GAG content micro-aggregates **DNA content micro-aggregates** 10 Nx Nx *** 0.4 Нx Нx 8 GAG content (µg) DNA content (µg) 0.3 6 0.2 4 0.1 2-0 0.0 SIN 0.548 control SIN 0.548 IN THE 1GF.B control 142148 164.⁸⁷ b а GAG content corrected for DNA GAG release micro-aggregates 50 30 Nx Nx Нx Нx 40 GAG/DNA (µg/µg) GAG release (µg) 20 30 20 10 10 0 0 SIN 0.5HB IN THE SINO.548 control IN THE control 16F.B 1GF.B d С

Supplementary File 3. The effect of human (s)Link-N on CD canine CLCs in hypoxia vs. normoxia

GAG and DNA content (mean + SD) of CD canine CLC micro-aggregates treated with basal culture medium (control), supplemented with 10 ng/mL TGF- β_1 (positive control), 1 µg/mL human Link-N (LN) or 0.5 µg/mL human sLink-N (sLN). The CLC micro-aggregates were cultured for 28 days in normoxia (Nx, 21% O₂) or hypoxia (Hx, 5% O₂). (a) GAG content (b) DNA content (c) GAG content corrected for DNA content (d) Total amount of GAGs released in the culture medium. * p < 0.05, ** p < 0.01, *** p < 0.001. \times , $\times\times$, $\times\times\times$: significantly different (p < 0.05, p < 0.01, p < 0.001 respectively) from all other conditions (growth factor treatment) in either Hx or Nx. n = 6 (in duplicates).