## SECOND EDITION

## Chemical Stability of Pharmaceuticals

A HANDBOOK FOR PHARMACISTS

Kenneth A. Connors Gordon L. Amidon Valentino J. Stella



NOT THE REAL PROPERTY AND VALUE OF THE PROPERTY AND ANY AND

Copyright © 1986 by John Wiley & Sons, Inc.
All rights reserved. Published simultaneously in Canada.

No part of this publication may be reproduced, stored in a retrieval system or transmitted in any form or by any means, electronic, mechanical, photocopying, recording, scanning or otherwise, except as permitted under Sections 107 or 108 of the 1976 United States Copyright Act, without either the prior written permission of the Publisher, or authorization through payment of the appropriate per-copy fee to the Copyright Clearance Center, 222 Rosewood Drive, Danvers, MA 01923, (978) 750-8400, fax (978) 750-4470. Requests to the Publisher for permission should be addressed to the Permissions Department, John Wiley & Sons, Inc., 111 River Street, Hoboken, NJ 07030, (201) 748-6011, fax (201) 748-6008.

Library of Congress Cataloging in Publication Data:

Connors, Kenneth A. (Kenneth Antonio), 1932-Chemical stability of pharmaceuticals.

"A Wiley-Interscience publication."
Includes bibliographies and index.

1. Drug stability. I. Amidon, Gordon L. II. Stella,
Valentino J., 1946— . III. Title. [DNLM: 1. Drug
Stability—handbooks. 2. Kinetics—handbooks.

QV 735 C752c]

RS424.C66 1986 615'.18 85-31455 ISBN 978-0-471-87955-8



## 5-Azacytidine

GENERAL

Names

5-AC; 5-azacytidine; 4-amino-1-b-D-ribofuranosyl-1,3,5-triazin-2-one.

Structure

Stability Summary

5-Azacytidine (5-AC) undergoes hydrolysis of the triazine ring in aqueous buffers. The hydrolysis takes place at the 5-6 double bond and the pH-rate profile exhibits both acid and base catalysis. The pH range of optimum stability is 6.5-7.0. Even at this pH range, however, drug degradation is rapid. Decomposition rate constants of 5-AC are also influenced by temperature and buffer concentration. It is possible that metal ions, for example, iron, act as catalysts in the hydrolysis so that the addition of EDTA to solutions of 5-AC in distilled water increases the stability.

