

## Common Markers of Bone and Soft Tissue Tumors

| Bone Tumor                                 | Immunohistochemical Markers                                              | Molecular Markers                                                                             |
|--------------------------------------------|--------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|
| <b>Osteosarcoma</b>                        | SATB2, MDM2 (in low-grade)                                               | RB1, TP53, CDK4, MDM2 (low-grade)                                                             |
| <b>Ewing Sarcoma</b>                       | CD99 (strong, diffuse), FLI1                                             | EWSR1-FLI1 (85%), EWSR1-ERG (10%)                                                             |
| <b>Chondrosarcoma</b>                      | S100                                                                     | IDH1/IDH2 mutations (central chondrosarcoma), EXT1/EXT2 mutations (peripheral chondrosarcoma) |
| <b>Giant Cell Tumor of Bone (GCTB)</b>     | H3.3G34W (specific)                                                      | H3F3A G34W mutation (majority of cases)                                                       |
| <b>Chordoma</b>                            | Brachyury (specific), S100, keratin                                      | Loss of CDKN2A, T (Brachyury) amplification                                                   |
| <b>Osteoid Osteoma</b>                     | SATB2                                                                    | No recurrent molecular alterations                                                            |
| <b>Osteoblastoma</b>                       | SATB2                                                                    | FOS gene rearrangements (specific)                                                            |
| <b>Chondroblastoma</b>                     | H3. 3 K36M                                                               | H3F3B K36M mutation (specific)                                                                |
| <b>Fibrous Dysplasia</b>                   | SMA, SATB2                                                               | GNAS1 mutations (GNAS R201H or R201C)                                                         |
| <b>Adamantinoma</b>                        | Cytokeratins , EMA                                                       | Associated with recurrent chromosomal changes, including gains of chromosome 7, 8, 12, and 19 |
| <b>Plasmacytoma/Multiple Myeloma</b>       | CD138, MUM1, kappa/lambda light chain restriction                        | IGH rearrangements, MYC alterations, RAS mutations                                            |
| <b>Langerhans Cell Histiocytosis (LCH)</b> | CD1a, Langerin (CD207), S100                                             | BRAF V600E mutation (50-60% cases), MAP2K1 mutations                                          |
| <b>Metastatic Carcinoma to Bone</b>        | CK7, CK20, TTF1 (lung), PSA (prostate), GATA3 (breast, urothelial), etc. | Depends on primary tumor site (e.g., TP53, EGFR, ALK in lung cancer)                          |

| Soft Tissue Tumor                                         | Immunohistochemical Markers | Molecular Markers                             |
|-----------------------------------------------------------|-----------------------------|-----------------------------------------------|
| <b>Liposarcoma (Well-Differentiated/Dedifferentiated)</b> | MDM2, CDK4, S100            | MDM2/CDK4 amplification (12q13-15)            |
| <b>Myxoid/Round Cell Liposarcoma</b>                      | S100 (variable), DDIT3      | FUS-DDIT3 (95%), EWSR1-DDIT3 (rare)           |
| <b>Pleomorphic Liposarcoma</b>                            | S100, MDM2 (negative)       | Complex karyotype, no specific translocations |
| <b>Leiomyosarcoma</b>                                     | SMA, Desmin                 | TP53, RB1, ATRX mutations                     |

|                                                        |                                                                      |                                                                   |
|--------------------------------------------------------|----------------------------------------------------------------------|-------------------------------------------------------------------|
| <b>Undifferentiated Pleomorphic Sarcoma (UPS)</b>      | No specific marker. Diagnosis of exclusion.                          | Complex karyotype, no specific translocation                      |
| <b>Rhabdomyosarcoma (Alveolar)</b>                     | MyoD1, Myogenin, Desmin                                              | PAX3-FOXO1 (70%), PAX7-FOXO1 (20%)                                |
| <b>Rhabdomyosarcoma (Embryonal)</b>                    | MyoD1, Myogenin, Desmin                                              | Loss of heterozygosity at 11p15, mutations in TP53, NRAS, FGFR4   |
| <b>Synovial Sarcoma</b>                                | TLE1, EMA                                                            | SS18-SSX1 (most common), SS18-SSX2                                |
| <b>Epithelioid Sarcoma</b>                             | EMA, CK, CD34                                                        | SMARCB1 (INI1) loss                                               |
| <b>Clear Cell Sarcoma</b>                              | S100, HMB45, MelanA, SOX10 (like melanoma)                           | EWSR1-ATF1 fusion                                                 |
| <b>Alveolar Soft Part Sarcoma</b>                      | TFE3, PAS-positive crystals                                          | ASPSCR1-TFE3 fusion                                               |
| <b>Dermatofibrosarcoma Protuberans (DFSP)</b>          | CD34 (strong, diffuse)                                               | COL1A1-PDGFB fusion                                               |
| <b>Malignant Peripheral Nerve Sheath Tumor (MPNST)</b> | S100 (focal or absent), SOX10 (variable), H3K27me3 loss (high-grade) | NF1 mutations, SUZ12/EED loss                                     |
| <b>Solitary Fibrous Tumor</b>                          | STAT6 (specific), CD34                                               | NAB2-STAT6 fusion                                                 |
| <b>Myxofibrosarcoma</b>                                | CD34 (variable)                                                      | Complex karyotype, no specific translocation                      |
| <b>Desmoid-Type Fibromatosis</b>                       | $\beta$ -catenin (nuclear), SMA (variable)                           | CTNNB1 mutations (common), APC mutations (rare, Gardner syndrome) |
| <b>Angiosarcoma</b>                                    | CD31, ERG, FLI1                                                      | MYC amplification (post-radiation cases), PTPRB, KDR mutations    |
| <b>Ewing-Like Sarcoma (BCOR-CCNB3, CIC-rearranged)</b> | BCOR-CCNB3: BCOR, SATB2; CIC-rearranged: WT1, ETV4                   | BCOR-CCNB3 fusion, CIC-DUX4 fusion                                |