Johan FENNETEAU Doctorant 2<sup>eme</sup> année Pharmacognosie D5-4

# Total synthesis of (+)-Ambrucitin

(+)-Ambrucitin is an natural antifungal isolated from fermentation extract of *Polyangium cellulosum* in 1977.

This compound has shown activity against various fungal strains, the mechanism of action is based on perturbations of the osmoregulation system.

1) Find structures from A to K. ( see scheme below)

### Synthesis of first fragment :

## Synthsesis of the second fragment:

## Completion of the total synthesis:

### Cat\* I

Cat\* II

2) Find the transition state for the formation of the adduct A or D

- 3) Explain the mechanism of formation of G, What is the name of this reaction?
  - Hydroformylation of double bond, see scheme below :

- 4) Explain the mechanism of formation of K, What is the name of this reaction? Julia-Kociensky olefination
- 5) Explain the selectivity into the step 8?
  - Carbocupration of propargylacohol

OR Syn addition H 
$$Cu(SnBu_3)(CN)Li_2$$
  $Cu(SnBu_3)(CN)Li_2$   $SnBu_3$ 

- 6) What is the name of the reaction 10?
  - Kumada cross coupling