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(71) Applicant *(for all designated States except US)*: **NOVARTIS AG** [CH/CH]; Lichtstrasse 35, CH-4056 Basel (CH).

(72) Inventors; and

(71) Applicants *(for US only)*: **CAPLAN, Shari L.** [US/US]; Novartis Institutes for BioMedical Research, Inc., 100 Technology Square, Cambridge, Massachusetts 02139 (US). **GOLOSOV, Andrei** [US/US]; 218 Thorndike Street, Apt. 204, Cambridge, Massachusetts 02141 (US). **GROSCHE, Philipp** [DE/CH]; Novartis Pharma AG, Postfach, CH-4002 Basel (CH). **GUIMARAES, Carla** [PT/US]; Novartis Institutes for BioMedical Research, Inc., 100 Technology Square, Cambridge, Massachusetts 02139 (US). **KANTER, Aaron** [US/US]; 20 Brastow Ave., Somerville, Massachusetts 02143 (US). **LOU, Changgang** [CN/US]; 3681 Fawn Cove Lane, Apt. 1, Portage, Michigan 49024 (US). **PARKER, David Thomas** [US/US]; Novartis Institutes for BioMedical Research Inc., 250 Massachusetts Avenue, Cambridge, Massachusetts 02139 (US). **PETERS, Eric C.** [US/US]; 4244 Corte Famosa, San Diego, California 92130 (US). **USERA, Aimee Richardson** [US/US]; Novartis Institutes for BioMedical Research, Inc., 100 Technology Square, Cambridge, Massachusetts 02139 (US). **YASOSHIMA, Kayo** [JP/US]; Novartis Institutes for BioMedical Research, Inc., 100 Technology Square, Cambridge, Massachusetts 02139 (US). **YUAN, Jun** [US/US]; Novartis Institutes for BioMedical Research, Inc., 400 Technology Square, Cambridge, Massachusetts 02139 (US). **ZECRI, Frederic** [US/US]; Novartis Institutes for BioMedical Research, Inc., 100 Technology Square, Cambridge, Massachusetts 02139 (US). **ZHAO, Hongjuan** [US/US]; Novartis Institutes for BioMedical Research, Inc., 250 Massachusetts Avenue, Cambridge, Massachusetts 02139 (US).

(74) Common Representative: **NOVARTIS AG**; Lichtstrasse 35, CH-4056 Basel (CH).

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(54) Title: BIOCONJUGATES OF SYNTHETIC APELIN POLYPEPTIDES

X1-X2-X3-X4-X5-X6-X7-X8-X9-X10-X11-X12-X13 (I')

(57) Abstract: The invention provides a bioconjugates comprising a synthetic polypeptide of Formula I': I' or an amide, an ester or a salt thereof, wherein X1, X2, X3, X4, X5, X6, X7, X8, X9, X10, X11, X12 and X13 are defined herein and a half-life extending moiety wherein the peptide and the half-life extending moiety are covalently linked or fuse, optionally via a linker. The polypeptides are agonist of the APJ receptor. The invention also relates to a method for manufacturing the bioconjugates of the invention, and its therapeutic uses such as treatment or prevention of acute decompensated heart failure (ADHF), chronic heart failure, pulmonary hypertension, atrial fibrillation, Brugada syndrome, ventricular tachycardia, atherosclerosis, hypertension, restenosis, ischemic cardiovascular diseases, cardiomyopathy, cardiac fibrosis, arrhythmia, water retention, diabetes (including gestational diabetes), obesity, peripheral arterial disease, cerebrovascular accidents, transient ischemic attacks, traumatic brain injuries, amyotrophic lateral sclerosis, burn injuries (including sunburn) and preeclampsia. The present invention further provides a combination of pharmacologically active agents and a pharmaceutical composition.

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