Glyoxalase 1 enhances bone marrow progenitor cell therapy for wound healing in diabetic mice.

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Supplementary Figure S1. Immunofluorescent staining of IRE1 α and CD31 of aortas from IRE1^{ECKO} and IRE1^{flox/flox} mice. White arrows indicate IRE1 α in endothelium of aorta isolated from IRE1^{flox/flox} mice. Bar = 200 µm.



 $\label{eq:constraint} \ensuremath{\mathbb{C}}\xspace{2019} American Diabetes Association. Published online at http://diabetes.diabetes.journals.org/lookup/suppl/doi:10.2337/db18-0933/-/DC1 and the state of the state o$

Supplementary Figure S2. Western blot analysis of GLO1 expression level in Ad-IRE1 α or Ad-GFP infected db/db BMPCs with 24 hours of MGO exposure. BMPCs from db/+ mice were culture for 7 days then transfected with adenovirus carrying human IRE1 α (Ad-IRE1 α , 100 MOI, 48 hours) using adenovirus carrying egfp (Ad-GFP, 100 MOI, 48 hours) as controls. GLO1 protein level was analyzed by Western blot assay. n = 5 per group.



 $@2019\ American\ Diabetes\ Association.\ Published\ online\ at\ http://diabetes.diabetes.journals.org/lookup/suppl/doi:10.2337/db18-0933/-/DC1$

Supplementary Figure S3. Glyoxalase 1 (GLO1) expression level in bone marrow-derived progenitor cells (BMPCs) form db/+ mice did not change after MGO exposure. BMPCs from db/+ mice were culture for 7 days then exposed to 10 μ M of MGO for 24 hours. GLO1 protein level was analyzed by Western blot assay, n = 5 per group.



Supplementary	Table	S1.	Key	functional	regions	and	potential	MGO	binding	sites	in	human
IRE1a.												

Feature Key	Position	Description	Sequence
Binding site	599	ATP binding	FDNRDVAV <mark>KR</mark>
Active site	688	Proton acceptor	SLNIVH <mark>R</mark> DL <mark>K</mark>
Binding site	577-585	Nucleotide binding, ATP required	FCP <mark>K</mark> DVLGHG AEGTIVY <mark>R</mark> GM
Mutagenesis	599	K->A: Loss of autophosphorylation & endoribonuclease activity	FDN <mark>R</mark> DVAV <mark>KR</mark>
Mutagenesis	907	K->A: Loss of endoribonuclease activity	L <mark>R</mark> AM <mark>R</mark> N <mark>KK</mark> HH
Modified site	724-729	Phosphorylation sites	GHSFS <mark>RR</mark> SG

Red AA: functional site; highlighted AA: potential MGO binding position.

Supplementary Table S2. Clinical and biochemical characteristics in lean healthy subjects, obese subjects and type 2 diabetic patients.

Characteristics	Lean healthy (LH)	Type 2 diabetes (T2D)
Age (years)	25 ± 2.70	61.5 ± 3.54
Gender (Male/Female)	3/1	1/1
BMI (kg/cm ²)	21.20 ± 2.24	24.65 ± 1.54
Fasting glucose (mg/dl)	86.95 ± 7.28	161.85 ± 85.05
OGTT (2h, mg/dl)	92.60 ± 21.06	215 ± 29.62
HbA1c (%) (mmol/mol))	5.4% (36mmol/mol) ± 0.20	7.90% (63mmol/mol) ± 3.11

Abbreviation: BMI - body mass index. OGTT - oral Glucose Tolerance Test. Values are percentages or mean \pm SD

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