BioPAL provides MRI products to the biomedical research community. We offer a wide range of iron-based (T2 imaging agent) and gadolinium-based (T1 imaging agent) contrast reagents to the life-science community. BioPAL is a leading provider of novel MRI contrast reagents directed toward the needs of the life science community.

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Molday ION^{TM} product line is a family of iron oxide-based superparamagnetic contrast reagents designed to label cells and mark the vascular space. These MRI contrast reagents have a colloidal size of 30-50 nm and are classified as darkening agents acting through the T2 relaxation process.

Molday ION comes with many chemical surfaces and is presented below organized by applications. These applications are as follows:

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BioPAL can also prepare custom nano-iron oxides. Please Inquire!

BioPAL is actively developing novel and innovative MRI products to assist the biomedical research community.



NOT FOR HUMAN USE.

Application 1: Vascular and Functional Imaging

Catalog Number		
CL-30Q02-2	Molday ION TM	275.00
CL-30Q02-A2	Molday ION TM Autoclaved	300.00
CL-50Q02-2	Molday ION TM \$ 2.0 ml of 46 nm iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 10 mg Fe/ml having a zeta potential of ~ -5mV. Molday ION (~50 nm) can be used to complement studies obtained with Molday ION (30 nm) or where a larger nanoparticle is needed. Applications: MRI vascular/functional imaging, Blood pool agent, EM, Magnetic cell-sorting.	275.00
CL-70Q02-2	Molday ION TM	275.00
CL-20Q02-3	Molday ION Aminodextran TM\$ 2.0 ml of 20 nm iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 10 mg Fe/ml. Applications: MRI vascular/functional imaging, Blood pool agent, EM, Magnetic cell-sorting.	275.00
CL-30Q02-5	Molday ION CLIOH TM \$ 2.0 ml of 30 nm iron-based superparamagnetic contrast agent having a neutral surface packaged in a 2 ml sealed serum bottle. 5 mg Fe/ml having a zeta potential of ~ 0mV. Molday ION CLIOH is formulated to provide a stable neutral inert surface.	475.00
CL-30Q02-7	Molday ION Carboxyl TM\$ 2.0 ml of 20 nm iron-based superparamagnetic contrast agent containing carboxyl groups packaged in a 2 ml sealed serum bottle. 10 mg Fe/ml having a zeta potential of ~ -20mV. Applications: MRI vascular/functional imaging, EM, Magnetic cell-sorting.	275.00
CL-30Q02-6	Molday ION (-) TM	300.00



CL-00-01	Poly-L-Lysine	60.00
CL-30Q02-6C	Molday ION Carboxyl Terminated TM\$ 2.0 ml of 30 nm iron-based superparamagnetic contrast agent containing carboxyl groups packaged in 2 ml sealed serum bottle. 2.5 mg Fe/ml, having a zeta potential of ~ -35 mV. Applications: MRI, EM, Magnetic cell-sorting, Cell targeting, Conjugation — Application Note #10.	395.00
CL-30Q02-CA	Molday ION Carboxyl/Amine Terminated TM\$ 2.0 ml of 30 nm iron-based superparamagnetic contrast agent containing amine and carboxyl groups designed to approximate the surface of a protein packaged in a 2 ml serum bottle. 5 mg Fe/ml, having a zeta potential of ~ +4 mV. Applications: MRI, EM, Magnetic cell-sorting, Cell labeling.	475.00
Application 2: Cell L	abeling	
Catalog <u>Number</u>		
CL-50Q02-6A-0	Molday ION TM Dye Free	375.00
CL-50Q02-A6A-0	Molday ION TM Dye Free Autoclaved	450.00
CL-50Q01-6A-50	Molday ION Rhodamine B TM (MIRB)\$ 0.5 ml of 35 nm rhodamine B labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~ +31 mV. A suggested procedure for labeling cells is provided as a PDF download on BioPAL's web site. PDF downloads depicting labeled cells, as well as presented posters, is also provided. Applications: Cell labeling, Cell labeling with MRI tracking, Live cell imaging (in vitro), Drug delivery, Theranostics, Fluorescent detection. For additional information, please review Application Note #3 on BioPAL Web Site.	140.00
CL-50Q02-6A-50	Molday ION Rhodamine B TM (MIRB)\$ 2.0 ml of 35 nm rhodamine B labeled iron-based superparamagnetic contrast agent	450.00

2.0 ml of 35 nm rhodamine B labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~ +31 mV.

A suggested procedure for labeling cells is provided as a PDF download on BioPAL's web site.

A suggested procedure for labeling cells is provided as a PDF download on BioPAL's web site. PDF downloads depicting labeled cells, as well as presented posters, is also provided. Applications: Cell labeling, Cell labeling with MRI tracking, Live cell imaging (*in vitro*), Drug delivery, Theranostics, Fluorescent detection.

For additional information, please review **Application Note #3** on BioPAL Web Site.

CL-50Q02-6A-51 Molday ION EverGreen™.....\$ 450.00

2.0 ml of 35 nm EverGreen labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of \sim +31 mV. Labeled cells may be visualized using a standard fluorescein filter set. Molday Ion EverGreen



has an excitation and emission maxima at 498 nm and 526 nm, respectively. Applications: Cell labeling, Cell labeling with MRI tracking, Live cell imaging (*in vitro*), Drug delivery, Theranostics, Fluorescent detection. For additional information, please review **Application Note #3** on BioPAL Web Site.

CL-50Q02-6A-52	Molday ION Coumarin TM \$ 2.0 ml of 35 nm Coumarin labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~ +30 mV. Labeled cells may be visualized using a standard DAPI filter set. Molday Ion Coumarin has an excitation and emission maxima at 356 nm and 456 nm, respectively. Applications: Cell labeling, Cell labeling with MRI tracking, Live cell imaging (in vitro), Drug delivery, Theranostics, Fluorescent detection. For additional information, please review Application Note #3 on BioPAL Web Site.	450.00
CL-50Q01-6A-53	Molday ION TM MI-750	400.00
CL-50Q02-6A-53	Molday ION Rose Bengal TM 2.0 ml of 35 nm Rose Bengal labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~ +31 mV. A suggested procedure for labeling cells is provided as a PDF download on BioPAL's web site. Applications: Cell labeling, Cell labeling with MRI tracking, Live cell imaging (<i>in vitro</i>), Drug delivery, Theranostics, Fluorescent detection. For additional information, please review Application Note #9 on BioPAL Web Site.	450.00
CL-50Q02-6S-51	Molday SION EverGreen TM	425.00
CL-30Q02-6	Molday ION (-) TM	300.00
CL-00-01	Poly-L-Lysine	60.00
CL-50Q02-71	Molday ION Spermidine TM 1.0 ml of 40 nm iron-based superparamagnetic contrast agent conjugated with spermidine packaged in a 2 ml sealed serum bottle. 1 mg Fe/ml having a zeta potential of ~ +35mV. A suggested procedure for labeling cells is provided as a PDF download on BioPAL's web site (Application Note #5). Applications: Cell labeling, Cell labeling with MRI tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	475.00



Application 3: Functional Chemistry for Conjugations

Catalog Number

Mol	Molday ION Containing Reactive Amines Molday ION C2Amine TM\$ 2.0 ml of 35 nm iron-based superparamagnetic contrast agent with amine groups packaged in a 2 ml sealed serum bottle. 5 mg Fe/ml having a zeta potential of ~ +48 mV. Applications: Conjugation with bifunctional agents, Click chemistry, NHS esters, Isothiocyanates.	475.00
CL-30Q02-CA	Molday ION Carboxyl/Amine Terminated TM\$ 475.00 2.0 ml of 30 nm iron-based superparamagnetic contrast agent containing amine and carboxyl groups designed to approximate the surface of a protein packaged in a 2 ml serum bottle. 5 mg Fe/ml, having a zeta potential of ~ +4 mV. Applications: Conjugation with bifunctional agents, Click chemistry, NHS esters, Isothiocyanates.	
CL-50Q02-15	Molday ION Aromatic Amine TM 2.0 ml of 40 nm iron-based superparamagnetic contrast agent containing aromatic amine groups packaged in a 2 ml sealed serum bottle. 2.5 mg Fe/ml having a zeta potential of ~ +48 mV. This nanoparticle is designed not to independently bind to or be internalized by cells. Therefore, cell labeling is achieved exclusively <i>via</i> your conjugated ligand. Applications: After activation to an isothiocyanate this nanoparticle will react with amine containing compounds.	395.00
CL-50Q02-161	Poly's L-Lysin Molday ION Amine Terminated TM\$ 2.0 ml of 40 nm iron-based 2.0 ml of 40 nm iron-based superparamagnetic contrast agent coated with L-lysine packaged in a 2 ml sealed serum bottle. 5 mg Fe/ml having a zeta potential of ~ +20 mV. The epsilon-amines groups of lysine are ideally suited for conjugation of ligands and macromolecules. This nanoparticle is designed not to independently bind to or be internalized by cells. Therefore, cell labeling is achieved exclusively via your conjugated ligand. Applications: Conjugation of amine containing nanoparticles, MRI, EM, Drug delivery, Theranostics. Magnetic cell-sorting.	475.00
Mol	Iday ION Containing Reactive Carboxyls	
CL-30Q02-6C	Molday ION Carboxyl Terminated TM	395.00
CL-30Q02-CA	Molday ION Carboxyl/Amine Terminated TM\$ 2.0 ml of 30 nm iron-based superparamagnetic contrast agent containing amine and carboxyl groups designed to approximate the surface of a protein packaged in a 2 ml serum bottle. 5 mg Fe/ml having a zeta potential of a +4 mV	475.00



in a 2 ml serum bottle. 5 mg Fe/ml, having a zeta potential of \sim +4 mV. <u>Applications:</u> Conjugation with bifunctional agents, Click chemistry, NHS esters, Isothiocyanates.

<u>Mold</u>	ay ION Containing Other Reactive Groups	
CL-30Q02-10	Molday ION Aldehyde TM	395.00
CL-50Q02-162	Poly's L-Tyrosine Molday ION Phenol Terminated TM\$ 2.0 ml of 40 nm iron-based superparamagnetic contrast agent coated with L-tyrosine packaged in a 2 ml sealed serum bottle. 5 mg Fe/ml having a zeta potential of ~ -5 mV. The phenol group of tyrosine is ideally suited for applications as a substrate for tyrosine kinases, iodination with 124-l (PET), 125-l (metabolic studies), and 127-l (CT). This nanoparticle is designed not to independently bind to or be internalized by cells. Therefore, cell labeling is achieved exclusively via your conjugated ligand. Applications: MRI, PET, CT, Metabolism studies, EM, Drug delivery, Theranostics.	475.00
Application 4: Recepto	or Targets	
Catalog <u>Number</u>		
CL-50Q02-6C-54	Molday ION Biotin TM	475.00
CL-50Q01-6C-54	Molday ION Biotin PEG TM 1.0 ml of 40 nm iron-based superparamagnetic contrast agent conjugated with Biotin packaged in a 2 ml sealed serum bottle. 1 mg Fe/ml. CL-50Q01-6C-54 reacts with streptavidin as shown by receptor double diffusion. Biotin Is conjugated to Molday ION through a PEG extended linker. Applications: Biotin-Streptavidin conjugation strategies, Cell labeling, Cell labeling with MRI tracking, Tumor tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	495.00
CL-160Q01-22	Molday ION StreptAvidin TM\$ 1.0 mg of Fe of 160 nm iron-based superparamagnetic contrast agent conjugated with Streptavin packaged in a 2 ml sealed serum bottle. CL-160Q01-22 reacts with biotin-fluorescein as shown by fluorescent measurements. Applications: Biotin-Streptavidin conjugation strategies, Cell labeling, Cell labeling with MRI tracking, Tumor tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	550.00
CL-100Q01-21	Molday ION GAM TM 1.0 ml of 100 nm iron-based superparamagnetic contrast agent conjugated with Goat anit-mouse IgG packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml. CL-100Q01-21 reacts with mouse IgG-fluorescein as shown by fluorescent measurements. Applications: Mouse-IgG conjugation strategies, Cell labeling, Cell labeling with MRI tracking, Tumor tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	550.00

 $Molday\ ION\ \textbf{Spermidine}^{\text{TM}}\\$ \ \ 475.00$

1.0 ml of 40 nm iron-based superparamagnetic contrast agent conjugated with spermidine packaged in a 2 ml sealed serum bottle. 1 mg Fe/ml having a zeta potential of \sim +35mV. CL-50Q02-71 enters cells via the polyamine receptor.

<u>Applications:</u> Cell labeling, Cell labeling with MRI tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.

CL-50Q02-71

PolyGalactose <i>Magnetite</i> ™\$	375.00
Packaged as 200 mg powder in a 10 ml sealed serum bottle.	
For additional information, please see Application Note #4 .	
	PolyGalactose Magnetite TM

Application 5: Fluorescent Labeled Iron Oxides

Catalog Number		
CL-50Q01-6A-50	Molday ION Rhodamine B TM (MIRB)\$ 0.5 ml of 35 nm rhodamine B labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~ +31 mV. A suggested procedure for labeling cells is provided as a PDF download on BioPAL's web site. PDF downloads depicting labeled cells, as well as presented posters, is also provided. Applications: Cell labeling, Cell labeling with MRI tracking, Live cell imaging (in vitro), Drug delivery, Theranostics, Fluorescent detection. For additional information, please review Application Note #3 on BioPAL Web Site.	140.00
CL-50Q02-6A-50	Molday ION Rhodamine B TM (MIRB)\$ 2.0 ml of 35 nm rhodamine B labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~+31 mV. A suggested procedure for labeling cells is provided as a PDF download on BioPAL's web site (Application Note #3). PDF downloads depicting labeled cells, as well as presented posters, is also provided. Applications: Cell labeling, Cell labeling combined with MRI and fluorescent tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	450.00
CL-50Q02-6A-51	Molday ION EverGreen TM 2.0 ml of 35 nm EverGreen labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~+31 mV. Labeled cells may be visualized using a standard fluorescein filter set. Molday Ion EverGreen has an excitation and emission maxima at 498 nm and 526 nm, respectively. Applications: Cell labeling, Cell labeling combined with MRI and fluorescent tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	450.00
CL-50Q02-6A-52	Molday ION Coumarin TM 2.0 ml of 35 nm Coumarin labeled iron-based superparamagnetic contrast agent packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~ +30 mV. Labeled cells may be visualized using a standard DAPI filter set. Molday Ion Coumarin has an excitation and emission maxima at 356 nm and 456 nm, respectively. Applications: Cell labeling, Cell labeling with MRI tracking, Live cell imaging (in vitro), Drug delivery, Theranostics, Fluorescent detection. For additional information, please review Application Note #3 on BioPAL Web Site.	450.00
CL-50Q01-6A-53	Molday ION TM MI-750	400.00
CL-50Q02-6C-50	Molday ION Rhodamine B Carboxyl TM\$ 2.0 ml of 35 nm rhodamine B labeled iron-based superparamagnetic contrast agent designed for conjugating a ligand, peptide, antibody, or protein using water soluble carbodiimide	450.00



packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of \sim -35 mV. This nanoparticle is designed not to independently bind to or be internalized by cells. Therefore, cell labeling is achieved exclusively *via* your conjugated ligand. A suggested procedure for conjugation of amine containing compounds is provided as a PDF download on BioPAL's web site (**Application Note #5**).

<u>Applications:</u> Conjugation of amine containing compounds using water soluble carbodiimide, MRI, EM, Drug delivery, Theranostics, Magnetic cell-sorting.

CL-50Q02-CA-50

2.0 ml of 50 nm rhodamine B labeled iron-based superparamagnetic contrast agent containing amine and carboxyl groups designed to approximate the surface of a protein packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml having a zeta potential of ~ +4 mV. <u>Applications:</u> MRI, EM, Fluorescent detection, Magnetic cell-sorting.

Application 6: Molecular Imaging

Catalog Number

CL-50Q02-6C-54	Molday ION Biotin TM	475.00
CL-50Q01-6C-54	Molday ION Biotin PEG TM 1.0 ml of 40 nm iron-based superparamagnetic contrast agent conjugated with Biotin packaged in a 2 ml sealed serum bottle. 1 mg Fe/ml. CL-50Q01-6C-54 reacts with streptavidin as shown by receptor double diffusion. Biotin Is conjugated to Molday ION through a PEG extended linker. Applications: Biotin-Streptavidin conjugation strategies, Cell labeling, Cell labeling with MRI tracking, Tumor tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	495.00
CL-160Q01-22	Molday ION StreptAvidin TM\$ 1.0 ml of 160 nm iron-based superparamagnetic contrast agent conjugated with Streptavin packaged in a 2 ml sealed serum bottle. 1.6 mg Fe/ml. CL-160Q01-22 reacts with biotin-fluorescein as shown by fluorescent measurements. Applications: Biotin-Streptavidin conjugation strategies, Cell labeling, Cell labeling with MRI tracking, Tumor tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.	550.00
CL-100Q01-21	Molday ION GAM TM 1.0 ml of 100 nm iron-based superparamagnetic contrast agent conjugated with Goat anit-mouse IgG packaged in a 2 ml sealed serum bottle. 2 mg Fe/ml. CL-100Q01-21 reacts with mouse IgG-fluorescein as shown by fluorescent measurements.	550.00

Application 7: Molday ION Gycoprotien

Catalog



Applications: Mouse-IgG conjugation strategies, Cell labeling, Cell labeling

with MRI tracking, Tumor tracking, EM, Magnetic cell-sorting, Drug delivery, Theranostics.

Number

CL-50Q01-OVA	Molday ION TM Ovalbumin\$ 1.0 ml of 40 nm iron-based superparamagnetic contrast agent conjugated with ovalbumin packaged in a 2 ml sealed serum bottle. 1 mg Fe/ml having a zeta potential of ~-11mV. Quality control by ovalbumin ELISA.	400.00
FIT-0615	Egg Ovalbumin ELISA Kit The kit contains ovalbumin concentrate, ovalbumin antiserum, goat anti-rabbit IgG-HRP, HRP substrate and stop reagents, an ovalbumin 96 well coated plate, two plate sealers, FIT-GFR Inulin kit manual and documentation.	\$ 200.00

Application 8: Helpful Products for Staining and Fixing Iron-Labeled Cells

Catalog Number

<u>ivumber</u>		
CL-01-50	Prussian Blue Reagent Pack	65.00
CL-01-51	$PBS++ \\ Phosphate buffered saline supplemented with (0.1 g/L CaCl_2) and (0.1 g/L MgCl_2). \\ Amount 100 ml, Store at 4°C. \\ Applications: Cell Fixation. \\ The procedure for fixing cells is provided as a separate PDF download on BioPAL's web site (Application Note #3). \\ \\ \$$	45.00
CL-01-52	25% Glutaraldehyde\$ Amount 5 ml, Store at 4°C. Applications: Cell Fixation. The procedure for fixing cells is provided as a separate PDF download on BioPAL's web site (Application Note #6).	30.00
CL-01-53	40% Formalin\$ Amount 5 ml, Store at 4°C. Applications: Cell Fixation. The procedure for fixing cells is provided as a separate PDF download on BioPAL's web site (Application Note #6).	30.00

References

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- 2. Lee J, Hirano Y, Fukunaga M, Silva A, and Dyun, JH: Investigating the sources of phase contrast: iron oxide nanoparticle study to exclude deoxyhemoglobin as a major source for the gray/white matter phase contrast. *NeuroImage* (2009) **47** Supplement 1, S39-S41.
- 3. Bogdanov AA, Jr., Martin C, Weissleder R, Brady TJ: Trapping of dextran-coated colloids in liposomes by transient binding to aminophospholipid: preparation of ferrosomes. *Biochim Biophys Acta* (1994) **1193**; 212-8.
- 4. Groman EV, Yang M, Reinhardt, CP, Weinberg, JS, Vaccaro, DE: Polycationic Nanoparticles: (1) Synthesis of a Polylysine-MION Conjugate and its Application in Labeling Fibroblasts. *J of Cardiovasc Trans Res* (2009) **2**; 30-38.



 $FeRex^{TM}$ is a superparamagnetic magnetite. This reagent is calassified as a darkening reagent acting through the T2 relaxation process. FeRex can be used as an MRI contrast reagent.

References

- 1. J Immunol Methods 1982 52(3):353-367.
- 2. FEBS Letters 1984 170(2):282-238.
- 3. J Aerosol Med Pulmonary Drug Delivery 2008 21(4):335-341.

NOT FOR HUMAN USE.

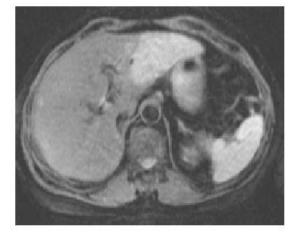
Superparamagnetic Magnetite

Catalog Number

MR-7200 Ferex.....\$ 400.00

2.0 ml sealed serum bottle at a concentration of 10 mg Fe/ml. NOT FOR HUMAN USE.

Assay Method: MRI and EM.







The twenty-first century began with the completion of the human genome project and the dawn of proteomics and systems-biology. These emerging fields will attempt to understand how newly discovered proteins impact human physiology and disease. As researchers progress from molecular- and cellular-based studies to *in vivo* systems, innovative technologies are needed to allow multiple tracking of probes and multiple measurements of functional parameters in *in vivo* models. One arena of research that has emerged due to recent discoveries has been the field of cellular transplantation.

BioPAL has developed a biodegradable 1µm superparamagnetic particles and a long-lived (non-biodegradable) superparamagnetic 1 µm microsphere for applications in cell labeling and drug delivery experiments for MRI technology. These reagents are classified as a darkening agent acting through the T2 relaxation process.

NOT FOR HUMAN USE.

1 µm Iron-Labeled Particles and Microspheres

Catalog		
Number CL-01Q02-B	Biodegradable MicroTRACK™ 1 µm biodegradable iron microparticles. 1 ml packaged in a 2 ml sealed serum bottle, containing approximately 8.4 billion particles. Particles are packaged in an aqueous suspension at 25 mg Fe/ml.	100.00
CL-01Q02-B-A	MicroTRACK [™] Amine-terminated	150.00
CL-01Q02-B-50	Micro TRACK™ Rhodamine B\$ 1 µm (± 0.5 µm) silane-coated, biodegradable iron microparticles labeled with rhodamine B. 1 ml packaged in a 2 ml sealed serum bottle (~8.4 billion particles). Particles are packaged in an aqueous suspension at 25 mg Fe/ml.	300.00
CL-01Q02-L	Non-Biodegradable MicroTRACK™ \$ 1 µm (± 0.5 µm) polystyrene, non-biodegradable iron microspheres. 1 ml packaged in a 2 ml sealed serum bottle, containing approximately 8.4 billion microspheres. Particles are packaged in an aqueous suspension at 5 mg Fe/ml.	100.00
CL-01Q02-L-50	MicroTRACK [™] Rhodamine B\$ 1 µm (± 0.5 µm) polystyrene, non-biodegradable iron microspheres labeled with rhodamine B. 1 ml packaged in a 2 ml sealed serum bottle (~8.4 billion microspheres). Particles are packaged in an aqueous suspension at 5 mg Fe/ml.	300.00



There has been increasing interest in modeling injection materials in *in vivo* systems using MRI technology. For example, one could model the fate of implanted cells, using a 15 μ m labeled microsphere or visualize an embolic region using a labeled 100 μ m microsphere. Therefore, BioPAL has developed a product-line specific for these applications. Upon request, BioPAL can provide different sized microspheres labeled with iron.

NOT FOR HUMAN USE.

Iron-Labeled Microspheres

Catalog

Number		
C-15Q10	Ferro <i>TRACK</i> [™] \$ 15 μm in diameter, iron-labeled microspheres and packaged in a 10 ml sealed serum bottle, containing approximately 25 million microspheres. 25 mg Fe/ml. Applications: Vascular imaging, Injection and injection-site visualization.	400.00
C-100Q10	Ferro TRACK [™] \$	400.00

100 µm in diameter, iron-labeled microspheres and packaged in a 10 ml sealed serum bottle, containing approximately 84,000 microspheres. 25 mg Fe/ml. Applications: Embolism studies and visualization.



Molday LION™ **Rhodamine B** is an ultrasmall mixed ferrite iron oxide-based superparamagnetic contrast reagent having a colloidal size of 35 nm. The core crystal is europium-doped magnetite surrounded by a cross-linked dextran to which a fluorescent dye, rhodamine B, is attached through a C6 amino group. This agent can be visualized by MRI and fluorescence, and can be quantified by neutron activation *via* BioPAL's short enhanced assay service. For neutron activation applications, please discuss your protocol with BioPAL in advance of purchase. For MRI, this agent is classified as a darkening agent acting through the T2 relaxation process. For fluorescence, the rhodamine B label excitation wavelength is 555 nm and the emission is at 565-620nm.

In addition to rhodamine B, alternative fluorescent dyes can be conjugated to CL-50KQ02-6A and we are pleased to discuss alternative conjugation procedures.

NOT FOR HUMAN USE.

Fluorescent Magnetite MRI Contrast Agent

Catalog
<u>Number</u>

CL-50KQ02-6A

Molday LION

2.0 ml of a 35 nm mixed ferrite iron oxide contrast agent containing europium having a positive surface packaged in a 2 ml sealed serum bottle. 1 mg Fe/ml. Applications: MRI, EM, Fluorescent detection, Neutron activation, Magnetic cell-sorting.

CL-50KQ02-6A-50

Molday LION Rhodamine B

1.0 ml of a 35 nm rhodamine B mixed ferrite iron oxide contrast agent

packaged in a 2 ml sealed serum bottle. 1 mg Fe/ml.

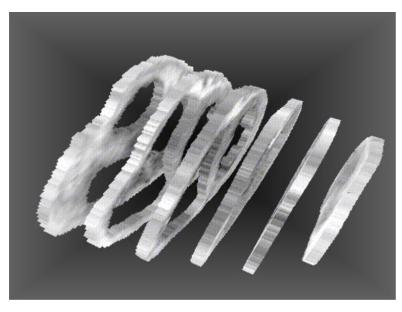
Applications: MRI, EM, Fluorescent detection, Neutron activation, Magnetic cell-sorting.

REF: Bioconjugate Chemistry 2007:18;1763-1771.



Magnetic resonance imaging (MRI) is an imaging technique used primarily in medical research to produce high quality images of *in vivo* systems. To track materials within *in vivo* systems, compounds-of-interest are tagged with materials that possess magnetic properties. These materials, as well as MRI contrast agents, generally fall into two classes. The first is soluble gadolinium compounds that act through the T1 relaxation process and are brightening agents. The second is particulate iron oxides that act through the T2 relaxation process and are darkening agents. Each type of contrast agent has its benefits.

BioPAL provides the research community with a comprehensive MRI product-line. This product-line includes standard MRI contrast agents, such as Gd-DTPA and Gd-DOTA, sold under our



brand name Gado-DTPATM and Gado-DOTATM, respectively. In addition, to aid researchers BioPAL offers immunoassay kits (ELISA) to measure both of these compounds in collected biological fluids.

Gadolinium-DTPA

Gadolinium-DTPA and gadolinium-DOTA are a brightening agent acting through the T1 relaxation process. BioPAL's Gado-DTPA is bottled at a concentration of 469 mg/ml and Gado-DOTA is bottled at a concentration of 279 mg/ml. NOT FOR HUMAN USE.

Catalog Number

MR-00P10	Gado-DTPA\$ 10.0 ml sealed serum bottle of gadolinium-DTPA at a concentration of 469 mg/ml. Applications: Perfusion.	50.00
MR-00P11	Gado-DOTA\$ 1.0 ml sealed serum bottle of gadolinium-DOTA at a concentration of 279 mg/ml. Applications: Perfusion.	250.00



BioPAL provides the research community with a comprehensive MRI product-line. BioPAL has applied its unique isotopic labeling technology to generate novel MRI reagents for specific applications. For example BioPAL offers gadolinium-labeled albumin and cholic acid. These products can be used for a wide range of applications.

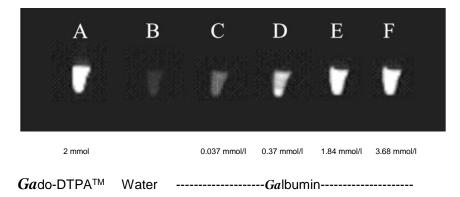


Figure: Using Spin Echo (SE) sequence with TR/TE = 100/14, images of gadolinium labeled solutions in microtubes were obtained. The average signal intensity within a region-of-interest (ROI) is listed below. The background intensity measured 0.7 ± 0.5 .

Gadolinium-Labeled Albumin

Cotolog

Gadolinium-labeled albumin is a brightening agent acting through the T1 relaxation process. BioPAL's Galbumin is gadolinium-labeled bovine albumin bottled at a concentration of 25 mg/ml. There are approximately 10 to 15 gadolinium atoms per albumin unit. NOT FOR HUMAN USE.

Catalog <u>Number</u>		
P-00P01	Galbumin TM \$ Gadolinium-labeled albumin. 1.0 ml at a concentration of 25 mg/ml, packaged in a 2 ml sealed serum bottle. Applications: Vascular imaging, Injection and injection-site visualization.	135.00
P-00P01-100	Concentrated <i>Ga</i> lbumin TM	600.00
P-00PK01	Trackable <i>Ga</i> lbumin TM	150.00
P-00PZ01	Glowing <i>Ga</i> lbumin TM - Fluorescein\$ Gadolinium-labeled albumin co-labeled with FITC for fluorescent imaging. 1.0 ml at a concentration of 25 mg/ml, packaged in a 2 ml sealed serum bottle. Applications: Vascular imaging, Injection and injection-site visualization.	175.00
P-00PY01	Glowing <i>Ga</i> Ibumin TM - Rhodamine B\$ Gadolinium-labeled albumin co-labeled with rhodamine B for fluorescent imaging. 1.0 ml at a concentration of 25 mg/ml, packaged in a 2 ml sealed serum bottle. Applications: Vascular imaging, Injection and injection-site visualization.	175.00



BioPAL provides the research community with a comprehensive MRI product-line. **Gd**-IgG[™] is a gadolinium-labeled IgG bottled at a concentration of 25 mg/ml. There are approximately 10 to 15 gadolinium atoms per IgG unit and the molecular weight of the product is approximately 160 kDa. This reagent is engineered to be used as a MRI contrast reagent and is classified as a brightening agent acting through the T1 relaxation process.

NOT FOR HUMAN USE.

Gadolinium-Labeled IgG

Cata	log
Numl	ber

Gadolinium-labeled IgG. 1.0 ml at a concentration of 25 mg/ml, packaged in a 2 ml sealed serum bottle. Applications: MRI, Vascular imaging, Injection and injection-site visualization

Gado-Inulin™ is a gadolinium-labeled inulin bottled at a concentration of 25 mg/ml. There are approximately 10 to 15 gadolinium atoms per inulin unit and the molecular weight of the product is approximately 160 kDa. BioPAL uses USP Grade inulin, which its availability was in part supported by the NIH-SBIR Program (DK088349). Inulin is classically recognized as the gold standard renal probe for the measurement of glomerular filtration rate (GFR). This reagent is engineered to be used as a MRI contrast reagent that can be used in nephrology research and is classified as a brightening agent acting through the T1 relaxation process.

NOT FOR HUMAN USE.

Gadolinium-Labeled IgG

Catalog <u>Number</u>

P-00P03 **Gado**-Inulin™\$ 175.00

Gadolinium-labeled Inulin. 1.0 ml at a concentration of 25 mg/ml, packaged in a 2 ml sealed serum bottle. Applications: MRI, Renal imaging, Injection and injection-site visualization



GadoluminateTM is the first commercially available ultra-small gadolinium oxide colloid for use in a wide range of MRI and MRA applications. This product is a 30 nm gadolinium colloid (CL-20P02-2). As shown in the figure below, this reagent provides a strong T1 signal that is comparable to the same molar concentration as Gd-DTPA. However, due to its colloidal size and neutral charge, this reagent is retained in the vascular space. The measured blood half-life ($t_{1/2}$) of Gadoluminate in a swine model is approximately 12.5 hours, as compared to ~11.9 minutes in mouse model.

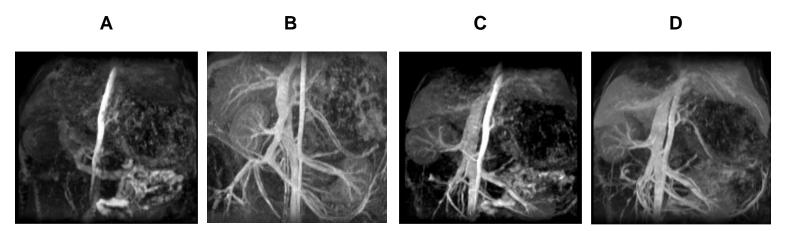


Figure 1: MIP reconstructed 3D images depicting pre and post administration of *Gado*luminate: **A** pre-contrast image, **B** 0.10 mmol Gd/kg, **C** 0.04 mmol Gd/kg, **D** 0.01 mmol Gd/kg. MRA images were acquired on a 3 T clinical scanner (Tio, Siemens) using a 7-inch volume coil and a 3D gradient echo (FLASH) sequence. Axial images were acquired to cover the abdominal region including liver and kidneys with acquisition parameters TR/TE = 19/4.9, FOV 180x124x128, matrix 128x176x256, effective slice thickness 1 mm, receiver bandwidth 120 Hzs/pixel, flip angle 30°.

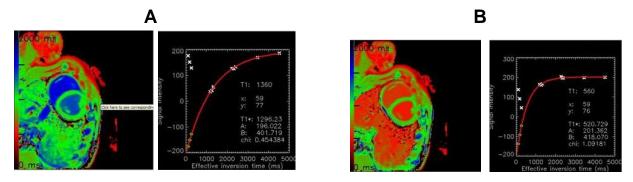


Figure 2: The change in T1 signal in blood following a 0.01 mmol Gd/kg dose of **Gado**luminate in a swine model. **A** is pre-dose and **B** is post dose administration. A Modified Look-Locked Inversion Recovery sequence with 11 different inversion times was performed. The signal intensity was measured in blood.

Ultra-Small Gadolinium Oxide Colloidal MRI Reagent

Catalog Number

CL-20P02-2 **Gado**luminateTM\$

 $2.0\ ml$ of $30\ nm$ gadolinium colloid (10 mg Gd/ml) packaged in a 2 ml sealed serum bottle. Neutral charge. Assay Method: MRI, MRA and EM.



450.00

GADOPOLYLYSINETM is a gadolinium-labeled polylysine. This reagent is engineered to be used as a MRI contrast reagent and is classified as a brightening agent acting through the T1 relaxation process. GADOPOLYLYSINE is designed for diffusion studies, for labeling cells, and to allow efficient labeling with ligands, peptides, proteins, and nucleic acid (RNAi, DNA, etc.), wherein a T1 weighted MRI contrast agent is desired. Other applications for these MRI contrast agents include use in drug delivery and use as a duel purpose colloid that combines drug delivery with imaging applications.

NOT FOR HUMAN USE.

Gadolinium-Labeled Polylysine

Catalog Number		
P-01P01-01	GADOPOLYLYSINE (10 kDa)\$ 1.0 ml of gadolinium-labeled 10 kDa polylysine packaged in a 2 ml sealed serum bottle. 10 mg polylysine per ml. Applications: MRI and EM.	175.00
P-01P01-02	GADOPOLYLYSINE (22 kDa)	175.00
P-01P01-05	GADOPOLYLYSINE (50 kDa)	175.00
P-01P01-11	GADOPOLYLYSINE (110 kDa)\$ 1.0 ml of gadolinium-labeled 110 kDa polylysine packaged in a 2 ml sealed serum bottle. 10 mg polylysine per ml. Applications: MRI and EM.	175.00
P-01P01-22	GADOPOLYLYSINE (225 kDa)\$ 1.0 ml of gadolinium-labeled 225 kDa polylysine packaged in a 2 ml sealed serum bottle. 10 mg polylysine per ml. Applications: MRI and EM.	175.00



GADO $Dextran^{\text{TM}}$ is a gadolinium-labeled dextran. This reagent is engineered to be used as a MRI contrast reagent and is classified as a brightening agent acting through the T1 relaxation process. Similar to GADO $POLYLYSINE^{\text{TM}}$, GADODextran is designed for diffusion studies, for labeling cells, and to allow efficient labeling with ligands, peptides, proteins, and nucleic acid (RNAi, DNA, etc.), wherein a T1 weighted MRI contrast agent is desired. Other applications for these MRI contrast agents include use in drug delivery and use as a duel purpose colloid that combines drug delivery with imaging applications.

NOT FOR HUMAN USE.

Gadolinium-Labeled Dextran

Catalog

<u>Number</u>		
P-02P01-01	GADODextran (10 kDa)	175.00
P-02P01-04	GADODextran (40 kDa)	175.00
P-02P01-07	GADODextran (70 kDa)\$ 1.0 ml of gadolinium-labeled 70 kDa dextran packaged in a 2 ml sealed serum bottle. 10 mg dextran per ml. Applications: MRI.	175.00



BioPAL has developed Gado $CELL\mathsf{Track}^\mathsf{TM}$ to aid researchers locate, track, and quantify implanted cells in *in vivo* systems using MRI technology. This product is a 50 nm gadolinium colloid (CL-50P02-6) that is negatively charged. As shown in the figure below, this reagent provides a strong T1 signal that is comparable to the same molar concentration as Gd-DTPA. Like all our colloidal markers, these reagents can also be measured by electron microscopy (EM).

BioPAL's *IVST* reagents, which contain both gadolinium for MRI imaging, as well as additional isotopes for quantification *via* neutron activation can be found in the Neutron Activation section of our catalog and web site.

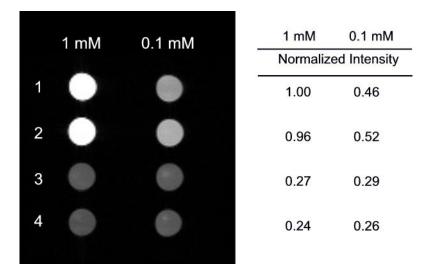


Figure: T1 weighted image from a series of compounds having the same molar concentration are shown on the left and normalized intensity values are tabulated on the right. Row 1 is Gd-DTPA, Row 2 is colloidal gadolinium, Row 3 is colloidal europium having the same size and matrix as colloidal gadolinium, Row 4 is PBS.

Colloidal Gadolinium Cell-Labeling Reagent

Catalog Number

CL-50P02-6	Gado <i>CELL</i> Track TM (-)\$ 2.0 ml of 50 nm gadolinium colloid (10 mg Gd/ml) packaged in a 2 ml sealed serum bottle. Negative charge. Assay Method: MRI and EM	450.00
CL-00-01	Poly-L-Lysine	60.00



BioPAL offers Evans Blue-DTPA-Gadolinium a novel functionalized MRI contrast agent that recognizes lesions in the vascular endothelium. Conventional MRI agents image the overall vasculature. However, these agents cannot directly detect and image endothelial lesions. Evans Blue-DTPA-Gadolinium is comprised of a detection unit (Evans-Blue dye) linked to an imaging unit (Gadolinium-DTPA). The dye component selectively adsorbs to a vascular endothelium-denuded region, and with the imaging component marks the site. Experimental work with this agent has been performed *in vivo* in apolipoprotein-E-deficient (ApoE-/-) mice of varying ages. In the mice experiments, MRI signal enhancement is seen only in the inner surface of the aorta wall, which means that the contrast agent does not permeate the tissue; this is important for MR imaging of blood vessels since rapid excretion is preferable. Signal intensity at the injured vascular site is 1.74 times as high as that of the normal site 10 minutes after injection; after 30 minutes the signal weakened and at 120 minutes there was no observed difference in signal intensity, indicating rapid excretion from the body. When administered intravenously, our agent effectively detected atherosclerotic plaques in the aorta, areas indicative of endothelial dysfunction. Interestingly, the signal intensity increased with age, consistent with the notion of age-related atherosclerosis. This agent is selective and specific for detecting vascular lesions and may be used to follow progression and regression of atherosclerosis and also perhaps aid in the management of cardiovascular diseases.

Evans Blue-DTPA-Gadolinium

Evans Blue-DTPA-Gadolinium is a brightening agent acting through the T1 relaxation process. This product is gadolinium-labeled Evans Blue dye bottled at a concentration of 10 mg. NOT FOR HUMAN USE.

Catalog <u>Number</u>		
GD-00P01	Evans Blue-DTPA-Gadolinium\$ Gadolinium-labeled Evens Blue dye. 10 mg packaged as a dry powder in a 2 ml sealed serum bottle. Applications: Vascular imaging, Endothelium lesion recognition.	400.00
GD-01P05	Buffered Saline Solution\$ Buffered saline solution used to reconstitute Evans Blue-DTPA-Gadolinium. 5 ml in a 10 ml sealed serum bottle.	50.00

Advantages:

- It is a targeted contrast agent that enhances the capability and specificity of MRI.
- The imaging agent does not permeate the tissue and is rapidly excreted out of the body within 120 min.
- It may aid in early intervention for both primary and secondary treatment of cardiovascular diseases.

Publications

Yasuda, S., Ikuta, K., et al. In vivo Magnetic Resonance Imaging of Atherosclerotic Lesions with a Newly Developed Evans Blue-DTPA-Gadolinium Contrast Medium in Apolipoprotein-E-Deficient Mice [J Vasc Res (2008) 45, 123-128]

Intellectual Property

JP 4569875, US 7,368,099



BioPAL's gadolinium protein labeling kit contents are specifically designed to support our FIT-MITT ELISA system. The labeling kit contains the bifunctional chelate (either DOTA or DTPA), gadolinium salt, required buffers, and instructions. The kit is designed to label up to 10 mg of protein. If planning to use your labeled compound for MRI experiments, the sensitivity of MRI detection is a function of many factors, including labeling efficiency and sample composition. Therefore, each researcher will need to validate MRI detection for their unique application. BioPAL also offers a custom protein labeling service. Customers should contact BioPAL if interested in custom labeling services.

The kits are nonradioactive.

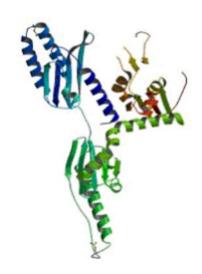
Catalog

PL-00P03

PROTrack - Gadolinium Protein Labeling Kits

PROTrack Gadolinium (DOTA).....\$ 675.00

Second generation kit designed to minimize potential precipitation during chelation. The kit contains the bifunctional chelate, gadolinium salt, required buffers, and instructions. The kit is designed to label 1 to 10 mg of protein.





FIT-MITT kits provide a superior analytical method to measure Gd-DTPA labeled compounds. BioPAL's FIT-MITT test relies on immunoassay as its analytical segment. Depending on the protocol chosen, the kit can support up to 42 unknown samples performed in duplicate. The kit does not contain the gadolinium-DTPA labeling kit. The labeling kit is sold separately. Reminder: Researchers MUST develop a standard curve using their gadolinium-labeled compound.

FOR RESEARCH USE ONLY. NOT FOR USE IN DIAGNOSTIC PROCEDURES.

Patents Pending.

Gd-DTPA FIT-MITT Catalog	Γ – Kit		
Number FIT-0115	FIT-MITT Kit (Gd-DTPA) The kit contains Gd-DTPA concentrate, rabbit Gd-DTPA antiserum, HRP-Gd-DTPA conjugate, HRP substrate and stop reagents, a goat anti-rabbit IgG coated 96 well plate, a plate sealer, FIT-MITT Gd-DTPA kit manual and documentation.	. \$ (600.00
Gd-DTPA FIT-MITT			
FIT-0101	Rabbit Anti-Gadolinium-DTPA	. \$ 2	250.00
FIT-0102	HRP-Gadolinium-DTPA 6.0 ml packaged in an 8 ml polyethylene screw-cap bottle.	. \$ 2	250.00
FIT-0103	Gadolinium-DTPA Concentrate	. \$	50.00
FIT-0104	Gd-DTPA Controls (High and Low)	. \$	50.00
FIT-MITT - Genera	l Kit Components		
FIT-0001	Standard and Sample Diluent	. \$	25.00
FIT-0002	HRP Substrate Reagent	. \$	35.00
FIT-0003	HRP Stop Reagent	. \$	25.00
FIT-0004	Plate Sealer	. \$	20.00
FIT-0005	Wash Buffer Concentrate	. \$	30.00
FIT-0006	Goat Anti-Rabbit IgG Coated 96 Well Plate		
FIT-0009	Long-Term Storage Diluent	. \$	35.00



FIT-MITT kits provide a superior analytical method to measure Gd-DOTA labeled compounds. BioPAL's FIT-MITT test relies on immunoassay as its analytical segment. Depending on the protocol chosen, the kit can support up to 42 unknown samples performed in duplicate. The kit does not contain the gadolinium-DOTA labeling kit. The labeling kit is sold separately. Reminder: Researchers MUST develop a standard curve using their gadolinium-labeled compound.

FOR RESEARCH USE ONLY. NOT FOR USE IN DIAGNOSTIC PROCEDURES.

Patents Pending.

Gd-DOTA FIT-MIT Catalog Number FIT-0215	FIT-MITT Kit (Gd-DOTA) \$600.00 The kit contains Gd-DOTA concentrate, rabbit Gd-DOTA antiserum, HRP-Gd-DOTA conjugate, HRP substrate and stop reagents, a goat anti-rabbit IgG coated 96 well plate, a plate sealer, FIT-MITT Gd-DOTA kit manual and documentation.
Gd-DOTA FIT-MIT	T – Kit Components
FIT-0201	Rabbit Anti-Gadolinium-DOTA
FIT-0202	HRP-Gadolinium-DOTA
FIT-0203	Gadolinium-DOTA Concentrate \$ 60.00 125.0 µl packaged in a 0.6 ml microtube.
FIT-0204	Gd-DOTA Controls (High and Low)
FIT-MITT - Genera	L Kit Components
FIT-0001	Standard and Sample Diluent
FIT-0002	HRP Substrate Reagent
FIT-0003	HRP Stop Reagent
FIT-0004	Plate Sealer \$20.00 Pack of 5.
FIT-0005	Wash Buffer Concentrate
FIT-0006	Goat Anti-Rabbit IgG coated 96 Well Plate

