**DETAILS** 

**RELATIONS** 

**<<** 

**(1)** 

0



**British Journal of Clinical** <u>Pharmacology</u> Volume 89, Issue 12

Dec 2023

Pages 3465-3772

### ARTICLE

Alkaline phosphatase to treat ischaemiareperfusion injury in living-donor kidney transplantation: APhIRI I feasibility pilot study

## View article page

Thei S. Steenvoorden, Robert E. Duin, Janneke A. J. Rood, Hessel Peters-Sengers, Azar ... See all authors

77 CITE

© 2023 The Authors. British Journal of Clinical Pharmacology published by John Wiley & Sons Ltd on behalf of British Pharmacological Society.

https://doi.org/10.1111/bcp.15871 🔀

0306-5251 ISSN

1365-2125 elSSN

30 June 2023 Accepted

9 June 2023 Revised

4 April 2023 Received

3629 - 3636 Pages

## Aims

Ischemia-reperfusion injury (IRI) during kidney transplant procedures is associated with adverse come. Alkaline phosphatase (AP) is an enzyme that as the potential to dampen IRI. Prior to this study, it

Received: 4 April 2023

Revised: 9 June 2023 | Accepted: 30 June 2023

DOI: 10.1111/bcp.15871

# ORIGINAL ARTICLE



# Alkaline phosphatase to treat ischaemia-reperfusion injury in living-donor kidney transplantation: APhIRI I feasibility pilot study

Thei S. Steenvoorden | Robert E. van Duin | Janneke A. J. Rood<sup>2</sup> Hessel Peters-Sengers 1,3 | Azam S. Nurmohamed 2 | Frederike J. Bemelman 1 Liffert Vogt 1 | Joost W. van der Heijden 2,4

<sup>1</sup>Dept. of Internal Medicine, Nephrology section, Amsterdam UMC, University of Amsterdam, Amsterdam, the Netherlands

<sup>2</sup>Dept. of Internal Medicine, Nephrology section, Amsterdam UMC, Vrije Universiteit, Amsterdam, the Netherlands

<sup>3</sup>Center of Experimental and Molecular Medicine, Amsterdam University Medical Centers, University of Amsterdam, Amsterdam, the Netherlands

<sup>4</sup>Dept. of Internal Medicine, Nephrology Section, Spaarne Gasthuis, Haarlem, the Netherlands

## Correspondence

Liffert Vogt, Dept. of Internal Medicine, Nephrology section, Amsterdam UMC, locatie Universiteit van Amsterdam, Meibergdreef 9, 1105AZ, Amsterdam, the Netherlands. Email: I.vogt@amsterdamumc.nl

### **Funding information**

This study was funded by Alloksys Life Sciences B.V.

Aims: Ischemia-reperfusion injury (IRI) during kidney transplant procedures is associated with adverse outcome. Alkaline phosphatase (AP) is an enzyme that has the potential to dampen IRI. Prior to this study, it had not been tested in the setting of kidney transplantation. This study aimed to evaluate the safety and feasibility of periprocedural AP administration in living donor kidney transplantation.

Methods: In this double blind, randomized, placebo-controlled, single-center pilot study, all eligible recipients of living donor kidneys were asked to give informed consent. AP (bRESCAP) or a placebo was administered intravenously over 24 hours after the transplantation procedure. The primary outcome—graft function at 1 year—was represented by iohexol measured glomerular filtration rate (mGFR). Serum and urine biomarkers within seven days after surgery were used as surrogate markers of kidney function and injury.

**Results:** Eleven patients were enrolled of whom five were treated with bRESCAP and six with placebo. After 1 year, mGFR was not different between groups. No specific adverse events were observed in the bRESCAP group. Urine expression of injury biomarkers CCL14, NGAL and Cystatin C was lower in the bRESCAP group at day seven. This was statistically significant.

Conclusion: This study illustrates that bRESCAP treatment is feasible in kidney transplantation, might have a dampening effect on IRI induced renal inflammation, and raises no safety concerns. Future research will evaluate the effects of bRESCAP

