(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau

(43) International Publication Date
11 June 2015 (11.06.2015)

WIPO | PCT

(10) International Publication Number
WO 2015/085044 A1

(51) International Patent Classification:
A61K 31/485 (2006.01) *A61P 3/04* (2006.01)
A61K 31/137 (2006.01)

(21) International Application Number:
PCT/US2014/068527

(22) International Filing Date:
4 December 2014 (04.12.2014)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:

61/913,216	6 December 2013 (06.12.2013)	US
61/914,938	11 December 2013 (11.12.2013)	US
61/984,580	25 April 2014 (25.04.2014)	US
14/322,810	2 July 2014 (02.07.2014)	US

(71) Applicant: **OREXIGEN THERAPEUTICS, INC.** [US/US]; 3344 North Torrey Pines Court, Suite 200, La Jolla, CA 92037 (US).

(72) Inventors: **KLASSEN, Preston**; 7762 Lookout Drive, La Jolla, CA 92037 (US). **TAYLOR, Kristin**; 15023 Palomino Mesa Road, San Diego, CA 92127 (US).

(74) Agent: **ALTMAN, Daniel, E.**; Knobbe Martens Olson & Bear, LLP, 2040 Main Street, 14th Floor, Irvine, CA 92614 (US).

(81) Designated States *(unless otherwise indicated, for every kind of national protection available)*: AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, JP, KE, KG, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

(84) Designated States *(unless otherwise indicated, for every kind of regional protection available)*: ARIPO (BW, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, ST, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

Declarations under Rule 4.17:

— *as to applicant's entitlement to apply for and be granted a patent (Rule 4.17(ii))*

— *as to the applicant's entitlement to claim the priority of the earlier application (Rule 4.17(iii))*

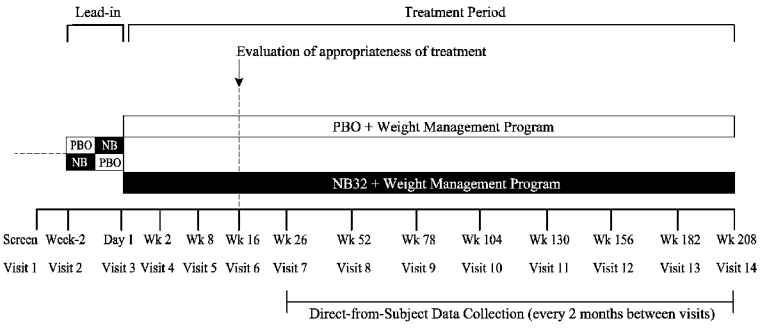
Published:

— *with international search report (Art. 21(3))*

— *before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments (Rule 48.2(h))*

(54) Title: COMPOSITIONS AND METHODS FOR REDUCING MAJOR ADVERSE CARDIOVASCULAR EVENTS

Figure 1. Study Design



(57) Abstract: The present application relates to compositions, kits, uses, systems and methods of using naltrexone and bupropion, or pharmaceutically acceptable salts thereof, for reducing the risk of adverse cardiovascular outcomes or events, including Major Adverse Cardiovascular Events (MACE) in subjects, preferably those at increased risk of adverse cardiovascular outcomes or MACE, that may be overweight or obese. The present application also relates to compositions, kits, uses, systems and methods of using naltrexone and bupropion or pharmaceutically acceptable salts thereof for treatment of overweight or obesity in subjects, preferably at increased risk of adverse cardiovascular outcomes or MACE, wherein the treatment reduces the risk of MACE.

WO 2015/085044 A1