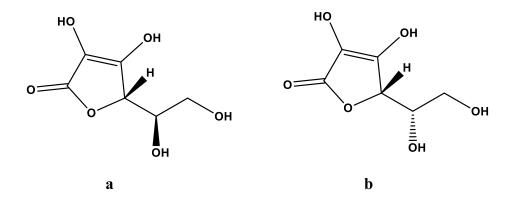
Vitamin C/ L-Ascorbic Acid

is a *cofactor* in several *hydroxylation* and *amidation* reactions by functioning as a *reducing agent*. As a result, vitamin C plays an important role in the *synthesis of collagen*, *carnitine*, *folinic acid*, *and norepinephrine*. It also influences the processing of hormones such as *oxytocin*, *antidiuretic hormone*, *and cholecystokinin*. Vitamin C *reduces* iron from the *ferric to the ferrous state* in the stomach, thereby *increasing* intestinal absorption of iron. Vitamin C may be involved in *steroidogenesis* in the adrenals. *Vitamin C also has a pro-oxidant effect in vivo which may occur in the setting of overdose*.

Some of Ascorbate-Dependent Enzymes

Enzyme	Its role
Procollagen-proline dioxygenase	Collagen synthesis
Procollagen-lysine 5-dioxygenase	Collagen synthesis
Procollagen-proline 3-dioxygenase	Collagen synthesis
Phytanoyl-CoA1 dioxygenase	Fatty acid metabolism
Dopamine beta-monooxygenase	Norepinephrine synthesis
Peptidylglycine monooxygenase	Peptide hormone synthesis

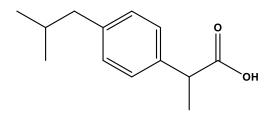


Structure of L-Ascorbic Acid (a) and D-Ascorbic Acid (b).

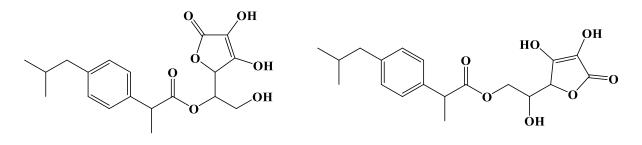
ASCORBIC ACID IN DRUG DESIGN

The intake of vitamin C (AA) in the body compartments of mammals is guaranteed by a class of Na⁺ dependent transporters recently characterized and called SVCT. In particular, SVCT₁ allows the absorption of vitamin C from the intestine and its recovery by the kidneys, while SVCT₂ allows accumulation of the vitamin C in the brain and eye, so that recent researches revealed that *AA can be used as tool to improve brain drug delivery*.

In order to achieve the satisfactory therapeutic effect, ibuprofen in the CNS needs to reach a higher concentration. However, because of the poor permeability of NSAIDs, the larger doses are required to achieve the desired therapeutic effect. This will cause a lot of gastrointestinal adverse effects and some toxicity, and may cause some damage to the body; so that conjugation of ibuprofen with ascorbic acid may resolve this problem.



Structure of Ibuprofen



Structures of ibuprofen prodrugs with ascorbic acid

- **1.** Of which type these prodrugs are?
- 2. Draw the mechanism for synthesis of these prodrugs.