

## SUPPLEMENTAL FIGURES

Primary CNS Tumors by WHO Classification Diagnosis	WHO Grade(s)	SWI/SNF Mutation(s)	Pathways Interacting with SWI/SNF	Human ERV(s) Expressed	Non-Human ERV(s) Expressed	References
<b>Gliomas, glioneuronal tumors, and neuronal tumors</b>						
<i>Adult-type diffuse gliomas</i>						
Astrocytoma, IDH-mutant	2-4	SMARCE1, SMARCA4, ATRX	TP53, MYC, RB1	-	-	[5,21–26,84]
Oligodendroglioma, IDH-mutant and 1p/19q-codeleted	2-3	BICRA (GLTSCR1); ARID1A	SETD2, MYC; RB1	-	-	[5,22,27,85–87]
Glioblastoma, IDH-wildtype	4	ATRX	EZH1P, SETD2; TP53; MYC; RB1	HERV1, HERVK, HERVL, ERV3, HML-6 (ERVK3-1)	-	[5,28,29,78,79,88]
<i>Pediatric-type diffuse low-grade gliomas</i>						
Diffuse astrocytoma, MYB- or MYBL1-altered	1	-	-	-	-	
Angiocentric glioma	1	-	-	-	-	
Polymorphous low-grade neuroepithelial tumour of the young	1	-	TP53/RB1 <sup>1</sup>	-	-	[5,89]
Diffuse low-grade glioma, MAPK pathway-altered	1	-	-	-	-	
<i>Pediatric-type diffuse high-grade gliomas</i>						
Diffuse midline glioma, H3 K27-altered	4	SMARCA4; ATRX	EZH1P, H3K27; TP53	-	-	[5,30,31,90,91]
Diffuse hemispheric glioma, H3 G34-mutant	4	ATRX	H3G34; TP53; MYCN	-	-	[5,32–34,92]
Diffuse pediatric-type high-grade glioma, H3-wildtype and IDH-wildtype	4	-	TP53, MYCN	-	-	[5,93]
Infant-type hemispheric glioma	NR	-	-	-	-	
<i>Circumscribed astrocytic gliomas</i>						
Pilocytic astrocytoma	1	ATRX <sup>2</sup>	-	-	-	[35]
High-grade astrocytoma with piloid features	3-4	ATRX	-	-	-	[35]
Pleomorphic xanthroastrocytoma	2-3	SMARCB1, ARID1A, ATRX	TP53 <sup>2</sup> , NOTCH <sup>2</sup>	-	-	[5,36–38]
Subependymal giant cell astrocytoma	1	-	-	-	-	
Chordoid glioma	2	-	-	-	-	

Astroblastoma, MN1-altered	NR	-	MN1	-	-	[94]
<i>Glioneuronal and neuronal tumors</i>						
Ganglioglioma	1	-	-	-	-	
Gangliocytoma	1	-	-	-	-	
Desmoplastic infantile ganglioglioma/desmoplastic infantile astrocytoma	1	ATRX <sup>1</sup>	TP53 <sup>1</sup>	-	-	[5,39–42]
Dysembryoplastic neuroepithelial tumor	1	-	-	-	-	
Diffuse glioneuronal tumor with oligodendroglioma-like features and nuclear clusters	NR	-	-	-	-	
Papillary glioneuronal tumor	1	-	-	-	-	
Rosette-forming glioneuronal tumor	1	-	-	-	-	
Myxoid glioneuronal tumor	1	-	-	-	-	
Diffuse leptomeningeal glioneuronal tumor	NR	-	-	-	-	
Multinodular and vacuolating neuronal tumor	1	-	-	-	-	
Dysplastic cerebellar gangliocytoma (Lhermitte-Duclos disease)	1	-	-	-	-	
Central neurocytoma	2	-	MYCN	-	-	[5,95]
Extraventricular neurocytoma	2	-	-	-	-	
Cerebellar liponeurocytoma	2	-	TP53	-	-	[5,96]
<i>Ependymal tumors</i>						
Supratentorial ependymoma (fusion-negative)	2-3	-	-	-	-	
Supratentorial ependymoma, ZFTA fusion-positive	2-3	-	ZFTA-RELA	-	-	[97]
Supratentorial ependymoma, YAP1 fusion-positive	2-3	-	YAP1	-	-	[98]
Posterior fossa ependymoma	2-3	-	EZH1P	-	-	[99]
Posterior fossa group A (PFA) ependymoma	2-3	-	EZH1P	-	-	[99]
Posterior fossa group B (PFB) ependymoma	2-3	-	-	-	-	
Spinal ependymoma	2-3	-	-	-	-	

Spinal ependymoma, MYCN-amplified	2-3	-	MYCN	-	-	[5,100–103]
Myxopapillary ependymoma	2	-	-	-	-	
Subependymoma	1	-	-	-	-	
<b>Choroid plexus tumors</b>						
Choroid plexus papilloma	1	-	TP53 (rare)	-	Roux Sarcoma Virus	[5,72,104]
Atypical choroid plexus papilloma	2	-	-	-	-	
Choroid plexus carcinoma	3	-	TP53; SHH, NOTCH	-	-	[5,104–106]
<b>Embryonal tumors</b>						
<i>Medulloblastoma</i>						
Medulloblastoma, WNT-activated	4	SMARCA4, SMARCB1, ARID1A, ARID2	WNT/CTNNB1, TP53, OCT4	-	-	[5,43,107]
Medulloblastoma, SHH-activated and TP53-wildtype	4	-	SHH, CREBBP; MYCN, MYCL, YAP1, OCT4	-	-	[5,43,44,107,109]
Medulloblastoma, SHH-activated and TP53-mutant	4	-	TP53, SHH, CREBBP, OCT4	-	-	[5,44,107,109]
Medulloblastoma, non-WNT/non-SHH (Group 3/4)	4	SMARCA4	MYC	-	-	[5,107,109,110]
Medulloblastoma, histologically defined	4	Not relevant	Not relevant	-	-	
<i>Other CNS embryonal tumors</i>						
Atypical teratoid/rhabdoid tumor	4	SMARCB1, SMARCA4	SHH, NRAS, MYC	HERV-K	-	[5,10,45–50]
Cribriform neuroepithelial tumor	NR	SMARCB1	-	-	-	[51]
Embryonal tumor with multilayered rosettes	4	-	LIN28A; TP53	-	-	[5,111]
CNS neuroblastoma, FOXR2-activated	4	-	-	-	-	
CNS tumor with BCOR internal tandem duplication	NR	SMARCA2	-	-	-	[52]
CNS embryonal tumor NEC/NOS	3-4	-	-	-	-	
<b>Pineal tumors</b>						
Pineocytoma	1	-	-	-	-	
Pineal parenchymal tumor of intermediate differentiation	2-3	-	-	-	-	



Solitary fibrous tumor	1-3	-	EP300; TP53	-	-	[120–122]
Soft tissue tumors: Vascular tumors						
Hemangiomas and vascular malformations	NR	-	-	-	ALV	[75]
Hemangioblastoma	1	-	-	-	-	
Soft tissue tumors: Skeletal muscle tumors						
Rhabdomyosarcoma	NR	-	-	ERV-9	Feline endogenous retrovirus	[77]
Soft tissue tumors: Tumors of uncertain differentiation						
Intracranial mesenchymal tumor, FET::CREB fusion-positive	NR	-	FET	-	-	[123]
CIC-rearranged sarcoma	4	-	NUTM1	-	-	[5]
Primary intracranial sarcoma, DICER1-mutant	NR	ATRX	DICER1; TP53	-	-	[68,112]
Ewing sarcoma	4	-	FET; TP53	Syncytin-1; ERV-L	-	[80,123–126]
Chondro-osseous tumors: Chondrogenic tumors						
Mesenchymal chondrosarcoma	NR	-	NOTCH, NCoA-2	-	-	[127]
Chondrosarcoma	1-3	-	RB1, H3K36	-	-	[128-130]
Notochordal tumors						
Chordoma	NR	SMARCB1	-	-	-	[69,70]
Melanocytic tumors						
Diffuse meningeal melanocytic neoplasms						
Melanocytosis and melanomatosis	NR	-	YAP1	-	-	[131]
Circumscribed meningeal melanocytic neoplasms						
Melanocytoma and melanoma	NR	-	YAP1	-	MMVL30	[81,131]
Hematolymphoid tumors involving the CNS						
Lymphomas: CNS lymphomas						
Primary diffuse large B-cell lymphoma of the CNS	NR	-	MYC	Variety - Unspecified	-	[82,132]
Immunodeficiency-associated CNS lymphomas	NR	-	-	-	-	
Lymphomatoid granulomatosis	1-3	-	-	-	-	
Intravascular large B-cell lymphoma	NR	-	-	-	-	
Lymphomas: Miscellaneous rare lymphomas in the CNS						

MALT lymphoma of the dura	NR	-	NOTCH	-	-	[5]
Other low-grade B-cell lymphomas of the CNS	NR	-	-	-	-	
Anaplastic large cell lymphoma (ALK+/ALK-)	NR	-	-	-	-	
T-cell and NK/T-cell lymphomas	NR	-	-	-	-	
<i>Histiocytic tumors</i>						
Erdheim-Chester disease	NR	-	NRAS	-	-	[133]
Rosai-Dorfman disease	NR	-	-	-	-	
Juvenile xanthogranuloma	NR	-	-	-	-	
Langerhans cell histiocytosis	NR	-	-	-	primate type D retroviruses, murine intracisternal A particles, Jaagsiekte sheep retrovirus, and murine long interspersed nuclear elements	[73]
Histiocytic sarcoma	NR	-	-	-	-	
<b>Germ cell tumors</b>						
Mature teratoma	NR	-	JMJD1C; RB1	ERVK24	-	[83,134,135]
Immature teratoma	NR	SMARCA4	JMJD1C; RB1	-	-	[71,134,135]
Teratoma with somatic-type malignancy	NR	-	JMJD1C	-	-	[134]
Germinoma	NR	-	JMJD1C, LIN28A; TP53; RB1	ERVK24	-	[83,134–136]
Embryonal carcinoma	NR	-	JMJD1C, LIN28A; RB1	ERVK24	-	[83,134,135]
Yolk sac tumor	NR	-	JMJD1C, LIN28A; TP53; RB1	ERVK24	-	[83,134-136]
Choriocarcinoma	NR	-	JMJD1C; RB1	ERVK24	-	[83,134,135]
Mixed germ cell tumor <sup>3</sup>	NR	-	JMJD1C; RB1	-	-	[134,135]
<b>Tumors of the sellar region</b>						
Adamantinomatous craniopharyngioma	1	-	SHH, CTNNB1 (B-catenin)	-	-	[137]
Papillary craniopharyngioma	1	-	-	-	-	

Pituicytoma, Granular cell tumor of the sellar region, Spindle cell oncocytoma	NR	-	-	-	-	
Pituitary adenoma/pituitary neuroendocrine tumor	NR	-	Ik1, Ik2; TP53 <sup>5</sup>	-	-	[138,139]
Pituitary blastoma	NR	-	DICER1; TP53	-	-	[140,141]

**Supplemental Table S1.** Database of primary central nervous system (CNS) tumors outlined by the 5<sup>th</sup> edition of the World Health Organization (WHO) classification scheme and stratified by tumor category, assigned WHO grade, known epigenetic and/or genetic mutations of interest, and documented aberrant endogenous retrovirus (ERV) expression. Abbreviations: ALK (Anaplastic lymphoma kinase); ALV (Avian leukosis virus); ARID1A (AT-Rich Interaction Domain 1A); ARID1B (AT-Rich Interaction Domain 1B); ARID2 (AT-Rich Interaction Domain 2); ATRX (Alpha-thalassemia/mental retardation, X-linked); BCOR (BCL6 Corepressor); BICRA (BRD4 Interacting Chromatin Remodeling Complex Associated Protein); CIC (Capicua transcriptional repressor); CREBBP (cAMP-response element binding protein); CTNNB1 (Catenin beta-1); EED (Embryonic Ectoderm Development); EP300 (E1A-associated protein p300); ERV (Endogenous retrovirus); ERV3 (Endogenous retrovirus group 3); ERV9 (Endogenous retrovirus group 9); ERVK24 (Endogenous retrovirus group K member 24); ERVK3-1 (Endogenous retrovirus group 3 member 1); EZHIP (Enhancer of Zeste Homologs Inhibitory Protein); FOXR2 (Forkhead Box R2); GLTSCR1 (Glioma tumor suppressor candidate region gene 1); H3K27 (Histone 3 on lysine 27); H3K36 (Histone 3 on lysine 36); H3G34 (Histone 3 on arginine 34); HERV1 (Human endogenous retrovirus group 1); HERVK (Human endogenous retrovirus group K); HERVL (Human endogenous retrovirus group L); HML-6 (Human endogenous MMTV-like 6); IDH (Isocitrate dehydrogenase); Ik1 (Ikaros 1); Ik2 (Ikaros 2); JMJD1C (Jumonji domain containing 1C); KLF4 (Kruppel-like factor 4); LIN28A (Lin-28 homolog A); MALT (Mucosa associated lymphoid tissue); MAPK (Mitogen-activated protein kinase); MMVL30 (Mouse murine leukemia virus group L member 30); MN1 (Meningioma (disrupted in balanced translocation) 1); MYBL1 (MYB proto-oncogene like 1); NCoA-2 (Nuclear receptor coactivator 2); NEC (Not elsewhere classified); NK (Natural Killer); NOS (Not otherwise specified); NOTCH (Neurogenic locus notch homolog protein); NR (Not Reported); NRAS (Neuroblastoma ras viral oncogene homolog); NUTM1 (NUT midline carcinoma family member 1); OCT4 (Octamer-binding transcription 4); PBRM1 (Polybromo-1); RB1 (Retinoblastoma Transcriptional Corepressor 1); RELA (V-rel reticuloendotheliosis viral oncogene homolog A); SETD2 (SET domain-containing 2); SHH (Sonic Hedgehog); SMARCA2 (SWI/SNF Related Matrix Associated, Actin Dependent Regulator of Chromatin Subfamily A, Member 2); ; SMARCA4 (SWI/SNF Related Matrix Associated, Actin Dependent Regulator of Chromatin Subfamily A, Member 4); SMARCB1 (SWI/SNF Related Matrix Associated, Actin Dependent Regulator of Chromatin Subfamily B, Member 1); SMARCE1 (SWI/SNF Related Matrix Associated, Actin Dependent Regulator of Chromatin Subfamily E, Member 1); SUZ12 (Zeste 12 homolog); SWI/SNF (SWItch/Sucrose Non-Fermentable); TP53 (Tumor protein 53); WNT (Wingless-related integration site); YAP1 (Yes-associated protein-1); ZFTA (Zinc finger translocation associated)

<sup>1</sup>reported in one recurrent case with glioblastoma-like histology

<sup>2</sup>reported in rare tumors with anaplastic features

<sup>3</sup>dependent on composition of tumor-subtypes contained within primary tumor

<sup>4</sup>reported in meningiomas without concomitant Merlin mutation

<sup>5</sup>reported only in pituitary carcinomas

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<b>Gliomas, glioneuronal tumors, and neuronal tumors</b>					
<i>Adult-type diffuse gliomas</i>					
Glioblastoma, IDH-wildtype	4	EZH1P, SETD2; TP53; MYC; RB1	HERV1, HERVK, HERVL, ERV3, HML-6 (ERV3-1)	-	[28,29,78,79,88]
<b>Choroid plexus tumors</b>					

Choroid plexus papilloma	1	TP53 (rare)	-	Roux Sarcoma Virus	[5,104–106]
<b>Embryonal tumors</b>					
<i>Other CNS embryonal tumors</i>					
Atypical teratoid/rhabdoid tumor	4	SHH, NRAS, MYC	HERV-K	-	[5,10,45–50]
<b>Cranial and paraspinal nerve tumors</b>					
Malignant peripheral nerve sheath tumor	NR	SUZ12, EED, H3K27; TP53	-	ALV	[61–66,115–117]
<b>Meningioma</b>					
Secretory meningioma	1-3	KLF4	HERV-K <sup>4</sup>	-	[59,118]
Atypical meningioma	2-3	TP53	HERV-K <sup>4</sup>	-	[59,119]
Anaplastic (malignant) meningioma	3	H3K27M; TP53	HERV-K <sup>4</sup>	-	[5,59,119]
<b>Mesenchymal, non-meningothelial tumors involving the CNS</b>					
<i>Soft tissue tumors: Tumors of uncertain differentiation</i>					
Ewing sarcoma	4	FET; TP53	Syncytin-1; ERV-L	-	[80,123–126]
<b>Melanocytic tumors</b>					
<i>Circumscribed meningeal melanocytic neoplasms</i>					
Melanocytoma and melanoma	NR	YAP1	-	MMVL30	[81,131]
<b>Hematolymphoid tumors involving the CNS</b>					
<i>Lymphomas: CNS lymphomas</i>					
Primary diffuse large B-cell lymphoma of the CNS	NR	MYC	Variety - Unspecified	-	[82,132]
<b>Germ cell tumors</b>					
Mature teratoma	NR	JMJD1C; RB1	ERV-K24	-	[83,134,135]
Germinoma	NR	JMJD1C, LIN28A; TP53; RB1	ERV-K24	-	[83,134–136]
Embryonal carcinoma	NR	JMJD1C, LIN28A; RB1	ERV-K24	-	[83,134,135]
Yolk sac tumor	NR	JMJD1C, LIN28A; TP53; RB1	ERV-K24	-	[83,134–136]
Choriocarcinoma	NR	JMJD1C; RB1	ERV-K24	-	[83,134,135]

**Supplemental Table S2.** Primary CNS tumors with known genetic and/or epigenetic altered pathways with direct interaction with SWI/SNF subunits and aberrant ERV expression implicated in their tumorigenesis.