	109.75	109.5
A0218-02	127.72	126.27	129.55	130.97	133	132	131.6	132.2	131.6	134.7	135.5	135.7	134.9	135.5	124.7	122.2	89.32	86.11	102.11	100.41	124.57	119.7
A0219-02	86.16	94.92	103.82	106.42	110.1	111.8	114.2	118.2	115.2	116.4	117.5	113.3	112	86.55	66.4	62.89	69.88	60.82	58.5	50.07	103.69	82.24
Mean	109.84	110.69	115.77	117.82	116.7	116.2	117.6	120.1	120.1	118.3	119.8	117.7	116.8	114.1	109.7	106.4	98.24	95.14	96.68	99.068	109.23	107.8
SD	13.069	11.54	8.6469	8.2282	9.683	10.89	7.984	8.037	8.017	9.549	7.54	9.128	9.737	14.56	19.49	19.57	20.91	21.9	21.536	21.506	9.791	12.34

Table 6b. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **0.3 mg/kg IQB-9302**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	109.1	111.23	112.67	115.93	114.9	116.5	115.9	119.2	120.8	106.9	115.4	118.9	118.4	118.5	111.2	112.8	115.6	117.7	112.75	118.43	117.85	102.9
A0311-01	113.46	112.57	110.27	111.51	114.9	114.4	114.1	114.9	114.7	113.7	115.3	116.7	111.5	109.2	114.3	112.9	116.6	117.4	114.43	116.13	115.83	115.8
A0309-01	114.61	114.34	118.27	119.85	121.1	121.2	122.9	117.4	115.6	120	117	118.7	120.5	117.7	120.3	122.8	117.5	117.9	115.86	119.69	120.53	122.7
A0304-02	117.8	116.41	118.28	117.5	119	121.1	122.5	118.4	123.3	121.2	120.4	120	121.3	121.7	120.1	120.7	122.6	128.3	127.91	125.23	124.34	121.6
A0302-02	107.75	100.32	113.17	104.69	112.2	99.64	119.4	118.1	117.2	120.2	111.6	106.6	109.3	108.9	108.9	112.4	109.6	105.7	114.15	77.72	79.99	93.42
A0216-03	110.77	109.96	115.76	112.7	112.3	113.1	115.5	116.3	112.5	116.7	114.5	111.4	115.6	113.2	113.1	111.1	109.3	110.6	110.27	113.04	111.49	113.6
A0218-02	107.82	110.25	116.14	114.74	121.1	113.4	115.9	115.5	111	113	105.9	118.3	101.5	103.9	93.16	95.08	106.6	122.8	88.75	86.67	98.33	120.1
A0219-02	48.74	51.26	69.96	94.64	106.4	109.7	108.9	110.5	113.1	116.6	119.3	121.5	124.7	125.1	121.6	120.9	115.7	118.4	99.05	112.11	82.81	54.14
Mean	103.76	103.29	109.32	111.45	115.2	113.6	116.9	116.3	116	116	114.9	116.5	115.3	114.8	112.8	113.6	114.2	117.4	110.4	108.63	106.4	105.5
SD	22.508	21.555	16.145	8.1757	5.039	6.903	4.609	2.774	4.204	4.753	4.577	4.996	7.582	7.21	9.206	8.781	5.267	6.9	11.77	16.985	17.265	23.08

Table 6c. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **1 mg/kg IQB-9302**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	102.92	105.56	109.64	111.48	112.7	114.4	115.7	115.1	114	112.5	118	113.3	115.6	112.1	111.7	111.5	111.3	113.6	91.1	111.41	88.1	81.42
A0311-01	115.83	116	114.06	107.89	111	110.3	113.9	112.9	113.3	115.8	115.9	116.6	118.5	117.1	117	117.7	117.9	115.4	115.99	115.86	115.33	114.6
A0309-01	122.65	122.9	121.19	121.33	123	123.8	127.6	127.8	128.1	130.5	131	132.7	125.9	127.3	124.4	119.5	119.9	120	121.27	124.1	131.51	115
A0304-02	121.59	117.14	106.87	104.37	107.5	108.5	110.7	112.4	112.6	114.6	115.1	118.4	119.8	118.6	122.8	118	116.8	114.7	115.09	104.11	119.16	81.7
A0302-02	93.42	95.95	105.59	99.99	101.5	100.8	102.5	100.2	101.6	95.27	95.74	96.73	97.95	96.99	98.49	100.2	101	101.1	104.71	105.54	105.84	114.3
A0216-03	113.41	93.74	91.74	103.24	111.6	114.8	117.8	117.1	117.2	117	118.1	119.8	120.3	122.6	118.8	115.8	120.5	116.3	116.89	117.44	114.08	114.5
A0218-02	114.23	113.18	116.66	117.76	117.3	120	119.2	119.6	121.1	123	124.4	124.8	126.7	124.6	120.4	121.1	93.78	88.92	82.77	113.53	101.77	121
A0219-02	50.08	47.44	55.37	88.25	88.14	92.18	96.23	99.38	102.2	104.5	108	111.9	112.5	114.3	114.6	111.3	109.1	101.9	107.17	86.7	55.44	50.37
Mean	104.27	101.49	102.64	106.79	109.1	110.6	113	113	113.8	114.1	115.8	116.8	117.2	116.7	116	114.4	111.3	109	106.87	109.84	103.9	99.11
SD	23.935	24.14	21.044	10.443	10.58	10.25	9.858	9.517	8.89	10.75	10.55	10.47	9.1	9.479	8.222	6.724	9.633	10.59	13.59	11.34	23.409	25.16

Table 6d. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **3 mg/kg IQB-9302**.

Rat	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75	2	2.5	3	3.5	4	4.5	5	6	7	8	10	15	20
A0316-01	96.17	101.09	107.82	115.48	102.4	96.04	45.22	21.52	69.49	120.4	137.2	131.7	125.3	127.1	127	126.1	122.6	119.3	120.47	117.56	116.01	96.47
A0311-01	114.55	119.12	117.43	121.16	121.3	114.2	107.6	111.1	109.1	109.1	111.1	109.3	109.4	110.1	110.5	112	113.2	112.1	115.34	112.85	111.54	108.9
A0309-01	132.35	134.07	119.7	129.68	138.4	140.6	142.4	141.3	141	141.2	137.8	135	131.3	128.8	127.8	127.4	129.5	132	134.73	133.8	125.44	129.4
A0304-02	81.7	78.66	91.86	98.43	100.5	104	103.5	107	108.1	106.9	111.1	113.2	111.1	111.9	114.1	112.8	117.3	118.2	120.03	112.67	106.83	108.5
A0302-02	117	114.97	115.26	113.61	110.4	119.1	107.9	114.9	110	114.2	112.9	115.1	115.1	117.3	116.2	112.4	110.3	110.3	110.14	110.1	122.99	115.8
A0216-03	113.36	112.33	114.23	120.97	126.5	127.2	125.5	122.4	124.7	124.1	123.4	119.8	116.4	117.5	114.9	116.2	112.5	109.1	106.22	103.05	73.98	68.67
A0218-02	121	119.23	124.84	120.07	117.4	108.3	101.6	96.31	84.6	77.54	74.66	82.83	78.59	81.28	81.72	76.06	76.55	80.72	84.88	113.85	123.31	122.2
A0219-02	50.37	48.49	48.63	61.57	62.92	82.65	94.76	98.95	101.5	100.5	105.5	106.2	107.2	105.5	105.3	104.6	101.4	78.3	77.86	93.17	48.81	45.83
Mean	103.31	103.5	104.97	110.12	110	111.5	103.5	101.7	106.1	111.8	114.2	114.1	111.8	112.4	112.2	110.9	110.4	107.5	108.71	112.13	103.61	99.46
SD	26.435	27.476	24.822	21.567	22.76	18.11	28.04	35.35	22.08	18.62	20.1	16.24	15.7	14.92	14.48	15.99	16.02	18.74	18.956	11.643	27.641	28.49

Table 6e. Mean arterial pressure (mmHg) recorded at different times (min) after the administration of **10 mg/kg IQB-9302**.

Rat*	-0.25	0	0.25	0.5	0.75	1	1.25	1.5	1.75
A0316-01	96.47	100.9	65.47	65.9	42.64	25.91	18.67	14.06	
A0311-01	108.88	110.16	92.15	58.12	33.04	18.01	73.65		
A0309-01	128.85	131.76	91.42	44.45	43.67	25.8	19.37	16.07	
A0302-02	115.24	116.93	112.46	111.26	56.24	24.91	12.33	10.35	9.64
A0216-03	65.69	70.26	73.65	70.13	69.1	24.34			
A0218-02	122.16	116.92	117.64	95.45	86.43	80.33	42.93	30.15	18.22
Mean	106.22	107.82	92.132	74.218	55.19	33.22	33.39	17.66	13.93
SD	22.77	20.991	20.576	24.715	19.76	23.27	25.34	8.659	6.067

* There are only 6 rats because by mistake rat A0304-02 received two consecutive doses of 3 mg/kg and did not receive the 10 mg/kg dose, and in rat A0219-02 we lost the record of the effects of the administration of 10 mg/kg.rat. All rats were dead at 2 min after the administration of the 10 mg/kg dose.

Table 7. Predose heart rate (HR) (bpm), minimum heart rate (HR min), difference of change (minimum HR minus predose HR) and time when minimum HR is reached (Tmin) (min) after the administration of a dose of 1 mg/kg or 3 mg/kg of bupivacaine or of IQB-9302.

Rat	Drug	Dose	Predose HR	HR min	Difference of change	Tmin (min)
A0316-02	Bupivacaine	1 mg/kg	403.2	413.5	10.3	0.25
A0311-02	Bupivacaine	1 mg/kg	304.1	289.5	-14.6	0.5
A0309-02	Bupivacaine	1 mg/kg	400.1	349.3	-50.8	0.75
A0304-01	Bupivacaine	1 mg/kg	426.6	370.3	-56.3	1
A0302-01	Bupivacaine	1 mg/kg	447	368.3	-78.7	0.75
A0216-02	Bupivacaine	1 mg/kg	332.5	11	-321.5	5
A0218-01	Bupivacaine	1 mg/kg	348.4	321.1	-27.3	2.5
A0219-01	Bupivacaine	1 mg/kg	377.1	325.3	-51.8	0.5
Mean					-73.8375	1.40625
SD					103.811807	1.60877893
A0316-02	Bupivacaine	3 mg/kg	440.5	434.1	-6.4	0.75
A0311-02	Bupivacaine	3 mg/kg	333.1	3.3	-329.8	1.5
A0309-02	Bupivacaine	3 mg/kg	308.5	274	-34.5	1
A0304-01	Bupivacaine	3 mg/kg	426.2	314.2	-112	0.5
A0302-01	Bupivacaine	3 mg/kg	336.8	311.6	-25.2	1
A0216-02	Bupivacaine	3 mg/kg	328.4	244.5	-83.9	0.25
A0218-01	Bupivacaine	3 mg/kg	376.1	2	-374.1	0.75
A0219-01	Bupivacaine	3 mg/kg	377.8	283.6	-94.2	0.5
Mean					-132.5125	0.78125
SD					140.694933	0.38816188
A0316-01	IQB-9302	1 mg/kg	418.3	388.3	-30	0.75
A0311-01	IQB-9302	1 mg/kg	422.6	353	-69.6	0.75
A0309-01	IQB-9302	1 mg/kg	386.6	336	-50.6	0.5
A0304-02	IQB-9302	1 mg/kg	378.7	346.4	-32.3	0.75
A0302-02	IQB-9302	1 mg/kg	318.7	289.9	-28.8	0.5
A0216-03	IQB-9302	1 mg/kg	358.7	326.3	-32.4	0.75
A0218-02	IQB-9302	1 mg/kg	376.1	328.4	-47.7	0.75
A0219-02	IQB-9302	1 mg/kg	371.3	336.7	-34.6	0.75
Mean					-40.75	0.6875
SD					14.2174943	0.11572751
A0316-01	IQB-9302	3 mg/kg	395.1	65.1	-330	1.5
A0311-01	IQB-9302	3 mg/kg	407.1	292.6	-114.5	1
A0309-01	IQB-9302	3 mg/kg	438.2	328.7	-109.5	0.5
A0304-02	IQB-9302	3 mg/kg	393.1	290.3	-102.8	0.5
A0302-02	IQB-9302	3 mg/kg	372.6	290.7	-81.9	0.75
A0216-03	IQB-9302	3 mg/kg	357.2	291.5	-65.7	0.75
A0218-02	IQB-9302	3 mg/kg	367.2	269.4	-97.8	1
A0219-02	IQB-9302	3 mg/kg	352.3	346.3	-6	0.25
Mean					-113.525	0.78125
SD					94.1805826	0.38816188

Table 8. Predose mean arterial pressure (MAP) (mmHg), maximum MAP (MAP max), difference of change (maximum MAP minus predose MAP) and time when maximum MAP is reached (Tmax) (min) after the administration of a dose of 1 mg/kg or 3 mg/kg of bupivacaine or of IQB-9302..

Rat	Drug	Dose	Predose MAP	MAP max	Difference of change	Tmax (min)
A0316-02	Bupivacaine	1 mg/kg	130.61	134.43	3.82	1.5
A0311-02	Bupivacaine	1 mg/kg	77.96	99.13	21.17	3.5
A0309-02	Bupivacaine	1 mg/kg	108.92	123.91	14.99	3
A0304-01	Bupivacaine	1 mg/kg	132.22	132.98	0.76	0.75
A0302-01	Bupivacaine	1 mg/kg	111.09	114.91	3.82	0.25
A0216-02	Bupivacaine	1 mg/kg	113.45	121.83	8.38	4
A0218-01	Bupivacaine	1 mg/kg	121.29	128.52	7.23	5
A0219-01	Bupivacaine	1 mg/kg	95.63	105.27	9.64	0.75
Mean					8.72625	2.34375
SD					6.642705855	1.76239235
A0316-02	Bupivacaine	3 mg/kg	124.73	125.68	0.95	0.25
A0311-02	Bupivacaine	3 mg/kg	105.51	102.1	-3.41	0.25
A0309-02	Bupivacaine	3 mg/kg	63.58	110.05	46.47	0.75
A0304-01	Bupivacaine	3 mg/kg	127.97	136.38	8.41	1
A0302-01	Bupivacaine	3 mg/kg	102.58	109.67	7.09	1.5
A0216-02	Bupivacaine	3 mg/kg	101.91	113.14	11.23	0.75
A0218-01	Bupivacaine	3 mg/kg	132.02	152.06	20.04	1
A0219-01	Bupivacaine	3 mg/kg	97.33	107.42	10.09	0.75
Mean					12.60875	0.78125
SD					15.35608622	0.41052014
A0316-01	IQB-9302	1 mg/kg	105.56	118	12.44	2.5
A0311-01	IQB-9302	1 mg/kg	116	118.51	2.51	3.5
A0309-01	IQB-9302	1 mg/kg	122.9	132.7	9.8	3
A0304-02	IQB-9302	1 mg/kg	117.14	122.82	5.68	4.5
A0302-02	IQB-9302	1 mg/kg	95.95	105.59	9.64	0.25
A0216-03	IQB-9302	1 mg/kg	93.74	122.6	28.86	4
A0218-02	IQB-9302	1 mg/kg	113.18	126.71	13.53	3.5
A0219-02	IQB-9302	1 mg/kg	47.44	114.57	67.13	4.5
Mean					18.69875	3.21875
SD					21.06973824	1.38510894
A0316-01	IQB-9302	3 mg/kg	101.09	137.16	36.07	2.5
A0311-01	IQB-9302	3 mg/kg	119.12	121.32	2.2	0.75
A0309-01	IQB-9302	3 mg/kg	134.07	142.35	8.28	1.25
A0304-02	IQB-9302	3 mg/kg	78.66	114.07	35.41	4.5
A0302-02	IQB-9302	3 mg/kg	114.97	119.11	4.14	1
A0216-03	IQB-9302	3 mg/kg	112.33	127.2	14.87	1
A0218-02	IQB-9302	3 mg/kg	119.23	124.84	5.61	0.25
A0219-02	IQB-9302	3 mg/kg	48.49	107.24	58.75	3.5
Mean					20.66625	1.84375
SD					20.4689709	1.49962793