

**Supplementary Table 1 The classic formula Zuo Gui Pill**

<b>Latin name</b>	<b>Chinese name</b>	<b>production batch number</b>
<i>Radix Rehmanniae Praeparata (24 g)</i>	Shu Di Huang	Lot.No. 2403063
<i>Rhizoma Dioscoreae Oppositae (12 g)</i>	Shan Yao	Lot.No. 2405091
<i>Fructus Lycii (12 g)</i>	Gou Qi Zi	Lot.No. 2404223
<i>Fructus Corni (12 g)</i>	Shan Zhu Yu	Lot.No. 2312032
<i>Radix Cyathulae (9 g)</i>	Chuan Niu Xi	Lot.No. 2402161
<i>Semen Cuscutae (12 g)</i>	Tu Si Zi	Lot.No. 2404161
<i>Colla Cornus Cervi (12 g)</i>	Lu Jiao Jiao	Lot.No. 2311132
<i>Colla Carapacis et Plastris (12 g)</i>	Gui Ban Jiao	Lot.No. 2402131

**Supplementary Table 2 The blood sample levels**

Indicators	ANOVA results (F value, <i>df</i> value, <i>P</i> value)	Effect size ( $\eta^2$ )	Levene's test <i>p</i> >0.05	Post hoc power analysis	Model vs. Control [mean diff (95%CI), <i>P</i> values]	ZGP vs model [mean diff (95%CI), <i>P</i> values]	ZGP + Fer-1 vs ZGP [mean diff (95%CI), <i>P</i> values]
TC	F (6, 14) = 199.6, <i>P</i> <0.0001	0.95	<i>p</i> >0.05	>0.99	[2.7121,3.1059], <i>p</i> <0.0001	[-0.9473,-0.5535], <i>p</i> <0.0001	[-0.6477,-0.2539], <i>p</i> <0.0001
TG	F (6, 14) = 714.8, <i>P</i> <0.0001	0.95	<i>p</i> >0.05	>0.99	[0.7725,0.8565], <i>p</i> <0.0001	[-0.3434,-0.2594], <i>p</i> <0.0001	[-0.2604,-0.1764], <i>p</i> <0.0001
LDL-C	F (6, 14) = 526.6, <i>P</i> <0.0001	0.95	<i>p</i> >0.05	>0.99	[0.8494,0.9296], <i>p</i> <0.0001	[-0.4867,-0.4066], <i>p</i> <0.0001	[-0.3754,-0.2953], <i>p</i> <0.0001
IFCC-HbA1c	F (6, 14) = 1250.9, <i>P</i> <0.0001	0.95	<i>p</i> >0.05	>0.99	[72.5419,77.3856], <i>p</i> <0.0001	[-11.3686,-6.5248], <i>p</i> <0.0001	[-47.7710,-42.9273], <i>p</i> <0.0001
Linsulin	F (6, 14) = 99.9, <i>P</i> <0.0001	0.95	<i>p</i> >0.05	>0.99	[17.3322,20.8617], <i>p</i> <0.0001	[-7.1396,-3.6101], <i>p</i> <0.0001	[-2.8675,0.6620], <i>p</i> >0.05
MDA	F (6, 14) = 2394.8, <i>P</i> <0.0001	0.95	<i>p</i> >0.05	>0.99	[0.6745,0.7068], <i>p</i> <0.0001	[-0.1764,-0.1442], <i>p</i> <0.0001	[-0.4065,-0.3742], <i>p</i> <0.0001

GSH/GSSH	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-1.8146,-1.7140],p<$	$[0.6945,0.7950],p<0.0$	$[0.2329,0.3335],p<0.0$
							0.0001	001	001
SOD	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-1.9808,-1.9485],p<$	$[0.0885,0.1208],p<0.0$	$[0.1319,0.1641],p<0.0$
							0.0001	001	001
BMD	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-0.1883,-0.1394],p<$	$[0.0379,0.0868],p<0.0$	$[0.0011,0.0500],p<0.0$
							0.0001	001	5
BV/TV	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-24.3725,-18.6209],p$	$[3.4775,9.2291],p<0.0$	$[2.9675,8.7191],p<0.0$
							$<0.0001$	001	5
BS/BV	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-14.0902,-11.6432],p$	$[4.4065,6.8535],p<0.0$	$[2.6298,5.0768],p<0.0$
							$<0.0001$	001	001
Tb.N	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-1.7116,-1.0342],p<$	$[0.1353,0.8127],p<0.0$	$[0.13680,0.8142],p<0.$
							0.0001	5	05
Tb.SP	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[0.1187,0.2150],p<0.$	$[-0.1424,-0.0461],p<0.$	$[-0.0679,0.0284],p>0.0$
							0001	05	5
Tb.Th	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-0.0953,-0.0784],p<$	$[0.0278,0.0447],p<0.0$	$[0.0213,0.0382],p<0.0$
							0.0001	001	001
BS/TV	F	(6, 14)	=	0.95	$p>0.05$	$>0.99$	$[-6.4507,-4.5626],p<$	$[0.7493,2.6374],p<0.0$	$[0.4626,2.3507],p<0.0$
							0.0001	5	5

PINP	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	[-27.0119,-21.6548], $p < 0.0001$	[2.3148,7.6719], $p < 0.0001$	[8.5081,13.8652], $p < 0.0001$
TRACP5b	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	[54.8952,61.1545], $p < 0.0001$	[-41.1247,-34.8654], $p < 0.0001$	[-12.8565,-6.5972], $p < 0.0001$
IL6	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	[15.8999,21.6170], $p < 0.0001$	[-11.7540,-6.0369], $p < 0.0001$	[-4.6857,1.0315], $p > 0.0001$
TNF $\alpha$	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	[28.2787,34.0423], $p < 0.0001$	[-12.3386,-6.5750], $p < 0.0001$	[-20.1781,-14.4145], $p < 0.0001$
CD86	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	[16.1222,26.8445], $p < 0.0001$	[-11.9845,-1.2622], $p < 0.05$	[-14.6145,-3.8922], $p > 0.05$
CD80	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	[15.1060,22.5007], $p < 0.0001$	[-10.6240,-3.2293], $p < 0.0001$	[-10.0940,-2.6993], $p > 0.05$
XCT/ $\beta$ -actin	F (6, 28)	= 0.95	$p > 0.05$	$> 0.99$	[-0.7769,-0.3909], $p < 0.0001$	[0.0244,0.4105], $p < 0.0001$	[-0.0439,0.3424], $p > 0.0001$
GPX4/ $\beta$ -actin	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	[-0.5120,-0.3111], $p < 0.0001$	[0.0157,0.2166], $p < 0.0001$	[-0.0547,0.1462], $p > 0.0001$
FTH1/ $\beta$ -actin	F (6, 21)	= 0.95	$p > 0.05$	$> 0.99$	[-0.8933,-0.5044], $p < 0.0001$	[0.0409,0.4297], $p < 0.0001$	[-0.0596,0.3292], $p > 0.0001$

HO1/ $\beta$ -actin	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	$[-0.6287, -0.3859], p < 0.0001$	$[-0.0086, 0.2341], p > 0.05$	$[0.0988, 0.3416], p < 0.05$
	17.1, $P < 0.0001$						
Nrf2/ $\beta$ -actin	F (6, 21)	= 0.95	$p > 0.05$	$> 0.99$	$[-0.1505, -0.0759], p < 0.0001$	$[0.0252, 0.0998], p < 0.05$	$[-0.0316, 0.0431], p > 0.05$
	7.1, $P < 0.0001$						
Rank1/ $\beta$ -actin	F (6, 35)	= 0.95	$p > 0.05$	$> 0.99$	$[0.1156, 0.2351], p < 0.0001$	$[-0.1751, -0.0556], p < 0.05$	$[-0.0758, 0.0437], p > 0.05$
	8.2, $P < 0.0001$						
Runx2/ $\beta$ -actin	F (6, 28)	= 0.95	$p > 0.05$	$> 0.99$	$[-0.3955, -0.2372], p < 0.0001$	$[0.0286, 0.1868], p < 0.05$	$[0.0228, 0.1810], p < 0.05$
	13.9, $P < 0.0001$						
XCT mRNA	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	$[-0.8896, -0.4052], p < 0.0001$	$[0.0208, 0.5051], p < 0.05$	$[-0.1674, 0.3169], p > 0.05$
	8.6, $P < 0.0001$						
GPX4 mRNA	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	$[-1.0493, -0.6793], p < 0.0001$	$[-0.0117, 0.3584], p > 0.05$	$[0.1523, 0.5224], p < 0.05$
	24.9, $P < 0.0001$						
Nrf2 mRNA	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	$[-1.0231, -0.7457], p < 0.0001$	$[-0.1093, 0.1681], p > 0.05$	$[0.1917, 0.4690], p < 0.05$
	42.8, $P < 0.0001$						
HO1 mRNA	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	$[-1.0632, -0.6332], p < 0.0001$	$[-0.1146, 0.3154], p > 0.05$	$[0.0867, 0.5167], p < 0.05$
	14.7, $P < 0.0001$						
Nrf2-IHC	F (6, 14)	= 0.95	$p > 0.05$	$> 0.99$	$[-42.3187, -38.7013], p < 0.0001$	$[2.1713, 5.7887], p < 0.001$	$[0.1246, 3.7420], p < 0.05$
	520.1, $P < 0.0001$						

HO1-IHC	F	(6, 14)	=	0.95	$p > 0.05$	$> 0.99$	[-32.5687,-29.8313],p	[4.3847,7.1220],p<0.0	[9.5313,12.2687],p<0.
				523.9,P<0.0001			<0.0001	001	0001
GPX4-IHC	F	(6, 14)	=	0.95	$p > 0.05$	$> 0.99$	[-27.9306,-24.7361],p	[2.6694,5.8639],p<0.0	[1.8361,5.0306],p<0.0
				254,P<0.0001			<0.0001	001	001

---