**Supplementary Table 1. Inter-group comparison of the different timepoints for MDA-MB-231 Spheroids.**

|  |  |  |  |
| --- | --- | --- | --- |
| **Timepoint** | **Comparison** | **p-value** | **Significance** |
| **Day 0** | Control vs. 1% Matrigel | 0.7 | ns |
| Control vs. 3.5% Matrigel | 0.07 | ns |
| 1%Matrigel vs. 3.5% Matrigel | 0.13 | ns |
| **Day 3** | Control vs. 1% Matrigel | 0.001 | \*\*\* |
| Control vs. 3.5% Matrigel | 0.0002 | \*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.5 | ns |
| **Day 7** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.05 | \* |
| **Day 10** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.34 | ns |
| **Day 14** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | 0.0001 | \*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.99 | ns |

ns: not significant, the significance level was indicated at \* p<0.05, \*\*\*p<0.001 and \*\*\*\*p<0.0001.

**Supplementary Table 2. Inter-group comparison of the different timepoints for Osteoblast Spheroids.**

|  |  |  |  |
| --- | --- | --- | --- |
| **Timepoint** | **Comparison** | **p-value** | **Significance** |
| **Day 0** | Control vs. 1% Matrigel | 0.9 | ns |
| Control vs. 3.5% Matrigel | 0.005 | \*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.04 | \* |
| **Day 3** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| **Day 7** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| **Day 10** | Control vs. 1% Matrigel | 0.0036 | \*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.09 | ns |
| **Day 14** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.0002 | \*\*\* |

ns: not significant, the significance level was indicated at \* p<0.05, \*\*p<0.01, \*\*\*p<0.001 and \*\*\*\*p<0.0001.

**Supplementary Table 3. Inter-group comparison of the different timepoints for Heterotypic Spheroids.**

|  |  |  |  |
| --- | --- | --- | --- |
| **Timepoint** | **Comparison** | **p-value** | **Significance** |
| **Day 0** | Control vs. 1% Matrigel | 0.39 | ns |
| Control vs. 3.5% Matrigel | 0.009 | \*\* |
| 1%Matrigel vs. 3.5% Matrigel | >0.99 | ns |
| **Day 3** | Control vs. 1% Matrigel | 0.0003 | \*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.956 | ns |
| **Day 7** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | >0.99 | ns |
| **Day 10** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | >0.99 | ns |
| **Day 14** | Control vs. 1% Matrigel | <0.0001 | \*\*\*\* |
| Control vs. 3.5% Matrigel | <0.0001 | \*\*\*\* |
| 1%Matrigel vs. 3.5% Matrigel | 0.09 | ns |

ns: not significant, the significance level was indicated at \*\*p<0.01, \*\*\*p<0.001 and \*\*\*\*p<0.0001.