

Heterocycles In Biological Chemistry

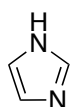
from chapter(s) _____ in the recommended text

A. Introduction

B. Names



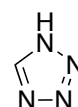
pyrrole



imidazole



pyrazole



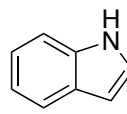
tetrazole



pyridine



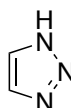
1,3-pyrimidine



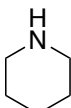
indole



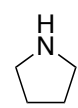
thiophene



1,2,3-triazole



piperidine



pyrrolidine



aziridine



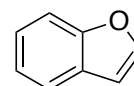
oxirane



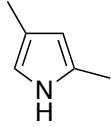
oxetane



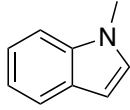
furan



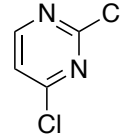
benzofuran



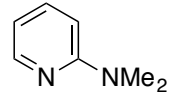
2,4-dimethylpyrrole



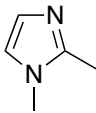
1-methylindole



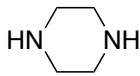
2,4-dichloropyrimidine



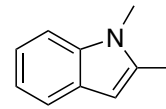
2-dimethylaminopyridine



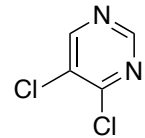
1,2-dimethylimidazole



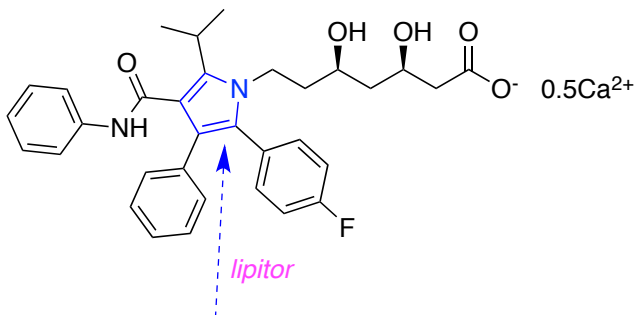
piperazine



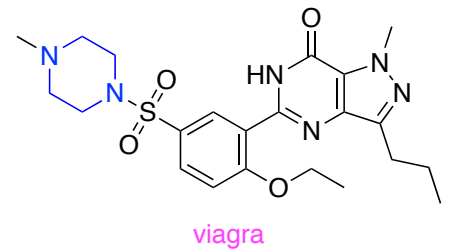
1,2-dimethylindole



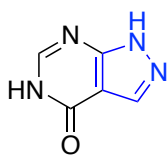
4,5-dichloropyrimidine



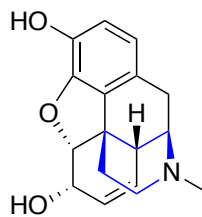
pyrrole



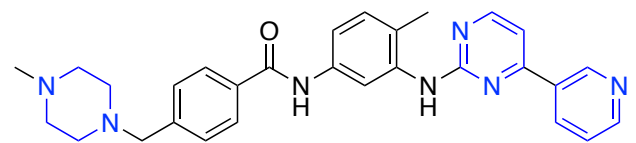
piperazine



allopurinol



morphine



piperazine and pyrimidine and pyridine

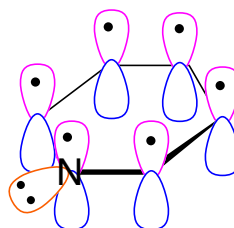
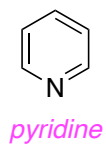
C. Aromaticity And Basicity Of Heterocycles

Pyridines And Pyrimidines

sp^2 hybridized with *a lone pair*

1 electron

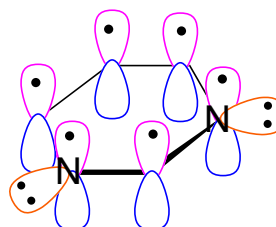
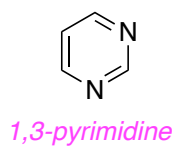
aromatic.



sp^2 hybridized with *a lone pair*

1 electron

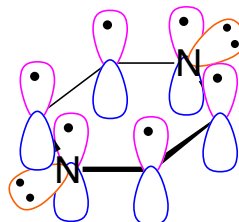
aromatic.



sp^2 hybridized with *a lone pair*

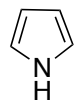
1 electron

aromatic.

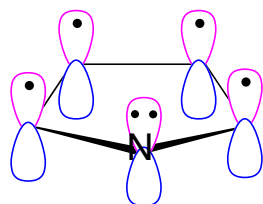


Pyrrole

sp^2 hybridized with 0
can
 aromatic.

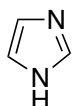


pyrrole

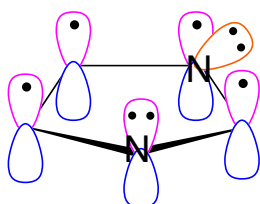


Imidazole

can
 are both sp^2 hybridized, and *one*
 is



imidazole



does influence

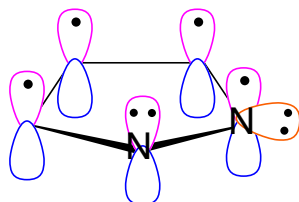
Pyrazole

cannot
 are
 one

Pyrazole *is*



pyrazole



aromatic stabilization.

1,3,4-Oxadiazole

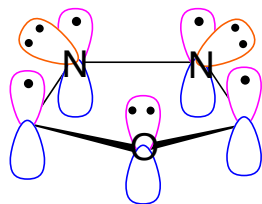
sp^2 hybridized and each contributes 1

sp^2 hybridized and contributes 2

aromatic.

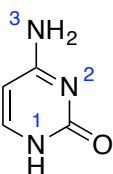


1,3,4-oxadiazole



does not
good base
is not lost.

Heterocycles In Nature

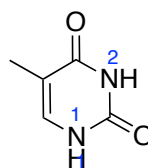


cytosine

N^3 : 0

N^2 : 1

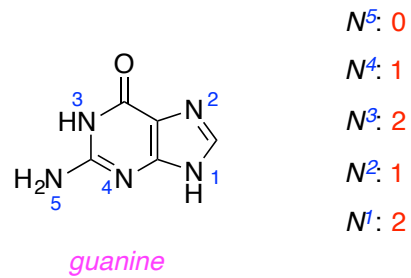
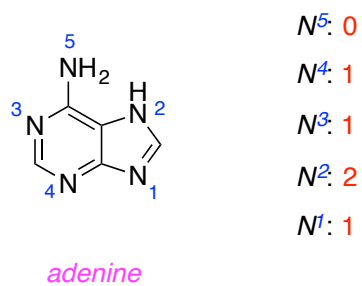
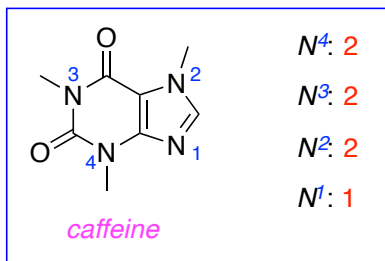
N^1 : 2



thymine

N^2 : 2

N^1 : 2

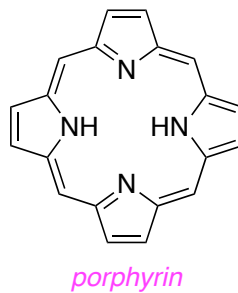


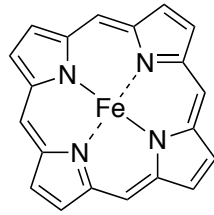
2 pyridine-like nitrogen atoms, 2

26 π -electrons

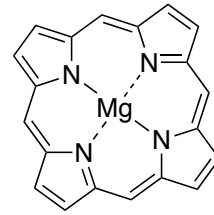
are aromatic.

2





Fe²⁺ complex overall charge 0



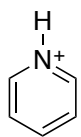
Mg²⁺ complex overall charge 0

Hemoglobin

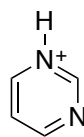
chlorophyll

): strongly UV absorbing / fluorescent / capable of redox chemistry.

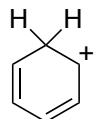
Aromatic Characteristics Of Protonated Heterocycles



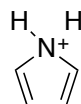
aromatic because it has
6 πe^- .



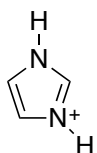
aromatic because it has
6 πe^- .



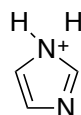
not aromatic because it has
4 πe^- .



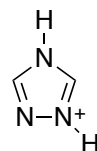
not aromatic because it has
4 πe^- .



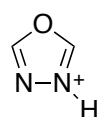
aromatic because it has
6 πe^- .



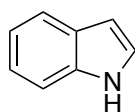
not aromatic because it has
4 πe^- .



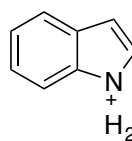
aromatic because it has
6 πe^- .



aromatic because it has
6 πe^- .

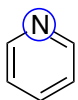


aromatic because it has
10 πe^- .

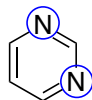


not aromatic because it has
8 πe^- .

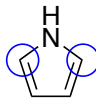
C^3



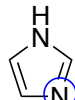
pyridine



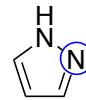
pyrimidine



pyrrole



imidazole



pyrrazole



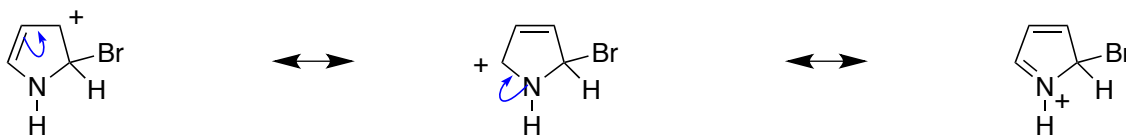
oxazole

D. Electrophilic Attack On Pyrrole And Indole Compared

Pyrrole

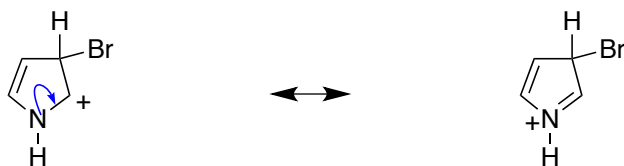
low

in the 2-position



complete diagrams and show arrows

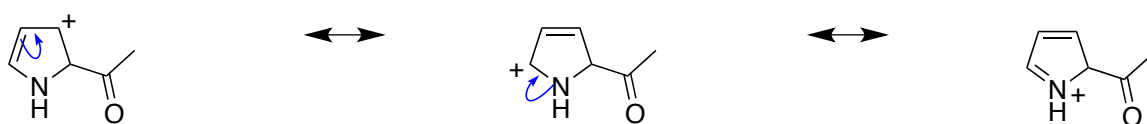
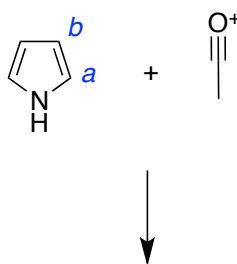
in the 3 position



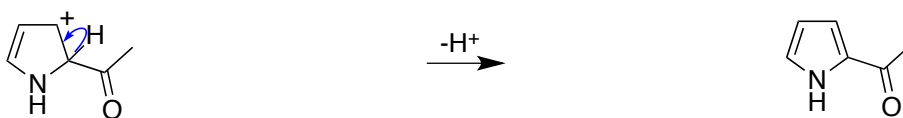
complete diagrams and show arrows

*2-position
thermodynamic*

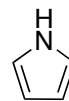
Hammond's postulate.



choose correct regiochemistry, show resonance structures, and electron flow that relates them using curly arrows



more electron rich than benzene, hence it reacts faster

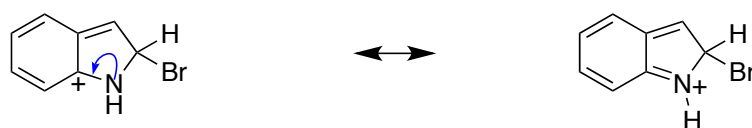
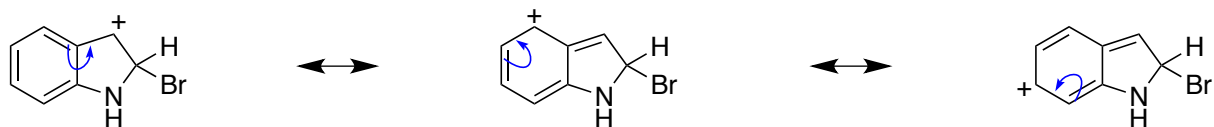


least reactive

most reactive

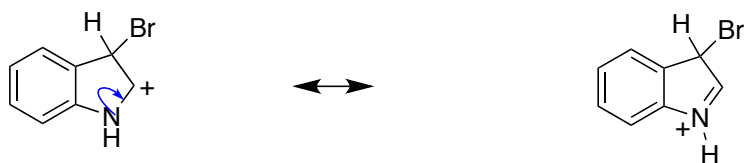
Indole

in the 2-position



*donation of the N-lone pair
does
disrupt aromaticity of the
benzene ring*

in the 3 position



*donation of the N-lone pair
need not
disrupt aromaticity of the
benzene ring*

3-position

because the positive charge can be delocalized onto the nitrogen without disrupting the aromaticity of the benzene, whereas for attack at the 2-position the aromaticity of the benzene must always be disrupted.