

Figure S1. Standard cytokine curve for interleukin 1  $\beta$  (IL-1  $\beta$ )

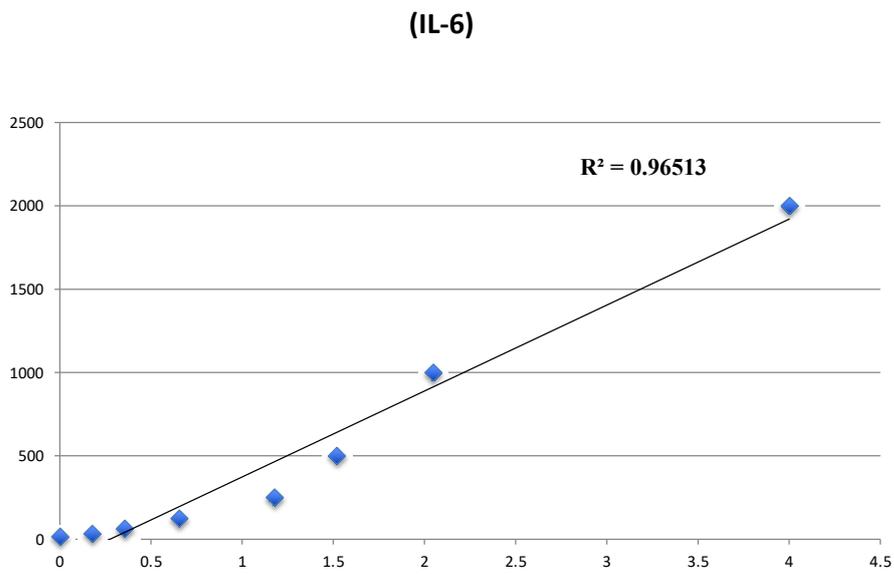


Figure S2. Standard cytokine curve for interleukin 6 (IL-6)

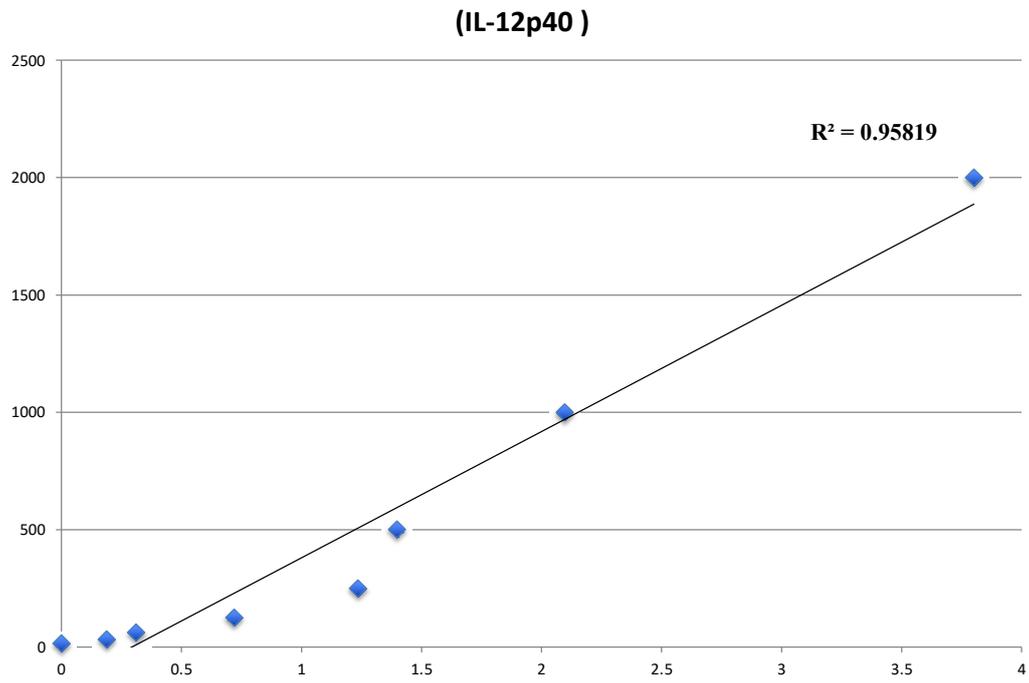


Figure S3. Standard cytokine curve for interleukin 12p40 (IL-12p40)

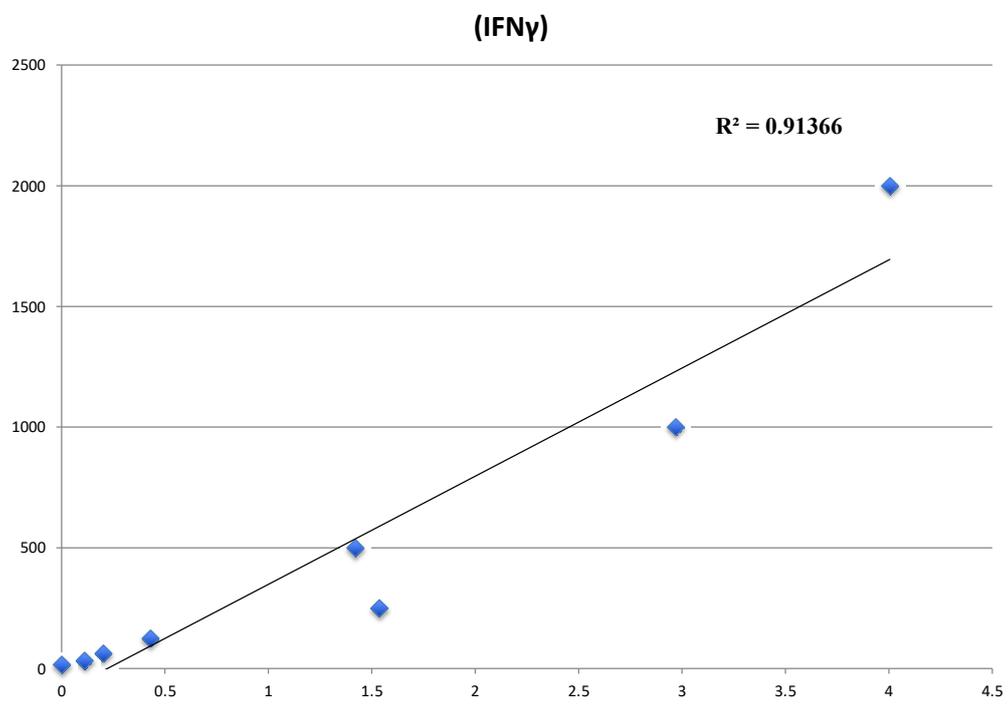


Figure S4. Standard cytokine curve for interferon gamma (IFN $\gamma$ )

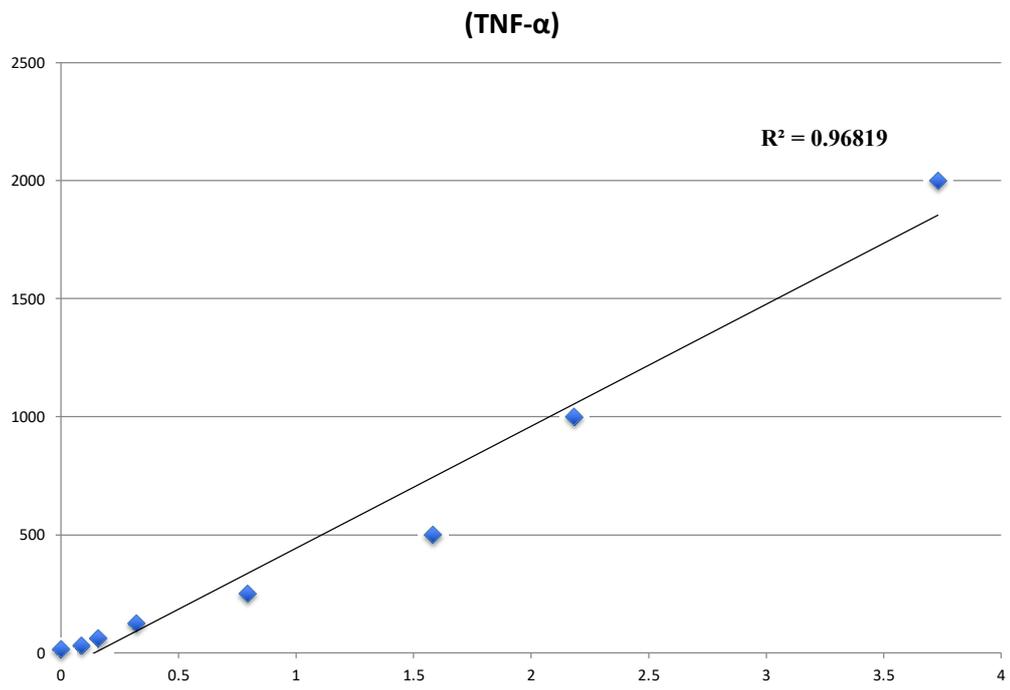


Figure S5. Standard cytokine curve for tumor necrosis factor alpha (TNF- $\alpha$ )

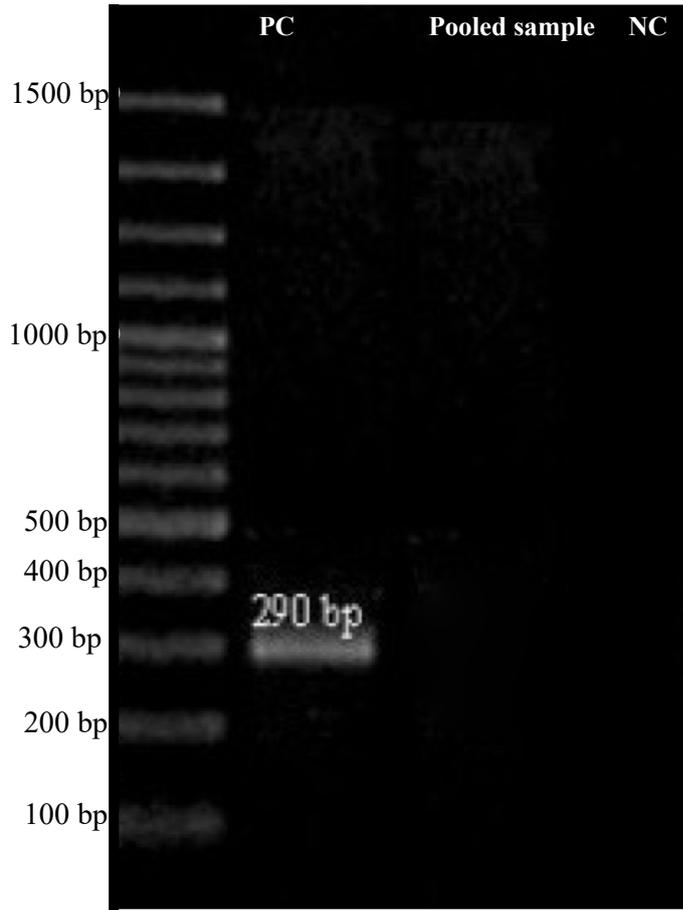


Figure S6. RT-PCR product visualized after agarose gel electrophoresis. The expected size of the PCR products was 290 bp. PC, Positive control sample, NC, Negative control, M= 100 bp ladder. RT-PCR was used to confirm the infection by BVD in the pooled buffy coat samples collected from the selected lambs. Cytopathic strain of BVDV genotype-2 (Strain 125) were obtained from the department of Rinderpest like Diseases, Veterinary Serum and Vaccine Research Institute, Cairo and used as a control strain for RT-PCR test. Pure RNA was extracted using viral RNA Purification Kit following the manufacturer protocol. PCR cycling conditions were performed using the following PCR primer sets: the forward primer (5'-ATGCCCTTAGTAGGACTAGCA-3')/reverse primer (5'- CAACTCCATGTGCCATGTACAGCAG -3').

Table S1. *P* values for the Pearson correlation test coefficient between clinical signs scores and the production of pro-inflammatory cytokines in lambs naturally infected with pneumonic pasteurellosis

Variables	CSS	IL-1 $\beta$	IL-6	IL-12p40	TNF- $\alpha$	IFN $\gamma$
CSS		0.57	0.70	0.36	0.44	0.80
IL-1 $\beta$	0.57		0.35	0.42	0.84	0.26
IL-6	0.70	0.35		0.12	0.81	0.93

<b>IL-12p40</b>	0.36	0.42	0.12		0.09	<b>0.004</b>
<b>TNF-<math>\alpha</math></b>	0.44	0.84	0.81	0.09		0.09
<b>IFN<math>\gamma</math></b>	0.80	0.26	0.93	<b>0.004</b>	0.09	

Abbreviations: CSS (clinical signs score), interleukin 1  $\beta$  (IL-1  $\beta$ ), interleukin 6 (IL-6), interleukin 12p40 (IL-12p40), interferon gamma (IFN $\gamma$ ) and tumor necrosis factor alpha (TNF- $\alpha$ ). Differences between means at  $P < 0.05$  were considered significant.

Table S2.  $P$  values for the Pearson correlation test coefficient between responses to the treatment by tulathromycin in lambs naturally infected with pneumonic pasteurellosis

Variables	CSS	IL-1 $\beta$	IL-6	IL-12p40	TNF- $\alpha$	IFN $\gamma$
<b>CSS</b>		0.25	<b>0.009</b>	0.05	0.41	0.63
<b>IL-1 <math>\beta</math></b>	0.25		0.06	0.64	0.13	<b>0.03</b>
<b>IL-6</b>	<b>0.009</b>	0.06		0.08	0.11	0.55
<b>IL-12p40</b>	0.05	0.64	0.08		0.23	0.93
<b>TNF-<math>\alpha</math></b>	0.41	0.13	0.11	0.23		0.52
<b>IFN<math>\gamma</math></b>	0.63	<b>0.03</b>	0.55	0.92	0.52	

Abbreviations: CSS (clinical signs score), interleukin 1  $\beta$  (IL-1  $\beta$ ), interleukin 6 (IL-6), interleukin 12p40 (IL-12p40), interferon gamma (IFN $\gamma$ ) and tumor necrosis factor alpha (TNF- $\alpha$ ). Differences between means at  $P < 0.05$  were considered significant.

Table S3.  $P$  values for the Pearson correlation test coefficient between responses to the treatment by tulathromycin combined with vitamin C in lambs naturally infected with pneumonic pasteurellosis

Variables	CSS	IL-1 $\beta$	IL-6	IL-12p40	TNF- $\alpha$	IFN $\gamma$
<b>CSS</b>		0.55	0.38	0.93	0.25	0.44

<b>IL-1 <math>\beta</math></b>	0.55		0.90	0.94	0.37	0.32
<b>IL-6</b>	0.38	0.90		0.34	0.17	0.61
<b>IL-12p40</b>	0.93	0.94	0.34		0.46	0.95
<b>TNF-<math>\alpha</math></b>	0.25	0.37	0.17	0.46		0.38
<b>IFN<math>\gamma</math></b>	0.44	0.32	0.61	0.95	0.38	

Abbreviations: CSS (clinical signs score), interleukin 1  $\beta$  (IL-1  $\beta$ ), interleukin 6 (IL-6), interleukin 12p40 (IL-12p40), interferon gamma (IFN $\gamma$ ) and tumor necrosis factor alpha (TNF- $\alpha$ ). Differences between means at  $P < 0.05$  were considered significant.