

Studies With Information Regarding Stability of Agents ALZET[®] Osmotic Pumps

This bibliography does not include references prior to 2010. To obtain the bibliographic list of earlier references, please contact ALZET Technical Services at (800) 692-2990 or email to alzet@durect.com.

| Agent 6E10 | <u>Stability Verified Through</u> 5 weeks | <u>References</u> P9867 |
|--|---|--|
| A-127722 | | P6953 |
| Acetylcholine HCl | 13 days 15 days | P0439 P1926 |
| ACTH | 7 days 3 days 7 days 81-120 days | P0449 P0446 S1087 P2344 |
| Adrenomedullin | 21 days | P4131 |
| Aldosterone | 3 days | P0612 |
| Aminooxyacetic acid | 14 days | P0886 |
| Aminolevulinic acid, 5- | 12 hrs | P3750 |
| Aminopyrine | 7 days | P3395 |
| Amitriptyline | 7 days | P2779 |
| Amyloid protein, beta | 2,4 weeks | P7068 |
| Angiostatin | 7 days | P5052 |
| Angiotensin II | 6 days 6 days 6 days 10-12 days 14 days 7 days | P0480 P1008 P3530 P1992 P3064 P4590 |
| Angiotensinogen | 7 days 7 days 6 days | P1047 P0699 P1008 |
| Antibody/Immunoglobulin | 15 days | P8266, P1136 |
| Antichymotripsin, alpha-1 Apomorphine | 28 days 20 days 14 days | Q0562 P5291 P1291 |



| Argatroban | 4 weeks | Q2300 |
|--|--------------|-------|
| Arginine, N(omega)-hydroxy-nor-l | 10 days | Q4902 |
| Atrial Natriuretic Factor | 14 days | P1724 |
| | 14 days | P1521 |
| | 13 days | P0911 |
| | 7-14 days | P1653 |
| | 6 days | P0871 |
| | 6 days | P1564 |
| | 7 days | P0990 |
| | 3 days | P1032 |
| | | |
| Atriopeptin III | 7 days | P0990 |
| Atropine, Methyl- | 5 weeks | P0683 |
| Azidothymidine | 7 days | P1644 |
| Bacteria, vibrio cholerae | 7 days | P1380 |
| Bafilomycin A1 | 2 weeks | P2976 |
| BDNF, recombinant human | 28 days | P5186 |
| | 2 weeks | P3290 |
| | 2 weeks | P3031 |
| | 2 weeks | P2933 |
| | 2 weeks | P2562 |
| | 30 days | P3430 |
| | 14 days | P9237 |
| | | |
| BE16627B | 18 days | P2788 |
| Benazeprilat | 2 weeks | P3939 |
| Benzylamine | 2 weeks | P4873 |
| | 2 weeks | P5688 |
| Benzylguanine, 06- | 14 days | P7702 |
| Bromodeoxyuridine | 1 week (80%) | P3208 |
| Buserelin | 2 weeks | P0457 |
| Buspirone HCl | 2 weeks | P0813 |
| C-reactive protein | 2 weeks | P9632 |
| C3 | >40 days | Q3859 |
| Calcitonin | 7 days | P1087 |
| Calcitonin-Gene Related Peptide (8-37) | 14 days | P5575 |





| Captopril | 4 weeks 2 weeks | P0910 |
|----------------------------|----------------------|----------------|
| | 2 weeks | P1036 |
| Carbachol | 6 days | P0628 |
| Carboplatin | 7 days | P5476 |
| | 4 weeks | Q0707 |
| Carboxyfullerine | >40 days | Q3959 |
| Catalase | 7 days 20 days | P2520 P1741 |
| | 20 days | P1/41 |
| CCX662 | 16 days | Q3365 |
| CART (42-89) | 10 days | P4764 |
| Celecoxib | 28 days | Q0379 |
| CGS-16949A | 4, 12 weeks | P2209 |
| | 6 weeks | P2208 |
| CGS-22652 | 2 weeks | P2678 |
| CHF-1024 | 1 month | P7942 |
| Chlordiazepoxide | "Half-time 8.8 days" | P9959 |
| Chlorpromazine | 2 weeks | P0385 |
| Cholecystokinin 8 | 5 days | P0879 |
| Chondroitinase ABC | 7 days | P7563 |
| Clozapine | 2 weeks | P0385 |
| CNP-22 | 7 days | P4017 |
| Cocaine HCl | 14 days | P2218 |
| Collagenase | 7 days | P1957 |
| Cortisol, 21-hemisuccinate | 18 days | P2997 |
| Cytochrome C | 14 days | P2256 |
| | 1 week | P4031 |
| DADLE (enkephalin analog) | 7 days | P2630 |
| Dexamethasone | 12-26 days | P3120 |
| | 26 days | Q2064 |
| | 7 days | Q4838 |





| Dopamine HCl | 14 days | P1291 |
|----------------------------------|-----------|-----------|
| | 13 days | P0439 |
| | 13 days | P0959 |
| | 10 days | P1733 |
| | 7 days | P0592 |
| | 7 days | P1313 |
| Dopamine, 6-hydroxy | 8 days | P2562 |
| | 7 days | P3514 |
| | 7 days | P0495 |
| | 7 days | P0606 |
| | 7 days | S0317 |
| | 7 days | P3932 |
| | 7 0895 | F 3 3 3 2 |
| Doxorubicin | 18 days | P2788 |
| DNA | 28 days | P6539 |
| DPAT, 7-hydroxy-U-99194A maleate | 14 days | P7137 |
| Dynorphin A (1-13) | 7 days | P1261 |
| Echistatin | 14 days | P5567 |
| | 14 00 / 3 | 13307 |
| Edaravone | 24 hours | P7529 |
| | | |
| Endomorphine Analog | >1 year | Q5105 |
| Endostatin | 32 days | P5030 |
| Endothelin | 3 days | P3586 |
| | 6 days | P1810 |
| | 6 days | P1803 |
| Endothelin-1 | 4 weeks | P3039 |
| | | |
| | 2 weeks | P4492 |
| | 2 weeks | P5690 |
| Endothelin-3 | 3 days | P3586 |
| | | |
| Endotoxin | 1 week | P1828 |
| | 1 week | P0425 |
| Faineabring | C dave | 02520 |
| Epinephrine | 6 days | P3530 |
| Estradiol, 17A- | 7 days | P0763 |
| Estradiol, 17B- | 7 days | P0763 |
| Estradiol, 2-hydroxy- | 7 days | P0763 |
| | | |





| Estrone, 2-hydroxy- | 7 days | P0763 |
|-------------------------------------|-----------------------------------|-------------------------|
| Fadrozole | 4 weeks 2 weeks | P2209 P2208 |
| Fibroblast Growth Factor | 4 days | P2490 |
| Fibroblast Growth Factor, basic | Up to 4 weeks | P2901 |
| Ficoll, 3H-carboxymethyl | 7 days | P6599 |
| Flurbiprofen | 14 days | Q0222 |
| Formoterol fumarate | 20 hrs | P5817 |
| ForsKolin | 7 days | P3932 |
| Fluorodeoxy Cytidine Fluvoxamine | 45 days 21 days | Q5174 P2641 |
| Furosemide | 28 days | P5336 |
| FMRFamide | | P6115 |
| GABA | 14 days 13 days | P0886 P0439 |
| Gabapentin | 14 days | P5337 |
| Gastrin-17 | 7 days | P7077 |
| Gene, HSV-Tk + liposome | 3 days | P3860 |
| Gene, LacZ + Liposome | 3 days | P3860 |
| Gentamicin | 7 days | Q2064 |
| Glibenclamide | 48 hours | Q2174 |
| Glycine | 13 days | P0959 |
| Gonadotropin-releasing hormone | 2 weeks | Q3663 |
| GP120 | 2 weeks | P3880 |
| Growth Hormone | 7 days 4 days 2 weeks (rat) | P1478 P3289 Q0258 |
| H7 | 24 hours | P2981 |
| H89 (Protein kinase A inhibitor) | 7 days | P5269 |





| Haloperidol | 2 weeks 4 weeks 2 weeks | P0813 P0385 P0556 |
|---|---|----------------------------------|
| Heregulin | 7 days | P3717 |
| Hexamethylphosphoramide | 4 weeks | P3737 |
| Horseradish Peroxidase | 48 hours | P0389 |
| HS024 | 4 weeks | P6559 |
| HU210 | 6 weeks | Q2667 |
| Human Placental Lactogen | 6 days | P2220 |
| Hydroxytryptamine Antagonist | 1,2,3 weeks | P1735 |
| Idazoxan | 7 days | P1270 |
| Imipramine | 14 days | P5337 |
| Insulin | 4 weeks 10 days 10 days >11 days | S0774 P1402 P1442 P5850 |
| Insulin-like Growth Factor I and II | 6 days 14 days | P0599 P4150 |
| Insulin-like growth factor I, receptor antagonist | 4 weeks | P7467 |
| Interferon, alpha | 8 weeks | Q1983 |
| Interferon-gamma | 7 days 8 weeks | Q1983 |
| Interleukin 1 | 14 days 7 days 7 days 14 days | P2542 P2140 P1957 P4586 |
| Interleukin 2 | 7 days 7 days 6 days | P0550 P2749 P1837 |
| Interleukin 3 | 7 days | P3567 |
| Interleukin 6-PE | 7 days | P1913 |
| IL13-PE38QQR | 7 days | P6373 |



| Interleukin B, recomb. human | 7 days 7 days | P2177 P2639 |
|--|---|---|
| lododeoxyuridine | 1 week (80%) | P3229 P3165 |
| JNJ-31020028 | 28 days | Q2631 |
| Ketamine | 14 days | P5337 |
| Kisspeptin-54 | 14 days | P7900 |
| КҮР-2047 | 1 month | Q3622 |
| Leptin | 14 days | P5391 |
| Lisuride | 14 days | P1320 |
| L-NAME | 4 or 8 days | P2585 P6953 |
| Marinobufagenin | 4 weeks | P7519 |
| Melanin Concentrating Hormone | 7+ days | P0985 |
| Melatonin | 14 days | P1111 |
| Methotrexate | 7 days | P0592 |
| Methylene Blue | | P1906 |
| MMP-2 | 28 days | P5819 |
| MK-771 (thyroxine analog) | 7 days | P0638 |
| MPP* | 28 days | P8284 |
| MTII (non-specific melanocortin agonist) | 28 days | P5207 |
| Morphine sulfate | 7 days 14 days | P2630 P5337 |
| Nerve Growth Factor | 15 days 1 month 1 month 14 days 14 days 14 days 14 days | P2348 P1927 P1658 P3260 P2933 P2256 P0923 |





| | 14 days | P0933 |
|--|---|---|
| | 14 days | S0632 |
| | 7 days | P0473 |
| | 14 days | P2758 |
| | | P2775 |
| | 14 days | P3877 |
| Neuropeptide Y | 14 days | P1983 |
| Neurotensin | 8 days | P0572 |
| Neurotrophin-4/5 | 14 days | P7844 |
| Neurotrophin-3 | 14 days | P8606 |
| Nicotine Bitartrate | 16 days | P0976 |
| Nogo receptor, soluble fragment | 1 month | P9513 |
| Norepinephrine | 14 days | P0632 |
| | 13 days | P0423 |
| | 13 days | P0439 |
| | 5 days | P0152 |
| NT-3 | 14 days | P3290 |
| | 14 days | P2562 |
| | 28 days | P3913 |
| NXY-059 | 3 days | P4957 |
| Olanzapine | 4 weeks | P9790 |
| | | 02024 |
| Oligodeoxynucleotide, antisense | 14 days | P3934 |
| Oligodeoxynucleotide, antisense Oligodeoxyn-, phosphorothioate | 14 days | P3934 P3809 |
| Oligodeoxyn-, phosphorothioate | 14 days | P3809 |
| | - | |
| Oligodeoxyn-, phosphorothioate | 14 days 14 days | P3809 P4902 |
| Oligodeoxyn-, phosphorothioate Osteogenic progein-1 | 14 days 14 days 14 days | P3809 P4902 P6921 |
| Oligodeoxyn-, phosphorothioate Osteogenic progein-1 Oxotremorine | 14 days 14 days 14 days 5 weeks 10 days 10 days | P3809 P4902 P6921 P0683 |
| Oligodeoxyn-, phosphorothioate Osteogenic progein-1 Oxotremorine | 14 days 14 days 14 days 5 weeks 10 days | P3809 P4902 P6921 P0683 P1976 |
| Oligodeoxyn-, phosphorothioate Osteogenic progein-1 Oxotremorine | 14 days 14 days 14 days 5 weeks 10 days 10 days | P3809 P4902 P6921 P0683 P1976 P2766 |
| Oligodeoxyn-, phosphorothioate Osteogenic progein-1 Oxotremorine Oxytocin P110-TAT | 14 days 14 days 14 days 5 weeks 10 days 10 days 10 days (44% destruction) 28 days | P3809 P4902 P6921 P0683 P1976 P2766 P3050 Q3188 |
| Oligodeoxyn-, phosphorothioate Osteogenic progein-1 Oxotremorine Oxytocin | 14 days 14 days 14 days 5 weeks 10 days 10 days 10 days (44% destruction) | P3809 P4902 P6921 P0683 P1976 P2766 P3050 |
| Oligodeoxyn-, phosphorothioate Osteogenic progein-1 Oxotremorine Oxytocin P110-TAT | 14 days 14 days 14 days 5 weeks 10 days 10 days 10 days (44% destruction) 28 days 13 days | P3809 P4902 P6921 P0683 P1976 P2766 P3050 Q3188 P0415 |





| Peptide YY (3-36) | 28 days 14 days | P8015 Q1576 |
|--|----------------------------------|----------------------------------|
| Phenylarsonous acid | 2 weeks | Q0586 |
| Phosphonovaleric acid | 7 days | P1597 |
| Physostigmine | 7 days | P1508 |
| Pioglitazone | 21 days 21 days | P8145 Q3340 |
| Platelet-activating factor | 12 days | P1926 |
| Plasminogen activator inhibitor-1 | | P6129 |
| Platelet factor-4 | 28 days | P5819 |
| PR-21 Prolactin | 14 days 2 days | Q0450 P1758 |
| Propranolol | 28 days 11 days 9 days | P1693 P5559 P0011 P2894 |
| Proline, l- | 7 days | P3740 |
| Proline, nitric acid | 7 days | P3740 |
| Proprionamide ditri-fluoroacetate | >2 hours | Q5308 |
| Prostaglandin $(A_2, D_2, E_2, F_{2A}, I_1)$ | 7 days | P3077 |
| Prostaglandin E1 | 7 days 1 week (70%) | P1917 P1092 P3028 |
| Quinpirole | 7 days 7 days | P2115 P1466 |
| Rapamycin | 14 days | P3013 |
| Reelin, recomb. | 2 weeks | P9766 |
| RG108 | 20 days | Q4616 |
| Rilmenidine | 30 days | P6378 |
| Rosiglitazone Maleate | 24-48 hours | Q5322 |





| Serotonin | 13 days | P0439 |
|--|--|----------------------------------|
| SHU-9119 | 7 days 21 days | P0671 Q2499 |
| Small Interfering RNA (LNA modified) | >48 hours | P8305 |
| Somatostatin | 28 days 7 days | P8852 P0967 |
| Strychnine | 13 days | P0959 |
| Substance P | 8 days 14 days | P0572 P2259 |
| Sulpiride | 7 days | P2115 |
| Superoxide Dismutase | 20 days 7 days | P1741 P2520 |
| TAT-peptide Theophylline | 2 weeks 7 days | Q1165 P0823 |
| THIP | 14 days | P0886 |
| Thymidine,5'amino5'deoxy- | 6 days | P3380 |
| Thyroid stimulating hormone | 7 days | P1126 |
| Thyrotropin Releasing Hormone | 8 days 7 days 7 days | P0572 P0967 P1126 |
| Thyroxine | 14 days 7 days | P1306 P1331 |
| Tiagabine | 7 days | P3415 |
| Toxin, TGF- $lpha$ chimeric- | 7 days (80-100%) | P3210 |
| Transforming Growth Factor | 7 days 7 days 21 days 2,4 weeks | P2590 P2583 P4770 P7068 |
| TGF-B20 soluble receptor | 14 days | P5046 |
| Thrombospondin-1, three type 1 repeats | 7 days | P8302 |
| TLQP-21, synthetic | 14 days | P7973 |
| Triazolam | 7 days | P2115 |
| Triiodothyronine | 7 days | P1331 |



| Tryptophan, L-5-hydroxy- | 1 week | P1882 |
|---------------------------------------|--------------------------------|-------------------------|
| ΤΤ-232 | 3 weeks 3 weeks >3 weeks | P8928 P6497 P6894 |
| Tumor Necrosis Factor | 14 days | P4672 |
| Urease | | 7 days |
| Uridine, Bromodeoxy- | 7 days 3-11 days | P3208 P7105 |
| Uridine, 5-fluoro-2'-deoxy- | 7 days 7 days | P3208 P1917 |
| Uridine, iododeoxy- | 7 days | P2917 |
| Valproic Acid | 14 days | P0886 |
| Valsartan | 2 weeks | P3939 |
| Vasoactive Intestinal Peptide | 17 days | P2858 |
| Vasopressin, arginine | 7 days | P3109 |
| Vasopressin, 1-desamino- 8-D-arginine | 3 days 7 days | P0668 P1556 |
| VEGF | 2 weeks | P5670 |
| Verapamil | 2 weeks | P5832 |
| Vitamin D3, 1,25-dihydroxy- | 4 weeks | P8289 |
| VX Y27632 | 1 week 4 weeks 13 days | P9487 P9438 P7559 |



Recent References (2010-Present) on the Stability of Agents Administered Using ALZET[®] Osmotic Pumps

Q10637: N. Orti-Casan, *et al.* A TNF Receptor 2 Agonist Ameliorates Neuropathology and Improves Cognition in an Alzheimer's Disease Mouse Model. Proceedings of the National Academy of Sciences 2022;119(37):e2201137119 **Agents:** NewStar2 **Vehicle:** PBS; **Route:** CSF/CNS (lateral ventricle); **Species:** Mice; **Pump:** 2006; **Duration:** 6 weeks; **ALZET Comments:** "Dose: (0.15 μL/h)Controls received mp w/ vehicle; animal info: C57BL/6 background; Male mice 6 mo of age behavioral testing: EPM; Y-Maze spontaneous alternation; MWM; stability of compound verified by cytotoxicity assay; Brain coordinates (anteroposterior,

0.05 mm; lateral, 0.1 mm; dorsoventral, 0.25 mm),); dental cement used; neurodegenerative (Alzheimer's disease); "

Q7293: A. T. Larsen, *et al.* The Dual Amylin and Calcitonin Receptor Agonist KBP-088 Induces Weight Loss and Improves Insulin Sensitivity Superior to Chronic Amylin Therapy. J Pharmacol Exp Ther 2019;370(1):35-43

Agents: Amylin, KBP-088 Vehicle: Saline; Route: SC; Species: Rat; Pump: 2ML4; Duration: 4 weeks;

ALZET Comments: Dose: KBP-088 (5 µg/kg/day), rat amylin (100, 300, 1000 µg/kg/day); Controls received mp w/ vehicle; animal info (Long Evans rats (10 weeks old); stability verified (a stability test of amylin in the concentrations used in the osmotic pumps shows that the peptide remains stable throughout the study); KBP-088 is a potent dual amylin and calcitonin receptor agonist (DACRA); obesity;

Q6978: R. D. Cole, *et al.* Adolescent and adult nicotine exposure differentially impacts oral nicotine and oral saccharin self-administration in mice. Behavioural Brain Research 2019;359(836-844

Agents: nicotine hydrogen tartrate **Vehicle:** Saline; **Route:** SC; **Species:** Mice; **Pump:** 1002; **Duration:** 2 weeks; **ALZET Comments:** Dose (3.0 mg/kg/d - adolescent, 6.3 mg/kg/d-adult); Controls received mp w/ vehicle; animal info (adolescent, adult); stability verified by (measuring the residual volume following surgical removal at the end of the exposure period for each time point); half-life 10 min (p.841); dependence;

Q7324: J. Yang, *et al.* Inhibition of the CD36 receptor reduces visceral fat accumulation and improves insulin resistance in obese mice carrying the BDNF-Val66Met variant. J Biol Chem 2018;293(34):13338-13348

Agents: Salvionolic acid B Vehicle: PBS; Route: SC; Species: Mice; Pump: 2004; Duration: 8 weeks;

ALZET Comments: Dose (50 mg/kg/day); Controls received mp w/ vehicle; animal info (C57BL/6, 6-weeks old, control 23.9 +/- 0.5 g, SAB treatment 23.3 +/- 0.6 g); pumps replaced after 4 weeks; stability verified by (measured effect of neutralized SAB on macrophage oxLDL uptake);

Q7540: S. Watanabe, *et al.* Intracerebroventricular administration of Cystatin C ameliorates disease in SOD1-linked amyotrophic lateral sclerosis mice. J Neurochem 2018;145(1):80-89

Agents: Cystatin C, Recombinant human Vehicle: PBS; Route: CSF/CNS (lateral ventricle); Species: Mice; Pump: 1004; Duration: 4 weeks;

ALZET Comments: Dose (66 ng/day); Controls received mp w/ vehicle; animal info (100-day-old, transgenic SOD1G93A, male≥30g and female≥20g); stability verified by (influenza hemagglutinin (HA)-tagged CysC administration for 1 week); CysC is an endogenous protease inhibitor; enzyme inhibitor (cathepsin); ALZET brain infusion kit used; neurodegenerative (ALS); "After 1 week of continuous HA-tagged CysC administration using an osmotic pump, the CysC was successfully delivered to the lumbar spinal cord and was predominantly distributed in the ventral horn neurons (Fig. 1b and c), whereas CysC was rarely found in neurons of the dorsal horn. These data indicate that intracerebroventricular administration was sufficient to deliver CysC to lower motor neurons in the lumbar spinal cord." pg.82; implanted pump remained on the back of mice until end-stage; "the disease end-stage was determined as the time when animals in a lateral position were unable to right themselves within 20s" p.81;Therapeutic indication (Bunina body formation and regulation of AMPK/PGC-1a pathway);



Q7312: N. Tsuburaya, *et al.* A small-molecule inhibitor of SOD1-Derlin-1 interaction ameliorates pathology in an ALS mouse model. Nat Commun 2018;9(1):2668

Agents: SOD1-Derlin-1 inhibitor #56-40, SOD1-Derlin-1 inhibitor #56-59 Vehicle: DMSO; Route: CSF/CNS (lateral ventricle); Species: Mice; Pump: 2006; Duration: 36 weeks;

ALZET Comments: Dose (1 mM #56-40 or 3 mM #56-59); Controls received mp w/ vehicle; animal info (22 weeks, male, C57BL/6); behavioral testing (rotarod performance); behavioral testing (rotarod performance); pumps replaced every 6 weeks until mouse showed paralysis onset; long-term study; stability verified by (in-vitro immunoprecipitation assay); 3-Amino-N-(4-pyridyl)-6-(3-pyridyl)thieno[2,3-b]pyridine-2-carboxamide aka #56-40;

N-Allyl-3-amino-N-phenyl-6-(pyridin-3-yl)thieno[2,3-b]pyridine-2-carboxamide aka #56-59; enzyme inhibitor (SOD1-Derlin-1 interaction); ALZET brain infusion kit 3 used; neurodegenerative (Amyotrophic lateral sclerosis);

Q7046: M. Shimamura, et al. Therapeutic Effects of Systemic Administration of the Novel RANKL-Modified Peptide, MHP1, for Ischemic Stroke in Mice. BioMed Research International 2018;2018(4637084

Agents: Microglial healing peptide 1 **Vehicle:** Saline; **Route:** SC; **Species:** Mice; **Pump:** 2001D; **Duration:** 24 hours; **ALZET Comments:** Dose (2 mg/mL); Controls received mp w/ vehicle; 24 hour stability verified by in vitro inhibitory activity of TLR4-induced inflammation. (); MHP1 is a Novel RANKL-Modified Peptide; Therapeutic indication (ischemic stroke);

Q5938: S. Zhu, *et al.* Recombinant Immunotoxin Therapy of Glioblastoma: Smart Design, Key Findings, and Specific Challenges. BioMed Research International 2017;2017(7929286

Agents: NZ-1-(scdsFv)-PE38KDEL Vehicle: Not Stated; Route: Not Stated; Species: Mice (SNG); Pump: Not Stated; Duration: 3 days;

ALZET Comments: cancer (glioblastoma);no stress (see pg. 10);stability verified by (33-98% of activity 3 days at 37C; verified via incubation);

Q6186: E. Tarasco, *et al.* Effect of AP102, a subtype 2 and 5 specific somatostatin analog, on glucose metabolism in rats. Endocrine 2017;58(1):124-133

Agents: AP102, Pasireotide Vehicle: Saline; Route: SC; Species: Rat; Pump: 2ML2; Duration: 4 weeks;

ALZET Comments: Dose: AP102 (3 or 10 µg/kg/h); pasireotide (3 or 10 µg/kg/h); Controls received mp w/ vehicle; animal info (male Sprague-Dawley rats); post op. care: antibiotic (Enrofloxacin; 5.7 mg/kg), and analgesics (Flunixin; 1 mg/kg s.c.); pumps replaced every 2 weeks; 13-days stability verified by pump incubation at 37 degrees and liquid chromatography-mass spectrometry analysis; AP102 is a subtype 2 and 5 specific somatostatin analog;

Q6119: Y. C. Shi, *et al.* Y5 receptor signalling counteracts the anorectic effects of PYY3-36 in diet-induced obese mice. J Neuroendocrinol 2017;29(10):

Agents: Peptide YY (3-36) Vehicle: Disodium hydrogen phosphate, NaCl, Tween 80; Route: SC; Species: Mice; Pump: 2004; Duration: 21 days;

ALZET Comments: animal info (diet-induced obese wild-type, Y5R knockout); stability verified by (Peptide YY "was stable and functional over the period of the experiment"); Obesity and diabetes;

Q5842: M. Kano. AMH/MIS as a contraceptive that protects the ovarian reserve during chemotherapy. Proceedings of the National Academy of Sciences 2017;114(9):E1688-E1697

Agents: Mullerian inhibiting substance, recombinant human Vehicle: Saline; Route: IP; Species: Mice; Pump: 1007D; Duration: 15 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (6-7 weeks old) ; functionality of mp verified by rhMIS activity; pumps replaced every 5 or 7 days; cancer; half-life is ~ 4 hours (p. E1689); post op. care (carprofen analgesic (2.5 mg/mL) by oral gavage (100 μL)); stability verified by bioassay ("rhMIS activity was remarkably stable, with the material recovered from pumps that had been implanted in mice for 1 wk conserving full biological activity in the rat urogenital ridge bioassay"); "To test the efficacy of rhMIS protein for the preservation of ovarian reserve, we elected to use osmotic pumps implanted i.p. in C57BL/6N female mice to allow very precise delivery of MIS" pg. E1691; Therapeutic indication (Oncofertility, cancer); Dose (1200 ug/mL);



Q5105: J. E. Zadina, *et al.* Endomorphin analog analgesics with reduced abuse liability, respiratory depression, motor impairment, tolerance, and glial activation relative to morphine. Neuropharmacology 2016;105(215-27 **Agents:** Morphine; endomorphine analog **Vehicle:** Saline; **Route:** CSF/CNS (intrathecal); **Species:** Rat; **Pump:** 2001; **Duration:** 7 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (male, Sprague Dawley, 250-400g); half-life (p. 219); behavioral testing (tail flick test; rotarod testing); stability verified by (internal testing; stability >1 year at 37C); pumps primed in 37C saline for 16 hours; used PE-8 IT catheter; Dose (2 ug/hr morphine; 0.056-0.075 ug/hr analog);

Q4902: C. Weber, *et al.* Macrophage Infiltration and Alternative Activation during Wound Healing Promote MEK1-Induced Skin Carcinogenesis. Cancer Research 2016;76(4):805-817

Agents: arginine, N(omega)-hydroxy-nor-l **Vehicle:** Not Stated; **Route:** SC; **Species:** Mice; **Pump:** 1004; **Duration:** 33 days; **ALZET Comments:** animal info (InvEE); functionality of mp verified by plasma levels; stress/adverse reaction: (see pg. 811); stability verified by (10 days see pg 811); immunology; "Continuous dosing at a rate of 0.25 mL per hour ensured constant compound levels. Successful ARG1 inhibition was confirmed in blood plasma and wounded skin samples taken 5 days after implantation" pg 811; nor-NOHA aka N(omega)-hydroxy-nor-l-arginine;

Q5174: Y. Pan, *et al.* Inhibition of DNA Methyltransferases Blocks Mutant Huntingtin-Induced Neurotoxicity. Sci Rep 2016;6(31022

Agents: Cytidine, fluorodeoxy- Vehicle: Saline; Route: CSF/CNS; Species: Mice; Pump: 2001; Duration: 1 week; ALZET Comments: ALZET brain infusion kit 3 used; neurodegenerative (Huntington's disease); stability verified by ("...FdCYd fully maintained its neuroprotective activity after 45 days of pre-incubation" see supplement 2); "...decitabine and FdCyd, are known to be degraded rapidly by cytidine deaminase in the liver (in vivo half life of decitabine <20 min)62, indicating that systemic administration may not be an effective strategy for drug delivery to the brain. We therefore chose intracerebroventricular (icv) administration using an Alzet osmotic pump, which provides continuous infusion of drug at a consistent rate from a subcutaneous pump" pg 8; FdCyd aka Cytidine, fluorodeoxy-; decitabine and FdCyd are similar in structure

Q5395: T. H. Lin, *et al.* NF-kappaB decoy oligodeoxynucleotide mitigates wear particle-associated bone loss in the murine continuous infusion model. Acta Biomaterialia 2016;41(273-81

Agents: Ultra-high molecular weight polyethylene particles; oligodeoxynucleotide, decoy; oligodeoxynucleotide, scrambled; Endotoxin, LPS; Brain-derived neurotropic factor; **Vehicle:** Saline; **Route:** In Vitro (cell culture); Bone (Femur); **Species:** Mice (nude); **Pump:** 2006; **Duration:** 4 weeks;

ALZET Comments: Controls received mp w/ vehicle; animal info (Male athymic nude mice, 10-15 weeks old); stability verified by (in vitro experiment); dose-response (pg. 277); good methods (pg. 276); tissue perfusion (bone); Dose (15 mg/ml UHMWPE, 50uM decoy, 1 ug/ml LPS); Therapeutic indication (Bone loss, chronic inflammation);

Q6165: C. K. Kandathil, *et al.* Effects of brain-derived neurotrophic factor (BDNF) on the cochlear nucleus in cats deafened as neonates. Hear Res 2016;342(134-143

Agents: Brain-derived neurotrophic factor, recomb. human Vehicle: Perilymph, artificial; Route: Ear (cochlea); Species: Cat; Pump: 1002, 2004; Duration: 10 weeks;

ALZET Comments: Dose (94 mg/ml; 0.25 ml/hr); pumps replaced after 2 and 4 weeks; BDNF stability verified by neuronal cell culture survival assay (28 days);

Q4838: H. JIA, *et al.* PREVENTION OF TRAUMA-INDUCED COCHLEAR FIBROSIS USING INTRACOCHLEAR APPLICATION OF ANTI-INFLAMMATORY AND ANTIPROLIFERATIVE DRUGS. neuroscience 2016;316(261-278

Agents: Dexamethasone; Ara-C Vehicle: Perilymph, artificial; Route: Ear (cochlea); Species: Rat; Pump: 2001; Duration: 7 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (Wistar, adult); animal info (Wistar, adult); stability verified by (incubation in 37C saline for 7 days see pg 268); one cochlea received vehicle only, while another received drug;



Q5158: M. Gujrati, *et al.* Multifunctional pH-Sensitive Amino Lipids for siRNA Delivery. Bioconjugate Chemistry 2016;27(1):19-35

Agents: RNA, small interfering/EHCO; PEGylated EHCO Vehicle: Not Stated; Route: Not Stated; Species: Mice (nude); Pump: Not Stated; Duration: 14 days;

ALZET Comments: Controls received treated with nonspecific PEGylated EHCO/siGFP nanoparticles (PEGGFP) and non-PEGylated EHCO/HIF-1α; cancer; gene therapy, RNA nanoparticle infusion; peptides; "These results indicate that PEGylation can significantly improve the stability of EHCO/siRNA nanoparticles during storage in solution, possibly by preventing the aggregation of the nanoparticles and providing better protection to the siRNA cargo from degradation" (pg 31);

Q4894: J. P. V. a. R. A. Gonzales. Chronic Intracerebroventricular Infusion of Monocyte Chemoattractant Protein-1 Leads to a Persistent Increase in Sweetened Ethanol Consumption During Operant Self-Administration But Does Not Influence Sucrose Consumption in Long-Evans Rats. Alcoholism Clinical and Experimental Research 2016;40(1):

Agents: Monocyte chemoattractant protein-1, recombinant rat Vehicle: CSF, artificial; water, distilled; albumin, rat serum; Route: CSF/CNS; Species: Rat; Pump: 1004; Duration: 5 weeks;

ALZET Comments: Controls received mp w/ vehicle; animal info (male, Long Evans, 350g); functionality of mp verified by ELISA and manufacturer's instructions see pg189; ALZET brain infusion kit 2 used; dose-response (pg.190); post op. care (bupivacaine intradermally; bupivacaine and gentamicin dripped into wound); behavioral testing (alcohol self-administration); stability verified by (ELISA pg 192); cyanoacrylate adhesive; "We chose the ICV method to simulate the presence of brain-induced MCP-1 and the method's ability to target the whole brain with molecules that do not readily cross the BBB" pg 188-189; pumps primed at 37C sterile saline; Dose (0.2, 20, 2000 ng/day);

Q5341: A. Drougard, *et al.* Central chronic apelin infusion decreases energy expenditure and thermogenesis in mice. Sci Rep 2016;6(31849

Agents: Apelin **Vehicle:** CSF, artificial; **Route:** CSF/CNS (lateral ventricle); **Species:** Mice; **Pump:** 2004; **Duration:** 2 weeks; **ALZET Comments:** Controls received mp w/ vehicle; animal info (C57Bl6/J mice, 13-15 week old); animal info (C57Bl6/J mice, 13-15 week old); "stability of apelin and the duration of the treatment were compatible with the stability of the molecule" (pg. 8); peptides; anesthetized mice with isoflurane; Brain coordinates – 1 mm lateral, – 0.2 mm anteroposterior from the bregma and – 1.7 mm deep; Dose (20 nM);

Q5339: Y. Dong, *et al.* Inhibition of SDF-1alpha/CXCR4 Signalling in Subchondral Bone Attenuates Post-Traumatic Osteoarthritis. Int J Mol Sci 2016;17(6):

Agents: AMD3100 Vehicle: PBS; Route: SC; Species: Mice; Pump: 1004; Duration: 30 days;

ALZET Comments: Controls received mp w/ vehicle and sham operation; animal info (C57BL/6J mice (30 males, 2 months old)); immunology; (AMD3100 is an immunostimulant); inhibitor of CXCR4; transected the anterior cruciate ligament on the right knee; caused joint instability; Dose (180 ug/day);

Q5308: A. Clermont, *et al.* Plasma Kallikrein Mediates Vascular Endothelial Growth Factor-Induced Retinal Dysfunction and Thickening. Invest Ophthalmol Vis Sci 2016;57(6):2390-9

Agents: Fluoroacetate, Propionamide ditri- Vehicle: PEG 400, PBS; Route: SC; Species: Rat, Mice; Pump: 1003D, 1007D; Duration: 1 day, 2 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (Male Sprague-Dawley (SD) rats 10 wks, plasma prekallikrein gene-deficient mice (KLKB1-/-)); functionality of mp verified by enzyme activity assays; 10% PEG 400 used; dose-response (pg 2394, 2398); stability verified by (single bolus subcutaneous injection); Fluoroacetate, Propionamide ditri- aka VA999272; enzyme inhibitor (PKal inhibitor); enzyme inhibitor (PKal inhibitor); Resultant plasma level (pg 2394);

Q5322: M. Bazargan, *et al.* Limited fetal metabolism of rosiglitazone: Elimination via the maternal compartment in the pregnant ewe. Reprod Toxicol 2016;61(162-8

Agents: Rosiglitazone Maleate Vehicle: Water, Ethanol; Route: SC; Species: Sheep (pregnant); Pump: 2ML1; Duration: 16 days;

ALZET Comments: animal info (Singleton pregnant sheep); functionality of mp verified by plasma level, amniotix fluid samples; 15% ethanol used; Multiple pumps per animal (4); stability verified by regular plasma level measurements (reached after day 5, tested through day 16; half-life of 24-48 hours in sheep); Catheters flushed with heparinized saline; Dose (2.7 mg/fetus/d);



Q7246: X. Bao, *et al.* Preclinical toxicity evaluation of a novel immunotoxin, D2C7-(scdsFv)-PE38KDEL, administered via intracerebral convection-enhanced delivery in rats. Invest New Drugs 2016;34(2):149-58

Agents: Immunotoxin, D2C7-(scdsFv)-PE38KDEL Vehicle: Not Stated; Route: CSF/CNS (right caudate nucleus); Species: Rat; Pump: 2ML1, 2001; Duration: 72 hours;

ALZET Comments: Dose (0, 0.05, 0.1, 0.35, 0.4 µg/rat/72 hrs); animal info (Sprague–Dawley; 13-17 weeks old; 360–460 g for males and 210–290 g for females); stability verified by (cytotoxicity assay); stability verified by (cytotoxicity assay); Brain coordinates (1 mm anterior to the bregma, 3 mm to the right of the cranium midline, and 5 mm into the caudate nucleus); cyanoacrylate adhesive (3 M Vetbond Tissue Adhesive); cancer (Glioblastoma); toxicology;

Q4646: B. A. Williams, et al. Multimodal Perineural Analgesia with Combined

Bupivacaine-Clonidine-Buprenorphine-Dexamethasone: Safe In Vivo and Chemically Compatible in Solution. PAIN MEDICINE 2015;16(186-198

Agents: Bupivacaine; clonidine; dexamethasone Vehicle: Saline; Route: CSF/CNS (sciatic nerve); Species: Rat; Pump: 2ML1; Duration: 7 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (male, albino, CD[SD]); no stress (see pg. 192); post op. care (IM butorphanol tartrate 0.05 mg/kg, ceftiofur sodium 5 mg/kg); stability verified by (pg. 195); used polyurethane catheter 0.5mm ID 0.9 mm OD; pumps removed after 1 week; dose (66.6 ug/mL)

Q4616: P. Tognini, *et al.* Experience-dependent DNA methylation regulates plasticity in the developing visual cortex. NATURE NEUROSCIENCE 2015;18(956-+

Agents: RG108 Vehicle: Cyclodextrin, 2-hydroxypropyl-b-; Route: CSF/CNS; Species: Mice; Pump: 1007D; Duration: 3 days; ALZET Comments: Controls received mp w/ vehicle; animal info (C57BL6J, P25); 5% cyclodextrin used; stability verified by (37C t1/2 of 20 days); enzyme inhibitor;

Q5007: C. Lahmann, *et al.* Systemic Administration of Glibenclamide Fails to Achieve Therapeutic Levels in the Brain and Cerebrospinal Fluid of Rodents. PLoS One 2015;10(7):e0134476

Agents: Glibenclamide Vehicle: CSF, artificial; DMSO; BSA; Route: CSF/CNS; Species: Rat; Mice; Pump: 2ML4; 2004; ALZET Comments: Controls received mp w/ vehicle; animal info (mice nV57M, 11-14 weeks old, 25-30g; rats male Lister-hooded, young adults, 200-300g); functionality of mp verified by plasma levels; rats ALZET brain infusion kit 2 used; mice ALZET brain infusion kit 3 used; 0.1% DMSO used; stability verified by (in vitro experimentation see pg 9); Cannula placement verified via histological analysis;

Q5406: J. K. Hu, *et al.* Murine Antibody Responses to Cleaved Soluble HIV-1 Envelope Trimers Are Highly Restricted in Specificity. J Virol 2015;89(20):10383-98

Agents: BG505 SOSIP/664 trimer Vehicle: Iscomatrix; Route: SC; Species: Mice; Pump: 1007D; 2001; 1002; 2002; Duration: 7 days; 14 days;

ALZET Comments: animal info (129/SvJ); Multiple pumps per animal (2); stability verified by (incubation and imaging; 14 day stability. See figure 7 of supplement); immunology; Dose (various, see supplement table 6);

Q3365: M. J. Walters, *et al.* Inhibition of CXCR7 extends survival following irradiation of brain tumours in mice and rats. British Journal of Cancer 2014;110(5):1179-1188

Agents: CCX662 Vehicle: Not Stated; Route: SC; Species: Rat (pregnant); Pump: 2004; Duration: 2 weeks; 4 weeks;; ALZET Comments: Controls received mp w/ vehicle; animal info (Sprague Dawley, 115 days old); functionality of mp verified by plasma levels sampled at 9 and 16 days post implantation; pumps replaced every 7 days; cancer (glioblastoma); stability verified by (IC90 value after 16 days); CCX662 is a CXCR7 inhibitor

Q3663: T. Takeda, *et al.* Maternal Exposure to Dioxin Imprints Sexual Immaturity of the Pups through Fixing the Status of the Reduced Expression of Hypothalamic Gonadotropin-Releasing Hormone. MOLECULAR PHARMACOLOGY 2014;85(1):74-82 **Agents:** Gonadotropin-releasing hormone **Vehicle:** Saline; HCl; **Route:** CSF/CNS; **Species:** Rat; **Pump:** 2002;

ALZET Comments: Controls received mp w/ vehicle; animal info (male, Wistar, PND63); ALZET brain infusion kit 2 used; behavioral testing (sexual behavior); stability verified by (incubation of GnRH at 37C for 2 weeks - half of GnRH remains in unchanged form pg.78); teratology; Cannula placement verified via 0.1% infusion of bromophenol blue; 0.1 M HCl



Q3622: M. H. Savolainen, *et al.* The beneficial effect of a prolyl oligopeptidase inhibitor, KYP-2047, on alpha-synuclein clearance and autophagy in A30P transgenic mouse. NEUROBIOLOGY OF DISEASE 2014;68(1-15

Agents: KYP-2047 **Vehicle:** DMSO; saline; **Route:** IP; **Species:** Mice (transgenic); **Pump:** Not Stated; **Duration:** 28 days; **ALZET Comments:** Controls received mp w/ vehicle; animal info (Snca tm(A30P) or WT, 12-13 months old); functionality of mp verified by decrease in PREP activity; 50% DMSO used; neurodegenerative (Parkinson's disease; Lewy body dementia); post op. care (buprenorphine); stability verified by (stability experiment one month); pumps primed overnight in 37C saline;

Q3616: C. W. Roman, *et al.* PAC1 receptor antagonism in the bed nucleus of the stria terminalis (BNST) attenuates the endocrine and behavioral consequences of chronic stress. Psychoneuroendocrinology 2014;47(151-165 Agents: Pituitary adenylate cyclase activating polypeptide Vehicle: Saline; BSA; Route: CSF/CNS; Species: Rat; Pump: 2002; Duration: 14 days;

ALZET Comments: Animal info (male, adult); behavioral testing (novel object recognition, open field, elevated plus maze); stability verified by (radioimmunoassays; 4% loss per day); cardiovascular; Cannula placement verified via crestyl violet staining and visual inspection; pituitary adenylate cyclase activating polypeptide aka PACP (6-38); delayed delivery; catheters filled with vehicle; pumps primed; PACAP (6-38) is a PAC1 receptor antagonist; stress; y-connector; PACAP (6-38)75uM;

Q3554: M. Oono, *et al.* Transglutaminase 2 accelerates neuroinflammation in amyotrophic lateral sclerosis through interaction with misfolded superoxide dismutase 1. Journal of Neurochemistry 2014;128(3):403-418

Agents: Cystamine Vehicle: Not Stated; Route: CSF/CNS (intrathecal); Species: Mice; Pump: 2006; Duration: 42 days; ALZET Comments: Controls received mp w/ vehicle; animal info (WT C57BL6 or mSOD1, 30 weeks old); neurodegenerative (amyotrophic lateral sclerosis); immunology; stability "Cystamine hydrochloride has previously been shown to be very stable for long time use in the osmotic minipump" pg 417;

Q3232: J. T. Ke, et al. Gliquidone decreases urinary protein by promoting tubular reabsorption in diabetic Goto-Kakizaki rats. Journal of Endocrinology 2014;220(2):129-141

Agents: Gliquidone Vehicle: Dimethylformamide; Route: IP; Species: Rat; Pump: Not Stated; Duration: 7 days; ALZET Comments: Animal info (male, SPF-grade, Goto-Kakizaki, 8 weeks old, 250-300g); functionality of mp verified by blood drug levels; 99.8% dimethylformamide used; comparison of intragastric administration vs mp (pg.133); stability verified by (plasma serum); diabetes;

Q3859: L. L. Dugan, *et al.* Carboxyfullerene Neuroprotection Postinjury in Parkinsonian Nonhuman Primates. Annals of Neurology 2014;76(393-402

Agents: C3 Vehicle: Not Stated; Route: SC; IP; Species: Monkey (macaque); Pump: 2ML4; Duration: 30 days; ALZET Comments: Controls received mp w/ placebo; animal info (Macaca facicularis); Multiple pumps per animal (2); neurodegenerative (Parkinson's disease); behavioral testing (locomotor activity); stability verified by (testing at 37C, C3 is stable for >40 days); Carboxyfullerine aka C3;

Q3426: F. Calcagnoli, *et al.* Chronic enhancement of brain oxytocin levels causes enduring anti-aggressive and pro-social explorative behavioral effects in male rats. Hormones and Behavior 2014;65(427-433

Agents: Oxytocin, synthetic; antagonist, peptidergic oxytocin Vehicle: Saline, sterile, pyrogen-free; Route: CSF/CNS; Species: Rat; Pump: 1007D; Duration: 7 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (male, Groningen, adult); ALZET brain infusion kit 2 used; behavioral testing (resident intruder test); stability verified by (pilot study; 44% destruction of OVT over 10 day period in vivo); peptides; Cannula placement verified via dye injection after sacrifice; Plastics One guide cannula; dental acrylic cement and two stainless steel screws; pumps and catheters incubated in room temperature saline overnight;

Q3417: O. Ben Menachem-Zidon, *et al.* Intra-Hippocampal Transplantation of Neural Precursor Cells with Transgenic Over-Expression of IL-1 Receptor Antagonist Rescues Memory and Neurogenesis Impairments in an Alzheimer's Disease Model. Neuropsychopharmacology 2014;39(2):401-414

Agents: Interleukin-1 receptor antagonist **Vehicle:** Not Stated; **Route:** CSF/CNS; **Species:** Mice (transgenic); **Duration:** 28 days; **ALZET Comments:** Controls received mp w/ vehicle; animal info (WT and Tg2576, 11 months old); pumps replaced every 2 weeks; ALZET brain infusion kit 3 used; neurodegenerative (Alzheimer's disease); behavioral testing (fear conditioning; morris water maze); stability verified by (IL-1-ra remains effective for 2 weeks in pumps); Used three anchoring screws;



Q2522: K. Yo, *et al.* Brown adipose tissue and its modulation by a mitochondria-targeted peptide in rat burn injury-induced hypermetabolism. American Journal of Physiology Endocrinology and Metabolism 2013;304(4):E331-E341 Agents: SS31 Vehicle: Saline; Route: IV (jugular); Species: Rat; Pump: Not Stated; Duration: 7 days; ALZET Comments: Control animals received mp w/ vehicle; animal info (Sprague Dawley, male, 16-19 wks old, 400-500 g); peptides; stability verified via stable isotope dilution method ("more than 7 days")

Q3188: X. Guo, *et al.* Inhibition of mitochondrial fragmentation diminishes Huntington's disease-associated neurodegeneration. Journal of Clinical Investigation 2013;123(12):5371-5388

Agents: P110-TAT **Vehicle:** Not Stated; **Route:** SC; **Species:** Mice (transgenic); **Pump:** Not Stated; **Duration:** 8 weeks; **ALZET Comments:** Controls received mp w/TAT control peptide; animal info (R6/2 HD model, 5 weeks old); functionality of mp verified by FITC positive signal in neurons pg 5379; pumps replaced every 28 days; neurodegenerative (Huntington's disease); no stress (see pg.5381, 5383); behavioral testing (motor function and behaviors); stability verified by (HPLC with UV detection for 28 days); peptides; P110-TAT is a Drp1-selective peptide inhibitor

Q3340: C. Grommes, *et al*. The PPARgamma agonist pioglitazone crosses the blood-brain barrier and reduces tumor growth in a human xenograft model. Cancer Chemotherapy and Pharmacology 2013;71(4):929-936

Agents: Pioglitazone Vehicle: Dulbecco's modified eagle medium; Route: CSF/CNS; Species: Mice (SCID); Pump: 2004; Duration: 21 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (Balb/CJHanHsd-Prkdc-SCID, 6 weeks old); ALZET brain infusion kit used; comparison of oral dosing vs mp; cancer (tumors); dose-response (CNS); stability verified by (p.934 - incubation at 37C for 21 days); "Intracerebral treatment with 1 IM pio prolonged survival significantly from 49 to 68 days... This defines the minimal effective dose for oral pio treatment at 240 PPM (20.2 mg/kg) and for intracerebral pio treatment at 1 IM (0.11 ug/kg)." pg 932;

Q2667: L. Geurts, *et al.* Chronic Endocannabinoid System Stimulation Induces Muscle Macrophage and Lipid Accumulation in Type 2 Diabetic Mice Independently of Metabolic Endotoxaemia. PLoS One 2013;8(2):U622-U631

Agents: HU210 Vehicle: Tween; saline; Route: SC; Species: Mice; Pump: 2006; Duration: 6 weeks;

ALZET Comments: Control animals received mp w/ vehicle; animal info (C57BL/6J, male, 9 wks old); HU210 is a potent agonist, a synthetic cannabinoid; functionality of mp verified via HPLC-HRMS in the brain tissue; stability verified (6 weeks, pg e55963);

Q2499: X. X. Zhu, *et al.* Hypothalamic signaling in anorexia induced by indispensable amino acid deficiency. American Journal of Physiology Endocrinology and Metabolism 2012;303(12):E1446-E1458

Agents: SHU-9119 Vehicle: Not Stated; Route: CSF/CNS; Species: Rat; Pump: 2004; Duration: Not Stated;

ALZET Comments: Control animals received mp w/ artificial CSF; animal info (Sprague Dawley, 7-8 wks old); ; brain infusion kit 2 used; stability verified (21 days, pg E1450)

Q1983: Y. Yuan, *et al.* Role of microRNA-15a in autoantibody production in interferon-augmented murine model of lupus. MOLECULAR IMMUNOLOGY 2012;52(2):61-70

Agents: Interferon, alpha; interferon, gamma **Vehicle:** PBS; BSA; **Route:** SC; **Species:** Mice; **Pump:** 2006; **Duration:** 16 weeks; **ALZET Comments:** Animal info (13 wks old, female, B/W); pumps replaced after 8 weeks; long-term study; stability verified after 8 weeks; "residue IFNs from the pumps at the end of treatment were tested on IFN and IFN responsive cell lines and demonstrated that the in vivo conditions in the pump did not affect bioactivity of both IFNs (data not shown)" pg 63

Q2174: J. M. Simard, et al. Glibenclamide-10-h Treatment Window in a Clinically Relevant Model of Stroke. Translational Stroke Research 2012;3(2):286-295

Agents: Glibenclamide Vehicle: DMSO; saline; NaOH; Route: SC; Species: Rat; Pump: 2001; Duration: 48 hours; ALZET Comments: Controls received mp w/ vehicle; animal info (Wistar, male, 250-275 g); 5.5 hour delayed delivery accomplished by connecting 5.5 ul volume PE60; functionality of mp verified by residual volume; 8% DMSO used; stability verified via spectrophotometry; ischemia (cerebral)



Q2631: J. C. Morales-Medina, *et al.* Chronic administration of the Y(2) receptor antagonist, JNJ-31020028, induced anti-depressant like-behaviors in olfactory bulbectomized rat. Neuropeptides 2012;46(6):329-334 Agents: JNJ-31020028 Vehicle: DMSO; saline; Route: CSF/CNS; Species: Rat; Pump: 2002; Duration: 14 days; ALZET Comments: Control animals received mp w/ saline; animal info (Sprague Dawley, male, 150-170 g, OBX); 1% DMSO used; stability verified (28 days) pg 330; JNJ-31020028 is a Y2 receptor antagonist

Q2300: K. M. Kassel, *et al.* Therapeutic Administration of the Direct Thrombin Inhibitor Argatroban Reduces Hepatic Inflammation in Mice with Established Fatty Liver Disease. American Journal of Pathology 2012;181(4):1287-1295 Agents: Argatroban Vehicle: Acetic acid, glacial; sodium acetate; PEG 400; Route: SC; Species: Mice; Pump: 2004; Duration: 4 weeks;

ALZET Comments: Control animals received mp w/ vehicle; animal info (LDLr -/-, male, 6 wks old); 20% PEG 400 used; stability verified (4 weeks); post op. care (buprenorphine HCl)

Q2064: E. Bas, *et al.* Efficacy of three drugs for protecting against gentamicin-induced hair cell and hearing losses. British Journal of Pharmacology 2012;166(6):1888-1904

Agents: Gentamicin; dexamethasone; melatonin Vehicle: Not Stated; Route: Ear (round window); Species: Rat; Pump: 2001; Duration: 7 days;

ALZET Comments: Controls received mp w/ saline; animal info (Wistar, male, 220-250 g); stability verified after 7 days (data not shown)

Q1165: J. Kim, *et al.* Sustained inhibition of PKC alpha; reduces intravasation and lung seeding during mammary tumor metastasis in an in vivo mouse model. Oncogene 2011;30(3):323-333

Agents: Peptide, TAT; peptide, alpha V5-3; pyrrolidine dithiocarbamate; antibody, anti-CXCR4 **Vehicle:** Saline; **Route:** SC; **Species:** Mice; **Pump:** 1002; **Duration:** 4 weeks;

ALZET Comments: Animal info (BALB/c, 6 wks old, female); pumps replaced every 2 weeks; stability verified (2 weeks, pg 331); incorrectly listed using Model 2001; "Pumps were replaced every 2 weeks, which corresponds to the stability of the peptides in the pump." pg 331; bioluminescence IVIS100

Q1576: S. L. Pedersen, *et al.* Peptide hormone isoforms: N-terminally branched PYY3-36 isoforms give improved lipid and fat-cell metabolism in diet-induced obese mice. JOURNAL OF PEPTIDE SCIENCE 2010;16(11):664-673

Agents: Peptide YY (3-36); peptide YY (3-36), isoform 2; peptide YY (3-36), isoform 3 Vehicle: Saline; Route: SC; Species: Mice; Pump: 1003D; 2002; Duration: 3 days;

ALZET Comments: Controls received mp w/ vehicle; animal info (male, DIO C57BL/6J, NMRI, 8-9 wks old); incorrectly listed Model 1003 pumps used; stability verified pg 667 (2, 7, 14 days via HPLC); wound clips used

Q1430: Y. L. Liu, *et al.* Subcutaneous oxyntomodulin analogue administration reduces body weight in lean and obese rodents. International Journal of Obesity 2010;34(12):1715-1725

Agents: OXM6421; oxyntomodulin Vehicle: Saline; Route: SC; Species: Rat; mice; Pump: 2001; Duration: 7 days; ALZET Comments: Controls received mp w/ vehicle; animal info (Wistar, male, 250-300 g; DIO, male); peptides; stability verified by HPLC (7 days); OXM6421 is a long-acting oxyntomodulin (OXM) analogue; obesity

Q0109: K. L. Chambliss, *et al*. Non-nuclear estrogen receptor-alpha signaling promotes cardiovascular protection but not uterine or breast cancer growth in mice. Journal of Clinical Investigation 2010;120(7):2319-2330

Agents: Estradiol; estrogen-dendrimer conjugate Vehicle: DMSO; Route: IP; Species: Mice (SCID); Pump: 1004; Duration: 72 hours; 28 days;

ALZET Comments: Controls received mp w/ empty dendrimer; cardiovascular; cancer (breast); animal info (female, ERE-Luc reporter, 10-13 weeks old; Ex3aERKO, 8-9 weeks old; C57BL/6 Apoe-/-, 6 weeks old; SCID, 8 weeks old); functionality of mp verified by serum agent levels; Estradiol Dose (6 ug/d); replacement therapy (ovariectomy; pumps replaced after 28 days); half-life (p.2321); half life of EDC = 28 hours; stability verified by (Serum evaluation of experimental and control mice); photon recording with light emission tomography (LET) system with a CCD camera; Research Diets D10001