

and unsystematic. Similar results have been obtained for *p*-xylene and bibenzyl.

Discussion

For the six polar liquids so far tested, a significant discontinuity in the slope of the static dielectric constant-temperature curve has been found in the neighborhood of the melting point in each case. On the other hand, no such effect has been observed with any non-polar or slightly polar liquids, suggesting that the *magnitude* of the change in slope may be related—at least in a qualitative way—to the permanent dipole moment. In Table II are listed the differences in slope between the lines for the normal and supercooled liquids, together with the respective dipole moments and values of ϵ_0 at the melting point. Examination of the table does reveal a qualitative relationship between the change in slope $d\epsilon_0/dT$ and the dipole

TABLE II

| | Change in slope $d\epsilon_0/dT$ | Dipole moment, $\mu \times 10^{18}$ | ϵ_0 | $\epsilon_0 - n^2$ |
|-------------------------------|-------------------------------------|--|--------------|--------------------|
| Azoxybenzene | 0.0010 | 3.39 | 5.2 | 2.4 |
| Salol | .0019 | 3.15 | 6.4 | 3.6 |
| Methyl <i>n</i> -nonyl ketone | .0036 | 2.69 | 8.3 | 6.3 |
| <i>p</i> -Bromotoluene | .0031 | 1.90 | 6.0 | 3.6 |
| Menthol | .0009 | 1.58 | 4.0 | 1.8 |
| Phenyl ether | .0006 | 1.15 | 3.6 | 1.1 |
| Diphenylmethane | .0 | <0.3 | 2.49 | 0.01 |
| Bibenzyl | .0 | <.3 | 2.42 | .05 |
| <i>p</i> -Xylene | .0 | <.2 | 2.23 | .00 |

moment for the four liquids of smallest dipole moment. This relationship does not hold, however, for azoxybenzene and salol, and one reason for this disagreement may be the existence of dipolar interaction or association which would be expected to have a strong influence on any process that might occur at the melting point. Thus it seems more legitimate to compare the magnitude of the

change in slope (change in $d\epsilon_0/dT$) with ϵ_0 itself, since ϵ_0 is a macroscopic measure of the polarization produced in the pure liquid, which includes the effects of dipolar interaction. If n^2 is regarded as the contribution toward the dielectric constant from electronic and atomic polarization, n being the index of refraction, then $(\epsilon_0 - n^2)$ is that part arising only from the orientation of permanent molecular dipoles. In Fig. 3, change in $d\epsilon_0/dT$ is shown plotted against $(\epsilon_0 - n^2)$ for each liquid tested and now the points are seen to lie reasonably well on a line through the origin.

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N-Trifluoroacetyl Derivatives of Carcinogenic Amines¹

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N-Acetyl derivatives of 2-aminofluorene,² 2,7-diaminofluorene,³ 2-naphthylamine,⁴ 4-aminobiphenyl,⁵ 3-aminodibenzofuran⁶ and 3-aminodibenzothiophene⁷ have been shown to have carcinogenic properties. The free amines, 2-anthramine,⁷ benzdine⁸ and 2',3-dimethyl-4-aminoazobenzene,⁹ have also been shown to be active.

In the fluorene series it has been shown that 2-aminofluorene is carcinogenic in the rat.¹⁰ The carcinogenic N-acetyl derivative is easily hydrolyzed *in vivo*.¹¹ On the other hand, 2-benzoylaminofluorene has little activity¹² and cannot be hydrolyzed readily.¹³ Similarly the non-carcinogenic¹² 2-*p*-tosylaminofluorene is stated to be very slightly hydrolyzed *in vivo*.¹⁴ On the basis of these facts it is probable that the free amines or their metabolites are the primary carcinogens. To test this hypothesis the N-trifluoroacetyl derivatives have been prepared. According to this hypothesis if the trifluoroacetyl derivatives cannot be hydrolyzed *in vivo*, then they should be non-carcinogenic. If these groups are hydrolyzed, then the compounds should be carcinogenic.

The physical properties and yields of the compounds are given in Table I.

General Procedure.—To 0.01 mole of the amine (or 0.005 mole of the diamine) in 20 ml. of hot benzene was carefully added through a reflux condenser 0.01 mole of tri-

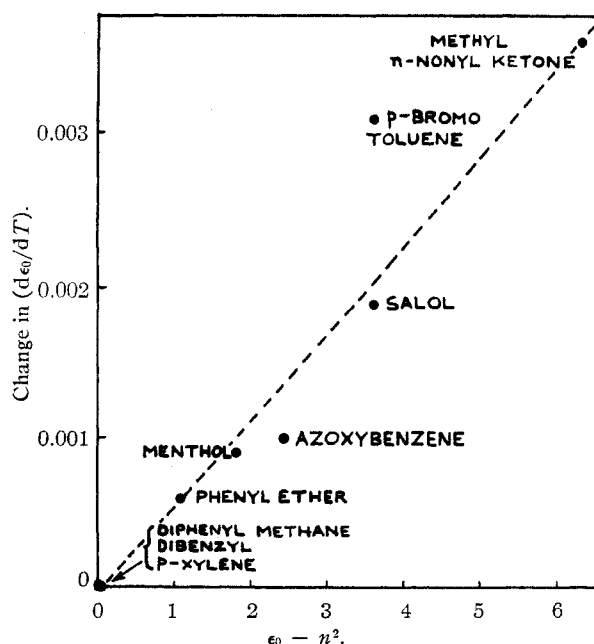


Fig. 3.—Correlation of change in $d\epsilon_0/dT$ with $(\epsilon_0 - n^2)$.

- (1) The work described in this paper was supported by a grant from the Sloan-Kettering Institute for Cancer Research.
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- (4) G. Bonser, *J. Path. Bact.*, **55**, 1 (1943).
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TABLE I

| Compound | M.p., ^a °C. | Yield, % | Nitrogen, % | | Sulfur, % | | Fluorine, % | |
|-------------------------------|------------------------|-------------|-------------|-------|-----------|-------|-------------|-------|
| | | | Calcd. | Found | Calcd. | Found | Calcd. | Found |
| 4-TFA ^b biphenyl | 200–201 | 95 | 5.28 | 5.42 | | | | |
| 4,4'-Di-TFA biphenyl | 324–325 | 98 | 7.45 | 7.13 | | | | |
| 2-TFA fluorene | 200–201 | 98 | | | | | 20.6 | 20.1 |
| 2,7-Di-TFA fluorene | 300–301 | 92 | 7.2 | 7.4 | | | | |
| 3-TFA dibenzofuran | 204–205 | 97 | 5.02 | 5.10 | | | | |
| 3-TFA dibenzothiophene | 186–187 | 98 | | | 10.85 | 10.7 | | |
| 2-TFA naphthalene | 146–147 | 95 | 5.86 | 5.58 | | | | |
| 2-TFA anthracene ^c | 245–246 | 87 | | | | | | |
| 4-TFA-2',3-dimethylazobenzene | 158–159 | 90 | 13.1 | 12.8 | | | | |

^a All melting points are uncorrected. ^b TFA = Trifluoroacetyl amino. ^c Anal. Calcd. for C₁₆H₁₀F₃NO: C, 66.44; H, 3.46. Found: C, 66.32; H, 3.82.

fluoroacetic anhydride.¹⁵ In the case of 2,7-diaminofluorene 100 ml. of hot xylene was used as the solvent. After the initial reaction had subsided, the mixture was refluxed for half an hour. The cooled mass was filtered and washed with 10 ml. of cold benzene. Further product was obtained by evaporating the mother liquid under reduced pressure at room temperature. The fairly soluble acyl derivative of 2-naphthylamine was obtained by a similar evaporation of the reaction mixture. The diamine derivatives were crystallized from alcohol. The azo dye and the 4-aminobiphenyl derivative were crystallized from heptane. The 2-naphthylamine derivative was crystallized from aqueous alcohol while 2-trifluoroacetylaminofluorene was crystallized from benzene. These derivatives are more soluble than the analogous acetyl derivatives in non-polar solvents, such as heptane, and polar solvents, such as alcohol. The strong electronegative trifluoroacetyl group confers acidic properties on the new molecules for they are all soluble to some extent in aqueous sodium hydroxide solution.

(15) Minnesota Mining and Manufacturing Co., St. Paul 6, Minnesota.

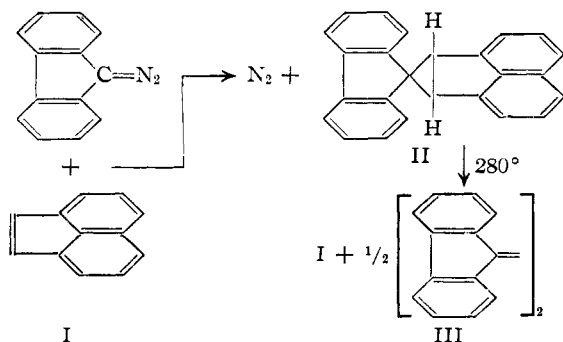
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Formation of a Spiro Compound by the Action of Diazofluorene on Acenaphthylene

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Though the action of diazomethanes on ethylenic compounds leading to three-membered rings has been observed frequently,¹ very little seems to be known about the formation of spiro products by this method. When acenaphthylene was treated with diazofluorene, the spiro product (II) was formed in good yield; in contrast with its generators and in agreement with the proposed formula,



(1) H. Wieland and O. Probst, *Ann.*, **530**, 277 (1937); P. C. Guha and S. Krishnamurthy, *Ber.*, **70**, 2113 (1937); W. Braren and E. Buchner, *ibid.*, **33**, 684, 3453 (1900).

it is colorless. There seems to be some strain in (II) since it decomposes at 280° into acenaphthylene and Δ^{9,9'}-bifluorene (III).

When acenaphthalene (I) was treated with an ethereal solution of diazomethane, it was recovered.

Experimental

Diazofluorene and Acenaphthylene.—A solution of 0.5 g. of acenaphthylene and 0.7 g. of 9-diazofluorene in 15 ml. of dry benzene was refluxed for 30 minutes. The mixture was allowed to stand overnight, concentrated and a few ml. of acetone was added to the residue. On standing, crystals separated which were recrystallized from benzene. 7,8-9',9'-Fluorenyleneacenaphthene (II) is colorless, m.p. 275° (red-brown melt), yield about 80%.

Anal. Calcd. for C₂₆H₁₆: C, 94.9; H, 5.0; mol. wt., 316. Found: C, 94.4; H, 5.1; mol. wt. (micro-Rast), 282.

Thermal Decomposition.—A 0.5-g. sample of II was heated for one hour at 280–300° (bath temp.) in a test-tube shaped vessel (Pyrex glass) which was connected during pyrolysis to a working oil vacuum pump. The reaction vessel was then allowed to cool in a vacuum; the oily drops on the upper part of the tube solidified and proved to be acenaphthylene (m.p. and mixed m.p.). The bottom of the reaction vessel contained a red substance which was extracted several times with boiling ether. After concentration of the ethereal extract to about 1 ml., light petroleum (b.p. 40–60°) was added dropwise at room temperature to turbidity. A reddish-brown substance separated and was dissolved in a very small amount of hot benzene. The addition of a saturated benzene solution of picric acid yielded orange crystals overnight; these proved to be the picrate of Δ^{9,9'}-bifluorene (III) (m.p. and mixed m.p.).

Diazomethane and Acenaphthylene.—An ethereal solution containing 1 g. of acenaphthylene and diazomethane² (prepared from 10 g. of nitrosomethylurea) was allowed to stand for 24 hours in the ice-chest. Acenaphthylene was recovered unchanged.

(2) *Org. Syntheses*, **15**, 3 (1935).

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σ-Constants of the Carboethoxyl and Hydroxyl Groups

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As part of another investigation,² we have had occasion to determine the σ-constants³ of the *m*- and *p*-carboethoxyl and *m*-hydroxyl groups. The

(1) U. S. Atomic Energy Commission Predoctoral Fellow, 1951–1952.

(2) J. D. Roberts and W. T. Moreland, Jr., *THIS JOURNAL*, **75**, 2167 (1953).

(3) L. P. Hammett, "Physical Organic Chemistry," McGraw-Hill Book Co., Inc., New York, N. Y., 1940, Chap. VII.