

Magnetofection™ *In vivo*

INSTRUCTION MANUAL

Magnetofection™ *In vivo* Kits

In vivo PolyMag
In vivo DogtorMag
In vivo ViroMag



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In vivo Magnetofection™

Instruction Manual

In vivo Magnetofection™ is an original, simple and efficient method to transfect target cells/tissue *in vivo*.

List of *In vivo* Magnetofection™ kits

Catalog Number	Product	Volume	Number of injections	Suitable for
IV-KC30210	<i>In vivo</i> PolyMag Starting kit ¹	500 µL	5-50 injections	500 µg
IV-KC30220	<i>In vivo</i> DogtorMag Starting kit ²	500 µL <i>In vivo</i> Dogtor + 500 µL <i>In vivo</i> CombiMag	5-50 injections	500 µg
IV-KC30230	<i>In vivo</i> ViroMag Starting kit ³	250 µL	10-25 injections	1-2 x 10 ⁷ pfu
IV-PN30500	<i>In vivo</i> PolyMag 500	500 µL	5-50 injections	500 µg DNA
IV-PN31000	<i>In vivo</i> PolyMag 1000	1000 µL	10-100 injections	1000 µg DNA
IV-DM30500	<i>In vivo</i> DogtorMag 500	500 µL <i>In vivo</i> Dogtor + 500 µL <i>In vivo</i> CombiMag	5-50 injections	500 µg DNA
IV-DM31000	<i>In vivo</i> DogtorMag 1000	1000 µL <i>In vivo</i> Dogtor + 1000 µL <i>In vivo</i> CombiMag	10-100 injections	1000 µg DNA
IV-VM30250	<i>In vivo</i> ViroMag 250	250 µL	10-25 injections	1-2 x 10 ⁷ pfu
IV-VM30500	<i>In vivo</i> ViroMag 500	500 µL	20-50 injections	2.5-5 x 10 ⁷ pfu
IV-MAG1	Magnets set ⁴	/	/	/
IV-MAG2	Square magnets set ⁵	/	/	/
IV-MAG3	Cylinder magnets set ⁶	/	/	/

¹ contains 1 vial of *In vivo* PolyMag and a Magnets set

² contains 1 vial of *In vivo* Dogtor, *In vivo* CombiMag and a Magnets set

³ contains 1 vial of *In vivo* ViroMag and a Magnets set

⁴ contains 1 extra small cylinder (ø 2 mm), 1 small cylinder (ø 5 mm), 1 cylinder (ø 10 mm) and 1 square (18x18 mm) magnets

⁵ contains 4 square magnets (18x18 mm)

⁶ contains 4 extra small cylinder (ø 2 mm), 4 small cylinder (ø 5 mm), 4 cylinder (ø 10 mm) magnets

Use the content of the list above to determine the appropriate catalog number for your needs. You can order these products by contacting us (phone, fax, email & website). For all other information, do not hesitate to contact our technical support (tech@ozbiosciences.com).

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1. Technology

1.1. Description

Congratulations on your purchase of the *In vivo* Magnetofection™ reagent!

In vivo Magnetofection™ is a rapid, simple and highly efficient method especially dedicated to transfect and/or transduce target cells/tissue *in vivo*. This original system combines magnetic nanoparticles and nucleic acid vectors that will be retained after injection at the magnetically targeted site. In this way, targeted delivery minimizes systemic distribution, decreases gene vectors inactivation and reduces toxicity. Furthermore, the magnetic forces enhance the uptake of magnetic nanoparticles by the target tissue, and thus improve the efficiency of transfection/transduction. This allows decreasing the required process time of delivery to few minutes which is crucial for improvement of *in vivo* nucleic acid delivery (principally in stringent conditions, refer to results for more information). All *In vivo* Magnetofection™ reagents have been especially designed by our R&D team to meet *in vivo* grade quality (reagents performed under high manufacturing and quality standards and tested by strict quality controls).

In vivo Magnetofection™ main advantages in comparison to standard procedures:

1. Increased transfection/transduction efficiency
2. Targeted process (magnetically-driven)
3. Reduction of the systemic dissemination of vectors during injection
4. Reduction of the vector doses (nucleic acid, virus...)
5. Work under non permissive conditions (hypothermia, physiological flow conditions)
6. Penetration of the vector into tissues
7. Minimized toxicity
8. Universal – suitable for all nucleic acids or viral vectors

In vivo Magnetofection™ magnetic nanoparticles are non-toxic, biodegradable and totally biocompatible. The toxicity of iron oxide-based magnetic nanoparticles has been extensively studied due to their widespread use in medical imaging such as MRI, cell tracking and hyperthermia or cancer therapies as well as their use in diagnosis and cell purification. The amount of iron oxide required for nucleic acid delivery is far below the doses inducing toxicity. The iron oxide contained in Magnetofection™ nanoparticles has been shown to degrade through natural iron metabolism pathway (for review, Gupta and Gupta, 2005; Laurent *et al.*, 2011; Plank *et al.*, 2011).

1.2. Available Reagents

OZ Biosciences offers three types of ready-to-use *In vivo* Magnetofection™ reagents:

1. ***In vivo* PolyMag** is a cationic polymer-based magnetic nanoparticles formulation. It can be used with any nucleic acid (plasmid DNA, antisense oligonucleotides, mRNA, shRNA and siRNA...)
2. ***In vivo* DogtorMag** is a cationic lipid-based magnetic nanoparticles formulation. It associates ***In vivo* DogtorMag**, a specific cationic lipid, and **CombiMag** magnetic nanoparticles. This reagent is suitable for any nucleic acid (plasmid DNA, antisense oligonucleotides, mRNA, shRNA and siRNA...)
3. ***In vivo* ViroMag** is an optimized nanoparticles formulation dedicated to viral vectors. It is particularly suitable for Lentiviral/Retroviral, Adenoviral and Adeno-Associated Viral (AAV) vectors.

1.3. Kit Contents, Stability and Storage

Kit contents differ according to the components and volumes:

1. ***In vivo* PolyMag.**
 - 1 tube containing 500 µL of *In vivo* PolyMag suitable for 5 to 50 injections
 - 1 tube containing 1 mL of *In vivo* PolyMag suitable for 10 to 100 injections
2. ***In vivo* DogtorMag.**
 - 1 tube containing 500 µL of *In vivo* CombiMag and 1 tube containing 500 µL of *In vivo* Dogtor suitable for 5 to 50 injections
 - 1 tube containing 1 mL of *In vivo* CombiMag and 1 tube containing 1 mL of *In vivo* Dogtor suitable for 10 to 100 injections
3. ***In vivo* ViroMag.**
 - 1 tube containing 250 µL of *In vivo* ViroMag suitable for 10 to 25 injections
 - 1 tube containing 500 µL of *In vivo* ViroMag suitable for 20 to 50 injections

Stability, Storage and shipping conditions:

Storage: Upon receipt and for long-term use, store all tubes at the indicated storage conditions.

- *In vivo* PolyMag: store at +4°C
- *In vivo* DogtorMag: store at -20°C and *In vivo* CombiMag: store at +4°C
- *In vivo* ViroMag: store at +4°C
- **Do not freeze the nanoparticles**
- **Do not add anything to the stock solution of magnetic particles**

Stability: Magnetofection™ kits are stable for at least 18 months at the recommended storage temperature.

Shipping condition: room temperature.

Magnets manipulation:

- Manipulate carefully the magnets. Danger of injury by strong magnetic attraction of ferromagnetic materials
- Keep away from electronic devices and magnetic storage devices
- Persons with cardiac pacemakers should not work with these magnets

2. Applications

2.1. Nucleic acids and vectors

In vivo Magnetofection has been developed for *in vivo* targeted transfection of various types of nucleic acids such as DNA, RNA, oligonucleotides (*In vivo* PolyMag / *In vivo* DogtorMag) or transduction with any viral vectors (*In vivo* ViroMag). DNA/nanoparticles or virus/nanoparticles can be easily administrated through

various injection routes such as systemic administration (intravenous, intra-artery) or local administration (intraperitoneal, intratumoral, intracerebroventricular, intramuscular).

The instructions given hereunder represent protocols that were successfully applied in several studies. Nevertheless, optimal conditions vary depending on the nucleic acid, viral vector, animal model, administration route and the target organ. Therefore, use the Table 1 as a starting point for DNA amount and volume of injection in mouse and rats and Table 3 for viral vectors amount.

Notes:

- Nucleic acids should be as pure as possible, endotoxins free and prepare in water
- For the complexes preparation and injection, prefer saline buffer (HBS, PBS, normal saline, Ringer's solution) or glucose 5% solution.

Table 1: Suggested amount of nucleic acid and volume of injection in mouse (20 g) and rat (250 g)

Mouse			
Route of injection	Amount of nucleic acid	Total volume of injection according to animal weight	Site of injection
<i>Intravenous</i>	40 µg	200 µL (10-25 µL/g)	Tail vein
<i>Intramuscular</i>	10 to 100 µg	100 µL (50 µL x 2 sites of injection)	Caudal thigh
<i>Subcutaneous</i>	10 µg	200 µL (10-40 µL/g)	Scruff
<i>Intraperitoneal</i>	100 µg	400 µL (20 µL/g)	Lower Ventral Quadrants
<i>Intratumoral</i>	10 to 50 µg	100 µL (0.5 µL/mm ³)	Tumor
<i>Intracerebroventricular</i>	0.5 µg	2 µL	Brain ventricle
Rat			
Route of injection	Amount of nucleic acid	Total volume of injection	Site of injection
<i>Intravenous</i>	150 µg	2.5 mL (10-20 µL/g)	Tail vein, saphenous vein
<i>Intramuscular</i>	50 to 300 µg	300 µL (100 µL x 3 sites of injection)	Triceps, Quadriceps, Gluteals
<i>Subcutaneous</i>	5 to 10 µg	1.25 mL (5-10 µL/g)	Scruff, Back, Abdomen
<i>Intraperitoneal</i>	200 µg	2.5 mL (10-20 µL/g)	Lower Ventral Quadrants
<i>Intratumoral</i>	10 to 50 µg	100 µL (0.5 µL/mm ³)	Tumor
<i>Intracerebroventricular</i>	1 µg	10 µL	Brain ventricle

Suggested volume of reagent:

- *In vivo* PolyMag: 1 µL per µg of DNA.
- *In vivo* DogtorMag and *In vivo* CombiMag: 1 µL each per µg of DNA.
- *In vivo* ViroMag: 10 to 20 µL per 1x10⁶ infectious viral particles.

2.2. Magnets

Several kinds of magnets are provided with the *In vivo* Magnetofection™ kit; use Table 2 to choose the best one adapted to your application.

Table 2: Examples of use of magnets:

Kind of magnet	Tissue
Extra Small Cylinder 2 mm (diameter) x 5 mm (height)	<ul style="list-style-type: none"> • Brain area • Endothelial cells • Small tumors • Lymph node • Ovary • Adrenal gland
Small Cylinder 5 mm (diameter) x 5 mm (height)	<ul style="list-style-type: none"> • Subcutaneous tumors • Salivary gland • Brain
Cylinder 10 mm x 5 mm (height)	<ul style="list-style-type: none"> • Subcutaneous tumors • Pancreas • Spleen
Square 17 mm (length) x 17 mm (length) x 5mm (height)	<ul style="list-style-type: none"> • Large organs • Large tumor • Muscle • Lung • Skin flap

OZ Biosciences is currently proposing only those magnets. If you need specific magnet in terms of shape and size, please contact our technical service for obtaining fundamental properties of the magnet to purchase.

3.2 Protocols

3.1. *In vivo* PolyMag

Please refer to **Table 1** to determine the required amount of DNA as well as volume injection. The DNA, *In vivo* PolyMag and saline solutions should be at room temperature. We recommend using **1 µL of *In vivo* PolyMag per µg of DNA**.

1. Reagents preparation.

- a. *In vivo PolyMag*. Before each use, vortex *In vivo* PolyMag vial. Add the required volume of *In vivo* PolyMag (according to DNA amount needed) to a sterile microtube.

Note: For DNA doses of less than 1 µg, pre-dilute an aliquot of *In vivo* PolyMag reagent with deionized water only, and use the volume required for your DNA dose. Discard the diluted *In vivo* PolyMag after use.

- b. *DNA solution*. Dilute DNA in the final injection volume in a sterile vial (subtract the *In vivo* PolyMag volume).

Note: The final concentration of DNA should not exceed 0.5 mg/mL.

2. **Complexes formation.** Add the DNA solution to the *In vivo* PolyMag and mix immediately by pipetting up and down. Incubate the complexes for 20 min at room temperature.

3. Injection.

- a. Place the magnet on your targeted tissue
- b. Slowly inject the complexes
- c. Let the magnet stand from 5 min to 1 h (see section 3.4 and Table 4)

Notes for intracerebroventricular or intra tumoral injections: Place the magnet few seconds after the complexes injection. Dye *e.g.* Fast Green FCF can be added to the complexes solution for a better monitoring of the injection.

4. **Monitor gene expression** at the appropriate time point.

Important notes:

- Do not inject more than 1 mL of *In vivo* PolyMag per animal.
- Do not inject complexes if precipitate has formed

3.2. *In vivo* DogtorMag

Please refer to **Table 1** to determine the required amount of DNA as well as volume injection. The DNA, *In vivo* CombiMag, *In vivo* DogtorMag reagents and saline solutions should be at room temperature. We recommend using **1 µL of *In vivo* DogtorMag and 1 µL of *In vivo* CombiMag per µg of DNA**.

1. Reagents preparation.

- a. *DNA solution*. Dilute DNA in half of the injection volume in a sterile vial (subtract the *In vivo* CombiMag volume).

Note: The final DNA concentration should not exceed 0.5 mg/mL. Dilute DNA in saline buffer (HBS, PBS, normal saline, Ringer's solution) or glucose 5% solution.

- b. *In vivo DogtorMag solution*. Gently mix the reagent before use. Dilute *In vivo* DogtorMag in half of the injection volume. Incubate for 5 minutes at room temperature.

- c. *In vivo CombiMag reagent*. Vortex the reagent before each use. Use 1 µL of CombiMag / µg DNA. Add the *In vivo CombiMag* reagent directly into a tube (do not dilute with any solution).

Note: For DNA doses of less than 1 µg pre-dilute an aliquot of *In vivo CombiMag* reagent with deionized water only and use the volume required for your DNA dose. Discard the diluted *In vivo CombiMag* after use.

2. **Complexes formation.** Combine the DNA solution with the Dogtor solution. Mix gently and incubate 5 min at RT. Combine the DNA/ *In vivo DogtorMag* mixture with the *In vivo CombiMag* reagent. Mix gently and incubate for 20 minutes at room temperature.

3. **Injection.**

- a. Place the magnet on your targeted tissue
- b. Slowly inject the complexes
- c. Let the magnet stand from 5 min to 1 h (see Table 6)

Notes for intracerebroventricular or intra tumoral injections: Place the magnet few seconds after complexes injection. Dye *e.g.* Fast Green FCF can be added to the complexes solution for a better monitoring of the injection.

4. **Monitor gene expression** at the appropriate time point.

Important notes:

- Do not inject more than 1 mL of *In vivo CombiMag* per animal.
- Do not inject complexes if precipitate has formed

Note: Magnetofectamine™ (MTX0750) has also been successfully tested *in vivo*. Please contact our technical support team at tech@ozbiosciences.com for more details.

3.3. *In vivo ViroMag*

Determine the required injection volume according to the **Table 1**. The amount of virus you need to inject is closely correlated to your viral preparation, the route of injection, the target tissue and your preliminary *in vitro* studies. **Table 3** indicates a starting point for your protocol optimization. The *In vivo ViroMag* and saline solutions should be at room temperature. You may use **10 to 20 µL of *In vivo ViroMag* per 1x10⁶ infectious viral particles**. Do not exceed recommended volumes for injection.

Table 3: Suggested quantity of viral particles and *In vivo ViroMag* injected per mouse

Virus type	Viral particles titer (pfu)	<i>In vivo ViroMag</i> quantity (µL)
Adenovirus	1x10 ⁶ to 5x10 ⁸	10-20
AAV	1x10 ⁶ to 1x10 ¹⁰	10-20
Lentivirus	1x10 ⁶ to 5x10 ⁸	10-20

1. **Reagents preparation.** Immediately before use, vortex *In vivo ViroMag* vial. Add suitable amount of *In vivo ViroMag* to a sterile microtube

Note: If required *In vivo ViroMag* can be diluted with deionized water only. Discard the diluted *In vivo ViroMag* after use.

2. **Complexes formation.** Add your virus preparation to the tube containing the *In vivo ViroMag* reagent and mix immediately by pipetting up & down. Incubate 15-30 min at room temperature.

3. **Injection.**

- a. Place the magnet on your targeted tissue
- b. Slowly inject the complexes
- c. Let the magnet stand from 5 min to 1 h (see Table 6)

Notes for intracerebroventricular or intratumoral injections: Place the magnet few seconds after complexes injection. Dye *e.g.* Fast Green FCF can be added to the complexes solution for a better monitoring of the injection.

4. **Monitor gene expression** at the appropriate time point.

Important notes:

- Do not inject more than 1 mL of *In vivo* ViroMag per animal.
- Do not inject complexes if precipitate has formed

3.4. Magnetic incubation

The magnetic incubation time depends on the animal and the targeted tissue:

- for tumor, from 20 min (mouse, rat) to 1 hour (hamster, cat)
- for endothelial cells, from 5 to 20 min for mouse and rat, from 20 min to 1 h for rabbit or pig
- for peripheral tissue (*e.g.* stomach, gut, heart), 20 min
- for intracerebroventricular injection, 5 min

See **Table 4**, for other magnetic incubation times depending on target tissue, route of injection and magnet type.

Table 4: Suggested magnetic incubation time for various tissue

Target tissue	Route of injection	Kind of magnet	Magnetic incubation
Tumor	Intravenous, Intra-arterial, Intratumoral	All kind	20 min to 1 h
Endothelial cells	Intravenous, Intra-arterial	Extra small Cylinder	5 min to 1 h
Heart	Intra-arterial	Cylinder	20 min
Liver	Intravenous	Cylinder, Square	10 min
Lung	Intravenous	Square	10 min
Pancreas	Intrapancreatic	Cylinder	20 min
Kidney	Intraperitoneal	Cylinder, Square	20 min
Gut	Ilea lumen	All kind	20 min
Stomach	Stomach lumen	Cylinder, Square	20 min
Brain	Intraventricular	Small Cylinder	5 min

Notes:

- For long incubation time, (*e.g.* intratumoral injection), the magnet could be attached to the animal using adhesive tape in order to create a strong magnetic field in the area of the injection.
- Magnets can be easily handled with any magnetic surgical instruments (forceps, clamps, needle holders).
- Magnets can be sterilized by heat (steam sterilization or dry heat sterilization) or chemical means (ethanol 70%).

4. Appendix

Our dedicated and specialized technical support team will be pleased to answer any of your requests and to help you with your experiments at tech@ozbiosciences.com. In addition, do not hesitate to visit our website www.ozbiosciences.com and the FAQ section.

4.1 Quality Controls

To assure the performance of each lot of *In vivo* PolyMag, *In vivo* DogtorMag, *In vivo* CombiMag and *In vivo* ViroMag produced, we qualify each component using rigorous standards. The following *in vitro* assays are conducted to qualify the function, quality and activity of each kit component.

Specification	Standard Quality Controls
<i>Sterility</i>	Thioglycolate assay. Absence of fungal and bacterial contamination shall be obtained for 14 days.
<i>Biological Activity</i>	<i>In vitro</i> transfection or infection efficiency. Every lot shall have an acceptance specification of > 80% of the activity of the reference lot.
<i>Magnetic nanoparticles</i>	1. Quality and size homogeneity of the magnetic nanoparticles. 2. Stability of the magnetic nanoparticles formulations.
<i>In vivo grade</i>	Endotoxin Quantification. Endotoxins level shall be < 0.1 EU/mL for every lot.

4.2. Troubleshooting

Problems	Comments and Suggestions
Low efficiency	<ol style="list-style-type: none"> Nucleic acid amount. Use different quantity of DNA or siRNA with the recommended transfection reagent / nucleic acid ratio. Magnetofection reagent / nucleic acid ratio. Optimize the reagent/nucleic acid ratio by varying the volume of reagent (<i>In vivo</i> PolyMag or <i>In vivo</i> DogtorMag) from 0.5 µL to 2 µL per µg of DNA. DNA quality. Nucleic acids should be as pure as possible, free of contaminants (proteins, phenol, ethanol...) and endotoxins free. It must be "transfection grade". Viral particles quality. Ensure that the ratio total/infectious particles number is correct Expression cassette. Make sure that the transgene is well expressed <i>in vivo</i>, meaning that the expression cassette (promoter, terminator...) is suited for <i>in vivo</i>. Route of Injection. In order to improve your transfection/transduction, prefer a route of injection close to your target tissue; <i>e.g.</i> inject your vector intra-arterially into the tissue supplying artery (use the same quantity as intravenous administration, be careful, intra-arterial manipulations enhance the risk of thrombosis) or directly into the tissue of interest together with magnet application. Medium used for preparing transfection complexes. Favour saline buffer (HBS, PBS, normal saline, Ringer's solution) or 5% glucose during the preparation of the complexes. Injection volume. Optimize the volume of injection to your application. Speed of injection. DNA expression could be dependent on the injection speed, adapt your speed of injection to your target tissue. Transfection reagent temperature. Reagents must be at room temperature and be vortexed prior to use. Old transfection reagent / DNA complexes. The transfection reagent / DNA complexes must be freshly prepared every time. Complexes prepared and stored for longer than 2 hours can be aggregated. Incubation time. Optimal time range between transfection and assay varies with tissue, promoter, expression product, etc. Transfection efficiency can be monitored after 4 – 48h by analyzing the gene product.

Toxicity	<ol style="list-style-type: none"> 1- Concentration of transfection reagent / nucleic acid too high. Decrease the amount of nucleic acid / reagent complexes injected by lowering the nucleic acid amount or the transfection reagent concentration. Complexes aggregation can cause some toxicity; prepare them freshly and never inject complexes where precipitate has formed. 2- DNA quality - Presence of contaminants. Ensure that nucleic acid is pure, contaminant-free and endotoxin-free.
Precipitate formation	<ol style="list-style-type: none"> 1- Concentration of transfection reagent / nucleic acid too high. Decrease the amount of nucleic acid / reagent complexes injected by lowering the nucleic acid amount or the transfection reagent concentration. Complexes aggregation can cause some toxicity; prepare them freshly and never inject complexes where precipitate has formed. 2- DNA quality - Presence of contaminants. Ensure that nucleic acid is pure, contaminant-free and endotoxin-free. Prefer water than buffer (TE, Tris-Cl) for your DNA preparation. 3- Injection solution. Prefer 5% glucose (final concentration) than saline buffer (HBS, PBS, normal saline, Ringer's solution).

Our dedicated and specialized technical support group will be pleased to answer any of your requests and to help you with your transfection experiments: tech@ozbiosciences.com. In addition, do not hesitate to visit our website www.ozbiosciences.com and the FAQ section.

4.3. Bibliographic references

Please refer to the results sheet and to our website for a more comprehensive list of bibliographic reference.

- **Gupta A.K and Gupta M** 2005 Synthesis and surface engineering of iron oxide nanoparticles for biomedical applications. *Biomaterials*. 26:3995-4021.
- **Laurent N, Sapet C, Le Gourrierc L, Bertosio E and Zelphati O** 2011 Nucleic acid delivery nanoparticles : the Magnetofection™ technology. *Therapeutic Delivery*. 2:471:482.
- **Plank C, Zelphati O, Mykhaylyk O.** 2011 Magnetically enhanced nucleic acid delivery. Ten years of magnetofection-progress and prospects. *Adv Drug Deliv Rev*. 63:1300-1331

5. Related Products

Description
MAGNETOFECTION TECHNOLOGY
Super Magnetic Plate (<i>standard size for all cell culture support</i>)
Mega Magnetic plate (<i>mega size to hold 4 culture dishes at one time</i>)
Transfection reagents:
PolyMag Neo (<i>for all nucleic acids</i>)
Magnetofectamine™ (<i>for all nucleic acids</i>)
NeuroMag (<i>dedicated for neurons</i>)
SilenceMag (<i>for siRNA application</i>)
Transfection enhancer:
CombiMag (<i>to improve any transfection reagent efficiency</i>)
Viral Transduction enhancers:
ViroMag (<i>to optimize viral transduction</i>)
ViroMag R/L (<i>specific for Retrovirus and Lentivirus</i>)
AdenoMag (<i>for Adenoviruses</i>)
LIPOFECTION TECHNOLOGY (LIPID-BASED)
Lullaby (<i>siRNA transfection reagent</i>)
DreamFect Gold (<i>Transfection reagent for all types of nucleic acids</i>)
VeroFect (<i>for Vero cells</i>)
FlyFectin (<i>for Insect cells</i>)
i-MICST TECHNOLOGY
Viro-MICST (<i>to transduce directly on magnetic cell purification columns</i>)
3D TRANSFECTION TECHNOLOGY
3Dfect (<i>for scaffolds culture</i>) / 3DfectIN (<i>for hydrogels culture</i>)
RECOMBINANT PROTEIN PRODUCTION
HYPE-5 Transfection Kit (<i>for High Yield Protein Expression</i>)
PROTEIN DELIVERY SYSTEMS
Ab-DeliverIN (<i>delivery reagent for antibodies</i>)
Pro-DeliverIN (<i>delivery reagent for protein in vivo and in vitro</i>)
PLASMIDS PVECTOZ
pVectOZ-LacZ / pVectOZ-SEAP / pVectOZ-GFP / pVectOZ-Luciferase
ASSAY KITS
Bradford – Protein Assay Kit
MTT cell proliferation kit
SEAP Assay kit
β-Galactosidase assay kits (CPRG/ONPG)
BIOCHEMICALS
D-Luciferin, K ⁺ and Na ⁺ 1g
X-Gal powder 1g / G-418, Sulfate 1g

6. Purchaser Notification

Limited License for *In vivo* Magnetofection™

The purchase of *In vivo* Magnetofection™ Reagents grants the purchaser a non-transferable, non-exclusive license to use the kit and/or its separate and included components (as listed in section 1, Kit Contents). This reagent is intended **for in-house research only** by the buyer. Such use is limited to the transfection and transduction of nucleic acids and virus as described in the product manual. In addition, research only use means that this kit and all of its contents are excluded, without limitation, from resale, repackaging, or use for the making or selling of any commercial product or service without the written approval of OZ Biosciences.

Separate licenses are available from OZ Biosciences for the express purpose of non-research use or applications of *In vivo* Magnetofection™ Reagents. To inquire about such licenses, or to obtain authorization to transfer or use the enclosed material, contact the Director of Business Development at OZ Biosciences.

Buyers may end this License at any time by returning all *In vivo* Magnetofection™ Reagents material and documentation to OZ Biosciences, or by destroying *In vivo* Magnetofection™ Reagents components. Purchasers are advised to contact OZ Biosciences with the notification that *In vivo* Magnetofection™ Reagents kit is being returned in order to be reimbursed and/or to definitely terminate a license for internal research use only granted through the purchase of the kit(s).

This document covers entirely the terms of *In vivo* Magnetofection™ Reagents research only license, and does not grant any other express or implied license. The laws of the French Government shall govern the interpretation and enforcement of the terms of this License.

Product Use Limitations for *In vivo* Magnetofection™

In vivo Magnetofection™ Reagents and all of its components are developed, designed, intended, and sold for research use only. They are not to be used for human diagnostic or included/used in any drug intended for human use. All care and attention should be exercised in the use of the kit components by following proper research laboratory practices.

For more information, or for any comments on the terms and conditions of this License, please contact:

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