

Supplementary Table 1. Seventeen differentially expressed genes between the tumorspheres relative to Br cell lines.

| Gene symbol | Mean difference (tumorsphere-MB231-Br) | P value | Mean difference (tumorsphere-CN34-Br) | P value | Description |
|-----------------|--|--------------------|---------------------------------------|--------------------|---|
| ADAMTS1 | 4.877 | 0.002 | 3.153 | 5X10 ⁻⁵ | ADAM metallopeptidase with thrombospondin type 1 motif, 1 |
| ARHGAP29 | 1.565 | 0.002 | 1.746 | 0.0015 | Rho GTPase activating protein 29 |
| COL12A1 | 1.482 | 0.004 | 1.648 | 0.0016 | collagen, type XII, alpha 1 |
| DUSP1 | 1.45 | 0.008 | 2.044 | 0.0009 | dual specificity phosphatase 1 |
| FGF5 | 3.486 | 0.008 | 1.684 | 0.0022 | fibroblast growth factor 5 |
| FRAS1 | 4.52 | 0.001 | 2.013 | 0.0014 | Fraser syndrome 1 |
| HSPG2 | 2.061 | 5X10 ⁻⁵ | 1.474 | 0.0075 | heparan sulfate proteoglycan 2 |
| IL18 | 3.111 | 0.0001 | 2.131 | 0.0011 | interleukin 18 (interferon-gamma-inducing factor) |
| LPAR1 | 3.241 | 0.0015 | 2.809 | 0.0003 | lysophosphatidic acid receptor 1 |
| LPHN2 | 1.855 | 0.0035 | 1.966 | 0.0013 | latrophilin 2 |
| MMP1 | 8.35 | 5X10 ⁻⁵ | 1.416 | 0.0093 | matrix metallopeptidase 1 (interstitial collagenase) |
| NMNAT2 | 3.032 | 0.0019 | 2.209 | 0.0067 | nicotinamide nucleotide adenylyltransferase 2 |
| PCDH7 | 5.859 | 0.0008 | 1.781 | 0.007 | protocadherin 7 |
| PTGFRN | 3.99 | 0.0011 | 4.929 | 0.0032 | prostaglandin F2 receptor negative regulator |
| SDPR | 1.757 | 0.0018 | 3.471 | 0.001 | serum deprivation response |
| TIE1 | 4.810 | 0.0081 | 2.105 | 0.0013 | tyrosine kinase with immunoglobulin-like and EGF-like domains 1 |
| STC1 | -1.385 | 0.0087 | -1.597 | 0.0053 | stanniocalcin 1 |

Supplementary Table 2. Expression of PCDH7 in brain metastases tissues of breast cancer patient.

| H score | Negative (0-9) | Low (10-100) | Medium (101-200) | High (201-300) |
|--|-------------------|-----------------|---------------------|-------------------|
| Primary breast tumors | 25 (73%) | 6 (18%) | 3 (9%) | 0 (0%) |
| Brain metastatic tumors of breast cancer | 0 (0%) | 0 (0%) | 2 (7%) | 27 (93%) |

Supplementary Table 3. Pairwise sequence comparison of PCDH7 EC1-EC2 domain to diverse cadherins with structures available in Protein Data bank.

| Protein | PDB code | Sequence Identity | Sequence Similarity | Residues | Description | Subfamily | Resolution (Å) |
|---------------------------------|-----------|-------------------|---------------------|----------|---|--------------|----------------|
| E cadherin | 2O72 (14) | 19.40% | 32.40% | 1-213 | EC1 and EC2; strand exchange trans dimer | TYPE I | 2.00 |
| C cadherin | 1L3W (10) | 20.80% | 35.50% | 1-540 | EC1 - EC5; strand exchange trans dimer | TYPE I | 3.08 |
| N cadherin | 2QVI | 20.60% | 35.50% | 1-215 | EC1 and EC2; strand exchange trans dimer | TYPE I | 3.01 |
| Desmoglein-2 | 2YQG | 10.40% | 15.00% | 1-123 | EC1, monomer | Desmoglein | Null* |
| Cadherin-6 | 3LND (15) | 22.90% | 36.90% | 1-207 | EC1 and EC2; X dimer, W4A mutation | TYPE II | 2.82 |
| VE-Cadherin | 3PPE (16) | 20.90% | 36.10% | 1-203 | EC1 and EC2; strand exchange trans dimer | TYPE II | 2.1 |
| Cadherin-8 | 2A62 (17) | 20.80% | 37.40% | 1-322 | EC1 - EC3; strand exchange trans dimer | TYPE II | 4.5 |
| Cadherin-11 | 2A4E (17) | 20.90% | 34.80% | 0-207 | EC1 and EC2; strand exchange trans dimer | TYPE II | 3.2 |
| MN-cadherin | 1ZVN (17) | 19.60% | 45.10% | 0-98 | EC1; strand exchange trans dimer | TYPE II | 2.16 |
| T-cadherin | 3K6D (18) | 9.20% | 14.30% | 1-99 | EC1, X dimer | nonclassical | 1.8 |
| Cadherin-23 Protocadherin 15 | 3MVS (19) | 19.00% | 36.20% | 1-210 | EC1 and EC2 | nonclassical | 1.1 |
| Protocadherin 4 | 1WUZ (20) | 28.90% | 44.70% | 1-103 | EC1, monomer | nonclassical | Null* |
| Protocadherin 7 | 2YST | 10.80% | 17.30% | 302-413 | EC3, monomer | nonclassical | Null* |
| E cadherin | 1EDH (21) | 16.10% | 32.60% | 3-213 | EC1 and EC2, X dimer; DW residues in N terminus were removed | TYPE I | 2 |