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| <b>(51) International Patent Classification <sup>6</sup>:</b><br><b>C07D 211/44, 211/46, 211/52, 211/58,</b><br><b>401/14, 417/14, 401/06, 491/10, 471/10,</b><br><b>405/14, 409/14, A61K 31/445</b>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             | <b>A1</b> | <b>(11) International Publication Number:</b> <b>WO 97/24324</b><br><b>(43) International Publication Date:</b> 10 July 1997 (10.07.97)                                                                                                                                                                                                                                                                                                                                                                                      |
| <b>(21) International Application Number:</b> PCT/EP96/05883<br><b>(22) International Filing Date:</b> 20 December 1996 (20.12.96)<br><b>(30) Priority Data:</b><br>95203651.5 27 December 1995 (27.12.95) HP<br><b>(34) Countries for which the regional or international application was filed:</b> DE et al.<br><br><b>(71) Applicant (for all designated States except US):</b> JANSSEN PHARMACEUTICA N.V. [BE/BE]; Turnhoutseweg 30, B-2340 Beerse (BE).<br><br><b>(72) Inventors; and</b><br><b>(75) Inventors/Applicants (for US only):</b> JANSSENS, Frans, Ed- uard [BE/BE]; Tinststraat 79, B-2820 Bonheiden (BE). SOM- MEN, François, Maria [BE/BE]; Langenberg 49, B-2323 Wortel (BE). SURLERAUX, Dominique, Louis, Nestor, Ghislaine [BE/BE]; Diegemstraat 48, B-1830 Machelen (BE). VAN ROOSBROECK, Yves, Emiel, Maria [BE/BE]; Zonderschotsesteenweg 26, B-2220 Heist-opden-Berg (BE).                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            |           | <b>(81) Designated States:</b> AL, AM, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HU, IL, IS, JP, KG, KR, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, TR, TT, UA, US, UZ, VN, ARIPO patent (KE, LS, MW, SD, SZ, UG), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).<br><br><b>Published</b><br>With international search report. |
| <b>(54) Title:</b> 1-(1,2-DISUBSTITUTED PIPERIDINYL)-4-SUBSTITUTED PIPERIDINE DERIVATIVES AS TACHYKININ RECEPTOR ANTAGONISTS<br><br><b>(57) Abstract</b><br><p>The present invention concerns compounds of formula (I), the N-oxide forms, the pharmaceutically acceptable addition salts and stereochemically isomeric forms thereof, wherein n is 0, 1 or 2; m is 1 or 2, provided that if m is 2, then n is 1; p is 0, 1 or 2; -Q is -O or -NR<sup>3</sup>; X is a covalent bond or a bivalent radical of formula -O-, -S-, -NR<sup>3</sup>-; R<sup>1</sup> is Ar<sup>1</sup>; Ar<sup>1</sup>C<sub>1</sub>-alkyl or di(Ar<sup>1</sup>)C<sub>1</sub>-alkyl, wherein each C<sub>1</sub>-alkyl group is optionally substituted with hydroxy, C<sub>1</sub>-alkyloxy, oxo or a ketalized oxo substituent of formula -O-CH<sub>2</sub>-CH<sub>2</sub>-O- or -O-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-O-; R<sup>2</sup> is Ar<sup>2</sup>; Ar<sup>2</sup>C<sub>1</sub>-alkyl; Het or HetC<sub>1</sub>-alkyl; R<sup>3</sup> is hydrogen or C<sub>1</sub>-alkyl; R<sup>4</sup> is hydrogen; C<sub>1</sub>-alkyl; C<sub>1</sub>-alkyloxyC<sub>1</sub>-alkyl; hydroxyC<sub>1</sub>-alkyl; carboxyl; C<sub>1</sub>-alkyloxycarbonyl or Ar<sup>3</sup>; R<sup>5</sup> is hydrogen; hydroxy; Ar<sup>3</sup>; Ar<sup>3</sup>C<sub>1</sub>-alkyloxy; di(Ar<sup>3</sup>)C<sub>1</sub>-alkyloxy; Ar<sup>3</sup>C<sub>1</sub>-alkylthio; di(Ar<sup>3</sup>)C<sub>1</sub>-alkylthio; Ar<sup>3</sup>C<sub>1</sub>-alkylsulfoxy; di(Ar<sup>3</sup>)C<sub>1</sub>-alkylsulfoxy; Ar<sup>3</sup>C<sub>1</sub>-alkylsulfonyl; di(Ar<sup>3</sup>)C<sub>1</sub>-alkylsulfonyl; -NR<sup>7</sup>R<sup>8</sup>; C<sub>1</sub>-alkyl substituted with -NR<sup>7</sup>R<sup>8</sup>; or a radical of formula (a-1) or (a-2); R<sup>4</sup> and R<sup>5</sup> may also be taken together; R<sup>6</sup> is hydrogen; hydroxy; C<sub>1</sub>-alkyloxy; C<sub>1</sub>-alkyl or Ar<sup>3</sup>C<sub>1</sub>-alkyl; Ar<sup>1</sup>, Ar<sup>2</sup> and Ar<sup>3</sup> are phenyl or substituted phenyl; Ar<sup>2</sup> is also naphthalenyl; and Het is an optionally substituted monocyclic or bicyclic heterocycle; as substance P antagonists; their preparation, compositions containing them and their use as a medicine.</p> <div style="text-align: center;"> </div> |           |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |