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International Bureau

INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 6 : C07D 401/14, 401/04, C07C 229/20 — A1

(11) International Publication Number: **WO 98/51685**

(43) International Publication Date: 19 November 1998 (19.11.98)

(21) International Application Number: PCT/DK98/00183

(22) International Filing Date: 7 May 1998 (07.05.98)

(30) Priority Data:

0536/97	9 May 1997 (09.05.97)	DK
60/046,011	9 May 1997 (09.05.97)	US

(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, 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).

Published
With international search report.

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(54) Title: METHOD OF MANUFACTURING SERTINDOLE

(57) Abstract

A process of manufacturing sertindole comprising preparation of N–(4–fluorophenyl)–N–(2–carboxy–4–chlorophenyl)glycine, by reacting an alkalimetal salt of 2,5–dichlorobenzoic acid with an alkalimetal salt of N–(4–fluorophenyl)glycine in an aqueous, alkaline environment in the presence of a copper catalyst; cyclisation of N–(4–fluorophenyl)–N–(2–carboxy–4–chlorophenyl)glycine to the corresponding 3–acetoxy–indole; reduction of the 3–acetoxy–indole and subsequent elimination of H_2O thereby obtaining 5–chloro–1–(4–fluorophenyl)indole which is reacted with 4–piperidone in a mixture of an acetic acid and concentrated HC1; reduction of the resulting 5–chloro–1–(4–fluorophenyl)–3–(1,2,3,6–tetrahydropyridin–4–yl)indole and reaction of this compound with 1–(2–chloroethyl)–2–imidazolidinon in order to obtain sertindole. Alternatively, 5–chloro–1–(4–fluorophenyl)–3–(1,2,3,6–tetrahydropyridin–4–yl)indole is first reacted with 1–(2–chloroethyl)–2–imidazolidinon followed by reduction thereby obtaining sertindole. This process uses reactants and solvents that are suitable and allowed in large scale manufacture. Furthermore good total yields are obtained.