Circulation Research

Loss of endothelial Endoglin promotes high output heart failure through peripheral arteriovenous shunting driven by VEGF signalling

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* Long In Vivo Checklist

Circulation Research - Preclinical Animal Testing: A detailed checklist has been developed as a prerequisite for every publication involving preclinical studies in animal models. **Checklist items must be clearly described in the manuscript; if the answer to a question is "No", an explanation should be provided both within the manuscript text and on the following screen.** If this information (checklist items and/or explanations) cannot be included in the main manuscript because of space limitations, please include it in an online supplement. If the manuscript is accepted, this checklist will be published as an online supplement. See the explanatory <u>editorial</u> for further information.

This study involves use of animal models:

Yes

been provided.

Study Design	
The experimental group(s) have been clearly defined in the article, including number of animals in each experimental arm of the study.	Yes
An overall study timeline is provided.	Yes
The protocol was prospectively written	Yes
The primary and secondary endpoints are specified	Yes
For primary endpoints, a description is provided as to how the type I error multiplicity issue was addressed (e.g., correction for multiple comparisons was or was not used and why). (Note: correction for multiple comparisons is not necessary if the study was exploratory or hypothesis-generating in nature).	N/A
A description of the control group is provided including whether it matched the treated groups.	Yes
Inclusion and Exclusion criteria	
Inclusion and exclusion criteria for enrollment into the study were defined and are reported in the manuscript.	Yes
These criteria were set a priori (before commencing the study).	Yes
Randomization	
Animals were randomly assigned to the experimental groups.	N/A
If random assignment was not used, adequate explanation has been provided.	Yes
Type and methods of randomization have been described.	N/A
Allocation concealment was used.	N/A
Methods used for allocation concealment have been reported.	N/A
Blinding	
Blinding procedures with regard to masking of group/treatment assignment from the experimenter were used and are described. The rationale for nonblinding of the experimenter has been provided, if such was not performed.	Yes
Blinding procedures with regard to masking of group assignment during outcome assessment were used and are described.	Yes

If blinding was not performed, the rationale for nonblinding of the person(s) analyzing outcome has

N/A

Sample size and power calculations

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Formal sample size and power calculations were conducted before commencing the s on a priori determined outcome(s) and treatment effect(s), and the data are reported.	tudy based Yes
If formal sample size and power calculation was not conducted, a rationale has been p	provided. N/A
Data Reporting	
Baseline characteristics (species, sex, age, strain, chow, bedding, and source) of anim reported.	als are Yes
The number of animals in each group that were randomized, tested, and excluded and reported. If the experimentation involves repeated measurements, the number of anima at each time point is provided is provided for all experimental groups.	
Baseline data on assessed outcome(s) for all experimental groups are reported.	Yes
Details on important adverse events and death of animals during the course of the expreported for all experimental groups.	periment are Yes
Numeric data on outcomes are provided in the text or in a tabular format in the main art supplementary tables, in addition to the figures.	ticle or as Yes
To the extent possible, data are reported as dot plots as opposed to bar graphs, espe small sample size groups.	ecially for No
In the online Supplemental Material, methods are described in sufficient detail to enable replication of the study.	e full Yes
Statistical methods	
The statistical methods used for each data set are described.	Yes
For each statistical test, the effect size with its standard error and ${\it P}$ value is presented. encouraged to provide 95% confidence intervals for important comparisons.	Authors are Yes
Central tendency and dispersion of the data are examined, particularly for small data s	sets. No
Nonparametric tests are used for data that are not normally distributed.	Yes
Two-sided P values are used.	Yes
In studies that are not exploratory or hypothesis-generating in nature, corrections for multiple testing and multiple comparisons are performed.	ultiple N/A
In "negative" studies or null findings, the probability of a type II error is reported.	N/A
Experimental details, ethics, and funding statements	
Details on experimentation including formulation and dosage of therapeutic agent, site administration, use of anesthesia and analgesia, temperature control during experimen postprocedural monitoring are described.	and route of Yes
Both male and female animals have been used. If not, the reason/justification is provide	tation, and
Both male and female animals have been used. If not, the reason/justification is provided. Statements on approval by ethics boards and ethical conduct of studies are provided.	ed. Yes
	ed. Yes

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