

In conclusion, the assessment of biopharmaceutical evaluation was successfully applied to the microcapsule formulations of verapamil hydrochloride. The rate of absorption appeared to be more sustained, resulting in a relatively more uniform plasma concentration profile of the drug about at least 12 h. About twice and more bioavailabilities were noted with sustained release formulations even though the drug has substantial first pass metabolism. The results indicate that it is possible to make a once a day oral controlled release dosage form for verapamil hydrochloride.

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Chemical Components of *Melia azedarach* Stems

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Melia azedarach stems have been shown to contain melianin B, sendanolactone, ohchinin acetate, surianol and 3 α -hydroxy-4,4,14 α -trimethyl-5 α -pregn-8-en-20-one. The first three have already been reported from this plant. The last two compounds have already been reported from *Suriana maritima* and *Fomes officinalis* respectively.

Melia azedarach, known as bakain, drek and mahanimb, belongs to the family Meliaceae. It resembles neem in many respects and possesses antimicrobial, insecticidal and nematicidal properties. An aqueous extract of its leaves increases haemoglobin¹. Its stems have been reported to contain melianin A, melianin

B, nimbolin A and nimbolin B^{2,3}. We have undertaken a reinvestigation of its stems.

The dry stems of *Melia azedarach* (12 kg) were procured from the Landscape, CCSHAU, Hisar and extracted with hot methanol. The methanolic extract (300 g) was subjected to silica gel (60-120 mesh) column chromatography. Elutions with petroleum ether, benzene, ethyl acetate, methanol and their mixtures in the order of their

* For Correspondence

increasing polarity afforded five compounds which were characterised on the basis of their spectral data. ¹H NMR was recorded on a Bruker AC-300F 300 MHz NMR spectrometer in CDCl₃ using TMS as the internal standard. The other instruments used are Hitachi 570 infrared spectrophotometer and VG-705 11-250J GCMS-DS Mass Spectrometer.

Surianol was crystallised from hexane as colourless solid (12 mg), m.p. 170° (lit⁴ m.p. 173-174.5°). The compound was acetylated with Ac₂O/Py; m.p. 131° (lit⁴ m.p. 132-134°). IR (cm⁻¹, KBr) 3463, 3060, 2960, 2360, 1604, 1095, 1026, 802, 702. ¹H NMR (δ, CDCl₃) 0.13 (d, 1H, *J* = 4 Hz, cyclopropane H), 0.46 (d, 1H, *J* = 4 Hz, cyclopropane H), 0.91 (s, 3H, CH₃), 0.99 (s, 6H, 2xCH₃), 1.03 (d, *J* = 6.5 Hz, 6H, 2 x CH₃), 3.08 (t, 1H, *J* = 9 Hz, CHOH), 3.39 (m, 1H, CHOH), 4.65 (br s, 1H, = CH-), 4.70 (br s, 1H, = CH).

MS (m/z, rel. int.) 429 (M⁺ + 1, 1.51), 428 (M⁺, C₂₉H₄₈O₂, 0.1), 274 (10.03), 259 (16.12), 244 (10.94), 231 (26.07), 121 (14.23), 109 (11.30), 107 (20.84), 93 (7.38), 55 (100), 43 (64.24), 41 (92.38).

Melianin B was crystallised from benzene as a colourless solid (13 mg), m.p. 195° (lit² mp 198°). It readily formed an acetate with Ac₂O/Py, m.p. 268° (lit² mp 268-270°). IR (cm⁻¹, KBr) 3463, 2954, 2360, 1712, 1380, 1026, 810. ¹H NMR (δ CDCl₃) 0.92 (s, 3H, CH₃), 1.03 (s, 9H, 3xCH₃), 1.15 (s, 6H, 2xCH₃), 1.27 (s, 3H, CH₃), 1.64 (s, 3H, OAc), 2.04 (s, 3H, OAc), 4.64 (m, 1H, H-1), 4.94 (m, 1H, H-3), 5.22 (m, 1H, H-7), 5.35 (m, 1H, H-15), 7.50 (m, 3H, Ar-H), 8.07 (m, 2H, Ar-H). MS (m/z, rel. int.) 694 (M⁺, C₄₁H₅₈O₉, 2.0), 680 (M⁺-14, 0.23), 472 (0.32), 438 (0.98), 424 (1.73), 409 (1.83), 299 (1.91), 245 (2.96), 161 (8.98), 133 (15.99), 107 (23.22), 95 (35.36), 69 (58.81), 55 (100), 43 (90.87).

Sendanolactone was crystallised from benzene as solid mass (15 mg), m.p. 208° (lit⁶ m.p. 208-209°). IR (cm⁻¹, KBr) 1780, 1712, 1674, 1380, 1049, 732. ¹H NMR (δ, CDCl₃) 0.98 (s, 3H, CH₃), 1.10 (s, 3H, CH₃), 1.33 (s, 3H, CH₃), 1.39 (s, 6H, 2xCH₃), 1.64 (s, 3H, CH₃), 1.70 (s, 3H, CH₃), 4.16 (m, 1H, H-17), 5.10 (m, 1H, = CH-) 5.34 (m, 1H, = CH-) MS (m/z, rel. int.) 466 (M⁺, C₃₀H₄₂O₄, 0.42), 438 (0.89), 410 (6.54), 396 (13.14), 313 (8.84), 269 (24.75), 253 (9.92), 229 (23.01), 147 (30.05), 124 (100), 107 (39.20).

3α-Hydroxy-4,4,14α-trimethyl-5α-pregn-8-en-20-one was crystallized from benzene as pale yellow solid (18 mg), m.p. 230° (lit⁶ m.p. 230-232°). Acetylation of the compound with Ac₂O/pyridine afforded acetate having m.p. 250° (lit⁶ m.p. 250-252°). IR (cm⁻¹, KBr) 3787, 3456, 1728, 1458, 1380, 1049, 810. ¹H NMR (δ, CDCl₃) 0.75 (s, 3H, Me), 0.90 (s, 3H, Me), 0.99 (s, 9H, 3 Me), 2.05 (s, 3H, Me). MS (m/z, rel. int.) 359 (M⁺+1, C₂₄H₃₈O₂ + 1, 53.40), 135 (39.52), 121 (69.42), 107 (65.70), 93 (67.14), 78 (66.62), 44 (100).

Ohchinin acetate was crystallised from ethyl acetate as a pale coloured compound (25 mg), m.p. 220° (lit⁷ m.p. 223-226°). IR (cm⁻¹, KBr) 1728, 1604, 1512, 1380, 1033, 810. ¹H NMR (δ, CDCl₃) 0.98 (s, 3H, Me), 1.20 (s, 3H, Me), 1.31 (s, 3H, Me), 1.68 (d, *J* = 1.4 Hz, 3H, Me), 1.95 (s, 3H, CH₃COO), 3.74 (s, 3H, COOMe), 1.70-5.51 (m, 16 H, 3 x CH, 2 x CH₂, 5 x CHO-, CH₂O-, CH₂COO), 6.32 - 7.70 (m, 10 H, furan, olefinic and phenyl protons). MS (m/z, rel. int.) 644 (M⁺, C₃₈H₄₄O₉, 27.59), 168 (6.83), 149 (6.21), 119 (9.39), 97 (13.96), 84 (15.28), 69 (24.93), 45 (100).

It may be mentioned here that there are several reports⁸⁻¹⁰ on the stem bark of this plant.

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