Supplement 1. The most upregulated differentially expressed protein in PDE-EVs of the H/A group

| Protein accession | Protein name | Fold of change(H/L) | p-value |
|-------------------|--------------|---------------------|---------|
| P32455            | Guanylate-binding protein 1 | 12.64               | 0.0276  |
| A0A024F8R8        | HLA-DRB1     | 7.73                | 0.0371  |
| P05164            | Myeloperoxidase | 7.72                | 0.0272  |
| P05107            | Integrin beta-2    | 6.91                | 0.0138  |
| A0A0S2Z3S6        | Cytochrome b-245 beta polypeptide isoform 1 | 5.87        | 0.0462  |
| B4DP27            | Transmembrane emp24 domain-containing protein 2 | 5.09        | 0.0048  |
| P12314            | High affinity immunoglobulin gamma Fc receptor I | 4.54        | 0.0435  |
| A0A024QZD1        | Ribosomal protein L18 | 4.33                | 0.0086  |
| A0A024R8U1        | Solute carrier family 16 (Monocarboxylic acid transporters), member 3, isoform CRA_a | 4.27        | 0.0266  |
| Q86UX7            | Fertin family homolog 3     | 4.11                | 0.0047  |
| A0A024R7M0        | Transmembrane emp24 protein transport domain containing 9, isoform CRA_a | 3.99        | 0.0316  |
| Q8N386            | Leucine-rich repeat-containing protein 25 | 3.97        | 0.0475  |
| Q99816            | Tumor susceptibility gene 101 protein | 3.83        | 0.0422  |
| A0A0A0MTS7        | Titin          | 3.70                | 0.0468  |
| B2R4C0            | 60S ribosomal protein L18a | 3.36        | 0.0119  |
| B2RBL3            | Thymidine phosphorylase    | 3.31                | 0.0347  |
| P62269            | 40S ribosomal protein S18  | 3.12        | 0.0252  |
| A0A024R705        | Vacuolar protein sorting-associated protein 4A | 3.09        | 0.0365  |
| A0A140VPJ5        | S-adenosylmethionine synthase isoform type-2 | 3.09        | 0.0363  |
| P28838            | Cytosol aminopeptidase     | 3.08                | 0.0117  |
| D6RGG3            | Collagen alpha-1(XII) chain | 3.00       | 0.0409  |
| P05109            | Protein S100-A8            | 2.94                | 0.0461  |
| Q15404            | Ras suppressor protein 1   | 2.83                | 0.0085  |
| P08311            | Cathepsin G               | 2.83                | 0.0286  |
| P18124            | 60S ribosomal protein L7   | 2.82                | 0.0049  |
| B7ZKY2            | Peripheral plasma membrane protein CASK | 2.79        | 0.0006  |
| A0A0UM2           | FOLR2 protein (Fragment)   | 2.78                | 0.0308  |
| A0A024RAS5        | Rho GDP dissociation inhibitor (GDI) beta, isoform CRA_a | 2.68        | 0.0357  |
| A0A024R9G4        | Family with sequence similarity 49, member B, isoform CRA_a | 2.61        | 0.0386  |
| A0A024RDT4        | Lymphocyte cytosolic protein 1 (L-plasmin), isoform CRA_a | 2.53        | 0.0403  |
| Q92982            | Ninjurin-1                | 2.48                | 0.0211  |
| A5PLK9            | Bone morphogenetic protein 1 | 2.46       | 0.0163  |
| B4DXC4            | Atlastin-3                | 2.39                | 0.0157  |
| Accession  | Description                                                                 | Fold Change | P-value |
|------------|-------------------------------------------------------------------------------|-------------|---------|
| A0A024R1V4 | 60S ribosomal protein L27                                                      | 2.38        | 0.0421  |
| A0A024R9N6 | EH-domain containing 4, isoform CRA_a                                       | 2.38        | 0.0318  |
| P26447     | Protein S100-A4                                                               | 2.32        | 0.0362  |
| L7RT22     | Integrin beta;Integrin beta-5                                                 | 2.28        | 0.0447  |
| Q1RMC9     | ERBB2IP protein                                                               | 2.24        | 0.0213  |
| K4RH61     | Matrix metalloproteinase-14                                                   | 2.21        | 0.0473  |
| A0A024R2F9 | Transmembrane protein 43                                                      | 2.19        | 0.0250  |
| A0A140VK6  | cDNA FLJ61307, highly similar to Netrin G1                                    | 2.18        | 0.0423  |
| A0A024R183 | Cyclin-dependent kinase 16, partial [Neogale vison]                           | 2.08        | 0.0143  |
| Q6FIG4     | RAB1B protein                                                                 | 2.06        | 0.0412  |
| A0A140VK27 | Leukotriene A(4) hydrolase                                                    | 2.06        | 0.0378  |
| A0A024R8S5 | Protein disulfide-isomerase                                                   | 2.05        | 0.0404  |
| Q15818     | Neuronal pentraxin-1                                                           | 1.98        | 0.0038  |
| P14625     | Glycoprotein 96                                                                | 1.97        | 0.0192  |
| Q9P1F3     | Costars family protein ABRACL                                                  | 1.96        | 0.0463  |
| P0CG39     | POTE ankyrin domain family member J                                           | 1.92        | 0.0470  |