

ported to possess appreciable hypoglycemic activity. Based on these observations, several indanamides like 1-*N*-alkylacetamidoindans⁴ and 3-oxo-1-*N*-alkylacetamidoindans have been synthesized to evaluate their hypoglycemic activity. None of these compounds, however, possessed any hypoglycemic activity.

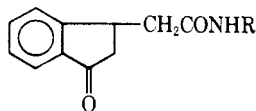
Experimental Section⁵

Methyl 3-Oxoindan-1-acetate.—3-Oxoindan-1-acetic acid⁶ (27 g) was esterified with dry MeOH (90 ml) in the presence of dry HCl (6 g) by refluxing on a steam bath for 8 hr. The crude ester was crystd from EtOAc-petr ether (bp 40–60°) in 90% yield, mp 67–68°. *Anal.* (C₁₂H₁₂O₃) C, H.

3-Oxo-1-*N*-alkylacetamidoindan. **A.**—A mixt of methyl 3-oxoindan-1-acetate (1 mole) and the appropriate alkylamine (2 moles) was heated in a sealed tube on steam bath for 6 hr. The reaction mass was poured into H₂O, acidified with 2 *N* HCl, either filtered or extd (PhH), and washed (H₂O). The crude product was crystd from PhH-petr ether (bp 40–60°) as shining crystals.

b.—SOCl₂ (5 ml) was added dropwise to a mixt of 3-oxoindan-1-acetic acid⁶ (3 g) and dry PhH (120 ml) with stirring till the evoln of HCl ceased. Approx 90 ml of PhH was distd off and the residual mass (3-oxoindan-1-acetyl chloride) was cooled in ice water. The cooled soln of 3-oxoindan-1-acetyl chloride (1 mole) was added dropwise under stirring to a soln of alkylamines (2.5 moles) in PhH (40 ml) with the simultaneous addn of 2 *N* NaOH to keep the mass alk. After stirring for 2 hr it was either filtered or extd (PhH), washed (H₂O), and purified by crystn from PhH-petr ether (bp 40–60°) as shining crystals (see Table I).

TABLE I
3-Oxo-1-*N*-alkylacetamidoindans



R	Mp, °C	Empirical formula ^c
Me ^a	144–146	C ₁₂ H ₁₃ O ₂ N
Et ^b	120–121	C ₁₃ H ₁₅ O ₂ N
<i>n</i> -Pr ^b	116–118	C ₁₄ H ₁₇ O ₂ N
<i>n</i> -Bu ^b	97–98	C ₁₅ H ₁₉ O ₂ N

^a Prep'd from ester. ^b Prep'd from acid chloride. ^c *Anal.* C, H, N.

Acknowledgment.—The authors' thanks are due to Bristol Laboratories, Syracuse, N. Y., for the hypoglycemic test report.

(4) A. U. De and B. Pathak, *J. Med. Chem.*, **13**, 152 (1970).

(5) Analytical results were within ±0.4% of the theoretical values. All melting points are uncorrected.

(6) R. H. Manske, *J. Amer. Chem. Soc.*, **53**, 1104 (1931).

Anti-*Trichinella spiralis* Activity of Some 1-Carbamoyl-3-methyl-2-pyrazolin-4,5-dione 4-Arylhydrazones

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Heterocyclic compounds containing a carbamoyl group have been reported to possess various activities¹ due to their ability to inhibit acetylcholinesterase,

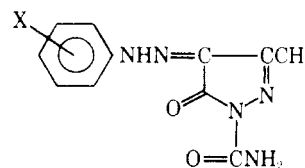
(1) I. T. Kay, D. J. Lovejoy, and S. Glue, *J. Chem. Soc.*, 445 (1970).

probably by the transfer of a carbamoyl group to an active site of the enzyme. This report includes the potencies against *Trichinella spiralis* of several 1-carbamoyl-3-methyl-2-pyrazolin-4,5-dione 4-arylhydrazones which were described earlier in connection with our work on potential antidiabetics.²

The compounds were prepared as described previously^{2,3} and were tested in mice and have shown the order of decreasing potency listed in Table I.

TABLE I

ANTI-*Trichinella* ACTIVITY^a



No.	X	Mp, °C	Mean worm count Control	Drug	% reduction ^a
1	2-Cl-4-NO ₂	210 ^b	396	326	17.7
2	2,5-Cl ₂	258–259 ^c	396	388	2.0
3	2-Cl-6-Me	226 ^c	396	394	0.5
4	4-NO ₂	257–258 ^c	495	536	0
5	2,6-Cl ₂	200 ^c	396	403	0

^a Drug administration was po in Charles River Mice. Compound effectiveness was calcd as a percentage reduction based on the following formula. % reduction = 100 - [(Mean of medicated group worm count)/(mean of unmedicated control group worm count)]. ^b Ref 2. ^c Ref 3.

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Modified Syntheses of 2,4,5-Trihydroxyphenylalanine, 2,4,5-Trihydroxyphenethylamine, and Analogs¹

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We are reporting new and more rewarding syntheses of 2,4,5-trihydroxyphenylalanine (I) (6-hydroxydopa),²

(1) This investigation was supported by the Psychopharmacology Research Branch, National Institute of Mental Health, Contract No. HSM-42-70-41.

(2) H. H. Ong, C. R. Creveling, and J. W. Daly, *J. Med. Chem.*, **12**, 458 (1969).