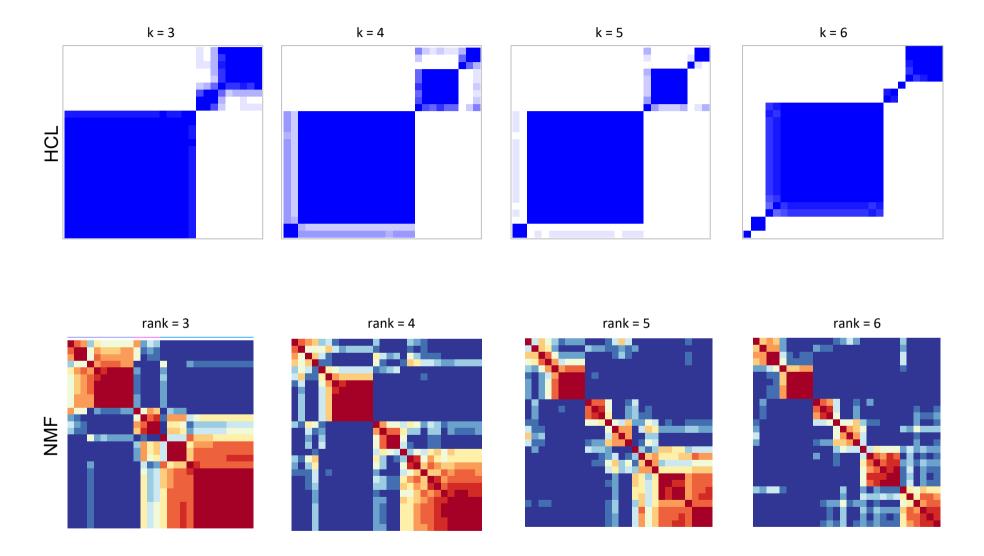
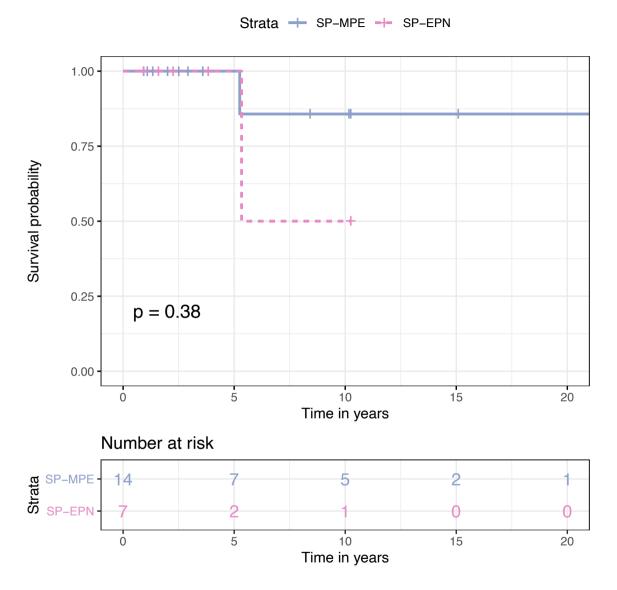


| Figure S1. Unsupervised analyses of 27 pediatric spinal ependymomas using DNA methylation profiles | .3 |
|--|----|
| Figure S2. Assignment of grade II and III pediatric spinal ependymomas using DNA methylation profiles | .5 |
| Figure S3. Assignment of grade II and III pediatric spinal ependymomas using DNA methylation profiles | .5 |
| Figure S4. Subgroup-associated gene signature | 6 |
| Figure S5. Differentially methylated distal probes between SP-MPE and SP-EPN subgroups of pediatric spinal ependymoma | 7 |
| Figure S6. DNA methylation CpG probe-gene pairs correlation between methylation and gene expression | 8 |
| Table S1. Clinical and histopathological data associated with all pediatric spinal ependymomas analysed in this study, along with molecular subgroup | |
| assignment | |

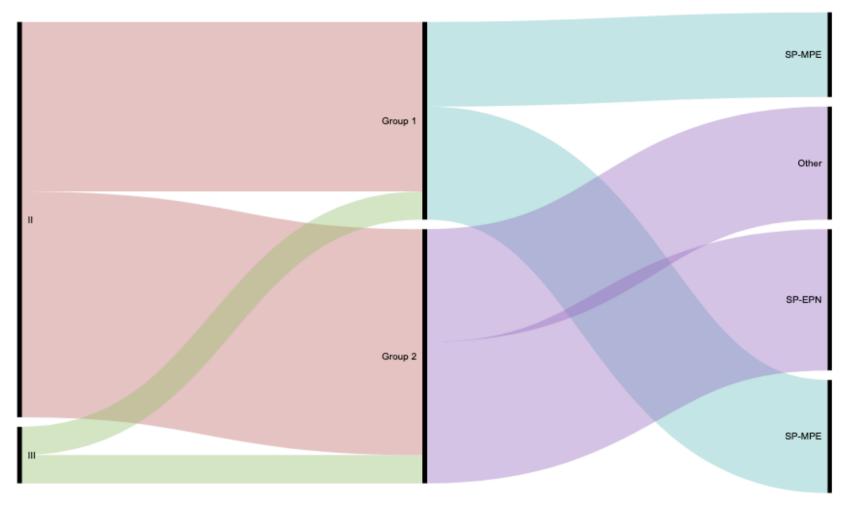
- **Table S2.** Summary of focal amplifications detected in spinal ependymoma by GISTIC2
- Table S3. Summary of focal deletions detected in spinal ependymoma by GISTIC2
- Table S4. List of significant pathways enriched by Gene Set Enrichment Analysis in SP-MPE subgroup
- Table S5. List of significant pathways enriched by Gene Set Enrichment Analysis in SP-EPN subgroup
- Table S6. List of significant pathways enriched in differentially methylated regions (DMR) that have high methylation levels in SP-MPE
- Table S7. List of significant pathways enriched in differentially methylated regions (DMR) that have low methylation levels in SP-MPE
- Table S8. List of CpG probe gene pairs with low methylation and high gene expression levels in SP-MPE
- Table S9. List of CpG probe gene pairs with high methylation and low gene expression levels in SP-MPE
- **Table S10.** Summary of significant transcription factor (TF) with motifs enriched among the list of CpG probe gene pairs with high methylation and low gene expression levels in SP-MPE



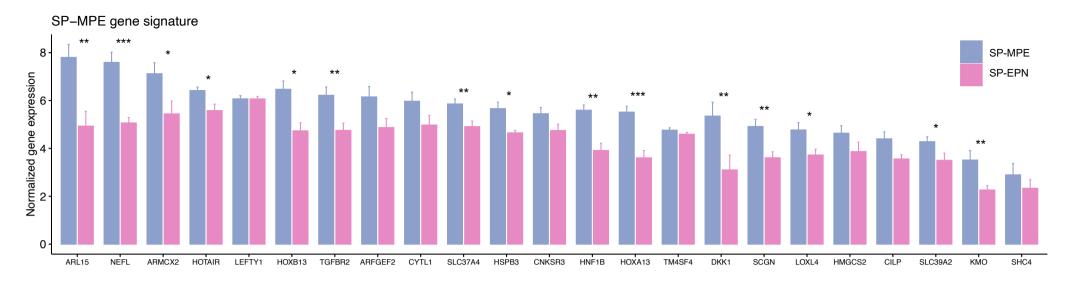
Supplementary figure S1. Unsupervised consensus hierarchical clustering (HCL) and non-negative matrix factorization (NMF) of 27 pediatric spinal ependymoma DNA methylation from a rank 2 to 6 classification.

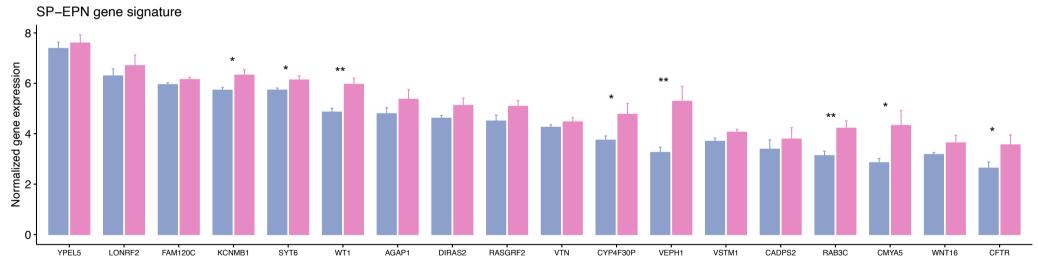


Supplementary figure S2. Kaplan-Meier curves of overall survival for our cohort when the patients are stratified by molecular subgroups, SP-MPE and SP-EPN. The median overall survival for the SP-EPN was 5.33 and was not reached for the SP-MPE. Significance levels were determined by log-rank test.

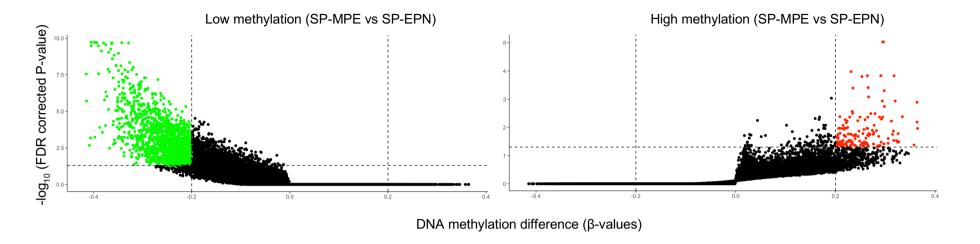


Supplementary figure S3. Assignment of grade II and III to molecular ependymoma subgroups based on DAN methylation profiles in our pediatric spinal ependymoma cohort.



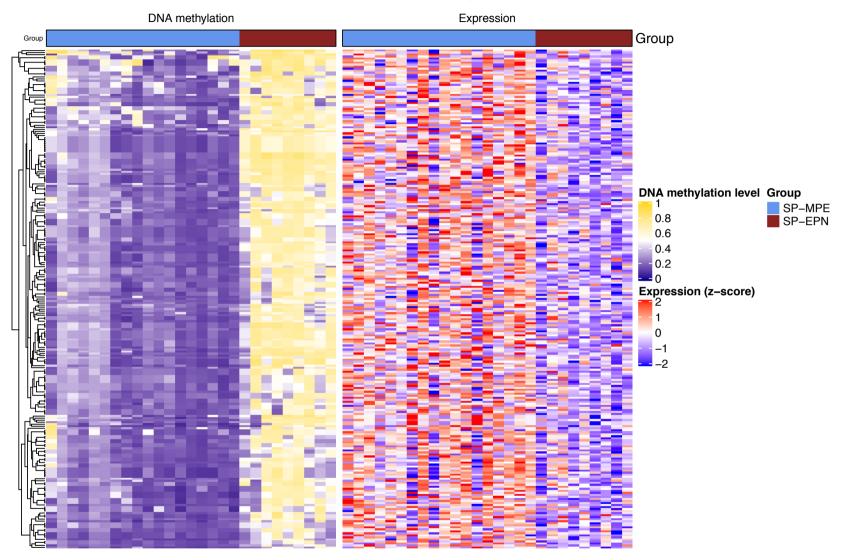


Supplementary figure S4. Subgroup-associated gene signature. Bar plot shows the expression levels of signature genes for SP-MPE (top) and SP-EPN subgroups (bottom). The subgroup-specific gene signatures were obtained from the previously published study by Pajtler et.al (2015). The Wilcoxon signed-rank test was used and p values (*: p < 0.05; **: p < 0.01; ***: p < 0.001) on the top indicate whether there was a significant difference in the distribution of expression values across the two subgroups, SP-MPE and SP-EPN. The p values for all SP-MPE signature genes remain significant after adjusting for multiple comparisons (false discovery rate, FDR) using the Benjamini-Hochberg adjustment whereas only three genes: *RAB3C*, *VEPH1* and *WT1* remain significant for the SP-EPN gene signature at the FDR < 0.05.



Supplementary figure S5. Differential methylated distal probes. Volcano plots showing DNA methylation differences (x-axis) and their statistical significance (y-axis) between SP-MPE and SP-EPN subgroups. Probes with low (left) and high (right) methylation levels in SP-MPE when compared to the SP-EPN subgroup are shown.

Correspondence between probe DNA methylation and distal gene expression



Supplementary figure S6. Probe-gene pairs showing inverse correlation between methylation and gene expression. Shown are the heatmaps for the significant DNA methylation probes (left) and gene expression (right) pairs at FDR < 0.05.