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MICHIGAN STATE UNIVERSITY

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THE SYNTHESIS OF DERIVATIVES OF
3-AMINO-4-H,1,2,4-TRIAZOLE

BY

Emerson A. Cooper

A THESIS

Submitted to the School for Advanced Graduate Studies of
Michigan State University of Agriculture and
Applied Science in partial fulfillment of
the requirements for the degree of

DOCTOR OF PHILOSOPHY

Department of Chemistry

1959

the following conditions are satisfied:

$$\begin{aligned} & \text{for } i = 1, 2, \dots, n-1 \\ & \quad \text{if } \alpha_i = 1, \text{ then } \alpha_{i+1} = 0; \end{aligned}$$

and

$$\alpha_n = 1, \quad \alpha_1 = 0.$$

where

the sequence $\{\alpha_i\}_{i=1}^n$ is defined by the recurrence relation

$$\alpha_{i+1} = \alpha_i + \alpha_{i-1}, \quad \alpha_1 = 0, \quad \alpha_2 = 1,$$

and the sequence $\{\beta_i\}_{i=1}^n$ is defined by the recurrence relation

$$\beta_{i+1} = \beta_i + \beta_{i-1}, \quad \beta_1 = 1, \quad \beta_2 = 0.$$

where α_i and β_i are defined by

$$\alpha_i = \frac{1}{2} (1 + (-1)^i), \quad \beta_i = \frac{1}{2} (1 - (-1)^i).$$

where

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9/27/62

To Marjorie, Roslyn and Stephen

1. The first part of the document is a letter from the author to the reader, explaining the purpose of the study and the methods used.

A C K N O W L E D G M E N T

I am indebted to Dr. Robert M. Herbst for his generous and invaluable counsel. His sage suggestions in the course of this investigation were indeed a source of much encouragement and inspiration. I am also thankful for the support given me by Oakwood College and the Danforth Foundation.

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AN ABSTRACT

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Department of Chemistry

Year 1959

Approved Robert M. Hecht

THE UNIVERSITY OF CHICAGO

PHYSICS DEPARTMENT

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It has been observed that rearrangements sometimes occur in certain heterocyclic systems containing the N-C-N arrangement where one of the nitrogens is an amino group. Dimroth (1) has described the thermal rearrangement of a number of 1-phenyl-5-amino-, and 1-phenyl-5-methylamino-1,2,3-triazoles to 5-phenylamino-, and 1-methyl-5-phenylamino-1,2,3-triazoles. Lieber et al (2,3) have investigated the rearrangement of 5-amino-1,2,3-triazoles. The rearrangement of 5-alkylaminotetrazoles to 1-alkyl-5-aminotetrazoles has also been studied (4).

The original purpose of this investigation was to determine whether 3-amino-4-phenyl-1,2,4-triazole and its homologs would undergo a similar rearrangement. After preliminary experimentation indicated the absence of such rearrangement in the 1,2,4-triazole series, attention was then directed to the synthesis of three different series of compounds having varied substituents in the fifth position of 3-amino-4-phenyl-4-H,1,2,4-triazole.

3-Amino-4-H,1,2,4-triazole is presently of considerable interest because of its remarkable effect on the physiological processes of plants and animals. It is able to inhibit the synthesis of chlorophylls and carotenoids in plants (5); it reduces the level of hepatic and renal catalase activity in rats thus producing an

effect similar to that observed in animals having malignant growth (6); and it is also capable of lowering the activity of δ -aminolevulinic acid dehydrase, an enzyme that catalyzes the conversion of δ -aminolevulinic acid to porphobilinogen (6). The last effect is also produced by tumors.

Since studies relating phytotoxicity to chemical structure indicate the importance of chloro, methyl, methoxy and phenoxy groups on herbicides (7), it was decided to prepare a representative group of substituted phenoxymethyltriazoles and some thiophenoxymethyltriazoles.

A number of dialkylaminomethyltriazoles was also prepared since the presence of two different heterocyclic nuclei in some, and the existence of two basic centers in all could possibly produce anesthetic action as well as some other useful physiological effect. It is of interest to note that 3-ethyl-4-cyclohexyl-1,2,4-triazole is similar to adrenaline in some of its actions on the body. It has been found to stimulate the heart and respiratory center (8).

The starting material for these syntheses, phenylaminoguanidinium bisulfate, was first prepared in the course of this work. This compound was allowed to react with a number of aliphatic acids (formic, acetic, glycolic, lactic and phenoxyacetic acid) to form the corresponding 3-amino-4-phenyl-5-alkyl-1,2,4-triazoles.

A procedure was perfected for the conversion of 3-amino-4-phenyl-5-hydroxymethyl-1,2,4-triazole to the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in good yield.

The 5-dialkylaminomethyltriazoles were prepared by allowing the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole to react with an excess of each of the following amines: piperidine, morpholine, pyrrolidine, dimethylamine, diethylamine, diallylamine, di-n-propylamine, di-isopropylamine, di-n-butylamine and di-isobutylamine.

The 5-phenoxymethyl-, and the 5-thiophenoxy-methyltriazoles were prepared by allowing the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole to react with an excess of the sodium salt of each of the following phenols and thiophenols: phenol, p-bromophenol, p-chlorophenol, p-methoxyphenol, p-cresol, o-cresol, m-cresol, thiophenol and p-chlorothiophenol.

Phenylthiourea derivatives of most of these new aminotriazoles were prepared. In a few cases, a number of the hydrochlorides and acetyl derivatives were prepared.

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References

- (1) Dimroth, Ann. Chem., 364, 183 (1909)
- (2) Lieber, E., Chao, T.S. and Rao, C.N.R., J. Org. Chem., 22, 654 (1957)
- (3) Lieber, E., Rao, C.N.R. and Chao, T. S., J. Am. Chem. Soc., 79, 5962 (1957)
- (4) Garbrecht, W.L. and Herbst, R.M., J. Org. Chem., 18, 1269 (1953)
- (5) Miller, C.S. and Hall, W.C., Weeds, 5, 304 (1957)
- (6) Tschudy, D.P. and Collins, A., Science, 126, 168 (1957)
- (7) Shaw, W.A. and Swanson, C.R., Weeds, 2, 43 (1953)
- (8) Lewenstein, M.J., U. S. Pat., 2,683,106 (1954)

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the 1990s, the number of people in the world who are undernourished has declined from 1.1 billion to 800 million. The number of people who are malnourished has declined from 1.5 billion to 1 billion. The number of people who are obese has increased from 100 million to 300 million. The number of people who are overweight has increased from 100 million to 300 million. The number of people who are obese and overweight has increased from 100 million to 300 million. The number of people who are obese and overweight has increased from 100 million to 300 million.

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the 1990s, the number of people in the world who are under 15 years of age is expected to increase from 1.1 billion to 1.5 billion. The number of people aged 65 and over is expected to increase from 200 million to 400 million. The number of people aged 15 and over is expected to increase from 3.5 billion to 4.5 billion. The number of people aged 15 and over is expected to increase from 3.5 billion to 4.5 billion. The number of people aged 15 and over is expected to increase from 3.5 billion to 4.5 billion.

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the 1990s, the number of people in the United States who are 65 years of age or older is projected to increase from 20 million to 30 million, and the number of people 75 years of age or older is projected to increase from 10 million to 15 million (U.S. Census Bureau, 1996). The number of people 85 years of age or older is projected to increase from 2 million to 4 million (U.S. Census Bureau, 1996). The number of people 90 years of age or older is projected to increase from 500,000 to 1 million (U.S. Census Bureau, 1996). The number of people 95 years of age or older is projected to increase from 100,000 to 200,000 (U.S. Census Bureau, 1996). The number of people 100 years of age or older is projected to increase from 10,000 to 20,000 (U.S. Census Bureau, 1996).

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Figure 1. Schematic diagram of the experimental setup.

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3-Amino-4-phenyl-5-di-n-propylaminomethyl-4-H,1,2,4-triazole	42
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1. The first part of the document discusses the importance of maintaining accurate records of all transactions and activities. It emphasizes that proper record-keeping is essential for transparency and accountability, particularly in financial matters. The text suggests that organizations should implement robust systems to track every detail, from small expenses to major investments.

2. The second section focuses on the role of technology in modern record-keeping. It highlights how digital tools can streamline the process, reduce errors, and provide real-time access to data. The author argues that embracing technology is not just a convenience but a necessity for staying competitive in today's fast-paced environment.

3. The third part of the document addresses the challenges of data security and privacy. It notes that as more information is stored digitally, the risk of breaches and unauthorized access increases. The text provides several recommendations for safeguarding sensitive data, including regular security audits, employee training, and the use of encryption techniques.

4. The fourth section discusses the importance of regular reviews and updates to record-keeping policies. It points out that what works today may not work tomorrow due to changing regulations and technological advancements. The author encourages organizations to conduct periodic assessments to ensure their systems remain effective and compliant.

5. The fifth part of the document explores the benefits of a well-maintained record-keeping system. It lists several advantages, such as improved decision-making, better risk management, and enhanced operational efficiency. The text also mentions that accurate records can be invaluable during audits and legal proceedings.

6. The sixth section provides a brief overview of common record-keeping practices across different industries. It compares various methods and tools used by businesses, government agencies, and non-profit organizations. This comparative analysis helps readers understand the best practices and adapt them to their own specific needs.

7. The seventh part of the document offers practical tips for implementing a successful record-keeping system. It covers topics such as setting clear goals, assigning responsibilities, and establishing a consistent workflow. The author stresses that success in record-keeping requires a commitment to discipline and attention to detail.

8. The eighth section discusses the future of record-keeping. It looks at emerging trends like cloud storage, artificial intelligence, and blockchain technology. The text speculates on how these innovations might further transform the way we manage and store information, potentially leading to even greater efficiency and security.

9. The ninth part of the document provides a summary of the key points discussed throughout the text. It reiterates the importance of accurate record-keeping and the need for continuous improvement. The author concludes by encouraging readers to take action and implement the strategies outlined in the document.

10. The final section of the document includes a list of references and resources for further reading. It cites several books, articles, and online resources that provide additional information on record-keeping and related topics. This section serves as a valuable guide for anyone looking to delve deeper into the subject.

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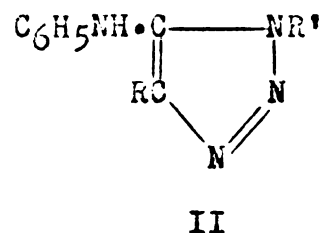
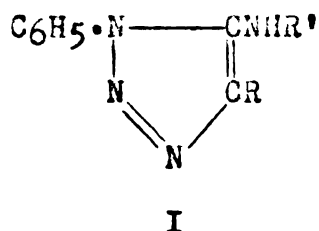
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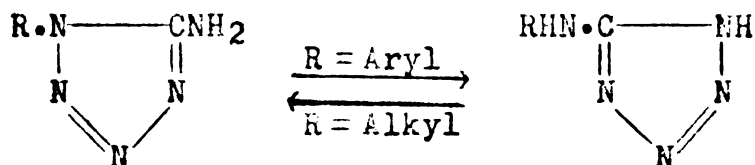
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Introduction

It has been observed that rearrangements sometimes occur in certain heterocyclic systems containing the N-C-N arrangement where one of the nitrogens is an amino group. Dimroth (1) has described the thermal rearrangement of a number of 1-phenyl-5-amino-, and 1-phenyl-5-methylamino-1,2,3-triazoles (I) to 5-phenylamino-, and 1-methyl-5-phenylamino-1,2,3-triazoles (II).



Lieber et al (2,3) have investigated the rearrangement of 5-amino-1,2,3-triazoles. The rearrangement of 5-alkylaminotetrazoles to 1-alkyl-5-aminotetrazoles has also been studied (4).



The original purpose of this investigation was to determine whether 3-amino-4-phenyl-1,2,4-triazole and its homologs would undergo a similar rearrangement. After preliminary experimentation indicated the absence of such rearrangement in the 1,2,4-triazole series, attention was then directed to the synthesis of three different series

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of compounds having varied substituents in the fifth position of 3-amino-4-phenyl-4-H,1,2,4-triazole.

3-Amino-4-H,1,2,4-triazole is presently of considerable interest because of its remarkable effect on the physiological processes of plants and animals. It is able to inhibit the synthesis of chlorophylls and carotenoids in plants (5); it reduces the level of hepatic and renal catalase activity in rats thus producing an effect similar to that observed in animals having malignant growth (6); and it is also capable of lowering the activity of δ -aminolevulinic acid dehydrase, an enzyme that catalyzes the conversion of δ -aminolevulinic acid to porphobilinogen (6). The last effect is also produced by tumors.

Since studies relating phytotoxicity to chemical structure indicate the importance of chloro, methyl, methoxy and phenoxy groups in herbicides (7), a representative group of substituted phenoxymethyltriazoles was prepared.

A number of dialkylaminomethyltriazoles was also prepared since the presence of two different heterocyclic nuclei in some, and the existence of two basic centers in all could possibly produce anesthetic action as well as some other useful physiological effect. It is of interest to note that 3-ethyl-4-cyclohexyl-1,2,4-triazole is similar to adrenaline in some of its actions on the body. It has been found to stimulate the heart and respiratory center (8).

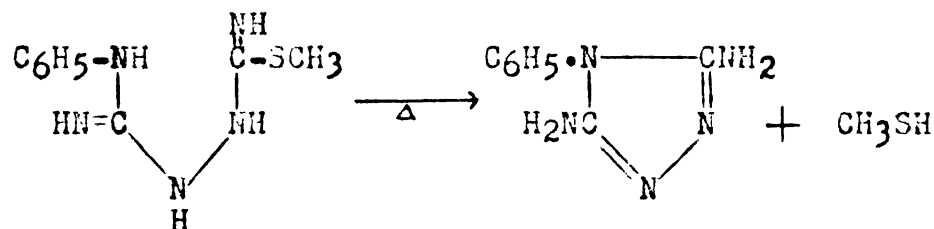
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Historical

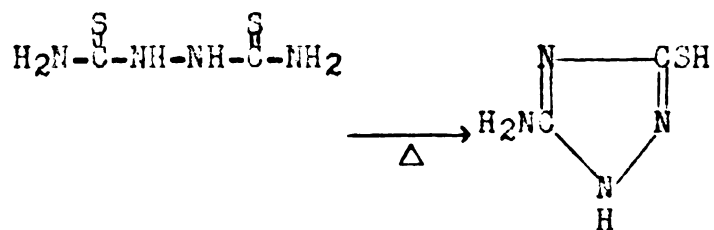
The first derivatives of 1,2,4-triazole were prepared by Bladin (9) in 1885. Six years later, in 1891, Bladin (10) and Andreocci (11), working independently, synthesized the parent compound, 1,2,4-triazole. In the same year, Thiele (12) discovered the first general method of preparing derivatives of 3-amino-1,2,4-triazole. He found that aminoguanidine reacts with acetic acid to form 3-amino-5-methyl-1,2,4-triazole. Later, Thiele and Manchot (13) showed the general nature of this reaction by preparing 3-amino-1,2,4-triazole from aminoguanidine and formic acid; and 3-amino-1,2,4-triazole-5-carboxylic acid from aminoguanidine and oxalic acid. This method has been termed the dehydration of acyl derivatives of aminoguanidines.

Some other methods that have been employed in the synthesis of derivatives of 3-amino-1,2,4-triazole are summarized below:

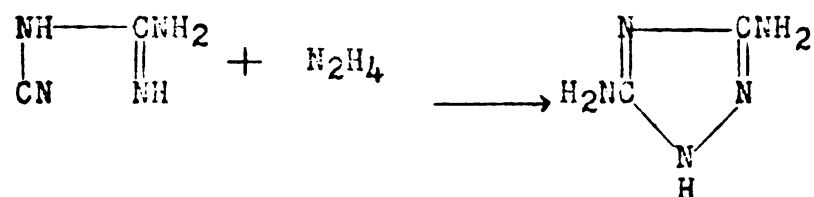
(1) The elimination of methylmercaptan from an S-methyl ether of phenylguanyllthiosemicarbazide (14):



(2) The fusion of a diamide of hydrazine-N:N'-dithiodicarboxylic acid (15):



(3) Reaction of dicyandiamide with hydrazine
(16):



A number of reviews on the chemistry of 1,2,4-triazoles are available (17,18,19,20).

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1. The first step is to identify the problem.

2. The second step is to define the problem.

3. The third step is to analyze the problem.

4. The fourth step is to plan the solution.

5. The fifth step is to implement the solution.

6. The sixth step is to evaluate the solution.

7. The seventh step is to monitor the solution.

8. The eighth step is to report the solution.

9. The ninth step is to conclude the solution.

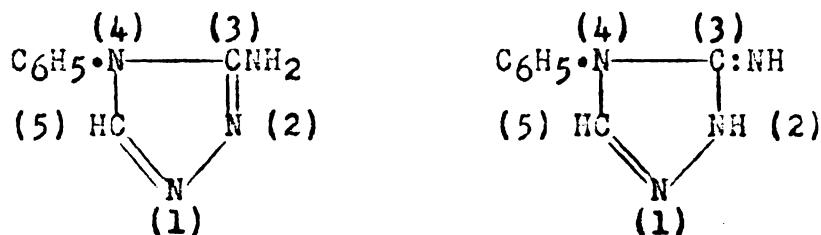
10. The tenth step is to reflect on the solution.

11. The eleventh step is to document the solution.

Discussion

Nomenclature and Properties

The nomenclature used in the discussion of derivatives of 3-amino-4-phenyl-4-H,1,2,4-triazole in this thesis, is based upon the following formulas for this compound:

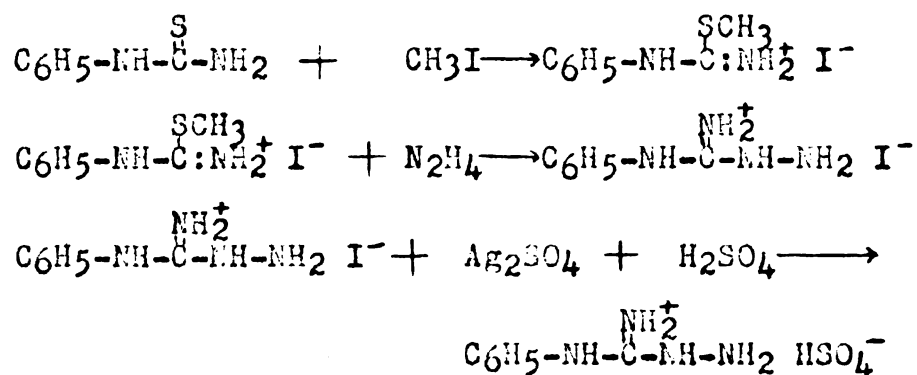


A survey of the literature shows that amino-triazoles exhibit many of the properties of aromatic amines. However, they also react both as weak bases and weak acids as is shown by the formation of hydrochlorides, nitrates, and picrates on the one hand, and of metal salts on the other. They are quite stable toward oxidizing agents, forming only azo derivatives. They diazotize to form diazonium salts which can couple with phenols and amines. These diazonium salts can be reduced to give C-hydrazino-1,2,4-triazoles and can react with hydrogen halides to form C-halogeno-1,2,4-triazoles (21). C-methyl groups can be oxidized to carboxyl groups, and N-phenyl groups can be oxidatively removed after nitration and reduction. The amino group can also react with aldehydes to form condensation products (22).

[illegible]

Phenylaminoguanidinium Bisulfate

Phenylaminoguanidinium bisulfate was an important intermediate in the synthesis of various derivatives of 3-amino-4-phenyl-1,2,4-triazole. It was prepared from S-methyl phenylisothiourea hydriodide by the method used by Kirsten and Smith (23) to prepare α -alkyl- α -aminoguanidines. Kirsten and Smith were unsuccessful in obtaining phenylaminoguanidine, but Finnegan *et al* (24) have prepared the hydriodide. The bisulfate was first prepared in the course of this investigation since the hydriodide was found to be unsuitable for the synthesis of 3-amino-1,2,4-triazoles. The following sequence of reactions illustrates the method:



The method of Finnegan, Henry and Lieber (24) was employed in preparing the S-methyl phenylisothiourea hydriodide. An alcoholic solution of the isothiourea was obtained by treating a slurry of phenylthiourea in alcohol at 5° C. with methyl iodide and then permitting the reaction to go to completion at room temperature. The second reaction was carried out without isolating the isothiourea.

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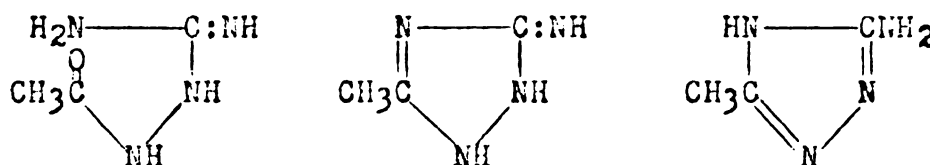
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In this reaction, the S-methyl phenylisothiourea reacts with hydrazine to form phenylaminoguanidinium iodide and methyl mercaptan. In the final reaction, the phenylaminoguanidinium iodide is converted to the bisulfate by a metathetical reaction with silver sulfate in the presence of sulfuric acid.

3-Amino-4-phenyl-5-alkyl-1,2,4-triazoles

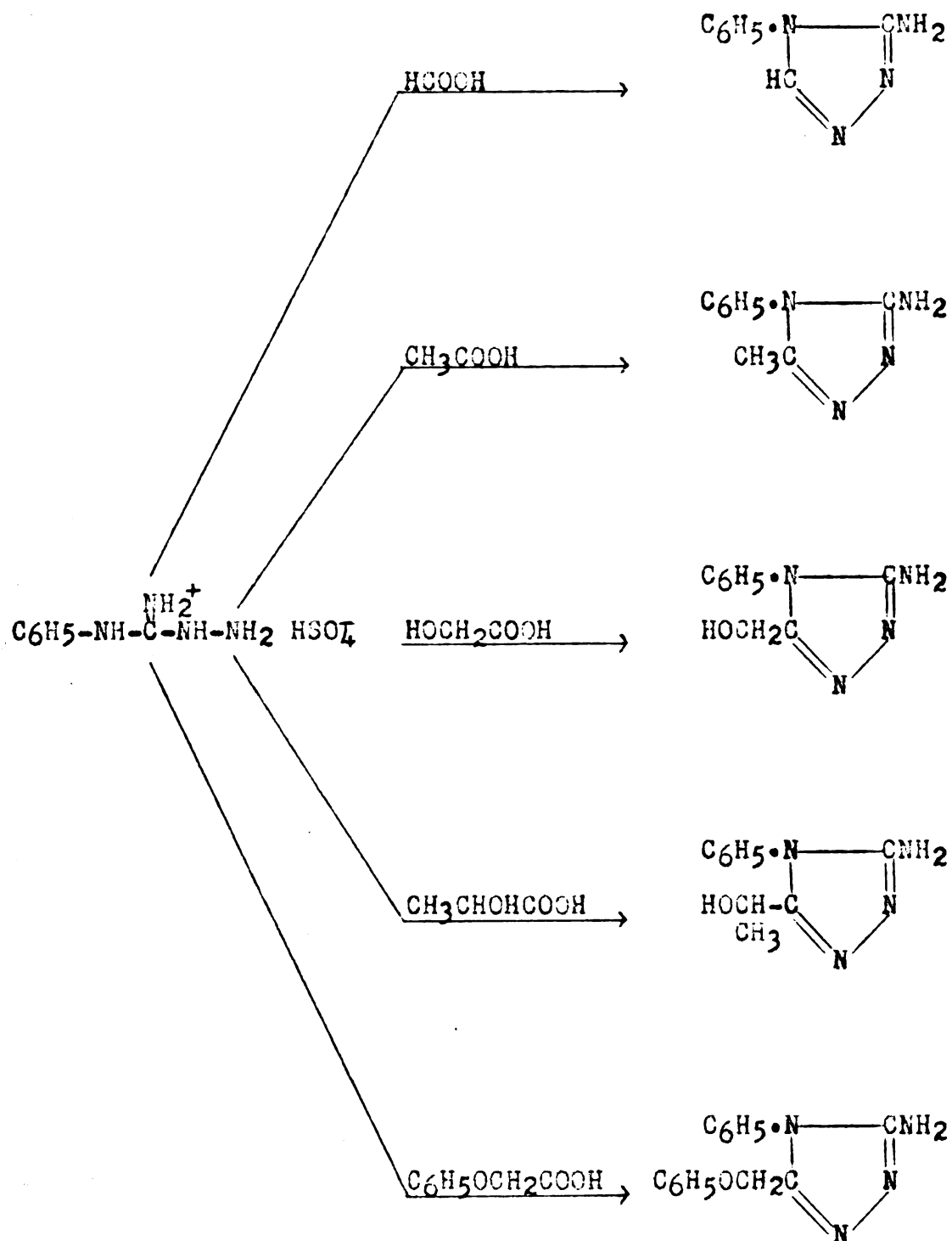
The method used in the synthesis of these compounds is basically the one discovered by Thiele (12) in 1891. He observed that aminoguanidine reacts with acetic acid to form acetylamino-guanidine which on warming or treatment with alkali loses a molecule of water and cyclizes to form an inner anhydro base:



Thiele and Heidenreich (25) found that this substance had the properties of an acid and that on oxidation an azomethyltriazole was formed; later work showed that an amino group capable of diazotization was present (21). Thiele and Manchot (13) showed the general nature of this reaction by preparing 3-amino-1,2,4-triazole from aminoguanidine and formic acid, and 3-amino-1,2,4-triazole-5-carboxylic acid from aminoguanidine and oxalic acid.

Reilly and Madden (26) have prepared the corresponding 5-ethyl-, 5-isopropyl-, and 5-isobutylaminotriazoles; and Reilly and Drumm (22) have prepared the 3-amino-5-n-propyl-1,2,4-triazole.

By the reaction of phenylaminoguanidinium bisulfate with an appropriate aliphatic acid or its derivative, it was possible to prepare 3-amino-4-phenyl-1,2,4-triazole and a number of 3-amino-4-phenyl-5-alkyl-1,2,4-triazoles. These syntheses are illustrated on the next page:



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3-Amino-4-phenyl-5-chloromethyl-1,2,4-triazole
Hydrochloride

Some difficulties were encountered in the preparation of this important intermediate. Several attempts to synthesize it by the reaction of phenylaminoguanidinium bisulfate and chloroacetic acid (Thiele's method) were fruitless. Bloom and Day (27) had succeeded in preparing 2-chloromethylbenzimidazole in good yields by using the method of Phillips (28). They heated o-phenylenediamine with chloroacetic acid in 4N hydrochloric acid for forty-five minutes, allowed the mixture to stand overnight at room temperature and then neutralized it in the cold with 6N ammonium hydroxide. When phenylaminoguanidinium bisulfate and chloroacetic acid were allowed to react under comparable conditions, 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole was obtained in poor yield. It was finally decided to attempt to prepare this compound from the corresponding hydroxymethyltriazole. After several experiments, a procedure was worked out for the smooth conversion of the hydroxymethyltriazole to the chloromethyltriazole in good yield.

The hydrochloride of 3-amino-4-phenyl-5-hydroxymethyl-1,2,4-triazole was prepared and then treated with an excess of thionyl chloride in chloroform. After refluxing for ten hours and allowing the mixture to stand overnight, the chloromethyltriazole could be isolated in good yield.

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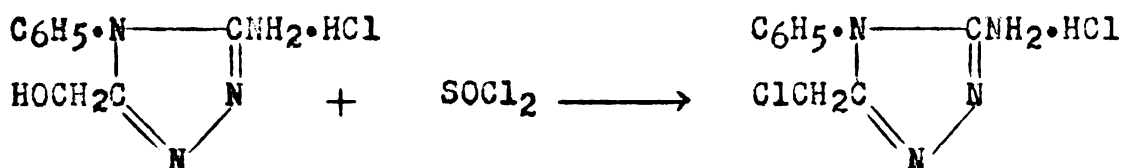
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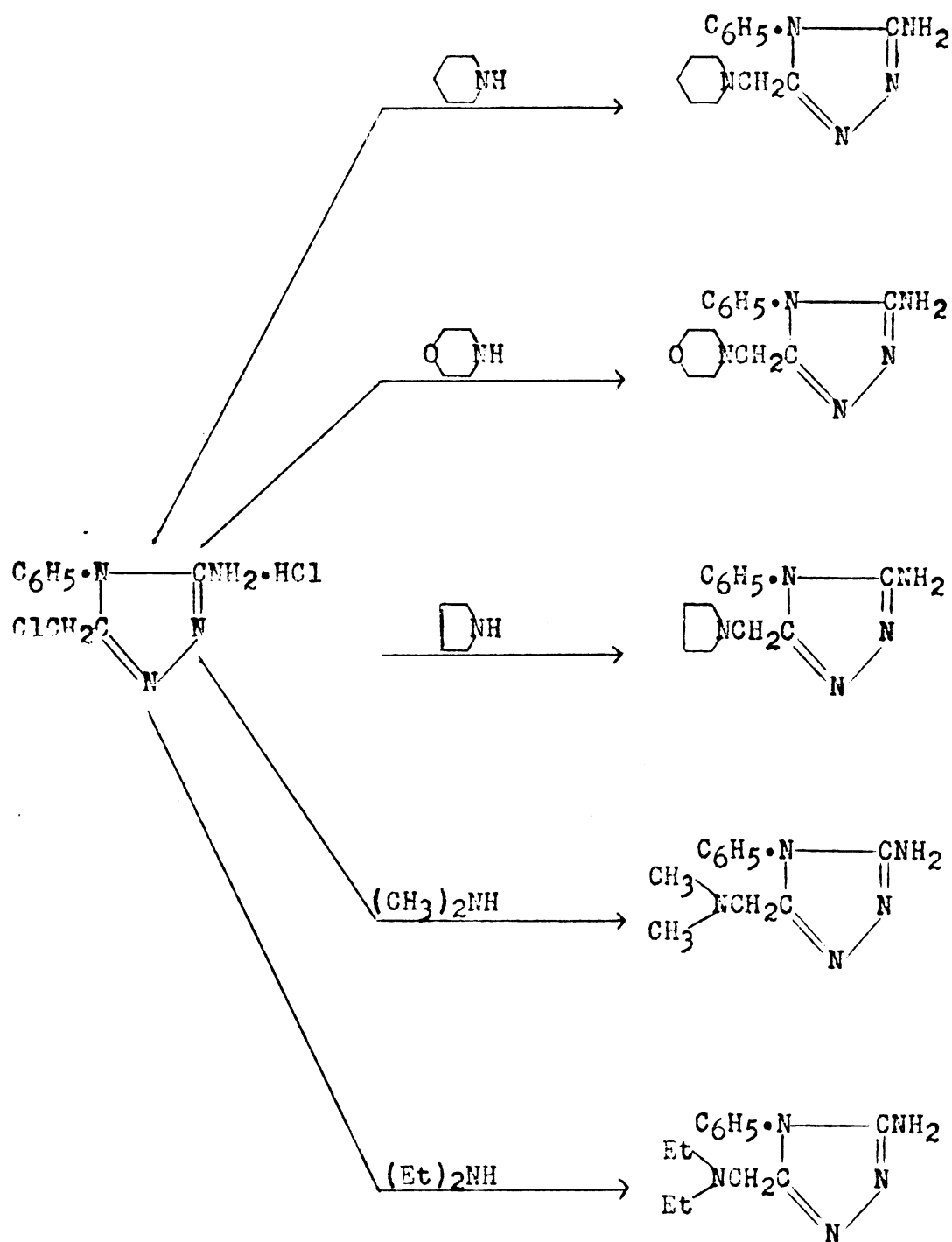
This intermediate was employed in the synthesis of the various dialkylaminomethyl- and phenoxymethyltriazoles.

Dialkylaminomethyltriazoles

The presence of both a reactive halogen and a moderately reactive amino group in the molecule of the intermediate was a source of some difficulty in the synthesis of these compounds.

In preparing these compounds, an alcoholic solution of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole was treated with the calculated amount of a secondary amine necessary to (1) liberate the amino group of the chloromethyltriazole, (2) react with the reactive halogen and (3) combine with the hydrogen chloride produced in the course of the reaction. Since the amino group of the secondary amine is much more basic than the amino group of the triazole, these reactions were allowed to proceed for a day or more at room temperature. These conditions were found to be most favorable for the reaction of the amino group of the secondary amine, and least favorable for the reaction of the amino group of the aminotriazole.

Sufficient sodium hydroxide was then added to liberate the excess secondary amine from its hydrochloride. The excess secondary amine was then removed along with the solvent by evaporating to dryness under reduced pressure. The product was then extracted from the sodium chloride with toluene. The best yields were obtained from those reactions involving the more basic secondary amines. These syntheses are illustrated on the following two pages:

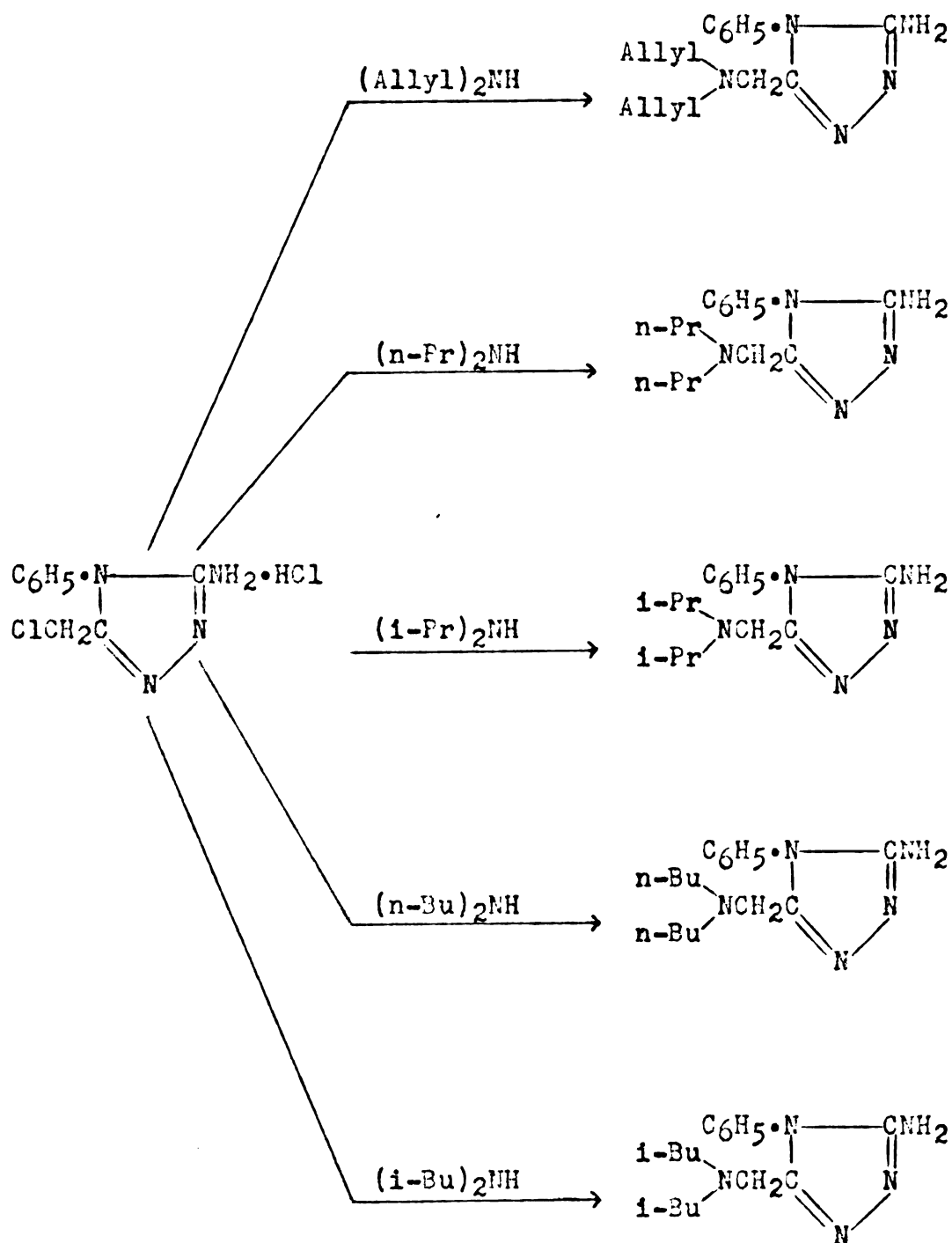


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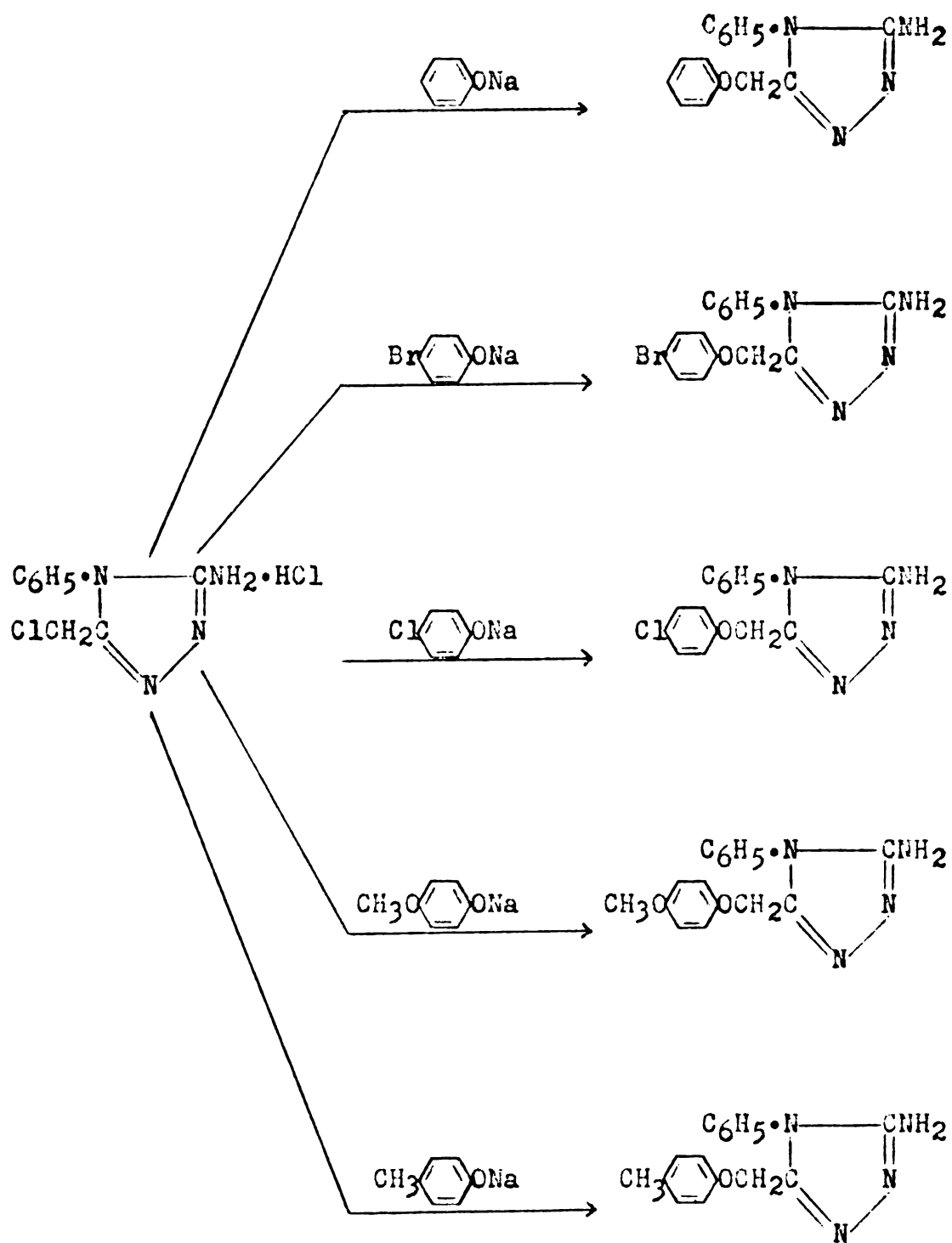




Phenoxymethyl- and thiophenoxymethyltriazoles

The same conditions employed in the synthesis of the dialkylaminomethyltriazoles were used in order to minimize the reaction of the amino group of the chloromethyltriazole.

In preparing the phenoxymethyltriazoles, an alcoholic solution of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole was treated with a solution containing at least the stoichiometric amount of the appropriately substituted sodium phenoxide in alcohol. These reactions were then allowed to proceed for a day or more at room temperature. In the case of those reactions involving sodium phenoxides with electron-repelling groups substituted in the ortho or para positions, precipitation of the product usually began promptly after the initial reaction had subsided. The best yields were usually obtained in these reactions. In those reactions in which no product precipitated at the end of the reaction time, the mixture was heated on the steam bath for about one hour to ensure completion of the reaction. Water was then added to the cooled solution to precipitate the product. These syntheses are illustrated on the following pages:



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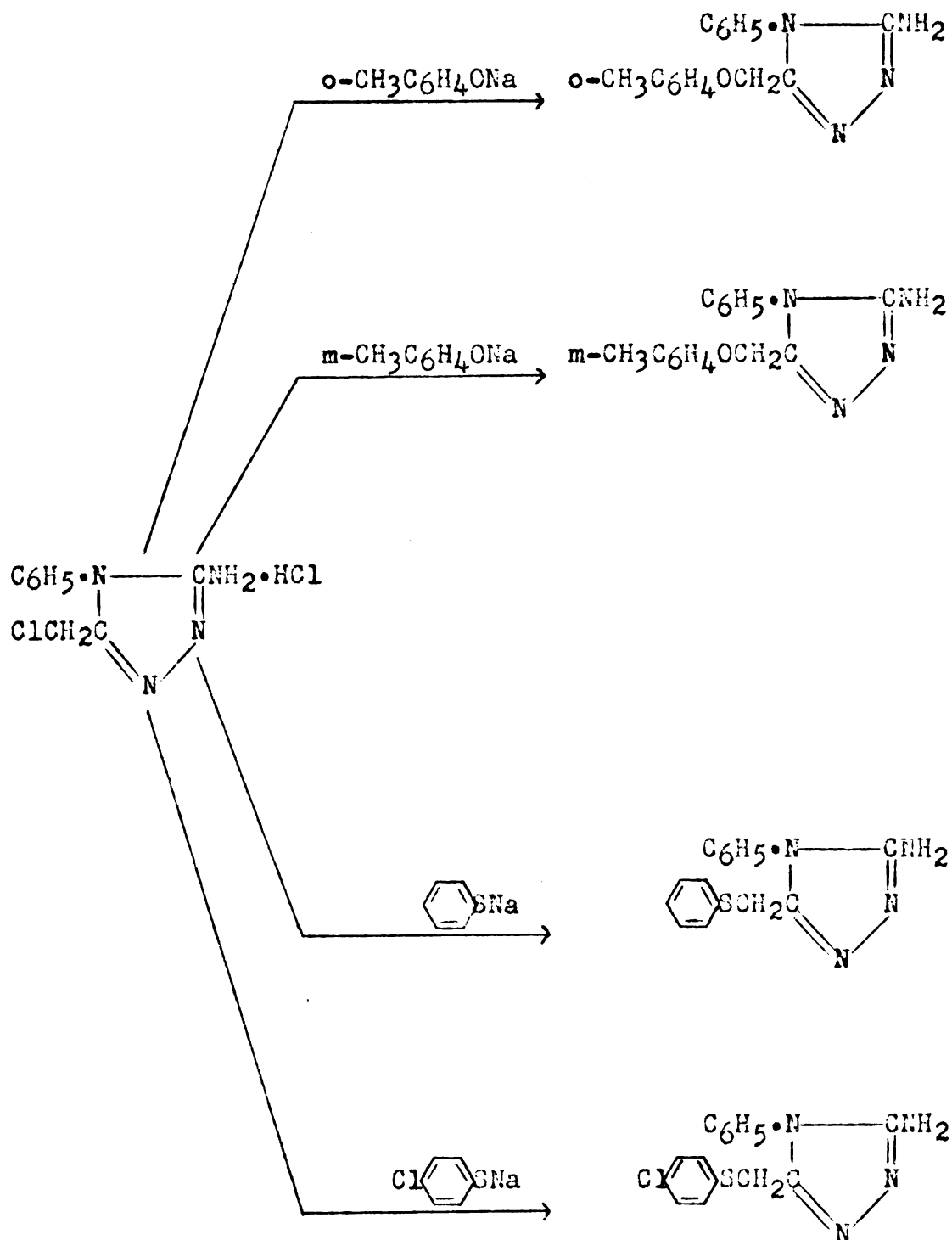
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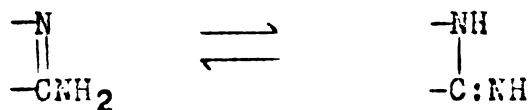
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Phenylthioureas of Aminotriazoles

Fromm (29) and other investigators were the first to note the remarkable fact that the phenylthioureas derived from a number of heterocyclic compounds containing the group:



occurred in two isomeric forms. The low melting isomer is obtained by allowing the heterocyclic amine to react with phenylisothiocyanate at room temperature. The higher melting isomer is formed when the reaction is carried out at a higher temperature. The low melting labile isomer can be converted into the high melting stable form by heating it above its melting point. The reverse of this change is not observed. Fromm suggested that the labile isomer is derived from a secondary amine and the stable form from a primary amine.

Fantl and Silbermann (30) have studied the behavior of aminotriazoles toward phenylisothiocyanate. They found that 3-amino-5-methyl-1,2,4-triazole reacted with phenylisothiocyanate in the cold to produce a substance melting at 137° C., which solidified at 140° C. and then melted at 197° C. They also obtained the higher melting compound directly by heating the reactants in a high boiling solvent (n-amyl alcohol) for two hours. They assigned the following formulas to their two products:

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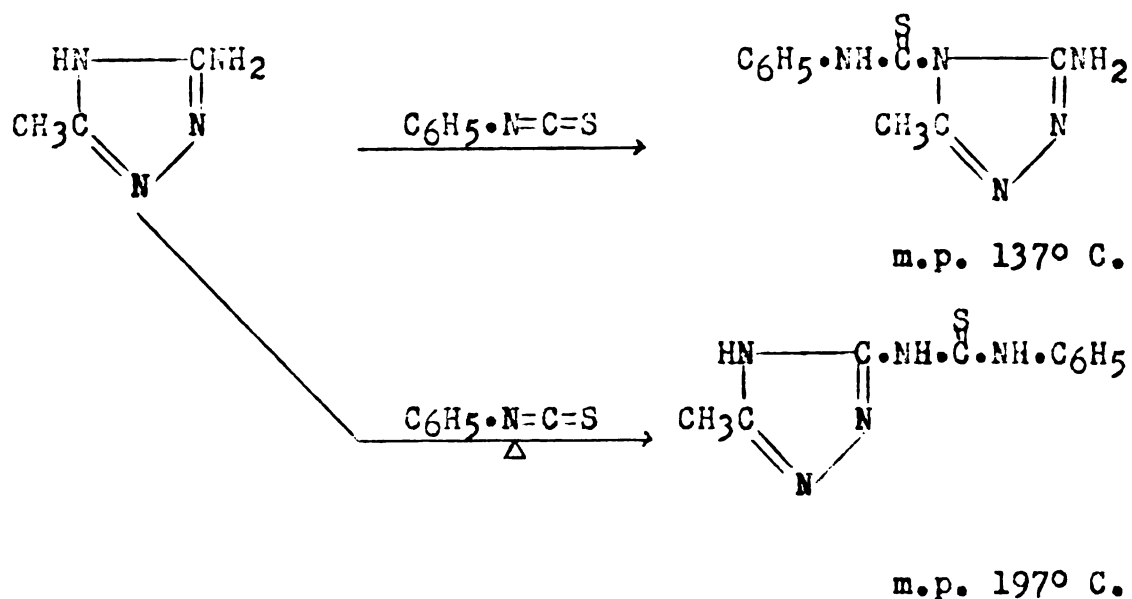
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Further studies showed that the presence of isomerism among the phenylthioureas of aminotriazoles was not a general occurrence since 2-phenyl-3-allylamino-1,2,4-triazole reacted with phenylisothiocyanate to yield only one stable phenylthiourea.

In preparing the phenylthioureas of the aminotriazoles which were synthesized in the course of this study, the author did not observe the presence of isomerism.

Biological Properties of Aminotriazoles

Interest in the study of the biological properties of aminotriazoles was strongly stimulated in 1952 when Hall et al (31) first noted that aminotriazole had defoliating and regrowth inhibiting properties on cotton during the course of tests conducted at the Texas Agricultural Experimental Station.

In 1953, Shaw and Swanson (7) reported their study of the relationship between phytotoxicity and chemical structure of herbicides. Aminotriazole was again found to have the desirable properties of a promising herbicide.

The following year, W.W. Allen (32) of the American Chemical Paint Co. was granted a patent on the use of aminotriazole as a herbicide and cotton defoliant. After a number of field tests, aminotriazole was marketed by the American Chemical Paint Co. as a herbicide under the trade names - "Amizol" and "Weedazol". The American Cyanamid Co. was also licensed to manufacture and sell aminotriazole.

This compound whose derivatives had been used only sparingly as fog inhibitors in photographic emulsions (33) was reported to be more effective against Canada thistle than any other known herbicide. Prior to the use of aminotriazole, Canada thistle could only be kept under control by dosing with large amounts of soil sterilants or repeated applications of 2,4-D. Other weeds that were

also susceptible to control by aminotriazole included the following: quackgrass, nutgrass, Bermuda grass, poison ivy, poison oak, scrub oak, sow thistle, milkweed, horse-tail rush, cattails, ash and red maple.

The fundamental reasons for the inherent herbicidal properties of aminotriazole are still obscure. It has been observed to cause chlorosis in plants. Sometimes the plant dies after the chlorosis. Generally, only new growth, in which pigment is being formed, becomes chlorotic. This has caused some (5) to suggest that chlorophyll synthesis is being affected. It has also been found that the degree of chlorosis is proportional to the amount of aminotriazole applied.

Kenneth A. Sund of the American Cyanamid Co. (34) has suggested that aminotriazole may affect plant physiological processes in three different ways: (1) as a precursor in the formation of some porphyrin similar to chlorophyll but unable to carry out the functions of chlorophyll; (2) it may bind by complex formation some metal or metals required for the development of chlorophyll; (3) it may upset some oxidation-reduction reactions within the plant.

By using radioactive 3-amino-1,2,4-triazole with the fifth position labeled with carbon-14, Miller and Hall (5) made a study to determine how aminotriazole effects chlorosis in plants. They suggested that aminotriazole in sub-lethal doses causes chlorosis by inhibiting the synthesis of chlorophylls and carotenoids. But when it is

applied in high concentrations, chlorosis is caused by the destruction of chlorophyll. From their studies they concluded that the possibility that aminotriazole acts as a precursor in the formation of an abnormal porphyrin is quite remote and that the complexing of an essential metal is unlikely. They agreed with Rogers (35) that aminotriazole inhibits plastid development.

While studying the depressing effect of aminotriazole on chlorophyll synthesis in plants, Heim et al (36) observed that the chemical also causes a great decrease in the catalase activity of plant tissue. This led them to study the effect of aminotriazole on catalase activity in rats. They found that aminotriazole reduced hepatic and renal catalase activity levels, but not that of red cells, thus producing an effect similar to that observed in tumor-bearing animals.

Tschudy and Collins (6) concluded that the ability of aminotriazole to affect two different porphyrin-containing compounds - catalase and chlorophyll - suggested a possible interference with porphyrin synthesis. They then studied the effect of aminotriazole on an enzyme known to be involved in porphyrin synthesis. They studied the effect of aminotriazole on δ -aminolevulinic acid dehydrase activity. This enzyme catalyzes the conversion of δ -aminolevulinic acid to porphobilinogen - an intermediate in the synthesis of porphyrins. They found

• The first step in the process of identifying a problem is to define the problem clearly.

• The second step is to identify the causes of the problem.

• The third step is to develop a plan to solve the problem.

• The fourth step is to implement the plan.

• The fifth step is to evaluate the results of the plan.

• The sixth step is to make adjustments to the plan if necessary.

• The seventh step is to document the process.

• The eighth step is to communicate the results.

• The ninth step is to review the process.

• The tenth step is to make improvements.

• The eleventh step is to monitor the results.

• The twelfth step is to report the results.

• The thirteenth step is to conclude the process.

• The fourteenth step is to reflect on the process.

• The fifteenth step is to learn from the experience.

• The sixteenth step is to apply the lessons learned.

• The seventeenth step is to share the results.

• The eighteenth step is to celebrate the success.

• The nineteenth step is to maintain the results.

• The twentieth step is to continue the process.

• The twenty-first step is to stay motivated.

• The twenty-second step is to stay focused.

• The twenty-third step is to stay organized.

• The twenty-fourth step is to stay positive.

that aminotriazole reduces the level of δ -aminolevulinic acid dehydrase activity in the kidneys. It is significant that this effect is also produced by tumors. Thus tumors and aminotriazole are capable of causing a decrease in activity of both δ -aminolevulinic acid dehydrase and catalase.

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1. The first step in the process of creating a new product is to identify a market need. This involves conducting market research to determine what consumers want and what problems they are trying to solve. Once a need is identified, the next step is to develop a concept for a product that addresses that need. This often involves brainstorming and sketching ideas.

2. After a concept is developed, the next step is to create a prototype. A prototype is a preliminary model of the product that allows designers to test and refine their ideas. This can be done using various materials and techniques, depending on the nature of the product.

3. Once a prototype is created, the next step is to conduct a feasibility study. This involves evaluating the technical, financial, and market viability of the product. This step is crucial for determining whether the product is worth developing further.

4. If the feasibility study is positive, the next step is to develop a business plan. This document outlines the company's strategy for producing, marketing, and selling the product. It also includes financial projections and a timeline for development.

5. The final step in the process is to launch the product. This involves manufacturing the product, distributing it to retailers or customers, and promoting it through marketing efforts. After launch, it's important to monitor sales and customer feedback to make any necessary adjustments.

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Experimental

Phenylaminoguanidinium Bisulfate

A slurry of 304 g. (2.0 moles) of phenylthiourea (Eastman Kodak Co.) in 600 ml. of absolute methanol was cooled to 5° C. and then treated with 132 ml., 301 g. (2.12 moles) of methyl iodide. The mixture was allowed to remain in an ice bath as it warmed up to room temperature and stand for forty-eight hours. At the end of this period, 100 ml. of methanol were removed by distillation and then replaced by 100 ml. of fresh methanol. The flask was then placed in an ice bath and 67.4 g. (2 moles) of 95% hydrazine was added. The ice bath was then removed and the solution refluxed gently for four hours in the hood. The methyl mercaptan which is a by-product of this reaction was collected in a trap which was cooled in an ice-salt bath. The alcohol was then removed under reduced pressure until a viscous liquid remained. A volume of water equal to the volume of the viscous liquid was then added. This solution was again evaporated to a viscous liquid under reduced pressure. The viscous liquid was used in preparing the phenylaminoguanidinium bisulfate.

The viscous solution of the phenylaminoguanidinium iodide was dissolved in a solution of 60 ml. of concentrated sulfuric acid in 300 ml. of water. This solution was then added to a slurry of 327 g. (2.05 moles) of

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silver sulfate in 200 ml. of water. The mixture became quite warm and yellow silver iodide precipitated. The mixture was stirred occasionally and permitted to stand overnight. The yellow silver iodide was collected on a filter and the filtrate was treated with hydrogen sulfide for about one hour until the precipitation of silver sulfide was complete. The solution was heated to coagulate the colloidal silver sulfide and then filtered. Next, it was treated with "Norit". After filtering, the colorless solution was evaporated to a viscous liquid under reduced pressure. The viscous liquid solidified on standing. It was dried in a vacuum desiccator and crushed while still soft to colorless granules. The yield of product was 400 g. (80% of theory based on the phenylthiourea). A sample for analysis was recrystallized from isopropyl alcohol and dried under a vacuum at 80° C. It melted at 94-96° C. (heated slowly) and near 85° C. when heated rapidly.

Anal. Calc'd. for $C_7H_{12}N_4O_4S$: N, 22.57; S, 12.92

Found: N, 22.31; S, 12.81

3-Amino-4-phenyl-4-H.1.2.4-triazole

A mixture of 25 g. (0.1 mole) of phenylamino-guanidinium bisulfate and 50 ml. of 88% formic acid was refluxed for six hours. The excess formic acid was removed under reduced pressure and 40 ml. of water was

added. The solution was evaporated under reduced pressure to a viscous liquid, and 35 ml. of water was added. The solution was then carefully neutralized with solid sodium carbonate. Colorless crystals precipitated; they were separated by filtration and recrystallized from methanol. The yield of triazole was 12 g. (75% of theory). A sample for analysis was recrystallized twice from methanol and melted at 218-219° C.

Anal. Calc'd. for $C_8H_8N_4$: C, 60.00; H, 5.00;
N, 35.00

Found: C, 59.99; H, 5.03; N, 34.98

3-Amino-4-phenyl-4-H,1,2,4-triazole Hydrochloride

A solution of 0.8 g. (0.005 mole) of 3-amino-4-phenyl-1,2,4-triazole in 5 ml. of concentrated hydrochloric acid was evaporated to dryness on the steam bath. The crystalline residue was recrystallized three times from an ethanol-ether solvent pair. The yield of product was 0.88 g. (90% of theory). The colorless crystals melted at 190-191° C.

Anal. Calc'd. for $C_8H_9ClN_4$: C, 48.86; H, 4.61;
Cl, 18.03; N, 28.49

Found: C, 48.55; H, 4.85; Cl, 18.03;
N, 28.74

3-Acetanido-4-phenyl-4-H,1,2,4-triazole

Two grams (0.012 mole) of 3-amino-4-phenyl-1,2,4-triazole was treated with 20 ml. of acetic anhydride and refluxed for three hours. The excess acetic anhydride was removed under reduced pressure and 40 ml. of water was added to precipitate the product. The crystalline product was removed by filtration and washed with water. Recrystallization of the product three times from ethanol gave 1.5 g. (60% of theory) of colorless crystals melting at 201-202° C.

Anal. Calc'd. for $C_{10}H_{10}N_4O$: C, 59.40; H, 4.98
N, 27.71

Found: C, 59.58; H, 5.15; N, 27.72

The hydrolysis of 3-acetamido-4-phenyl-1,2,4-triazole by heating with concentrated hydrochloric acid produced a product which was identical with 3-amino-4-phenyl-1,2,4-triazole as shown by a mixture melting point determination. This suggests the absence of rearrangement of the original triazole in the process of acetylation. The original triazole also did not rearrange when heated above its melting point.

3-Amino-4-phenyl-5-methyl-4-H,1,2,4-triazole

A mixture of 25 g. (0.1 mole) of phenylaminoguanidinium bisulfate and 50 ml. of glacial acetic acid was refluxed for twelve hours. The excess acetic acid was removed under reduced pressure and 40 ml. of water was

added. The solution was evaporated under reduced pressure to a viscous liquid, and 35 ml. of water was added. The solution was then neutralized with solid sodium carbonate. Colorless crystals precipitated; they were removed by filtration and extracted with hot methanol. This solution was filtered while hot and cooled in an ice bath. Precipitation of the crystalline product took place. The yield of crude triazole was 13 g. (75% of theory). A sample for analysis was recrystallized twice from methanol and melted at 161-162° C.

Anal. Calc'd. for $C_9H_{10}N_4$: C, 62.10; H, 5.75;
N, 32.20

Found: C, 62.05; H, 5.79; N, 32.17

3-Amino-4-phenyl-5-methyl-4-H,1,2,4-triazole Hydrochloride

A solution of 0.87 g. (0.005 mole) of 3-amino-4-phenyl-5-methyl-1,2,4-triazole in 5 ml. of concentrated hydrochloric acid was evaporated to dryness on the steam bath. The crystalline residue was recrystallized three times from an ethanol-ether solvent pair. The yield was 0.97 g. (92% of theory) of colorless crystals melting at 223-225° C.

Anal. Calc'd. for $C_9H_{11}ClN_4$: C, 51.31; H, 5.26
Cl, 16.83; N, 26.60

Found: C, 51.09; H, 5.44; Cl, 16.92; N, 26.76

1-(4'-Phenyl-5'-methyl-4'-H,1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.003 mole) of 3-amino-4-phenyl-5-methyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for two hours and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration; washed with 50 ml. of ligroin, and recrystallized twice from methanol to yield 0.78 g. (90% of theory) of product melting at 186-188° C.

Anal. Calc'd. for $C_{16}H_{15}N_5S$: N, 22.64; S, 10.36

Found: N, 22.82; S, 10.28

3-Amino-4-phenyl-5-hydroxymethyl-4-H,1,2,4-triazole

A mixture of 25 g. (0.1 mole) of phenylaminoguanidinium bisulfate, 10 ml. of 70% glycolic acid, and 12 ml. of water was refluxed for twenty-two hours. The solution was then cooled and carefully neutralized with 7.5 N ammonia. Fine, cream-colored crystals precipitated. They were removed by filtration and washed with methanol. The yield of crude triazole was 14 g. (75% of theory). A sample for analysis was recrystallized three times from methanol and melted at 239-240° C.

Anal. Calc'd. for $C_9H_{10}N_4O$: C, 56.84; H, 5.26;

N, 29.47

Found: C, 56.98; H, 5.18; N, 29.57

1. The first part of the paper is devoted to the study of the properties of the function $f(x)$ defined by the equation

$$f(x) = \int_0^x \frac{1}{1+t^2} dt$$
for $x \in \mathbb{R}$. It is shown that $f(x)$ is an odd function, i.e., $f(-x) = -f(x)$, and that it is strictly increasing on $\mathbb{R}$. Moreover, it is proved that $f(x)$ is bounded on $\mathbb{R}$, with $\lim_{x \rightarrow -\infty} f(x) = -\frac{\pi}{2}$ and $\lim_{x \rightarrow \infty} f(x) = \frac{\pi}{2}$.

2. In the second part, we consider the function $g(x)$ defined by the equation $g(x) = \int_0^x \frac{1}{1+t^4} dt$ for $x \in \mathbb{R}$. It is shown that $g(x)$ is an even function, i.e., $g(-x) = g(x)$, and that it is strictly increasing on $[0, \infty)$. Moreover, it is proved that $g(x)$ is bounded on $\mathbb{R}$, with $\lim_{x \rightarrow -\infty} g(x) = 0$ and $\lim_{x \rightarrow \infty} g(x) = \frac{\pi}{4}$.

3. The third part of the paper is devoted to the study of the function $h(x)$ defined by the equation $h(x) = \int_0^x \frac{1}{1+t^6} dt$ for $x \in \mathbb{R}$. It is shown that $h(x)$ is an even function, i.e., $h(-x) = h(x)$, and that it is strictly increasing on $[0, \infty)$.

Moreover, it is proved that $h(x)$ is bounded on $\mathbb{R}$, with $\lim_{x \rightarrow -\infty} h(x) = 0$ and $\lim_{x \rightarrow \infty} h(x) = \frac{\pi}{6}$. Finally, we show that the function $h(x)$ is concave down on $[0, \infty)$, i.e., $h''(x) < 0$ for $x > 0$.

4. The fourth part of the paper is devoted to the study of the function $k(x)$ defined by the equation

$$k(x) = \int_0^x \frac{1}{1+t^8} dt$$

for $x \in \mathbb{R}$. It is shown that $k(x)$ is an even function, i.e., $k(-x) = k(x)$, and that it is strictly increasing on $[0, \infty)$.

3-Amino-4-phenyl-5-hydroxymethyl-4-H.1,2,4-triazole
Hydrochloride

A solution of 9.5 g. (0.05 mole) of 3-amino-4-phenyl-5-hydroxymethyl-1,2,4-triazole in 10 ml. of concentrated hydrochloric acid was evaporated to dryness on the steam bath. The crystalline residue was washed with acetone. A sample for analysis was recrystallized three times from an ethanol-ether solvent pair. The yield of hydrochloride was 10.6 g. (94% of theory). The colorless crystals melted at 218-220° C.

Anal. Calc'd. for $C_9H_{11}ClN_4O$: C, 47.69; H, 4.89
 Cl, 15.64; N, 24.72

Found: C, 47.66; H, 4.97; Cl, 15.65;
 N, 24.77

3-Amino-4-phenyl-5-(α -hydroxyethyl)-4-H.1,2,4-triazole

A mixture of 25 g. (0.1 mole) of phenylaminoguanidinium bisulfate, 12 ml. of 85% lactic acid and 24 ml. of water was refluxed for sixteen hours. The solution was then carefully neutralized with solid sodium carbonate. Colorless crystals precipitated, were removed by filtration and recrystallized from methanol. The yield of triazole was 7 g. (35% of theory). A small sample was further recrystallized once from ethanol and twice from isopropyl alcohol and melted at 205-207° C.

Anal. Calc'd. for $C_{10}H_{12}N_4O$: C, 58.81; H, 5.92;
 N, 27.44

Found: C, 58.88; H, 6.09; N, 27.26

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3-Acetamido-4-phenyl-5-(α -acetoxyethyl)-4-H,1,2,4-triazole

Two grams (0.01 mole) of 3-amino-4-phenyl-5-(α -hydroxyethyl)-1,2,4-triazole was treated with 20 ml. of acetic anhydride and refluxed for three hours. The excess acetic anhydride was removed under reduced pressure and 40 ml. of water was added to precipitate the product. The crystalline product was removed by filtration and washed with water. Recrystallization of the product three times from ethanol gave 1 g. (35% of theory) of colorless crystals melting at 201-203° C.

Anal. Calc'd. for $C_{14}H_{16}N_4O_3$: C, 58.32; H, 5.60;
N, 19.43

Found: C, 58.26; H, 5.73; N, 19.22

3-Amino-4-phenyl-5-phenoxyethyl-4-H,1,2,4-triazole

A mixture of 7.6 g. (0.05 mole) of phenoxyacetic acid, 12.4 g. (0.05 mole) of phenylaminoguanidinium bisulfate, and 25 ml. of water was refluxed for twenty-two hours. The solution was then carefully neutralized with solid sodium carbonate; a gummy product precipitated. The aqueous layer was decanted and the product was recrystallized four times from toluene to yield 2.5 g. (20% of theory) of colorless crystals melting at 195-196° C.

Anal. Calc'd. for $C_{15}H_{14}N_4O$: C, 67.65; H, 5.30;
N, 21.04

Found: C, 67.81; H, 5.44; N, 21.06

[illegible]

3-Amino-4-phenyl-5-phenoxyethyl-4-H,1,2,4-triazole
Hydrochloride

A solution of 1.3 g. (0.005 mole) of 3-amino-4-phenyl-5-phenoxyethyl-1,2,4-triazole in 5 ml. of concentrated hydrochloric acid was evaporated to dryness on the steam bath. The crystalline residue was recrystallized three times from an ethanol-ether solvent pair. The yield of product was 1 g. (70% of theory), m.p. 198-199° C.

Anal. Calc'd. for $C_{15}H_{15}ClN_4O$: Cl, 11.71;
 N, 18.51

Found: Cl, 11.84; N, 18.35

1-(4'-phenyl-5'-phenoxyethyl-4'-H,1',2',4'-triazolyl-3')-
3-phenylthiourea

A mixture of 1 g. (0.004 mole) of 3-amino-4-phenyl-5-phenoxyethyl-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.6 g. (37% of theory) of product melting at 165-167° C.

Anal. Calc'd. for $C_{22}H_{19}N_5OS$: N, 17.44; S, 7.98

Found: N, 17.63; S, 7.94

3-Amino-4-phenyl-5-chloromethyl-4-H,1,2,4-triazole
Hydrochloride

A slurry of 10.8 g. (0.048 mole) of the hydrochloride of 3-amino-4-phenyl-5-hydroxymethyl-1,2,4-triazole

in 25ml. of chloroform was treated with 10 g. (6 ml., 0.08 mole) of thionyl chloride in 15 ml. of chloroform. After thirty minutes at room temperature, the mixture was refluxed for six hours. Another addition of 6 g. (4 ml., 0.053 mole) of thionyl chloride in 10 ml. of chloroform was made and refluxing continued for four hours. The mixture became dark red and was permitted to stand overnight. The mixture was then cooled and stirred; a cream-colored solid precipitated. An additional crop of crystals was obtained by the addition of a small amount of ether to the filtrate. The product was washed with ether. The yield of chloromethyl triazole hydrochloride was 10.4 g. (86% of theory). A sample for analysis was recrystallized twice from a methanol-acetone solvent pair and melted at 180-190° C.

Anal. Calc'd. for $C_9H_{10}Cl_2N_4$: C, 44.10; H, 4.11;
Cl, 28.93; N, 22.86

Found: C, 44.26; H, 4.23; Cl, 28.98;
N, 22.59

3-Amino-4-phenyl-5-piperidinomethyl-4-H,1,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol was treated with 12.75 g. (14.8 ml., 0.15 mole) of piperidine and allowed to stand for twenty-four hours. The piperidine hydrochloride was precipitated by the addition of ether. It was removed by filtration

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and the filtrate was evaporated under reduced pressure until only a viscous liquid remained. The residue was treated with 25 ml. of concentrated hydrochloric acid, and the solution was again evaporated under reduced pressure until only a viscous liquid remained. Upon cooling, this liquid solidified to a colorless mass. The amine hydrochloride was washed with acetone and treated with a large excess of concentrated ammonia. Lustrous crystals appeared which were washed with concentrated ammonia and then with water. The yield of product was 11 g. (84% of theory). The amine was recrystallized twice from toluene and melted at 153-154° C.

Anal. Calc'd. for $C_{14}H_{19}N_5$: C, 65.33; H, 7.44;
N, 27.22

Found: C, 65.43; H, 7.47; N, 27.39

When the free amine was refluxed in acetic anhydride for five hours, no product could be isolated. When acetic acid was used in place of the anhydride, the free amine was recovered unchanged.

1-(4'-Phenyl-5'-piperidinomethyl-4'-H,1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-piperidinomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product began. The

phenylthiourea derivative was removed by filtration, washed with ligroin, and recrystallized twice from methanol to yield 0.6 g. (77% of theory) of product melting at 184-186° C.

Anal. Calc'd. for $C_{21}H_{24}N_6S$: N, 21.41; S, 8.17

Found: N, 21.39; S, 8.03

3-Amino-4-phenyl-5-morpholinomethyl-4-H,1,2,4-triazole

A solution of 24.5 g. (0.1 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 100 ml. of methanol was treated with 26.13 g. (26 ml., 0.3 mole) of morpholine and allowed to stand for twenty-four hours. The morpholine hydrochloride was precipitated by the addition of ether and removed by filtration. The filtrate was evaporated under reduced pressure until only a viscous liquid remained. The residue was treated with 25 ml. of concentrated hydrochloric acid, and the solution was again evaporated under reduced pressure until only a viscous liquid remained. Upon cooling, this liquid solidified to a colorless mass. The amine hydrochloride was washed with acetone and treated with a large excess of concentrated ammonia. Lustrous crystals appeared which were washed with concentrated ammonia and then with water. The yield of base was 13 g. (50% of theory). The product was recrystallized twice from methanol and melted at 204-205° C.

Anal. Calc'd. for $C_{13}H_{17}N_5O$: C, 60.20; H, 6.61;
N, 27.01

Found: C, 59.91; H, 6.69; N, 26.89

1-(4'-Phenyl-5'-morpholinomethyl-4'-H,1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-morpholinomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with ligroin, and recrystallized twice from methanol to yield 0.6 g. (77% of theory) of product melting at 176-178° C.

Anal. Calc'd. for $C_{20}H_{22}N_6OS$: N, 21.31; S, 8.13

Found: N, 21.40; S, 8.15

3-Amino-4-phenyl-5-pyrrolidinomethyl-4-H,1,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol was treated with 10.65 g. (12.2 ml., 0.15 mole) of pyrrolidine and allowed to stand for forty-eight hours. The pyrrolidine hydrochloride was precipitated as a dark heavy oil by the addition of 200 ml. of anhydrous ether. The clear upper layer of the mixture was decanted and evaporated under reduced pressure until

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only a viscous liquid remained. The residue was treated with 25 ml. of concentrated hydrochloric acid, and the solution was again evaporated under reduced pressure until only a viscous liquid remained. Upon cooling, the amine hydrochloride precipitated as a hygroscopic, colorless solid. After triturating with acetone, it was treated with a large excess of concentrated ammonia. Lustrous crystals appeared which were washed with concentrated ammonia and then with water. The yield of base was 6 g. (50% of theory). It was recrystallized three times from toluene and melted at 158-160° C. A sample was further recrystallized once from toluene, three times from acetone and treated with "Morit"; it also melted at 158-160° C.

Anal. Calc'd. for $C_{13}H_{17}N_5$: C, 64.17; H, 7.05;
N, 28.79

Found: C, 64.34; H, 7.17; N, 28.75

1-(4'-Phenyl-5'-pyrrolidinomethyl-4'-H,1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-pyrrolidinomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with ligroin, and recrystallized twice from

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1. *Chlorophyll a* and *Chlorophyll b* were determined by the method of Arar and Collins (1971) using a Shimadzu 1601 UV-Visible Spectrophotometer.

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methanol to yield 0.4 g. (53% of theory) of product melting at 178-180° C.

Anal. Calc'd. for $C_{20}H_{22}N_6S$: N, 22.21; S, 8.47

Found: N, 22.43; S, 8.40

3-Amino-4-phenyl-5-dimethylaminomethyl-4-H,1,2,4-triazole

A solution of dimethylamine was prepared by treating 24.5 g. (0.3 mole) of dimethylamine hydrochloride with a solution of 12 g. (0.3 mole) of sodium hydroxide in 10 ml. of water and 50 ml. of methanol. After cooling the mixture, the sodium chloride which had precipitated was removed by filtration. This solution of dimethylamine was then treated with a solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol contained in an Erlenmeyer flask. After the initial reaction had subsided, the flask was stoppered and permitted to stand for ninety-six hours. It was then treated with a solution of 4 g. (0.1 mole) of sodium hydroxide in 10 ml. of water and 50 ml. of methanol. The sodium chloride which precipitated was removed by filtration and the filtrate was evaporated to dryness under reduced pressure. A yellow solid remained in the flask. The solid was extracted with hot toluene. A colorless product precipitated from the toluene extract upon cooling. The product was recrystallized from toluene five times to yield 4 g. (36% of theory) of colorless

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Figure 1. The effect of the number of trials on the number of correct responses. The number of correct responses was significantly higher than the number of incorrect responses in all cases. The number of correct responses was significantly higher than the number of incorrect responses in all cases. The number of correct responses was significantly higher than the number of incorrect responses in all cases.

crystals melting at 139-140° C.

Anal. Calc'd. for $C_{11}H_{15}N_5$: C, 60.80; H, 6.96;
N, 32.23

Found: C, 60.60; H, 7.07; N, 32.04

1-(4'-Phenyl-5'-dimethylaminomethyl-4'-H,1',2',4'-
triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-dimethylaminomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with 50 ml. of ligroin, and recrystallized twice from methanol to yield 0.7 g. (87% of theory) of product melting at 182-183° C.

Anal. Calc'd. for $C_{18}H_{20}N_6S$: N, 23.85; S, 9.10

Found: N, 23.65; S, 8.96

3-Amino-4-phenyl-5-diethylaminomethyl-4-H,1,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of absolute ethanol was placed in an Erlenmeyer flask and cooled. It was then treated with 62 ml. (0.6 mole) of diethylamine. Twenty-five milliliters of methanol was added; cooling was discontinued and the flask was stoppered and permitted to stand for sixty-eight hours. A solution of 4 g. (0.1 mole) of sodium hydroxide in

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25 ml. of water was diluted with 50 ml. of methanol and then added to the reaction mixture. The sodium chloride which precipitated was removed by filtration and the excess solvent and liberated diethylamine were removed under reduced pressure. A pinkish-white solid remained in the flask. The solid was extracted with hot toluene. A colorless product precipitated from the toluene extract upon cooling. It was recrystallized three times from toluene and melted at 148-150° C. The yield of product was 8 g. (75% of theory).

Anal. Calc'd. for $C_{13}H_{19}N_5$: C, 63.65; H, 7.81;
N, 28.55

Found: C, 63.65; H, 8.00; N, 28.40

1-(4'-Phenyl-5'-diethylaminomethyl-4'-H,1',2',4'-
triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-diethylaminomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with ligroin, and recrystallized twice from methanol to yield 0.7 g. (90% of theory) of product melting at 171-172° C.

Anal. Calc'd. for $C_{20}H_{24}N_6S$: N, 22.09; S, 8.43

Found: N, 21.90; S, 8.22

3-Amino-4-phenyl-5-diallylaminomethyl-4-H,1,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol was placed in an Erlenmeyer flask and treated with 29 g. (38 ml., 0.3 mole) of diallylamine. After the initial reaction had subsided, the flask was stoppered and permitted to stand for seventy-two hours. The reaction mixture was then treated with a solution of 4 g. (0.1 mole) of sodium hydroxide in 10 ml. of water and 50 ml. of methanol. The sodium chloride which precipitated was removed by filtration and the filtrate evaporated to dryness under reduced pressure. A yellow solid remained in the flask. The solid was recrystallized five times from ligroin to yield 3.4 g. (25% of theory) of product as yellow crystals melting at 92-93° C.

Anal. Calc'd. for $C_{15}H_{19}N_5$: C, 66.88; H, 7.11;
N, 26.00

Found: C, 66.56; H, 7.37; N, 26.16

1-(4'-Phenyl-5'-diallylaminomethyl-4'-H,1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-diallylaminomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration,

washed with ligroin and recrystallized twice from methanol to yield 0.5 g. (71% of theory) of product melting at 144-145° C.

Anal. Calc'd. for $C_{22}H_{24}N_6S$: N, 20.78; S, 7.93

Found: N, 20.76; S, 7.95

3-Amino-4-phenyl-5-di-n-propylaminomethyl-4-H.1,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol was placed in an Erlenmeyer flask and treated with 15 g. (20.4 ml., 0.15 mole) of di-n-propylamine. After the initial reaction had subsided, the flask was stoppered and permitted to stand for seventy-two hours. The reaction mixture was then treated with a solution of 4 g. (0.1 mole) of sodium hydroxide in 10 ml. of water and 50 ml. of methanol. The sodium chloride which precipitated was removed by filtration and the filtrate was evaporated to dryness under reduced pressure. The yellow solid that remained in the flask was recrystallized four times from toluene and twice from an ethanol-water solvent pair to yield 4 g. (26% of theory) of faintly yellow crystals melting at 142-143° C.

Anal. Calc'd. for $C_{15}H_{23}N_5$: C, 65.90; H, 8.48;

N, 25.62

Found: C, 65.72; H, 8.60; N, 25.45

3-Amino-4-phenyl-2-di-n-propylaminomethyl-5-H-I,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-2-chloromethyl-1,2,4-triazole in 50 ml. of methanol was placed in an Erlenmeyer flask and treated with 15 g. (20.4 ml., 0.15 mole) of di-n-propylamine. After the initial reaction had subsided, the mixture was stirred for 24 hours. The mixture was then poured into 100 ml. of water and the solid was removed by filtration and the filtrate was evaporated to dryness under reduced pressure. The residue was recrystallized from benzene and twice from an ethanol-water mixture. Yield 3.5 g. (28.6%).

1-(4'-Phenyl-5'-di-n-propylaminomethyl-4'-H,1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-di-n-propylaminomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with ligroin, and recrystallized twice from methanol to yield 0.6 g. (82% of theory) of product melting at 157-158° C.

Anal. Calc'd. for $C_{22}H_{28}N_6S$: N, 20.58; S, 7.85

Found: N, 20.46; S, 7.68

3-Amino-4-phenyl-5-di-isopropylaminomethyl-4-H,1,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol was placed in an Erlenmeyer flask and treated with 15 g. (21 ml., 0.15 mole) of di-isopropylamine. After the initial reaction had subsided, the flask was stoppered and permitted to stand for one week. The reaction mixture was then treated with a solution of 4 g. (0.1 mole) of sodium hydroxide in 10 ml. of water and 50 ml. of methanol. The sodium chloride which precipitated was removed by filtration and the filtrate evaporated to dryness under reduced pressure. The yellow solid which remained in the flask was recrystallized four times from

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toluene and twice from methanol to yield 4 g. (26% of theory) of faintly yellow crystals melting at 192-193° C.

Anal. Calc'd. for $C_{15}H_{23}N_5$: C, 65.90; H, 8.48;
N, 25.62

Found: C, 65.71; H, 8.45; N, 25.92

1-(4'-Phenyl-5'-di-isopropylaminomethyl-4'-H,1',2',4'-
triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-di-isopropylaminomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with ligroin and recrystallized twice from methanol to yield 0.66 g. (90% of theory) of product melting at 170-171° C.

Anal. Calc'd. for $C_{22}H_{28}N_6S$: N, 20.58; S, 7.85

Found: N, 20.64; S, 7.79

3-Amino-4-phenyl-5-di-n-butylaminomethyl-4-H,1,2,4-
triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of absolute methanol was placed in an Erlenmeyer flask and treated with 19.35 g., 26 ml. (0.15 mole) of di-n-butylamine. After the initial reaction had subsided,

the flask was stoppered and permitted to stand for seventy-two hours. The reaction mixture was then treated with a solution of 4 g. (0.1 mole) of sodium hydroxide in 10 ml. of water and 50 ml. of methanol. The sodium chloride which precipitated was removed by filtration and the filtrate was evaporated to dryness under reduced pressure. The yellow solid which remained in the flask was extracted with hot toluene. A colorless product precipitated from the toluene extract upon cooling. The yield of product was 13.5 g. (90% of theory). A small sample was recrystallized three times from toluene and melted at 135-136° C.

Anal. Calc'd. for $C_{17}H_{27}N_5$: C, 67.74; H, 9.03;
N, 23.24

Found: C, 67.48; H, 9.22; N, 23.07

1-(4'-Phenyl-5'-di-n-butylaminomethyl-4'-H,1',2',4'-
triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-di-n-butylaminomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with ligroin and recrystallized twice from methanol to yield 0.63 g. (90% of theory) of product melting at 129-130° C.

Anal. Calc'd. for $C_{24}H_{32}N_6S$: N, 19.25; S, 7.34

Found: N, 19.12; S, 7.36

• The first part of the document discusses the importance of maintaining accurate records of all transactions and the role of the accounting department in ensuring the integrity of the financial statements.

• The second part of the document outlines the various methods used to collect and analyze data, including the use of statistical techniques and the application of mathematical models.

• The third part of the document describes the results of the study, highlighting the key findings and the implications for the field of accounting.

• The fourth part of the document discusses the limitations of the study and the need for further research in this area.

• The fifth part of the document provides a conclusion and a summary of the main points discussed throughout the document.

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3-Amino-4-phenyl-5-di-isobutylaminomethyl-4-H.1,2,4-triazole

A solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of absolute methanol was placed in an Erlenmeyer flask and treated with 19.35 g., 26 ml. (0.15 mole) of diisobutylamine. After the initial reaction had subsided, the flask was stoppered and permitted to stand for five days. The reaction mixture was then treated with a solution of 4 g. (0.1 mole) of sodium hydroxide in 10 ml. of water and 50 ml. of methanol. The sodium chloride which precipitated was removed by filtration and the filtrate evaporated to dryness under reduced pressure. The yellow solid which remained in the flask was extracted with hot toluene. A colorless product precipitated from the toluene extract upon cooling. The yield of product was 10 g. (66% of theory). A small sample was recrystallized three times from toluene and melted at 189-191° C.

Anal. Calc'd. for $C_{17}H_{27}N_5$: C, 67.74; H, 9.03;

N, 23.24

Found: C, 67.88; H, 9.03; N, 23.19

1-(4'-Phenyl-5'-di-isobutylaminomethyl-4'-H.1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-di-isobutylaminomethyl-1,2,4-triazole and 2 ml. of phenylisothiocyanate was heated on the steam bath for one hour and then cooled in an ice bath. The flask was

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Figure 1. The effect of the number of trials on the number of correct responses. The number of correct responses was significantly higher than the number of incorrect responses in all cases. The number of correct responses was significantly higher than the number of incorrect responses in all cases. The number of correct responses was significantly higher than the number of incorrect responses in all cases.

scratched until precipitation of the product occurred. The phenylthiourea derivative was removed by filtration, washed with ligroin and recrystallized twice from methanol to yield 0.6 g. (85% of theory) of product melting at 168-169° C.

Anal. Calc'd. for $C_{24}H_{32}N_6S$: N, 19.25; S, 7.34

Found: N, 19.13; S, 7.50

3-Amino-4-phenyl-5-phenoxyethyl-1,2,4-triazole

A solution of 4.7 g. (0.05 mole) of phenol in 25 ml. of methanol was treated with a solution of 4.5 g. (0.11 mole) of sodium hydroxide in 10 ml. of water. This solution of sodium phenoxide was then added to a solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for twenty-four hours. The precipitated product was removed by filtration, and additional crystals were obtained by adding water to the filtrate. Both crops of crystals were combined and washed three times with 200 ml. of water. The product was recrystallized four times from toluene to yield 8 g. (60% of theory) of colorless crystals melting at 195-196° C. The product obtained in this reaction was identical with that obtained from the reaction of phenoxyacetic acid with phenylaminoguanidinium bisulfate as shown by a mixture melting point determination.

• The first step in the process of creating a new product is to identify a market need. This is often done through market research, which involves gathering information about the target market and its needs. Once a market need has been identified, the next step is to develop a concept for a product that meets that need. This is often done through brainstorming and prototyping. Once a concept has been developed, the next step is to create a business plan. This involves determining the costs of production, the pricing strategy, and the marketing strategy. Once a business plan has been created, the next step is to secure funding. This can be done through a variety of methods, including bank loans, venture capital, and crowdfunding. Once funding has been secured, the next step is to manufacture the product. This involves sourcing materials, hiring workers, and setting up a production line. Once the product has been manufactured, the next step is to distribute it. This can be done through a variety of methods, including retail stores, online marketplaces, and direct sales.

• The second step in the process of creating a new product is to develop a concept for a product that meets that need. This is often done through brainstorming and prototyping. Once a concept has been developed, the next step is to create a business plan. This involves determining the costs of production, the pricing strategy, and the marketing strategy. Once a business plan has been created, the next step is to secure funding. This can be done through a variety of methods, including bank loans, venture capital, and crowdfunding. Once funding has been secured, the next step is to manufacture the product. This involves sourcing materials, hiring workers, and setting up a production line. Once the product has been manufactured, the next step is to distribute it. This can be done through a variety of methods, including retail stores, online marketplaces, and direct sales. The third step in the process of creating a new product is to create a business plan. This involves determining the costs of production, the pricing strategy, and the marketing strategy. Once a business plan has been created, the next step is to secure funding. This can be done through a variety of methods, including bank loans, venture capital, and crowdfunding. Once funding has been secured, the next step is to manufacture the product. This involves sourcing materials, hiring workers, and setting up a production line. Once the product has been manufactured, the next step is to distribute it. This can be done through a variety of methods, including retail stores, online marketplaces, and direct sales. The fourth step in the process of creating a new product is to secure funding. This can be done through a variety of methods, including bank loans, venture capital, and crowdfunding. Once funding has been secured, the next step is to manufacture the product. This involves sourcing materials, hiring workers, and setting up a production line. Once the product has been manufactured, the next step is to distribute it. This can be done through a variety of methods, including retail stores, online marketplaces, and direct sales. The fifth step in the process of creating a new product is to manufacture the product. This involves sourcing materials, hiring workers, and setting up a production line. Once the product has been manufactured, the next step is to distribute it. This can be done through a variety of methods, including retail stores, online marketplaces, and direct sales. The sixth step in the process of creating a new product is to distribute it. This can be done through a variety of methods, including retail stores, online marketplaces, and direct sales.

3-Amino-4-phenyl-5-(p-bromophenoxymethyl)-4-H,1,2,4-triazole

A solution of 8.6 g. (0.05 mole) of p-bromophenol in 10 ml. of methanol was treated with a solution of 3 g. (0.075 mole) of sodium hydroxide in 6 ml. of water. This solution of sodium p-bromophenoxide was then added to a solution of 6 g. (0.025 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 25 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for forty hours. Precipitation of the product began within five minutes. The product was removed by filtration and additional product was obtained by adding water to the filtrate. Both crops of crystals were combined and washed three times with 200 ml. of water. The product was then recrystallized three times from methanol to yield 3 g. (35% of theory) of colorless crystals melting at 182-183° C.

Anal. Calc'd. for $C_{15}H_{13}BrN_4O$: C, 52.19; H, 3.79;
Br, 23.15; N, 16.23

Found: C, 52.26; H, 3.82; Br, 23.24;
N, 16.28

1-[4'-Phenyl-5'-(p-bromophenoxymethyl)-4'-H,1',2',4'-triazolyl-3'7-3-phenylthiourea

A mixture of 1 g. (0.003 mole) of 3-amino-4-phenyl-5-(p-bromophenoxymethyl)-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for two hours. It was then cooled in an ice bath and treated

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with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 1 g. (69% of theory) of product melting at 188-190° C.

Anal. Calc'd. for $C_{22}H_{18}BrN_5O_3$: Br, 16.63;

N, 14.58; S, 6.67

Found: Br, 16.47; N, 14.80; S, 6.67

3-Amino-4-phenyl-5-(p-chlorophenoxymethyl)-4-H.1,2,4-triazole

A solution of 6.5 g. (0.05 mole) of p-chlorophenol in 25 ml. of methanol was treated with a solution of 5 g. (0.125 mole) of sodium hydroxide in 10 ml. of water. This solution of sodium p-chlorophenoxide was then added to a solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for one week. The precipitated product was removed by filtration. Additional crystals were obtained by adding water to the filtrate. Both crops of crystals were combined and washed three times with 200 ml. of water. The product was recrystallized three times from methanol to yield 6 g. (55% of theory) of colorless crystals melting at 197-198° C.

Anal. Calc'd. for $C_{15}H_{13}ClN_4O$: C, 59.90;

H, 4.35; Cl, 11.79; N, 18.63

Found: C, 59.81; H, 4.43; Cl, 12.08; N, 18.86

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Figure 1. The effect of the number of trials on the number of correct responses. The number of correct responses was significantly higher than the number of incorrect responses for all conditions. The number of correct responses was significantly higher than the number of incorrect responses for all conditions. The number of correct responses was significantly higher than the number of incorrect responses for all conditions.

Journal of Management Studies, 20(6), 791-807.

[illegible]

1. *Journal of the American Medical Association*, 1997; 277: 1033-1037.

1-[4'-Phenyl-5'-(p-chlorophenoxy)methyl]-4'-H,1',2',4'-
triazolyl-3']-3-phenylthiourea

A mixture of 1 g. (0.003 mole) of 3-amino-4-phenyl-5-(p-chlorophenoxy)methyl-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.6 g. (44% of theory) of product melting at 185-186° C.

Anal. Calc'd. for $C_{22}H_{18}ClN_5OS$: Cl, 8.13;

N, 16.07; S, 7.36

Found: Cl, 8.03; N, 16.22; S, 7.28

3-Amino-4-phenyl-5-(p-methoxyphenoxy)methyl-4-H,1,2,4-
triazole

A solution of 7 g. (0.056 mole) of p-methoxyphenol in 25 ml. of methanol was treated with a solution of 5 g. (0.125 mole) of sodium hydroxide in 10 ml. of water. This solution of sodium p-methoxyphenoxide was then added to a solution of 12.25 g. (0.05 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 50 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for twenty-four hours. The solution became intensely red and after five hours beautiful, colorless needles precipitated. The precipitated

product was removed by filtration and additional crystals were obtained by adding water to the filtrate. Both crops of crystals were combined and washed three times with 200 ml. of water. The product was recrystallized three times from methanol to yield 9 g. (60% of theory) of colorless needles melting at 179-180° C.

Anal. Calc'd. for $C_{16}H_{16}N_4O_2$: C, 64.85; H, 5.44;
N, 18.91.

Found: C, 64.82; H, 5.42; N, 18.85

1-[4'-Phenyl-5'-(p-methoxyphenoxyethyl)-4'-H,1',2',4'-
triazolyl-3']-3-phenylthiourea

A mixture of 1 g. (0.003 mole) of 3-amino-4-phenyl-5-(p-methoxyphenoxyethyl)-4-H,1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.7 g. (50% of theory) of product melting at 177-178° C.

Anal. Calc'd. for $C_{23}H_{21}N_5O_2S$: N, 16.23; S, 7.43

Found: N, 16.43; S, 7.11

Figure 1. The effect of the concentration of the *Agrobacterium* suspension on the transformation efficiency of *Agrobacterium* strains. The *Agrobacterium* strains were grown in YEA medium for 24 h at 28 °C. The cell concentration of the strains was adjusted to 10⁸ cells/ml. The cell suspension was then diluted to 10⁶, 10⁷, 10⁸, 10⁹, and 10¹⁰ cells/ml. The cell suspension was then inoculated into the plant tissue. The transformation efficiency was determined by the number of transformants per 10⁶ cells. The data were expressed as the mean ± SD of three independent experiments. The asterisk (*) indicates a significant difference (p < 0.05) between the control and the treatment.

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3-Amino-4-phenyl-5-(p-methylphenoxymethyl)-4-H,1,2,4-triazole

A solution of 5.4 g. (5.4 ml., 0.05 mole) of p-cresol in 10 ml. of methanol was treated with a solution of 3 g. (0.075 mole) of sodium hydroxide in 6 ml. of water. This solution of sodium p-methylphenoxide was then added to a solution of 6 g. (0.025 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 25 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for twenty-four hours. Precipitation of the product began within five minutes. The precipitated product was removed by filtration and additional crystals were obtained by adding water to the filtrate. Both crops of crystals were combined and washed three times with 200 ml. of water. The product was recrystallized four times from methanol to yield 5 g. (71% of theory) of colorless crystals melting at 163-169° C.

Anal. Calc'd. for $C_{16}H_{16}N_4O$: C, 68.55; H, 5.75; N, 19.99

Found: C, 68.41; H, 5.92; N, 19.99

1-[4'-Phenyl-5'-(p-methylphenoxymethyl)-4'-H,1',2',4'-triazolyl-3']-3-phenylthiourea

A mixture of 1 g. (0.0035 mole) of 3-amino-4-phenyl-5-(p-methylphenoxymethyl)-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for

two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.7 g. (50% of theory) of product melting at 168-169° C.

Anal. Calc'd. for $C_{23}H_{21}N_5OS$: N, 16.85; S, 7.71

Found: N, 17.07; S, 7.56

3-Amino-4-phenyl-5-(o-methylphenoxymethyl)-4-H,1,2,4-triazole

A solution of 5.4 g. (5.4 ml., 0.05 mole) of o-cresol in 10 ml. of methanol was treated with a solution of 3 g. (0.075 mole) of sodium hydroxide in 6 ml. of water. This solution of sodium o-methylphenoxide was then added to a solution of 6 g. (0.025 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 25 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for twenty-four hours. Precipitation of the product began within thirty minutes. Finally the mixture was heated for thirty minutes on the steam bath, cooled and treated with water. The colorless, crystalline product was removed by filtration and recrystallized three times from ethanol to yield 2 g. (30% of theory) of colorless crystals melting at 186-187° C.

Anal. Calc'd. for $C_{16}H_{16}N_4O$: C, 68.55; H, 5.75;

N, 19.99

Found: C, 68.61; H, 5.76; N, 19.87

• The first step in the process of creating a new product is to identify a market need. This can be done through market research, which involves gathering information about the target market and its needs. Once a market need has been identified, the next step is to develop a concept for a new product that meets this need. This concept should be based on the market research and should be unique and innovative. The concept should also be feasible, meaning that it can be developed and marketed within the company's resources and capabilities. Once the concept has been developed, the next step is to create a prototype of the product. This prototype should be used to test the concept and to gather feedback from potential customers. Finally, once the concept has been tested and refined, the product can be developed and marketed to the target market.

• The second step in the process of creating a new product is to develop a business plan. This plan should outline the company's goals, objectives, and strategies for developing and marketing the new product. It should also include a detailed financial plan, which shows the expected costs and revenues of the product. The business plan is a critical document that provides a roadmap for the company's development and marketing efforts. It is also a key tool for securing financing from investors and lenders. Once the business plan has been developed, the next step is to secure financing for the product. This can be done through a variety of sources, including venture capitalists, angel investors, and banks. Once financing has been secured, the next step is to develop a marketing plan for the product. This plan should outline the company's strategies for promoting the product and reaching its target market. It should also include a detailed budget for the marketing efforts. Finally, once the marketing plan has been developed, the product can be launched into the market. The company should continue to monitor the product's performance and make adjustments as needed to ensure its success.

• The third step in the process of creating a new product is to launch the product into the market.

1-[4'-Phenyl-5'-(o-methylphenoxyethyl)-4'-H,1',2',4'-
triazolyl-3']-3-phenylthiourea

A mixture of 1 g. (0.0035 mole) of 3-amino-4-phenyl-5-(o-methylphenoxyethyl)-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.6 g. (43% of theory) of product melting at 182-183° C.

Anal. Calc'd. for $C_{23}H_{21}N_5OS$: N, 16.85; S, 7.71

Found: N, 17.00; S, 7.41

3-Amino-4-phenyl-5-(m-methylphenoxyethyl)-4-H,1,2,4-
triazole

A solution of 5.4 g. (5.4 ml., 0.05 mole) of m-cresol in 10 ml. of methanol was treated with a solution of 3 g. (0.075 mole) of sodium hydroxide in 6 ml. of water. This solution of sodium m-methylphenoxide was then added to a solution of 6 g. (0.025 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 25 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for forty-eight hours. The solution was then heated on the steam bath for thirty minutes, cooled and treated with water. An oil separated which crystallized upon cooling in an ice bath. The product

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1. The first step in the process of the scientific method is to ask a question.

2. The second step is to do background research.

- The third step is to form a hypothesis, which is a prediction about the outcome of the experiment.
- The fourth step is to design an experiment to test the hypothesis.
- The fifth step is to conduct the experiment and collect data.
- The sixth step is to analyze the data and draw conclusions.
- The seventh step is to communicate the results of the experiment.

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- The sixth step is to analyze the data and draw conclusions.
- The seventh step is to communicate the results of the experiment.

was recrystallized three times from toluene to yield 2 g. (30% of theory) of colorless needles melting at 155-157° C.

Anal. Calc'd. for $C_{16}H_{16}N_4O$: C, 68.55; H, 5.75; N, 19.99

Found: C, 68.55; H, 5.90; N, 20.23

1-[4'-Phenyl-5'-(m-methylphenoxyethyl)-4'-H,1',2',4'-triazolyl-3']-3-phenylthiourea

A mixture of 1 g. (0.0035 mole) of 3-amino-4-phenyl-5-(m-methylphenoxyethyl)-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.5 g. (35% of theory) of product melting at 166-167° C.

Anal. Calc'd. for $C_{23}H_{21}N_5OS$: N, 16.85; S, 7.71

Found: N, 17.15; S, 7.53

3-Amino-4-phenyl-5-thiophenoxyethyl-4-H,1,2,4-triazole

A solution of 5 g. (5 ml., 0.05 mole) of thiophenol in 10 ml. of methanol was treated with a solution of 3 g. (0.075 mole) of sodium hydroxide in 6 ml. of water. This solution of sodium thiophenoxide was then added to a solution of 6 g. (0.025 mole) of the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in 25 ml. of

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• The second step is the development of the (10-11)
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• The ninth step is the development of the (24-25)
• The tenth step is the development of the (26-27)

• The eleventh step is the development of the (28-29)
• The twelfth step is the development of the (30-31)

• The thirteenth step is the development of the (32-33)
• The fourteenth step is the development of the (34-35)

• The fifteenth step is the development of the (36-37)
• The sixteenth step is the development of the (38-39)

• The seventeenth step is the development of the (40-41)

• The eighteenth step is the development of the (42-43)
• The nineteenth step is the development of the (44-45)

• The twentieth step is the development of the (46-47)
• The twenty-first step is the development of the (48-49)

• The twenty-second step is the development of the (50-51)
• The twenty-third step is the development of the (52-53)

• The twenty-fourth step is the development of the (54-55)

methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for seventy-two hours. The mixture was then heated on the steam bath for one hour, cooled and treated with 200 ml. of water. The product which precipitated was removed by filtration and washed with water. It was recrystallized three times from toluene to yield 4 g. (57% of theory) of colorless crystals melting at 130-132° C.

Anal. Calc'd. for $C_{15}H_{14}N_4S$: C, 63.80; H, 4.97;
N, 19.84; S, 11.35

Found: C, 63.72; H, 5.14; N, 19.86; S, 11.47

1-(4'-Phenyl-5'-thiophenoxymethyl-4'-H,1',2',4'-triazolyl-3')-3-phenylthiourea

A mixture of 1 g. (0.0035 mole) of 3-amino-4-phenyl-5-thiophenoxymethyl-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath for two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.5 g. (43% of theory) of product melting at 141-143° C.

Anal. Calc'd. for $C_{22}H_{19}N_5S_2$: N, 16.77; S, 15.36

Found: N, 16.70; S, 15.50

3-Amino-4-phenyl-5-(p-chlorothiophenoxymethyl)-4-H,1,2,4-triazole

A solution of 7.2 g. (0.05 mole) of p-chlorothiophenol in 25 ml. of methanol was treated with a solution of 3 g. (0.075 mole) of sodium hydroxide in 6 ml. of water. This solution of sodium p-chlorothiophenoxide was then added to a solution of 6 g. (0.025 mole) of the hydrochloride of 3-amino -4-phenyl-5-chloromethyl-1,2,4-triazole in 25 ml. of methanol. After the initial reaction had subsided, the flask was stoppered and permitted to stand for sixty hours. The solution was then heated on the steam bath for thirty minutes, cooled and treated with 200 ml. of water. An oil separated which crystallized upon cooling in an ice bath. The product was recrystallized three times from toluene to yield 4 g. (50% of theory) of colorless crystals melting at 121-122° C.

Anal. Calc'd. for $C_{15}H_{13}ClN_4S$: C, 56.86; H, 4.13;
Cl, 11.19; N, 17.69; S, 10.12;
Found: C, 57.13; H, 4.25; Cl, 11.22;
N, 17.90; S, 9.92

1-[4'-Phenyl-5'-(p-chlorothiophenoxymethyl)-4'-H,1',2',4'-triazolyl-3']-3-phenylthiourea

A mixture of 0.5 g. (0.002 mole) of 3-amino-4-phenyl-5-(p-chlorothiophenoxymethyl)-1,2,4-triazole and 3 ml. of phenylisothiocyanate was heated on the steam bath

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for two hours. It was then cooled in an ice bath and treated with 50 ml. of ligroin to precipitate the phenylthiourea derivative. The product was removed by filtration and recrystallized twice from methanol to yield 0.4 g. (57% of theory) of product melting at 167-168° C.

Anal. Calc'd. for $C_{22}H_{18}ClN_5S_2$: Cl, 7.84;

N, 15.50; S, 14.19

Found: Cl, 7.96; N, 15.52; S, 14.06

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- The first step in the process of creating a new product is to identify a market need. This involves conducting market research to determine what consumers want and what problems they are trying to solve.
- Once a market need is identified, the next step is to develop a concept for a product that addresses that need. This involves brainstorming ideas and creating a prototype.
- The third step is to conduct a feasibility study to determine if the product is viable. This involves assessing the technical, financial, and market feasibility of the product.
- If the feasibility study is positive, the next step is to develop a business plan. This involves outlining the marketing, sales, and distribution strategy for the product.
- The final step is to launch the product and monitor its performance. This involves tracking sales, customer feedback, and market trends to ensure the product is successful.

Summary

In the course of this investigation, three different series of compounds having varied substituents in the fifth position of 3-amino-4-phenyl-4-H,1,2,4-triazole were synthesized. The starting material for these syntheses, phenylaminoguanidinium bisulfate, was first prepared in the course of this work.

A procedure was perfected for the conversion of 3-amino-4-phenyl-5-hydroxymethyl-1,2,4-triazole to the hydrochloride of 3-amino-4-phenyl-5-chloromethyl-1,2,4-triazole in good yield. The latter was a required intermediate for the synthesis of the 5-dialkylamino-methyl-, the 5-phenoxy-methyl-, and the 5-thiophenoxy-methyltriazoles.

Phenylthiourea derivatives of most of these new aminotriazoles were prepared. In a few cases, a number of the hydrochlorides and acetyl derivatives were prepared.

References Cited

- (1) Dimroth, Ann. Chem., 364, 183 (1909)
- (2) Lieber, E., Chao, T.S. and Rao, C.N.R., J. Org. Chem., 22, 654 (1957)
- (3) Lieber, E., Rao, C.N.R. and Chao, T.S., J. Am. Chem. Soc., 79, 5962 (1957)
- (4) Garbrecht, W.L. and Herbst, R.M., J. Org. Chem., 18, 1269 (1953)
- (5) Miller, C.S. and Hall, W.C., Weeds, 5, 304 (1957)
- (6) Tschudy, D.P. and Collins, A., Science, 126, 168 (1957)
- (7) Shaw, W.A. and Swanson, C.R., Weeds, 2, 43 (1953)
- (8) Lewenstein, M.J., U.S. Pat., 2,683,106 (1954)
- (9) Bladin, J.A., Ber. deut. chem. Ges., 18, 1544 (1885)
- (10) Bladin, J.A., Ber. deut. chem. Ges., 25, 741 (1892)
- (11) Andreocci, A., Ber. deut. chem. Ges., 25, 225 (1892)
- (12) Thiele, J., Ann. Chem., 270, 1 (1892)
- (13) Thiele, J. and Manchot, W., Ann. Chem., 303, 33 (1898)
- (14) Arndt, F. and Tschenschner, F., Ber. deut. chem. Ges., 56, 1984 (1923)
- (15) Guha, P.C. and Mehta, D.R., Quart. J. Indian Inst. Sci., 1938, 21 (A), 42; C.A. 33, 598 (1939)
- (16) Pellizzari, G., Gazz. chim. ital., 1894, 24, I, 481
- (17) Meyer, V. and Jacobson, P., Lehrbuch der Organischen Chemie, Zweiter Band - Cyclische Verbindungen - Naturstoffe Dritter Teil - Heterocyclische Verbindungen, Walter DeGruyter & Co., Berlin (1923)
- (18) Grignard, V., Du Pont, G. and Locquin, R., Traite De Chemie Organique, chapter XXI, Masson et C., Paris (1953)

- (19) Rodd, F.H., Chemistry of Carbon Compounds, Vol. 4A, p. 452, Elsevier Publishing Co., Amsterdam (1957)
- (20) Lieber, E. and Smith, G.B.L., Chem. Rev., 25, 213 (1939)
- (21) Manchot, W. and Noll, R., Ann. Chem., 343, 1 (1905)
- (22) Reilly, J. and Drumm, P., J. Chem. Soc., 1926, 1729
- (23) Kirsten, G.W. and Smith, G.B.L., J. Am. Chem. Soc., 58, 800 (1936)
- (24) Finnegan, W.G., Henry, R.A. and Lieber, E., J. Org. Chem. 18, 779 (1953)
- (25) Thiele, J. and Heidenreich, K., Ber. deut. chem. Ges., 26, 2598 (1893)
- (26) Reilly, J. and Madden, D., J. Chem. Soc., 1929, 815
- (27) Bloom, A. and Day, A.R., J. Org. Chem., 4, 14 (1939)
- (28) Phillips, M.A., J. Chem. Soc., 2393 (1928)
- (29) Fromm, Ann. Chem., 447, 295 (1926)
- (30) Fantl, P. and Silbermann, H., Ann. Chem., 467, 278 (1928)
- (31) Hall, W.C., Truchelut, G.B. and Lane, H.C., Texas Agr. Exp. Sta. Bul. 759 (1953)
- (32) Allen, W.W., U. S. Pat., 2,670,282 (1954)
- (33) Heimbach, N. and Kelly, W., U. S. Pat., 2,449,225 and 2,449,226 (1948)
- (34) Sund, K.A., Chem. Eng. News, 33, 1508 (1955)
- (35) Rogers, B.J., Weeds, 5, 5 (1957)
- (36) Heim, W.G., Appleman, D. and Pyfrom, H.T., Science, 122, 693 (1955)

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