Short Communications

A Short Stereospecific Synthesis of (\pm) -Deplancheine *

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A new indole alkaloid (+)-deplancheine 1 has recently been isolated from the New Caledonian plant Alstonia deplanchei van Heurck et Mueller Arg. (Apocynaceae). Three syntheses of (\pm) -deplancheine have been described in the literature, 1-4 one of which appeared 2,3 before deplancheine was known as a natural product.

In the present communication we present a new and much shorter synthesis of (±)-deplancheine.

The reduction of substituted amides with LiAlH₄ to amines proceeds by an initial reduction to a geminal amino alcohol derivative, followed by elimination and subsequent reduction of the resulting iminium salt.⁵ Considering the fact that 1,2,6,7,-12,12b-hexahydro-3-acetylindolo[2,3-a]quinolizine 2 ⁶ is a vinylogous amide, we thought it should be ideally suited for the rapid and short preparation of (\pm)-deplancheine 1 (E-isomer) and/or its Z-isomer (Scheme 1).

In fact, we found that when 1,2,6,7,12,12b-hexahydro-3-acetylindolo[2,3-a]quinolizine 2 was treated with LiAlH₄ in THF, (\pm)-deplancheine 1 was obtained in 20% yield. [MS (70 eV, 130 °C) m/e 252 (100%) (M⁺), 251 (98%), 237 (15%), 223 (20%), 170 (37%), 169 (58%), 156 (29%); ¹H NMR (CDCl₃) δ 1.61 (3H, d, J = 6.5 Hz, CH_3 – CH =), 5.42 (1H, q, J = 6.5 Hz, CH_3 – CH =), 7.0 – 7.5 (4H, m, aromatic protons), 7.90 (1H, br s, NH), no aliphatic proton signals beyond δ 3.65 (eliminating the Z-configuration); ⁴ IR (CHCl₃) Bohlmann bands 2815 and 2760 cm⁻¹; (cf. Ref. 3)]. No traces of the other possible isomer (Z-isomer) could be found among the reaction products.

The present method provides an extremely easy route to create an E configurational ethylidene side chain at the 3-position (corresponding to the 20-position in the biogenetic numbering) of the indolo [2,3-a] quinolizine skeleton. This configuration is present in the majority of indole alkaloids having a similar unsaturation. The applicability of the method for the synthesis of other alkaloids of a similar nature is under investigation.

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Scheme 1.

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