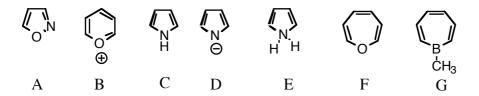
Exercises KJ 5220

Nov. 4, 2009

1. Indicate which of the heterocycles listed below can be formally regarded as aromatic, explain



2. A standard protocol for silylation of hydroxy groups is shown below. Explain the role of DMAP

$$ROH \xrightarrow{ClSiMe_2Bu^t} ROSiMe_2Bu^t$$

$$ROH \xrightarrow{CH_2Cl_2} ROSiMe_2Bu^t$$

3. Suggest a synthesis of the Ca-channel blocking drug Nifedipine

$$O_2N$$
 H_3CO_2C
 CO_2CH_3
 H_3C
 CH_3
 CH_3
 CH_3
 CH_3
 CH_3

4. Suggest mechanism for the reaction below.

5. $C_9H_8N_2O_2$ is formed when 2-methyl-5-nitropyridine is reacted with bromoacetone followed by NaHCO₃. Explain.

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(a) The structure of two antimalaria drugs are shown below. Discuss the possibilities of synthesizing these from dichloroquinolines

(b) Show how Mefloquine can be synthesized from 4-bromo-2,8-di(trifluoromethyl)quinoline.

(c) Suggest a synthesis of 4-bromo-2,8-di(trifluoromethyl)quinoline from 2-(trifluoromethyl)aniline.

7.

Explain th reaction below

8.

Suggest structures for the deuterated apigenines A and B

HO OH O
$$D_2O/Aceton$$
 A $\frac{1) D_3PO_4 \cdot BF_3 / D_2O}{2) H_2O}$ B $(C_{15}H_5D_5O_5)$ Apigenin

mefloquine J Med Chem 1971

apigenin Tetrahedron 2000, Pages 913-916