MONITOR synthesis

## Enantiospecific route to tirandamycin B

Tirandamycin B **XII** is a dienoyl tetramic acid antibiotic which was originally isolated from the culture broth of *Streptomyces flaveolus*. This compound possesses antimicrobial activity as well as inhibitory activity against bacterial DNA-directed RNA polymerase.

Shiratani, T. and coworkers [Chem. Commun. (1996) 21–22] describe the stereocontrolled enantiospecific synthesis of an enal intermediate from (S)-3-benzyloxy-2-methylpropanol using a highly regio- and stereo-selective methylation step.

# Total synthesis of angucyclinone antibiotics

Larsen, D.S., O'Shea, M.D. and Brooker, S. [*Chem. Commun.* (1996) 203–204] describe the first asymmetric synthesis of the angucyclinone antibiotics emycin A **XIII** and ochromycinone **XIV** from 5-hydroxy-1,4-naphthoquinone using a chiral Lewis acid derived from (*S*)-3,3'-diphenyl-1,1'-

XIV

binaphthalene-2,2'-diol to promote the kinetic resolution of a racemic diene in a Diels-Alder reaction.

### Regiospecific synthesis of acronycine

The acronycine alkaloid **XV** has been shown to have antineoplastic activity against C-1498 myelogenous leukemia, a tumour which is unresponsive to other antitumour agents. Anand, R.C. and Selvapalam, N. [Chem. Commun. (1996) 199–200] describe a the synthesis of acronycine and related alkaloids using a highly regiospecific prenylation of 3,5-dimethoxyacetanilide and cyclization of 2-[3,5-dimethoxy-2-(3-methylbut-2-enyl)]aminobenzoic acid under mild conditions.

#### Synthesis of L-734,217

ΧV

L-734,217 **XXI** is a nonpeptide, orally active fibrinogen receptor antagonist that is presently undergoing clinical trials as a potential inhibitor of vascular occlusion by thrombus formation associated with cerebral and cardiovascular diseases. Chung, J.Y.L. and coworkers [*J. Org. Chem.* (1996) 61, 215–222] report a six step synthesis of L-734,217 suitable for large-scale

production (Scheme 3). The synthesis involves the conversion of 2-piperidone **XVI** to ethyl (2-oxopiperidin-1-yl)acetate **XVII** followed by a novel chemoselective silyl-mediated conjugate addition of **XVII** to 4-vinylpyridine. The hydrolysis and kinetic resolution of product **XVIII** with quinine yielded **XIX** which was was coupled with benzyl 3-(*R*)-aminobutyrate in a biphasic system to give **XX**. Concomitant hydrogenation of the pyridine ring and debenzylation yielded L-734,217 **XXI** in 20% overall yield.

#### Nitric oxide sensor

Nitric oxide is now widely accepted as having a wide range of biological roles. However, the detection of nitric oxide, particularly in vivo is problematic. Leung, E. and coworkers [Chem. Commun. (1996) 23-24] report the development of novel in vivo nitric oxide sensor based on graphite-epoxy electrodes modified with N,N'-O-phenylenebis(salicylidineiminato)iron(III). The electrode has been successfully used by the group to monitor in vivo levels of nitric oxide in the muscle and liver of anaethetized rats which increased on perfusion with L-arginine, the biological precursor to nitric oxide and decreased on administration of L-ω-N-monomethyl arginine (L-MMA), anitric oxide synthase inhibitor. Such an electrode will have application in the investigation of the physiological role of nitric oxide in vivo and may also have utility in the screening of molecules that modulate the release of nitric oxide in vivo.