

**Adult effects of fetal IGF1 therapy for FGR**

1 **Effects of intrauterine insulin-like growth factor-1 therapy**  
2 **for fetal growth restriction on adult metabolism and body**  
3 **composition are sex-specific**

4 ***Supplementary material***

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11 **Running title:** Adult effects of fetal IGF1 therapy for FGR

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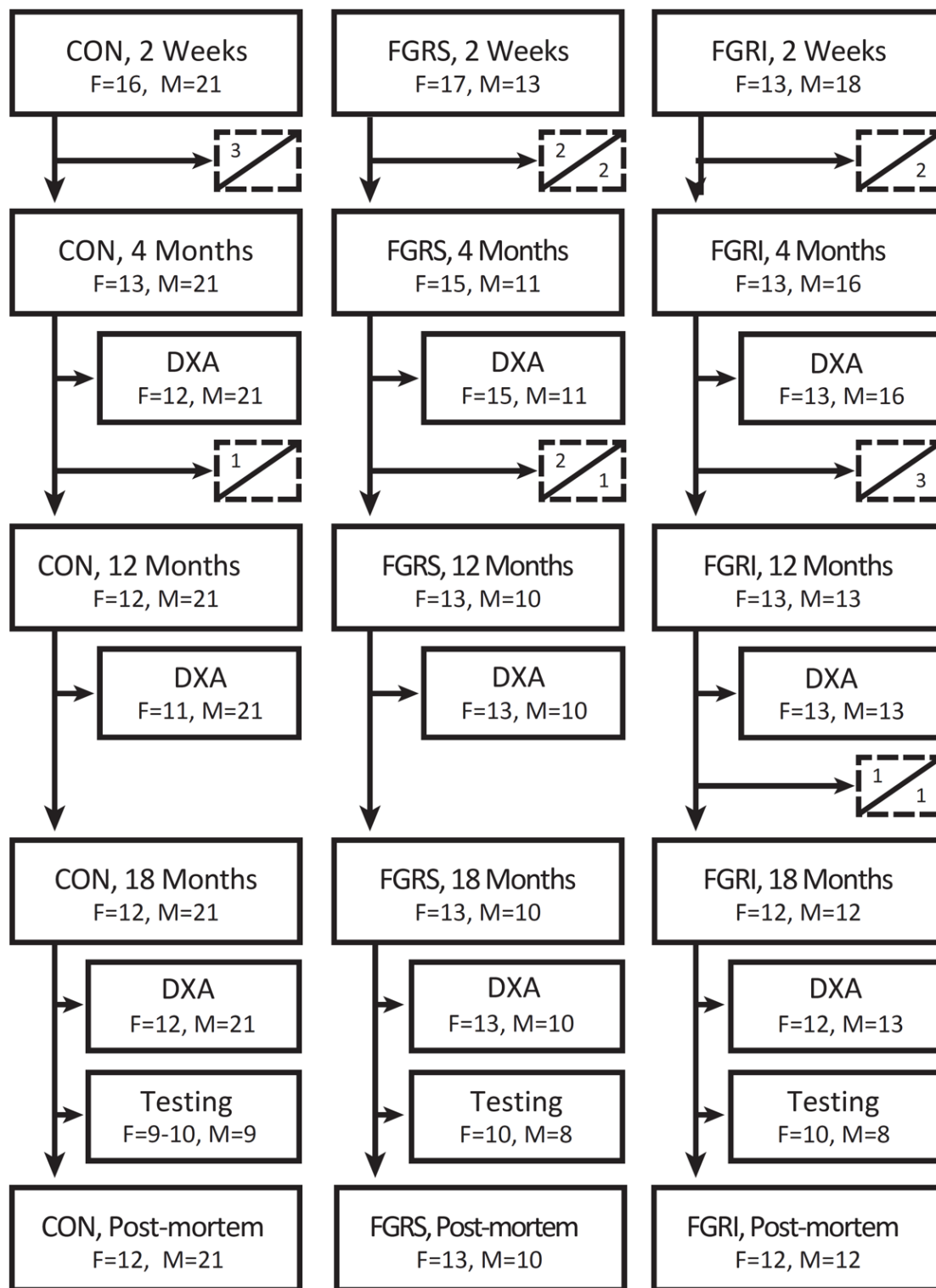
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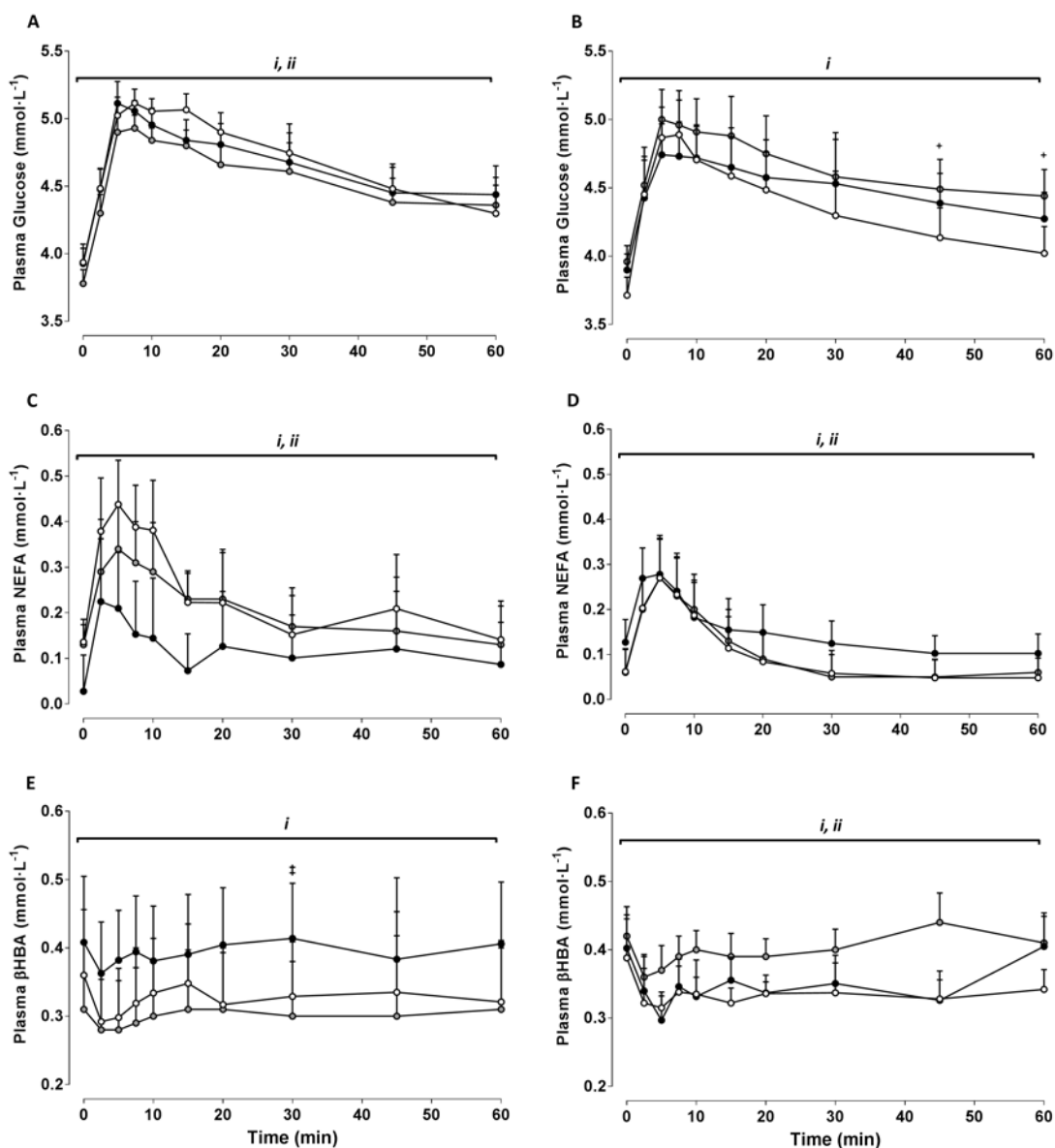
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## Adult effects of fetal IGF1 therapy for FGR



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 23 **Supplementary Figure 1. Animals utilised during the project according to**  
 24 **experimental group and sex.** Dashed boxes report exclusions, deaths and  
 25 euthanasia. Split boxes reporting postnatal deaths indicate the number of females  
 26 (top left) and males (bottom right) for each experimental group. Circumstances  
 27 constituting exclusion from the project and/or euthanasia were due to postnatal  
 28 illness or paddock deaths.

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 30 **Supplementary Figure 2. Plasma metabolite response to intravenous**  
 31 **epinephrine injection.** Plasma glucose, non-esterified fatty acids (NEFA), and β-  
 32 hydroxybutyric acid (βHBA) response to an intravenous epinephrine injection in  
 33 female (A, C, E: CON, white,  $n=9$ ; FGRS, grey,  $n=10$ ; FGRI, black,  $n=10$ ) and male  
 34 (B, D, F: CON, white,  $n=9$ ; FGRS, grey,  $n=9$ ; FGRI, black,  $n=8$ ) sheep. Data are  
 35 mean  $\pm$  SEM. Roman numerals denote the significant difference between  
 36 experimental groups (RM ANOVA: *i*, effect of time,  $p<0.05$ ; *ii*, time\*experimental  
 37 group interaction  $p<0.05$ ). Symbols denote significant differences amongst  
 38 experimental groups upon post hoc testing: ‡FGRS vs. FGRI  $p<0.05$ .

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	CON	FGRS	FGR <sub>39</sub>
Postnatal illness	4 <sup>°</sup>	4 <sup>°</sup> , 1 <sup>•</sup>	4 <sup>°</sup> 40 <sup>††</sup>
Paddock death			1 <sup>+</sup>

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45 **Supplementary Table 1. Animal losses during experimental procedures.** Sheep  
 46 excluded, euthanised, or found dead postnatally due to: <sup>°</sup>infection-related deaths  
 47 (coccidiosis, n=2; eosinophilic gastroenteritis and hepatocellular atrophy, n=1;  
 48 listeriosis, n=2; lymphocytic meningoencephalitis, n=1; pneumonia, n=4;  
 49 rhododendron poisoning, n=1; umbilical infection, n=1), <sup>•</sup>non-specific pulmonary  
 50 edema, n=1; <sup>+</sup> fight wounds, n=1.

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	Female			Male			Significance (treatment effect)	
	CON n=11	FGRS n=13	FGRI n=13	CON n=21	FGRS n=10	FGRI n=13	Female	Male
<b>BMC (g)</b>	745±45	703±42	698±45	826±83	760±87	827±86	ns	ns
<b>BMD (g·cm<sup>-2</sup>)</b>	1.38±0.05	1.33±0.04	1.26±0.13	1.40±0.10	1.37±0.12	1.42±0.11	ns	ns
<b>Chest LM (kg)</b>	12.3±0.6	12.0±0.6	12.1±0.6	15.8±0.6	14.6±0.7	15.1±0.7	ns	ns
<b>Chest FM (kg)</b>	2.5±0.6	2.1±0.6	2.2±0.6	1.2±0.2	0.7±0.2	0.7±0.2	ns	ns
<b>Abdominal LM (kg)</b>	20.3±1.2	19.5±1.2	19.2±1.2	24.8±0.5	22.8±0.8	23.0±0.8	ns	ns
<b>Abdominal FM (kg)</b>	4.1±1.0	3.4±0.9	3.5±1.0	1.9±0.4	1.1±0.4	1.1±0.4	ns	ns
<b>Rump LM (kg)</b>	8.9±0.4	8.6±0.4	8.9±0.4	10.7±0.4	9.8±0.5	10.2±0.4	ns	ns
<b>Rump FM (kg)</b>	1.8±0.4	1.5±0.4	1.6±0.4	0.8±0.1	0.5±0.2	0.5±0.2	ns	ns
<b>Total LM (%)</b>	41.4±2.2	40.2±2.1	40.2±2.2	51.4±1.4	47.3±1.8	48.4±1.6	ns	ns
<b>Total FM (%)</b>	8.4±2.0	7.0±1.9	7.2±2.0	3.9±0.7	2.3±0.8	2.2±0.8	ns	ns
<b>LM:WT (%)</b>	70.0±1.5	70.5±1.3	73.1±1.7	75.8±1.0	76.7±1.3	76.2±1.2	ns	ns
<b>FM:WT (%)</b>	14.1±2.7	12.4±2.6	12.1±2.6	5.7±1.1	3.4±1.2	3.5±1.2	ns	ns
<b>Chest FM:WT (%)</b>	4.2±0.8	3.7±0.8	3.6±0.8	1.7±0.3	1.0±0.4	1.1±0.4	ns	ns
<b>Abdominal FM:WT (%)</b>	6.8±1.3	6.0±1.2	5.8±1.3	2.7±0.6	1.6±0.6	1.7±0.6	ns	ns
<b>Rump FM:WT (%)</b>	3.0±0.6	2.7±0.5	2.6±0.6	1.2±0.2	0.7±0.2	0.7±0.2	ns	ns
<b>Abdominal FM:Total FM (%)</b>	48.8±0.6	48.5±0.5	47.3±0.7	48.2±0.6	48.3±0.6	48.2±0.6	ns	ns

**Supplementary Table 2. Body composition at 12 months of age.** Bone mineral content (BMC), bone mineral density (BMD) compartmental and total lean mass (LM) and fat mass (FM), absolute and, where appropriate, relative to weight (WT). Data are mean ± SEM.