

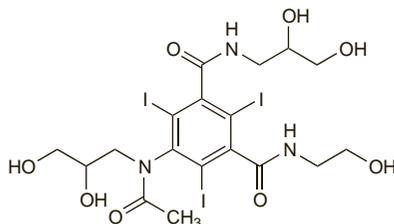
Hexabrix; **Norw.:** Hexabrix; **NZ:** Hexabrix; **Port.:** Hexabrix; **Spain:** Hexabrix; **Swed.:** Hexabrix; **Switz.:** Hexabrix; **UK:** Hexabrix; **USA:** Hexabrix; **Venez.:** Hexabrix.

Ioxilan (USAN, rINN)

Ioxilán; Ioxilane; Ioxilanum. *N*-(2,3-Dihydroxypropyl)-5-[*N*-(2,3-dihydroxypropyl)acetamido]-*N'*-(2-hydroxyethyl)-2,4,6-triiodo-sophthalamide.

Йоксилан

$C_{18}H_{24}I_3N_3O_8 = 791.1$.
CAS — 107793-72-6.
ATC — V08AB12.
ATC Vet — QV08AB12.



Description. Ioxilan contains about 48.1% of I.

Pharmacopeias. In *US*.

USP 31 (Ioxilan). A white to off-white, practically odourless, powder. Soluble in water and in methyl alcohol. pH of a 10% solution in water is between 5.0 and 7.5. Store at a temperature of 25°, excursions permitted between 15° and 30°. Protect from light.

Adverse Effects, Treatment, and Precautions

See under the amidotrizoates, p.1475.

Pharmacokinetics

After intravascular use, ioxilan is rapidly eliminated unchanged in the urine; about 94% of a dose is excreted within 24 hours. Protein binding is reported to be very low. Ioxilan is dialysable.

Uses and Administration

Ioxilan is a nonionic monomeric iodinated radiographic contrast medium (see p.1474). It is given intra-arterially or intravenously for procedures including angiography and urography; it is also used for contrast enhancement during computed tomography.

Ioxilan is usually available as solutions containing 62.3 or 72.7% of ioxilan (equivalent to 300 or 350 mg/mL of iodine). The dose and strength used vary according to the procedure and route.

Preparations

USP 31: Ioxilan Injection.

Proprietary Preparations (details are given in Part 3)

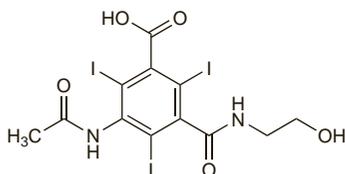
Jpn: Imagenil; **USA:** Oxilan.

Ioxitalamic Acid (rINN)

Acide Ioxitalamique; Ácido Ioxitalámico; Acidum Ioxitalamicum; AG-58107; Ioxitalamic Acid; Joksitalaamihappo; Joxitalamsyra. 5-Acetamido-*N*-(2-hydroxyethyl)-2,4,6-tri-iodoisophthalamic acid.

Йокситаламвая Кислота

$C_{12}H_{11}I_3N_2O_5 = 643.9$.
CAS — 28179-44-4.
ATC — V08AA05.
ATC Vet — QV08AA05.



Description. Ioxitalamic acid contains about 59.1% of I.

Pharmacopeias. In *Fr*:

Meglumine Ioxitalamate (rINNM)

Ioxitalamate de Méglumine; Ioxitalamate Meglumine; Ioxitalamate de meglumina; Meglumini Ioxitalamas. The *N*-methylglucamine salt of ioxitalamic acid.

Меглумина Йокситаламат

$C_{12}H_{11}I_3N_2O_5 \cdot C_7H_{17}NO_5 = 839.2$.
CAS — 29288-99-1.
ATC — V08AA05.
ATC Vet — QV08AA05.

Description. Meglumine ioxitalamate contains about 45.4% of I.

The symbol † denotes a preparation no longer actively marketed

Sodium Ioxitalamate (rINNM)

Ioxitalamate de Sodium; Ioxitalamate Sodium; Ioxitalamato sodico; Natrii Ioxitalamas.

Натрий Йокситаламат

$C_{12}H_{10}I_3N_2NaO_5 = 665.9$.
CAS — 33954-26-6.
ATC — V08AA05.
ATC Vet — QV08AA05.

Description. Sodium ioxitalamate contains about 57.2% of I.

Profile

Ioxitalamic acid is an ionic monomeric iodinated radiographic contrast medium (p.1474) with actions similar to those of the amidotrizoates (p.1475). It is given intravenously or by instillation into body cavities for procedures including angiography, cholangiography, cystography, hysterosalpingography, and urography; it may be given orally or rectally for imaging of the gastrointestinal tract. It is also used for contrast enhancement in computed tomography.

Ioxitalamic acid is usually available as a solution containing 21% of the sodium salt (equivalent to 120 mg/mL of iodine), 55.1 to 66% of the meglumine salt (equivalent to 250 to 300 mg/mL of iodine), or as a mixture of both salts. The dose and strength used vary according to the procedure and route.

Monoethanolamine ioxitalamate has also been used.

Preparations

Proprietary Preparations (details are given in Part 3)

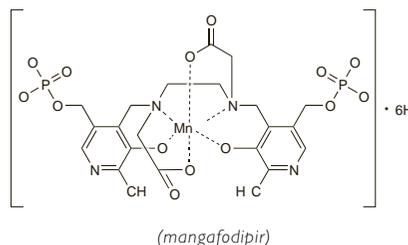
Arg.: Telebrix 30; Telebrix 38; Telebrix Coronario; Telebrix Hystero; **Belg.:** Telebrix; Telebrix Gastro; Telebrix Hystero; **Braz.:** Telebrix†; **Canad.:** Telebrix; **Chile:** Telebrix 30; Telebrix 35; **Cz.:** Telebrix 30; Telebrix 35; Telebrix Gastro; **Fr.:** Telebrix 12; Telebrix 30; Telebrix 35; Telebrix Gastro; Telebrix Hystero; **Ger.:** Telebrix Gastro; Telebrix N 180 and 300; **Gr.:** Telebrix Gastro; Telebrix Hystero†; **Hung.:** Telebrix; Telebrix Gastro; **Israel:** Telebrix; Telebrix Gastro; **Mex.:** Telebrix; **Neth.:** Telebrix 12; Telebrix 30; Telebrix 35; Telebrix Gastro; Telebrix Hystero; **Port.:** Telebrix 12; Telebrix 30; Telebrix 35; Telebrix Gastro; Telebrix Hystero; **Switz.:** Telebrix 12; Telebrix 30; Telebrix 35; Telebrix Gastro; Telebrix Hystero; **Venez.:** Telebrix 30; Telebrix 35; Telebrix Hystero.

Mangafodipir Trisodium (BANM, USAN, rINNM)

Mangafodipir trisódico; Mangafodipir Trisodique; Mangafodipirum Trinatricum; MnDPDP (mangafodipir); S-095 (mangafodipir); Win-59010; Win-59010-2 (mangafodipir). Trisodium trihydrogen (OC-6-13)-{[*N,N'*-ethylenebis(*N*-[3-hydroxy-5-(hydroxymethyl)-2-methyl-4-pyridyl]methyl)glycine] 5,5'-bis(phosphato)}(8-) manganate(6-); Trisodium trihydrogen (OC-6-13)-*N,N'*-ethane-1,2-diybis[*N*-(2-methyl-3-oxido-κO-5-(phosphonatoxy)methyl)-4-pyridylmethyl]glycinato(*O,N*)]manganate(II).

Тринатрий Мангафодипир

$C_{22}H_{27}MnN_4Na_3O_{14}P_2 = 757.3$.
CAS — 155319-91-8 (mangafodipir); 140678-14-4 (mangafodipir trisodium).
ATC — V08CA05.
ATC Vet — QV08CA05.



Pharmacopeias. In *US*.

USP 31 (Mangafodipir Trisodium). Pale yellow crystals or crystalline powder. Freely soluble in water; very slightly soluble in alcohol and in acetone; slightly soluble in chloroform; sparingly soluble in methyl alcohol. pH of a 1% solution in water is between 5.5 and 7.0. Store at a temperature not exceeding 8°.

Adverse Effects and Precautions

The most common adverse effects of mangafodipir are injection site discomfort, feelings of warmth or flushing, headache, nausea, vomiting, abdominal pain, and taste disturbances. Hypersensitivity reactions, including anaphylactoid reactions, may occur. Transient increases in bilirubin and liver transaminase concentrations and decreases in plasma-zinc concentrations have been reported.

Mangafodipir should be used with caution in patients with hepatic or renal impairment and should be avoided if impairment is severe. It should not be given to patients with phaeochromocytoma.

◇ References.

1. Federle MP, et al. Safety and efficacy of mangafodipir trisodium (MnDPDP) injection for hepatic MRI in adults: results of the U.S. multicenter phase III clinical trials (safety). *J Magn Reson Imaging* 2000; **12**: 186-97.

Pharmacokinetics

After intravenous injection, mangafodipir is dephosphorylated and manganese is exchanged for zinc leading to the release of free manganese ions and the formation of 2 inactive metabolites. Manganese is rapidly taken up by the liver, pancreas, kidney and spleen; about 15 to 20% is excreted in the urine within 24 hours, with most of the remainder excreted in the faeces over about 4 days. The metabolites are almost entirely excreted in the urine within 24 hours.

Uses and Administration

Mangafodipir is a manganese chelate that is used as a magnetic resonance contrast medium (p.1474) for imaging of the liver and pancreas. Manganese has paramagnetic properties that increase the relaxivity of hydrogen ions, leading to signal enhancement. Free manganese is released from mangafodipir in the body and is taken up by normal liver and pancreatic tissue, increasing the degree of contrast.

Mangafodipir is given intravenously as the trisodium salt.

In the UK, a solution containing mangafodipir trisodium 7.57 mg/mL (10 micromol/mL) is used. Usual doses for imaging are:

- liver: 0.5 mL/kg (5 micromol/kg) given by intravenous infusion at a rate of 2 to 3 mL/minute
- pancreas: 0.5 mL/kg (5 micromol/kg) given by intravenous infusion at a rate of 4 to 6 mL/minute

In the USA, a more concentrated preparation is used, containing mangafodipir trisodium 37.9 mg/mL (50 micromol/mL). Usual doses are:

- liver: 0.1 mL/kg (5 micromol/kg), given by slow intravenous injection to a maximum dose of 15 mL

Preparations

USP 31: Mangafodipir Trisodium Injection.

Proprietary Preparations (details are given in Part 3)

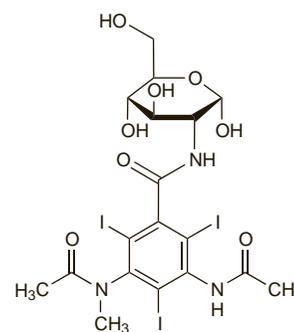
Austria: Teslscan; **Belg.:** Teslscan; **Cz.:** Teslscan; **Denm.:** Teslscan†; **Fin.:** Teslscan†; **Fr.:** Teslscan; **Ger.:** Teslscan; **Gr.:** Teslscan; **Hung.:** Teslscan; **Ital.:** Teslscan; **Neth.:** Teslscan; **Norw.:** Teslscan; **NZ:** Teslscan; **Port.:** Teslscan; **Spain:** Teslscan; **Swed.:** Teslscan; **Switz.:** Teslscan; **UK:** Teslscan; **USA:** Teslscan.

Metrizamide (BAN, USAN, rINN)

Metritsamidi; Metrizamid; Metrizamida; Métrizamide; Metrizamidum; Win-39103. 2-[3-Acetamido-2,4,6-tri-iodo-5-(*N*-methylacetamido)benzamido]-2-deoxy-D-glucose.

Метризамид

$C_{18}H_{22}I_3N_3O_8 = 789.1$.
CAS — 31112-62-6 (metrizamide); 55134-11-7 (metrizamide, glucopyranose form).
ATC — V08AB01.
ATC Vet — QV08AB01.



Description. Metrizamide contains about 48.2% of I.

Profile

Metrizamide is a nonionic monomeric iodinated radiographic contrast medium (p.1474) that has been used in myelography, angiography, intravenous urography, and arthrography, and also for contrast enhancement during computed tomography.

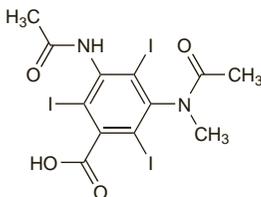
Breast feeding. No adverse effects have been seen in breast-feeding infants whose mothers were receiving metrizamide and the American Academy of Pediatrics considers¹ that it is therefore usually compatible with breast feeding.

1. American Academy of Pediatrics. The transfer of drugs and other chemicals into human milk. *Pediatrics* 2001; **108**: 776-89. Correction. *ibid.*; 1029. Also available at: <http://aappolicy.aappublications.org/cgi/content/full/pediatrics%3b108/3/776> (accessed 27/03/06)

Metrizoic Acid (BANM, rINNM)

Acide Métrizoïque; Ácido metrizoico; Acidum Metrizoicum; Metritsoinihappo; Metrizoisynra. 3-Acetamido-2,4,6-tri-iodo-5-(N-methylacetamido)benzoic acid.

Метризоевая Кислота
 $C_{12}H_{11}I_3N_2O_4 = 627.9$.
 CAS — 1949-45-7.
 ATC — V08AA02.
 ATC Vet — QV08AA02.



Description. Metrizoic acid contains about 60.6% of I.

Meglumine Metrizoate (BANM, rINNM)

Meglumini Metrizoas; Métrizoate de Méglumine; Metrizoate Meglumine; Metrizoato de meglumina. The N-methylglucamine salt of metrizoic acid.

Меглумина Метризоат
 $C_{12}H_{11}I_3N_2O_4 \cdot C_7H_{17}NO_5 = 823.2$.
 CAS — 7241-11-4.
 ATC — V08AA02.
 ATC Vet — QV08AA02.

Description. Meglumine metrizoate contains about 46.3% of I.

Sodium Metrizoate (BAN, rINN)

Métrizoate de Sodium; Metrizoate Sodium (USAN); Metrizoato de sodio; Natrii Metrizoas; NSC-107431.

Натрия Метризоат
 $C_{12}H_{10}I_3N_2NaO_4 = 649.9$.
 CAS — 7225-61-8.
 ATC — V08AA02.
 ATC Vet — QV08AA02.

Description. Sodium metrizoate contains about 58.6% of I.

Profile

Metrizoic acid is an ionic monomeric iodinated radiographic contrast medium (p.1474) with actions similar to those of the amidotrizoates (p.1475). It has been used as the meglumine and sodium salts, often with calcium metrizoate and magnesium metrizoate, for a variety of diagnostic procedures including angiography, cholangiography, and hysterosalpingography.

Breast feeding. No adverse effects have been seen in breast-feeding infants whose mothers were receiving metrizoate and the American Academy of Pediatrics considers¹ that it is therefore usually compatible with breast feeding.

1. American Academy of Pediatrics. The transfer of drugs and other chemicals into human milk. *Pediatrics* 2001; **108**: 776-89. Correction. *ibid.*: 1029. Also available at: <http://aapolicy.aappublications.org/cgi/content/full/pediatrics%3b108/3/776> (accessed 27/03/06)

Preparations

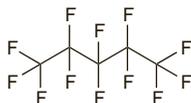
Proprietary Preparations (details are given in Part 3)

UK: Isopaque Cysto†.

Perflenanpent (USAN, rINN)

Dodecafluoropentanam; Dodekafluoropentaani; Dodekafluoropentani; Perflénapent; Perflenanpentum. Dodecafluoropentane.

Перфленапент
 $C_5F_{12} = 288.0$.
 CAS — 678-26-2.
 ATC — V08DA03.
 ATC Vet — QV08DA03.

**Profile**

Perflenanpent is a liquid perfluorocarbon that has been used as an ultrasound contrast medium (p.1474) for echocardiography. It has been given intravenously as an emulsion containing droplets of perflenanpent; on warming to body temperature the droplets

form microbubbles of perflenanpent gas that provide echo-enhancement. A small amount of perflisopent (below) was also included in the formulation.

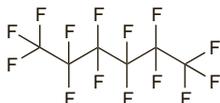
References.

- Robbin ML, Eisenfeld AJ. Perflenanpent emulsion: a US contrast agent for diagnostic radiology—multicenter, double-blind comparison with a placebo. *Radiology* 1998; **207**: 717-22.
- Kitzman DW, Wesley DJ. Safety assessment of perflenanpent emulsion for echocardiographic contrast enhancement in patients with congestive heart failure or chronic obstructive pulmonary disease. *Am Heart J* 2000; **139**: 1077-80.

Perflexane (USAN, rINN)

Perflexano; Perflexanum. Tetradecafluorohexane.

Перфлексан
 $C_6F_{14} = 338.0$.
 CAS — 355-42-0.

**Profile**

Perflexane is a perfluorocarbon gas that has been used as an ultrasound contrast medium (p.1474) for echocardiography. Dry microspheres containing the gas are reconstituted immediately before use, leading to the formation of microbubbles of perflexane that provide echo-enhancement; lipids are included in the microspheres to stabilise the bubbles when they form.

Preparations

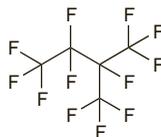
Proprietary Preparations (details are given in Part 3)

USA: Imagent.

Perflisopent (USAN, rINN)

Perflisopentum. Nonafluoro-2-(trifluoromethyl)butane.

Перфлизопент
 $C_5F_{12} = 288.0$.
 CAS — 594-91-2.

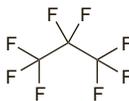
**Profile**

Perflisopent is a perfluorocarbon that has been used as an ultrasound contrast medium (p.1474) with perflenanpent (above) for echocardiography.

Perflutren (USAN, rINN)

DMP-115; FS-069; MRX-115; Octafluoropropanum; Oktafluoropropaani; Oktafluoropropan; Perfluoropropane; Perflutrène; Perflutreno; Perflutrenum. Octafluoropropane.

Перфлутрен
 $C_3F_8 = 188.0$.
 CAS — 76-19-7.

**Profile**

Perflutren is a perfluorocarbon gas used as either albumin- or lipid-coated microspheres as an ultrasound contrast medium (p.1474) for echocardiography.

The albumin-coated microspheres are suspended in 1% albumin solution immediately before use and are given in a dose of 0.5 to 3 mL by intravenous injection, repeated if necessary up to a total dose of 8.7 mL.

The lipid-coated microsphere suspension is formed by agitating the gas with a lipid solution immediately before use and is given in a dose of 10 microlitre/kg by intravenous injection, repeated once after 30 minutes if required, or as intravenous injections of 100 to 400 microlitres repeated as required up to a maximum total dose of 1.6 mL. Alternatively, 1.3 mL of the suspension may

be diluted in 50 mL of sodium chloride 0.9% and given by intravenous infusion at an initial rate of 4 mL/minute, adjusted as required, to a maximum rate of 10 mL/minute.

Serious cardiopulmonary reactions, including fatalities, have been reported with perflutren and it should be used with extreme caution in patients with pulmonary hypertension or unstable cardiopulmonary conditions. The safety of perflutren has not been established in patients with right-to-left cardiac shunts; it should also be used with extreme caution or avoided in such patients.

Perflutren has also been given by intra-ocular injection to provide tamponade in the management of retinal detachment.

Adverse effects. References.

- Herzog CA. Incidence of adverse events associated with use of perflutren contrast agents for echocardiography. *JAMA* 2008; **299**: 2023-5.

Preparations

USP 31: Perflutren Protein-Type A Microspheres Injectable Suspension.

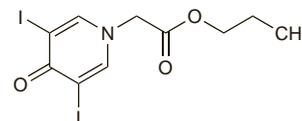
Proprietary Preparations (details are given in Part 3)

Arg.: Optison†; **Austria:** Optison; **Canada:** Definity; **Chile:** Definity; **Cz.:** Luminity; Optison; **Denm.:** Optison; **Fr.:** Luminity; **Ger.:** Optison; **Gr.:** Optison; **Hung.:** Optison; **Israel:** Definity; **Ital.:** Optison; **Neth.:** Optison; **Port.:** Luminity; Optison; **Spain:** Optison; **Swed.:** Optison; **Switz.:** Optison; **UK:** Luminity; Optison; **USA:** Definity; Luminity; Optison.

Propylidone (rINN)

Propilidona; Propylidonom; Propyljodon; Propyljiodoni. Propyl 1,4-dihydro-3,5-di-iodo-4-oxo-1-pyridylacetate.

Пропилидон
 $C_{10}H_{11}I_2NO_3 = 447.0$.
 CAS — 587-61-1.
 ATC — V08AD03.
 ATC Vet — QV08AD03.



Description. Propylidone contains about 56.8% of I.

Pharmacopoeias. In *Int.* and *US*.

USP 31 (Propylidone). A white or almost white, crystalline powder. Is odourless or has a faint odour. Practically insoluble in water; soluble in alcohol, in acetone, and in ether. Store in airtight containers at a temperature of 25°, excursions permitted between 15° and 30°. Protect from light.

Profile

Propylidone is an iodinated radiographic contrast medium (p.1474) that has been used for bronchography as either an aqueous or oily suspension.

Preparations

USP 31: Propylidone Injectable Oil Suspension.

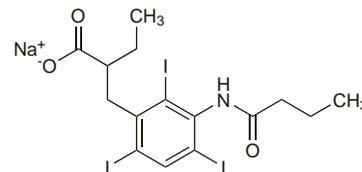
Proprietary Preparations (details are given in Part 3)

Israel: Dionosil†.

Sodium Tyropanoate (BAN, rINN)

Natrii Tyropanoas; NSC-107434; Tiropanoate de sodio; Tyropanoate de Sodium; Tyropanoate Sodium (USAN); WVin-8851-2. Sodium 2-(3-butyramido-2,4,6-tri-iodobenzyl)butyrate.

Натрия Тиропаноат
 $C_{15}H_{17}I_3NNaO_3 = 663.0$.
 CAS — 27293-82-9 (tyropanoic acid); 7246-21-1 (sodium tyropanoate).
 ATC — V08AC09.
 ATC Vet — QV08AC09.



Description. Sodium tyropanoate contains about 57.4% of I.

Profile

Sodium tyropanoate is an ionic monomeric iodinated radiographic contrast medium (p.1474) with actions similar to those of iopanoic acid (p.1484) and has been used for cholecystography and cholangiography.