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## Selective Reduction of $\beta$ -Hydroxy Ketones

The reduction of  $\beta$ -hydroxy ketones to give the corresponding 1,3-*anti*-diols has been accomplished by internal hydride delivery. Reduction occurs intramolecularly as a result of the attenuated reactivity of the reducing agent (**Scheme 1**).

$$R_1$$
 $R_2$ 
 $(CH_3)_4NBH(OAc)_3$ 
 $R_1$ 
 $R_2$ 
 $R_2$ 

Scheme 1: Internal hydride delivery.

The synthesis of Oasomycin A by Evans utilised a reduction of a  $\delta$ -hydroxy- $\beta$ -ketoimide (**Scheme 2**). There are countless other examples of this methodology in synthesis.

Scheme 2: Internal hydride delivery in synthesis.

Access to the analogous 1,3-syn-diols has been achieved by external hydride delivery. Both chelating metal hydrides<sup>3</sup> and alkoxydialkylboranes in the presence of sodium borohydride<sup>4</sup> will perform this transformation *via* axial attack (**Scheme 3**).

$$\begin{array}{c|c} OH & O \\ \hline \vdots \\ R_1 \end{array} \xrightarrow{R_2} \begin{array}{c} Zn(BH_4)_2 \text{ or} \\ \hline Et_2BOMe/NaBH_4 \end{array} \xrightarrow{QH} \begin{array}{c} OH & OH \\ \hline \vdots \\ R_2 \end{array}$$

Scheme 3: External hydride delivery.

This methodology was utilised in the Sammakia synthesis of oxopolyene macrolide RK-397 (**Scheme 4**).<sup>5</sup>

Scheme 4: External hydride delivery in synthesis.

## References:

- 1. Tetrahedron Lett. 1986, 27, 5939-5942.
- 2. Angew. Chem. Int. Ed. 2007, 46, 541-544.
- 3. Tetrahedron Lett. 1983, 24, 5385-5386.
- 4. Tetrahedron Lett. 1987, 28, 155-158.
- 5. Angew. Chem. Int. Ed. 2007, 46, 1066-1070.