TREATMENT OF RHEUMATOID ARTHRITIS WITH YODOXIN

A Preliminary Report from the

Desert Arthritis Medical Clinic

by Robert Bingham, M. D.

July 14, 1984

Yodoxin is a brand of Iodoquinol, also known generically as
Diiodohydroxyquinon. Iodine is the active ingredient, used for
generations as a treatment for amoebic infections and similar
large cell organisms, fungi, algae and protozoa.

About two years ago the Searle company, manufacturers of Diodoquin, another brand of Iodoquinol, ceased its distribution in the United States for two reasons - the poor demand for this medicine due to the rarity of amoebic disentery in this country and the fear that it would be used for the common bacillary disenteries, especially in children, with possible ophthalmic complications and the resulting legal liabilities.

We needed a drug of this type to treat patients where Flagyl was poorly tolerated or where the rheumatoid disease appeared resistant to this medication.

Fortunately, we were able to find another distributor with the product, Yodoxin, which we have used with considerable success as the primary treatment of rheumatoid arthritis.

Prior to this meeting I requested the Clinic Staff to compile a short series of consecutive cases in which Yodoxin, was prescribed and in which we had follow-up laboratory confirmation of the clinical improvement we obtained in these active disease.

These results were all obtained within two or three weeks, the length of time we consider necessary to verify the value of Yodoxin on the patient.

We have found that an elevated sedimentation rate, an elevated white blood cell count with a total increase in eosinophils, and a relative low hemoglobin and hematocrit are significant and verifying laboratory findings in rheumatoid diseases. Shown here are findings before and /after ten days of treatment with Yodoxin at the usual adult dose: 650mg. three times a day, for ten days, taken with meals

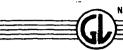
Patient	Sedimentation	Rate White Blood Count	Eosinophils	Hemoglobin or Hematocrit
·s.c.	61 / 30	7.5 M / 6.5	1% / 0	38 % / 37 %
P.J.	16 / 14	4.7 / 5.5 *	4 / 3	13.5 / 14.9 Gm.
B.J.	63 / 44	7.0 / 6.0	2 / 0	12.5 / 13.0
J.H.	100 / 48	4.3 / 5.8 *	1 / 0	14.0 / 14.5
L.G.	110 / 64	9.6 '/ 6.3	5 / 2	38.7 / 37.0
M.P.	23 / 49 *	14.9 / 8.6	2 / 5 *	53.0 / 51.0
C.A.	28 / 6	6.8 / 7.2 *	6 / 2	16.6 / 14.0
R.R	107 / 83	7.3 / 5.4	2 / 2	40.0 / 38.0
J.R.	130 / 97	14.0 / 5.6	0 / 0	12/2 / 13/6
M.R.	38 / 16	11.0 / 9.5	0 / 2 *	40.0 / 38.0
U.W.	95 / 72	9.0 / 7.6	1 / 0	41.0 / 40.0
E.S.	56 / 54	8.4 / 8.0	1 / 0	8.4 / 9.0
v.s.	104 / 68	5.7 / 4.4	5 / 3	34.0 / 36.0
G.M.	126 / 34	8.7. / 6.1	1 / 2 *	35.0 / 34.3
J.C.	125 / 70	17.0 / 9.8	12 / 2	12.5 / 14.3

^{*} Note: Occasionally there is seen a reversal of the usual trend in the clinical laboratory findings, clinical improvement not withstanding.

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7750 Katella Avenue STANTON, CA 90680

(714) 891-4443



U.S.P. ...650 mg

SEE PACKAGE INSERT FOR DOSAGE INFORMATION.

NDC 0516 - 0093 - 10

650 mg. Tablets

YODOXIN®

(Iodoquinal Tablets, U.S.P.)

1000 Tablets

CAUTION: Federal law prohibits dispensing without prescription.

Mtg for GLENWOOD INC., Tenafly, N.J. 07670 by: The Vitarine Co., Springheld Gardens, N.Y. 11413 Store at Controlled Room Temperature - 15:30°C (59-86°F).
PROTECT FROM HEAT AND MOISTURE WARNING: Keep out of the reach of children.
PACKARS MAT CHILD RESISTANT

YODOXIN (iodoquinol Tablets, U.S.P.) 210 mg. and 650 mg. Tablets

Composition: Each tablet contains: Iodoquinol, U.S.P. 210 mg. or 650 mg.

Description: Iodoquinol is of a light yellowish to tan color, nearly odorless and stable in air. The compound is practically insoluble in water, and sparingly soluble in most other solvents. It con-

tains 64 per cent organically bound iodine.

Action: Iodoquinol is amebicidal against Entamoeba histolytica and is considered effective against the trophosoite and cyst forms.

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Indications: Indication is used in the treatment of intestinal amebiasis.

Contraindications: Known hypersensitivity to iodine and 8-hydroxyquinolines. Contraindicated in patients with hepatic damage.

Warnings: Optic neuritis, optic atrophy, and peripheral neuropathy have been reported following prolonged high dosage theraphy with halogenated 8-hydroxyquinolines. Long term use of this drug should be avoided.

Use in Pregnancy: Safety for use in pregnancy or during lactation has not been established.

Precautions: Iodoquinol should be used with caution in patients with thyroid disease.

Protein-bound serum iodine levels may be increased during treatment with iodoquinol and therefore interfere with certain thyroid function tests. These effects may persist for as long as six months after discontinuation of therapy. Discontinue the drug if hypersensitivity reactions occur. Adverse Reactions: Skin: various forms of skin eruptions (acneiform papular and pustular; ballae; vegetating of tuberous iododerma), urticaria and pruritus. Gastrointestinal: nauses, vomiting, abdominal cramps, diarrhes, and pruritus ani. Fever, chilis, headache, vertigo and enlargement

Fever, chills, headache, vertigo and enlargement of thyroid have been reported. Optic neuritis, optic atrophy and peripheral neuropathy have been reported in association with prolonged high-dosage 8-hydroxyquinoline therapy.

Dosage and Administration: Usual adult dose: (210 mgm. each) 3 tablets three times daily, after meals for 20 days. Children 6 to 12 years: (210 mgm. each) 2 tablets, t.i.d. Children under 6: (210 mgm. each) one tablet per 15 pounds of body weight. Usual adult dose: (650 mgm. each) One tablet three times a day for twenty days, to be taken after meals. Children (650 mgm. each). For twenty days, 40 mg. per Kg. of body weight daily divided into 2 doses.

divided into 3 doses.

How Supplied: 210 mgm. NDC-00516-0092-01 bottle of 100 tablets and NDC-00516-0092-10 bottle of 1,000 tablets. 650 mgm. NDC-00516-0093-01 bottle of 100 tablets and NDC-00516-0093-10 bottle of 1,000 tablets.

Storage: Store at Controlled Room Temperature 15-30° C. (59-86°F.)

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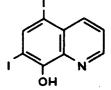
Mfg. for:
GLENWOOD INC.
Tenafly, New Jersey 07670
by VITARINE CO., Springfield Gardens, N.Y.
11413

Made in U.S.A

diiodohydroxyquin (introduced in 1936) are effective against organisms in the bowel lumen but not against trophozoites in the intestinal wall or extraintestinal tissues.

Chemistry

Three synthetic halogen-substituted 8-hydroxy-quinolines have had extensive clinical use—chiniofon (8-hydroxy-7-iodoquinoline-5-sulfonic acid), iodochlorhydroxyquin (clioquinol) (5-chloro-8-hydroxy-7-iodoquinoline), and diiodohydroxyquin (iodoquinol) (8-hydroxy-5,7-diiodoquinoline). Iodochlorhydroxyquin contains approximately 40% iodine and 12% chlorine, and diiodohydroxyquin contains approximately 64% iodine.



Iodochlorhydroxyquin

Diiodohydroxyquin

Absorption, Metabolism, & Excretion

Knowledge is incomplete on the pharmacokinetics of the hydroxyquinolines. Iodochlorhydroxyquin is more readily absorbed than diiodohydroxyquin. Metabolic studies in humans using ¹⁴C-iodochlorhydroxyquin indicated that maximal plasma concentrations were reached at 4 hours after administration of a single dose and then decreased, with an apparent half-life of between 11 and 14 hours. Approximately 25% of a single 750-mg oral dose was excreted in the urine over 72 hours. Use of radioactive iodochlorhydroxyquin in animals showed high uptake of the drug in visceral tissues.

The drugs may interfere with certain thyroid function tests by increasing protein-bound serum iodine levels, leading to a decrease in ¹³¹I uptake.

Antiamebic Effects

The mechanism of action of diiodohydroxyquin and iodochlorhydroxyquin against amebas is not known. Opinions vary on whether the drugs act only against trophozoites or against cysts as well.

Clinical Uses

A. Intestinal Amebiasis: Diiodohydroxyquin and iodochlorhydroxyquin are alternative drugs for the treatment of asymptomatic or mild to moderate intestinal amebiasis. However, until the question of the association of iodochlorhydroxyquin with the SMON syndrome is resolved, only diiodohydroxyquin should be used in therapy. The drugs are not effective in the initial treatment of severe intestinal disease but are used in the subsequent eradication of the infection. They are not effective against amebomas or extraintestinal forms of the disease, including hepatic amebiasis,

THE HALOGENATED
HYDROXYQUINOLINES
Diiodohydroxyquin (Iodoquinol),
Iodochlorhydroxyquin (Clioquinol)

The halogenated hydroxyquinolines were among the first synthetic drugs active in amebiasis. Iodochlorhydroxyquin (introduced in 1931) and