Waters of Application of Application

LAH 0134 10/83 AN/LS,ES/MD/TX/OT

SEPARATION OF PYRIDINIUM ALDOXIMES ON BARE SILICA USING REVERSED-PHASE ELUENTS

In an earlier Lab Highlight, the novel use of silica in reversed-phase eluents was presented (1,2). This recent report is an application of that approach.

Pyridinium aldoximes are a class of compounds which are cholinesterase reactivators. Because of this property of reactivating the enzyme cholinesterase, one of the compounds, pyralidoxime chloride (Structure 1 in Table I), is used as an antidote for nerve gas by removing the phosphoryl groups from the enzyme. The structures of the compounds studied are shown in Table I.

The compounds have been separated on silica based ion-exchange columns (e.g. a Radial-PAKTM SCX type material). However, it is generally accepted that these ion-exchangers are not "rugged" and often require special attention to insure long life.

Another approach is to separate these compounds on bare silica using a typical reverse phase eluent. This was done by Kientz, Verweij and Boter of Prins Mauritis Laboratory TNO in the Netherlands (3) and is shown in Figure 1. Clearly a good separation is available with all of the benefits of using silica (1). They also did a similar separation on a silica based ion exchange packing.

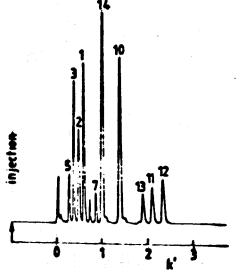


FIGURE 1: Separation of pyridinium aldoximes and P_2 -amide on bare silica 5 μ m. Solvent - MeOH (80%): H_2O (20%), 0.5 \underline{M} potassium formate, pH 6.3. UV at 290 nm.

| | I (mono-quaternary) | | | | | | |
|------|---------------------------|-------------------------------|----------------|---------------------|---------------------|-----------------|-----|
| Code | Compound | R | R ₂ | R ₃ | R4 | τ - | |
| 1 | P ₂ s | CH ₃ | CH=NOH | H | H | CH 3503 | |
| 2 | ethyl-P ₂ A | с ₂ н ₅ | CK=NOH | H | H | C1 | |
| 3 | n-propyl-P ₂ A | С ₃ н ₇ | CH=NOH | H | Ħ | cı_ | |
| 4 | n-pentyl-P ₂ A | C5H11 | CH=NOH | H | H | cı ⁻ | |
| 5 | benzyl-P ₂ A | C6H5CH2 | CH=NOH | H | H | CH3503 | |
| 6 | n-heptyl-P2A | C7H15 | CH-NOH | H | H | C1 | |
| 7 | P ₂ -amide | CH ₃ | C(0)NH2 | H | H | C1 | |
| 8 | P ₃ AM | CH ₃ | H | CH=NOH | H | C1 ⁻ | |
| 9 | P ₄ AM | CH ₃ | H | H | CH=NOH | Cl | |
| | | 1 | I (bis-q | uaternar | y) | | |
| | | R | R 2 · | R ₃ | R ₄ | R ₅ | x |
| 10 | TNB-4 | Ħ | CH=NOH | CH=NOH | Н | H | CH, |
| 11 | HI-6 | CH=NOH | H | C(0)NH ₂ | H | H | 0 |
| 12 | HS-6 | CH=NOH | H | H | C(0)NH ₂ | H | 0 |
| 13 | 2,2-LüH-6 | CH=NOH | H | H | H | CH=NOH | 0 |
| 14 | Toxogonin | H | CH-NOH | CH=NOH | H | H | 0 |

^{1.} LH0095, 0096

^{2.} Bidlingmeyer, B. A., Del Rios, J. K. and Korpi, J. Anal. Chem <u>54</u>, 442 (1982).

^{3.} Drs. Kientz, Verweij and Boter (of Prins Mauritis Laboratory TNO), The Netherlands, presented at 7th Symposium of LC at Baden-Baden, 4 May, 1983.