

Supplementary table 1. State of the art diagnostic methods for selected tumor types previously diagnosed as CNS PNET

Tumor diagnosis	Immunohistochemistry markers	Genetic techniques	Other molecular techniques
Astroblastoma, <i>MN1</i> -altered	GFAP positive EMA positive OLIG2 at least focally positive	FISH with break-apart <i>MN1</i> probe to detect fusion	
Atypical teratoid/rhabdoid tumor	EMA positive Vimentin positive SMA and cytokeratin frequently positive Nuclear loss of Ini-1 or Brg1 protein	<i>SMARCB1</i> or <i>SMARCA4</i> mutations can be identified by DNA sequencing. Larger deletions can be detected by FISH with specific probes or array-based methods (CGH, MIP)	
Choroid Plexus Carcinoma	Cytokeratin positive EMA positive p53 nuclear accumulation in TP53 mutant cases	Detection of chromosomal losses by array-based methods (CGH, MIP)	
CNS neuroblastoma, <i>FOXR2</i> -activated	OLIG2 positive Synaptophysin positive TTF-1 frequently positive GFAP negative Vimentin negative	Almost all cases show gain of chromosome 1q, loss of 16q is also frequent. Detection by FISH with specific probes or array-based methods (CGH, MIP)	Detection of mRNA overexpression of <i>FOXR2</i> and <i>NKX2.1</i> by quantitative RT-PCR or Nanostring technology
CNS tumor with <i>BCOR</i> internal tandem duplication	Vimentin positive CD56 (NCAM) often positive <i>BCOR</i> nuclear positive	Internal tandem duplication of <i>BCOR</i> detected by PCR	
Embryonal tumor with multilayered rosettes	LIN 28 positive Vimentin positive Synaptophysin positive in neuropil-like matrix	<i>C19MC</i> amplification detected by FISH or array-based methods (CGH, MIP) For <i>C19MC</i> non-amplified cases, DNA sequencing for <i>DICER1</i> mutations	
Ependymoma <i>ZFTA</i> fusion-positive	EMA positive Vimentin positive Nuclear accumulation of p65-RelA in most cases	FISH with break-apart <i>ZFTA</i> probe RT-PCR or Nanostring technology to detect <i>ZFTA</i> fusion transcripts	
HGG (different types)	GFAP positive OLIG2 positive (all but H3-G34 mutant HGG) Vimentin positive H3-K27me3 negative and H3K27M mostly positive and in diffuse midline gliomas H3G34R/V positive in hemispheric gliomas with H3-G34 mutation BRAFV600E often positive in anaplastic PXA /epithelioid GBM	DNA (Pyro)sequencing for mutations of H3F3A, other Histone genes, IDH1, IDH2, BRAF (V600) FISH with break-apart probes to detect fusions of <i>NTRK1</i> , -2, -3, <i>ALK</i> , <i>ROS</i> , <i>MET</i> in Infantile hemispheric gliomas. RNA-based methods to detect fusion transcripts (RT-PCR, Nanostring technology, RNA-Seq)	
Pineoblastoma	OT2 and OTX 3 positive Synaptophysin positive Vimentin negative		

GFAP, glial fibrillary acidic protein; EMA, epithelial membrane antigen; OLIG2, oligodendrocyte transcription factor 2; SMA, smooth muscle actin; TTF-1, Thyroid transcription factor-1; HGG, high grade glioma