

Diastereocontrol in Intermolecular Diels-Alder Reactions of Allenic Lactones: Synthetic Approach to the Plaunols

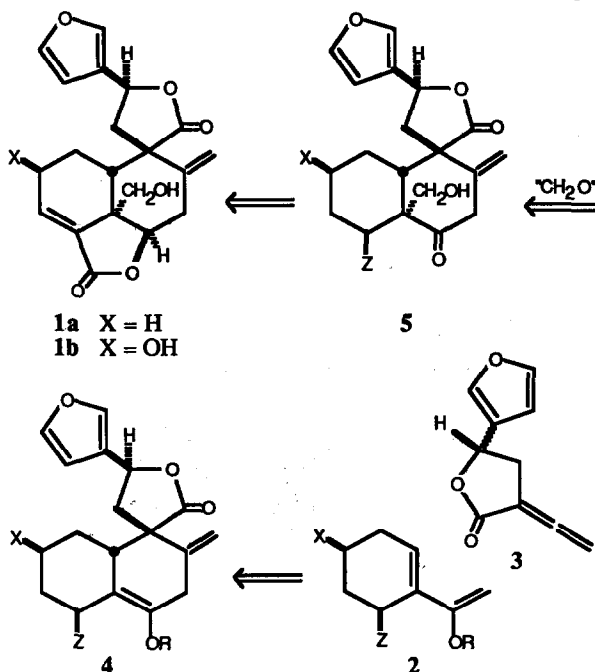
Michael E. Jung,*¹ Craig N. Zimmerman, Gregory T. Lowen, and Saeed I. Khan

Department of Chemistry and Biochemistry, University of California,

Los Angeles, California 90024

Summary: Diels-Alder cycloaddition of the allenic lactone **11** with 1-(1-(*t*-butyldimethylsilyloxy)vinyl)cyclohexene **6b** produces the desired cycloadduct **12** with good endo selectivity (4:1) and excellent diastereoselectivity.

Plaunol B and C, **1ab**, are diterpenes of the clerodane family² isolated from the stems of *Croton sublyratus* Kurz which were found to exhibit significant inhibitory activity against ulcers in Shay rats.³ This activity appears to result from the ability of the plaunols to depress gastric secretions.⁴ Several other structurally related diterpenes have also been isolated, including the insect antifeedants, the ajugarins.⁵ Because of their interesting structures and biological activity, these molecules have engendered a lot of work aimed at their synthesis.⁶ We proposed a synthetic approach



References and Notes

- 1) UCLA McCoy Award recipient, 1991-92; UCLA Hanson-Dow Teaching Award recipient, 1992.
- 2) Hanson, J. R. *Nat. Prod. Rep.* **1988**, *5*, 211.
- 3) a) Kitazawa, E.; Sato, A.; Takahashi, S.; Kuwano, H.; Ogiso, A. *Chem. Pharm. Bull.* **1980**, *28*, 227 and references therein. b) Ogiso, A.; Kitazawa, E.; Takahashi, S.; Kurabashi, M.; Sato, A.; Kuwano, H.; Tamura, C. *Tetrahedron Lett.* **1979**, 1117.
- 4) Shay, H.; Komorov, S. A.; Fels, S.; Meranze, D.; Gluentein, M.; Siple, H. *Gastroenterology* **1945**, *5*, 43.
- 5) Kubo, I.; Klocke, J. A.; Miura, I.; Fukuyama, Y. *J. Chem. Soc. Chem. Commun.* **1982**, 61.
- 6) a) Ley, S. V.; Simpkins, N. S.; Whittle, A. J. *J. Chem. Soc. Chem. Commun.* **1983**, 503. b) Jones, P. S.; Ley, S. V.; Simpkins, N. S.; Whittle, A. J. *Tetrahedron* **1986**, *42*, 6519. c) Ley, S. V. *Pestic. Sci. Biotechnol. Proc. Int. Congr. Pestic. Chem., 6th* **1986** (Pub. 1987), 25; *Chem. Abstr.* **1988**, *108*, 187002r. d) Kende, A. S.; Roth, B.; Kubo, I. *Tetrahedron Lett.* **1982**, *23*, 1751. e) Takahashi, S.; Kusumi, T.; Kakisawa, H. *Chem. Lett.* **1979**, 515. f) Bouchard, H.; Lallemand, J. Y. *Tetrahedron Lett.* **1990**, *31*, 5151.
- 7) a) Hansen, H. J.; Lang, R. W. *Helv. Chim. Acta* **1980**, *63*, 1204. b) Trippett, S.; Walker, D. M. *J. Chem. Soc.* **1959**, 3874. c) Hamlet, Z.; Barker, W. D. *Synthesis* **1970**, 543. d) Abell, A. D.; Trent, J. O.; Whittington, B. I. *J. Org. Chem.* **1989**, *54*, 2762.
- 8) The stereochemistry of the major isomer **9a** was unambiguously assigned by x-ray analysis of one of its hydrolysis products, the conjugated enone with a cis ring juncture.
- 9) All questions concerning the x-ray structures should be directed to Dr. Kahn.
- 10) a) Fotiadu, F.; Michel, F.; Buono, G. *Tetrahedron Lett.* **1990**, *31*, 4863. b) Fotiadu, F.; Archavlis, A.; Buono, G. *Tetrahedron Lett.* **1990**, *31*, 4859.
- 11) Jung, M. E.; Zimmerman, C. N. *J. Am. Chem. Soc.* **1991**, *113*, 7813.
- 12) The stereochemistry of the phenyl substituent in the minor isomer **13** is assumed based on steric arguments but has not been confirmed.

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