



# Granulocyte colony-stimulating factor improves early remodeling in isoproterenol-induced cardiac injury in rats

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## Abstract:

**Background:** Granulocyte colony-stimulating factor (G-CSF) has been used in some animal models and humans with well-established cardiovascular diseases. However, its effects in the initial stage of progressive non-ischemic heart failure are unknown.

**Methods:** Wistar rats (260–300 g) were divided into three groups: control (without any intervention), ISO (150 mg/kg isoproterenol hydrochloride *sc*, once a day for two consecutive days), and ISO-GCSF (50 µg/kg/d G-CSF for 7 days beginning 24 h after the last administration of ISO). Echocardiography was performed at baseline and after 30 days of follow-up. Subsequently, animals were anesthetized for hemodynamic analysis. The left ventricle was removed for analysis of interstitial collagen deposition and cardiomyocyte hypertrophy.

**Results:** Isoproterenol led to left ventricular dilation (control,  $7.7 \pm 0.14$  mm; ISO,  $8.7 \pm 0.16$  mm; ISO-GCSF  $7.8 \pm 0.09$  mm;  $p < 0.05$ ), myocardial fibrosis (control,  $2.0 \pm 0.18\%$ ; ISO,  $9.1 \pm 0.81\%$ ; ISO-GCSF  $5.9 \pm 0.58\%$ ;  $p < 0.05$ ) and cardiomyocyte hypertrophy (control,  $303 \pm 10 \mu\text{m}^2$ ; ISO,  $356 \pm 18 \mu\text{m}^2$ ; ISO-GCSF  $338 \pm 11 \mu\text{m}^2$ ;  $p < 0.05$ ). However, G-CSF partially prevented collagen deposition and left ventricular enlargement, with a slight effect on hypertrophy. Characterizing a compensated stage of disease, hemodynamic analysis did not change.

**Conclusion:** G-CSF administered for 7 days was effective in preventing the onset of ventricular remodeling induced by high-dose isoproterenol with decreased collagen deposition and chamber preservation.

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**Key words:** isoproterenol, heart failure, G-CSF, collagen

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