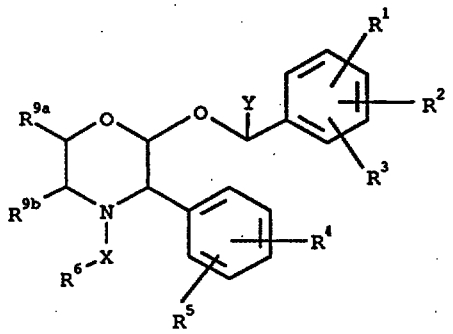
ABSTRACT

SUBSTITUTED MORPHOLINE DERIVATIVES AND THEIR USE AS THERAPEUTIC AGENTS

The present invention relates to compounds of the formula (I):



(I)

wherein

R^1 is hydrogen, halogen, C_{1-6}alkyl, C_{1-6}alkoxy, CF_3, NO_2, CN, SR^a, SOR^a, SO_2R^a, CO_2R^a, $CONR^aR^b$, C_{2-6}alkenyl, C_{2-6}alkynyl or C_{1-4}alkyl substituted by C_{1-4}alkoxy, where R^a and R^b are hydrogen or C_{1-4}alkyl;

R^2 is hydrogen, halogen, C_{1-6}alkyl, C_{1-6}alkoxy substituted by C_{1-4}alkoxy or CF_3;

R^3 is hydrogen, halogen or CF_3;

R^4 is selected from the definitions of R^1;

R^5 is selected from the definitions of R^2;

R^6 is a 5-membered or 6-membered heterocyclic ring containing 2 or 3 nitrogen atoms optionally substituted by =O, =S or a C_{1-4}alkyl group, and optionally substituted by an aminoalkyl group;

R^{9a} and R^{9b} are hydrogen or C_{1-4}alkyl, or R^{9a} and R^{9b} are joined to form a C_{5-7} ring;

X is C_{1-4}alkylene optionally substituted by oxo; and

Y is a C_{1-4}alkyl group optionally substituted by hydroxyl;

with the proviso that if Y is C_{1-4}alkyl, R^6 is substituted at least by an aminoalkyl group;

and pharmaceutically acceptable salts and prodrugs thereof.

The compounds are of particular use in the treatment of pain, inflammation, migraine and emesis.