



lopamiro®
lopamidol

The product lopamiro® (lopamidol) was discovered and developed in the research laboratories of Bracco s.p.a. of Milan (Italy). Chemically the substance is: (S)-N,N'-bis [2-hydroxy-1-(hydroxymethyl)-ethyl]-2,4,6-triiodo-5-lactamido-isophthalamide.

lopamiro® is a x-ray contrast medium of the new generation of non ionic compounds, which are water soluble because the molecular structure incorporates hydrophilic groups. This new class of contrast media differs significantly from other compounds currently used in connection with radiological procedures, all of which are soluble only when the radiopaque molecule is ionised by forming a salt with sodium and/or meglumine. While such products are remarkably well tolerated, their aqueous solutions show an inherently high osmolality and this is responsible for a number of side effects that may occur after administration. The discovery of non ionic contrast agents has afforded, along with a sizable reduction of general toxicity, a considerable improvement in local and tissue tolerability, even by the more delicate structures of the human body such as vascular endothelia and the central nervous system. The product is available as preconstituted solution in four different concentrations, with the following physical properties:

Concentration		Viscosity mPa.s		Relative density d ₂₀ ²⁰	Osmometric values at 37°C		
mg iodine/ml	g lopamidol/100ml	20°C	37°C		Osmolality (osmol.kg ⁻¹)	Osmotic pressure (atm)	pH
150	30.6	2.3	1.5	1.17	0.342	8.7	7 ± 0.5
200	40.8	3.3	2.0	1.22	0.413	10.5	7 ± 0.5
300	61.2	8.8	4.7	1.33	0.616	15.7	7 ± 0.5
370	75.5	20.9	9.4	1.41	0.796	20.3	7 ± 0.5

The low osmotic pressure of solutions, the nonionic nature of the molecule and its inherently low chemotoxicity contribute to the exceptionally good local and systemic tolerability of lopamiro®. The favourable results of extended biological and clinical investigations confirm the suitability of lopamiro® as a contrast medium for intrathecal, intra-arterial and intravenous administration.

GENERAL INDICATIONS

NEURORADIOLOGY: myelogram, cisternography and ventriculography. **ANGIOGRAPHY:** cerebral arteriography, coronary arteriography, thoracic aortography, abdominal aortography, angiocardiology, selective visceral arteriography, peripheral arteriography, venography, digital subtraction angiography (DSA), DSA of cerebral arteries, DSA of peripheral arteries, DSA of abdominal arteries. **UROGRAPHY:** intravenous urography. **CONTRAST ENHANCEMENT IN CT SCANNING. ARTHROGRAPHY. FISTULOGRAPHY.** lopamiro® 150 is indicated in paediatric radiology and in DSA.

Contraindications

There are no definite or absolute contraindications to the use of lopamiro®, with the possible exception of Waldenström's macroglobulinemia, multiple myeloma, and severe liver and kidney diseases.

General warnings

The use of organic iodine compounds may cause untoward side effects and manifestations of anaphylaxis. The symptoms include nausea, vomiting, widespread erythema, generalized heat sensation, headache, coryza or laryngeal edema, fever, sweating, asthenia, dizziness, pallor, dyspnoea and moderate hypotension. Skin reactions may occur in the form of various types of rash or diffuse blister formation. More severe reactions involving the cardiovascular system such as peripheral vasodilation with pronounced hypotension, tachycardia, dyspnoea, agitation, cyanosis and loss of consciousness, may require emergency treatment. For these reasons the use of organic iodine contrast media must be limited to cases for which the diagnostic procedure is definitely indicated, as suggested by the patient's clinical status with special attention to existing pathology of the cardiovascular, urinary or hepatobiliary system. In particular, contrast media designed for cardioangiographic procedures should be used in hospitals or clinics equipped and staffed for intensive care in emergencies. For other more common diagnostic procedures calling for the use of iodinated contrast media, the public or private institutions, where such procedures take place, should be supplied at all times with all equipment and drugs that experience has shown to be suitable in case of an accident: the Ambu balloon, oxygen bottles, antihistamines, vasopressor drugs, cortisones. Never mix other drugs with contrast medium solutions. When examining small children or babies, do not limit fluid intake before administering a hypertonic contrast solution; also, correct any existing water and electrolyte imbalance. Pregnant women and patients with hyperthyroidism should receive iodinated contrast media only if the attending physician finds it absolutely necessary.

In patients scheduled for thyroid examination with a radioactive iodine tracer, bear in mind that iodine uptake in the thyroid gland will be reduced for several days (sometimes up to 2 weeks) after dosing with an iodinated contrast medium that is eliminated through the kidneys.

Keep lopamiro® solutions away from strong light. Exceptionally, the event of crystallization of lopamiro® solutions could occur. It has been shown that such a phenomenon is caused by a damaged or defective container and therefore the product should not be used in this case. The bottle, once opened, must be used immediately. Any residue of contrast medium must be discarded. lopamiro®, as other iodinated contrast media, can react with metallic surfaces containing copper (e.g. brass), therefore the use of equipment, in which the product comes into direct contact with such surfaces, should be avoided.

NEURORADIOLOGY

	Concentration (mg iodine/ml)	Recommended dosage (ml)
Myelographic	200-300	5-15
Cisternography and ventriculography	200-300	5-15

Warnings and side effects

In the event of CSF blockade, remove as much of the administered contrast solution as possible. The use of organic iodine contrast media may be contraindicated for patients with a history of epilepsy. Also the presence of blood in the CSF contraindicates the use of lopamiro®: in such cases, the operator should carefully assess the need for the diagnostic procedure against possible risk to the patient. Patients receiving treatment with anticonvulsant drugs must continue such treatment before and after the procedure. Should a convulsive seizure develop during the examination, administer diazepam or sodium phenobarbital intravenously. Neuroleptics must be absolutely avoided because they lower the seizure threshold. The same applies to analgesics, antiemetics, antihistamines and sedatives of the phenothiazine group. Whenever possible, treatment with such drugs should be discontinued at least 48 hours before administration of the contrast medium and not be resumed less than 12 hours after completion of the procedure. Clinical trials have shown that lopamiro® is readily tolerated by the vast majority of patients, both in general terms and, particularly, in terms of CNS tolerability. Reported side effects include headache (sometimes with a delayed onset), nausea, vomiting and pain at the site of injection, all these manifestations being generally mild and of short duration. More rarely or indeed exceptionally, patients have developed dizziness, neck rigidity, lumbar pain, or sciatic pain - the latter often as a transient exacerbation of pre-existing symptoms. Fever was reported in a few cases. Exceptionally, patients dosed with lopamiro® have developed muscular spasms or generalized convulsions, sometimes representing a recurrence of established epilepsy or attributable to accidental overdosage of neuroleptic drugs. Very few cases of transient mental confusion have been reported.

ANGIOGRAPHY

	Concentration (mg iodine/ml)	Recommended dosage (ml)
Cerebral arteriography	300	5-10 (bolus)
Coronary arteriography	370	8-15 (bolus)
Thoracic aortography	370	1.0 - 1.2/kg
Abdominal aortography	370	1.0 - 1.2/kg
Angiocardiography	370	1.0 - 1.2 /kg
Selective visceral arteriography	300-370	depending on examination
Peripheral arteriography	300-370	40-50
Digital subtraction angiography	150-370	depending on examination
Venography	300	30-50

Warnings and side effects

The use of lopamiro® as a contrast medium for cerebral angiography, may cause side effects, which are usually mild and of short duration. Many patients report a sensation of heat in the face and neck; a few complain of headache. A fairly frequent cardiovascular reaction to dosing with lopamiro® is bradycardia associated with systemic hypotension. The reaction is transient and requires no treatment. Severe neurological sequelae may arise as direct complications of pre-existing pathology in the individual patient. Such reactions are diverse and may include tonic/clonic convulsions, aphasia, fainting, transient narrowing of visual field, hemiparesis and coma. The risk associated with a particular investigation involved may be increased by conditions such as advanced arteriosclerosis, hypertension, heart failure, major systemic diseases and recent cerebral embolism or thrombosis.

In patients undergoing angiocardiographic procedures special attention should be paid to the status of the right heart and pulmonary circulation. Right heart insufficiency and pulmonary hypertension may precipitate bradycardia and systemic hypotension, when the organic iodine solution is injected. In paediatric roentgenology, one should proceed with great caution when injecting the contrast medium into the right heart chambers of cyanotic neonates with pulmonary hypertension and impaired cardiac function. In examinations of the aortic arch the tip of the catheter should be positioned carefully to avoid hypotension, bradycardia and CNS injury due to excess pressure transmitted from the injector pump to the brachiocephalic branches of the aorta.

Likewise, in abdominal aortography, excess pressure from the pump may cause renal infarction, spinal cord injury, retroperitoneal bleeding, intestinal infarction and necrosis. In peripheral arteriography lopamiro® 370 may sometimes cause a painful reaction in the involved limb.

This is usually not the case with the less concentrated solution lopamiro® 300.

A property of nonionic contrast media is the extremely low interference with normal physiological functions. As a consequence of this, nonionic contrast media have less anticoagulant activity in vitro than ionic media. Medical personnel performing vascular catheterisation procedures should be aware of this and pay meticulous attention to the angiographic technique and catheter flushing so as to minimize the risk of procedure-related thrombosis and embolism.

UROGRAPHY

lopamiro® 300 and 370 should be used; the dose recommended for this type of examination is 30 to 50 ml. The less marked osmotic diuresis induced by the nonionic agent makes lopamiro® 370 especially suitable for patients with mild or moderately severe renal insufficiency and for neonates. The new contrast medium affords diagnostically useful nephrography even in patients with major renal insufficiency.

Warnings and side effects

The side effects that may arise in connection with intravenous urography are those described in the paragraph on General warnings.

OTHER DIAGNOSTIC PROCEDURES

	Concentration (mg iodine/ml)	Recommended dosage (ml)
Contrast enhancement in CT scanning	300-370	0.5 - 2.0/kg
Arthrography	300	depending on examination
Fistulography	300	depending on examination

For the enhancement of contrast in CT scans lopamiro® may be injected intravenously as a bolus, as a drip infusion or by a combination of the two methods.

Warnings and side effects

The reactions reported in cases of arthrography and fistulography usually represent irritative manifestations superimposed on existing tissue inflammation.

Storage

Store in cool (15-25) °C and dry place. Keep away from light. The vial, once opened, has to be used immediately. Any residue of contrast medium must be discarded.

Composition & Registration Number

lopamiro® 300 - Each vial (30 ml) contains: 18.37 g of lopamidol
lopamiro® 300 - Each vial (50 ml) contains: 30.62 g of lopamidol
lopamiro® 300 - Each bottle (100 ml) contains: 61.24 g of lopamidol
lopamiro® 370 - Each vial (30 ml) contains: 22.66 g of lopamidol
lopamiro® 370 - Each vial (50 ml) contains: 37.76 g of lopamidol
lopamiro® 370 - Each bottle (100 ml) contains: 75.53 g of lopamidol

Reg No : DK10678200543A1
Reg No : DK10678200543A1
Reg No : DK10678200543A2
Reg No : DK10678200543B1
Reg No : DK10678200543B1
Reg No : DK10678200543B2

How supplied

lopamiro® 300 - Boxes of 1 bottle, 30 ml - Boxes of 1 bottle, 50 ml - Boxes of 1 bottle, 100 ml
lopamiro® 370 - Boxes of 1 bottle, 30 ml - Boxes of 1 bottle, 50 ml - Boxes of 1 bottle, 100 ml



HARUS DENGAN RESEP DOKTER

Manufactured by

Patheon Italia S.p.A. - Italy for Bracco S.p.A. - Italy

Imported by

P.T. Dipa Pharmalab Intersains
Jakarta-Indonesia