Repeated administration of the opioids
Analgesia time-course

Maximum Possible Effects, %

Interval post-Morphine, min

Interval post-DAMGO, min

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First dose; + 3h Second dose; + 6h Third dose; + 24h Fourth dose

Membrane Internalized

MOR P-Ser375

MOR levels

Dose: 1st 2nd 3rd

Sampled 3 h after each dose

10% sucrose

40% sucrose

EEA1 (early endosomes)

Rab 5 (very early endosomes)

Rab 4 (early endosomes)

Rab11 (recycling endosomes)

Lamp-1 (lysosomes)

Control

kDa 50

MOR

1st dose, + 30 min

MOR

P-Ser375

4th dose, + 30 min

MOR

P-Ser375

10% sucrose

40% sucrose

EEA1 (early endosomes)

Rab 5 (very early endosomes)

Rab 4 (early endosomes)

Rab11 (recycling endosomes)

Lamp-1 (lysosomes)

IgG hc