

**New England
Medical Center Hospitals**

1796 Boston Dispensary
1894 Floating Hospital for Infants and Children
1938 Pratt Diagnostic Clinic
1948 New England Center Hospital
1958 Rehabilitation Institute

The principal teaching
hospitals for Tufts University
School of Medicine

750 Washington Street
Boston, Massachusetts 02111
Telephone: (617) 956-5000

February 29, 1988.

Medical Director
Blood Products Laboratory
Lestree Herts.
United Kingdom WD6 3BX

Dear

We are writing in regard to methods of preparation of Factor VIII concentrates. As you may be aware, several pharmaceutical companies here in the United States have recently introduced methods of preparation of Factor VIII which entail the use of several potentially toxic materials. Specifically, TPNB (Tri-n-butylphosphate) has been coupled with the detergent Triton X-100 by one manufacturer, while another has introduced sodium cholate in their process, and a third the use of N-heptane. Despite disclaimer made, it would be all but impossible to remove all traces of these materials from the Factor VIII preparations, and would inevitably amount to a potential problem over the lifetime of patients receiving it on a regular basis.

These methods you know to be in contrast to your own process of heating of lyophilized Factor VIII to temperatures as high as 70 to 80 degrees Centigrade, without recourse to those potentially toxic additives. Accordingly, we would kindly request that the Food and Drug Administration (FDA) of the United States approve your process. In fact, 80 degrees for a full 72 hours may be more than is absolutely required; 80 degrees for some ten to twenty hours would likely, we believe, prove adequate for inactivation of all virus.

The persons to contact at the FDA for approval and distribution of your method in this country are:

and/or

FDA
National Institutes of Health (all at same address)
Building 29 Rm. 302
9000 Rockville Pike
Bethesda, MD 20892 USA

or call, to expedite matters, at (301) 496-4833

We'd like to facilitate or help to get FDA approval in any way

possible. Such approval should present no problem; a previous method of heating to 60 degrees has already been approved. Request for approval of your method, must, of course, come from you.

We do believe your Factor VIII to be the safest concentrate available. It simply avoids the addition of any toxins and should be the product of choice available to all patients.

Hoping for your action in this very important matter, we are,

Sincerely yours,

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