

THCA - Thyroid carcinoma

Subtype	Biology & Expression	Genomic Alterations	Clinical Features
BRAF-like	<ul style="list-style-type: none">• High MAPK pathway output; low thyroid differentiation score• Enrichment of proliferation and EMT signatures	<ul style="list-style-type: none">• BRAF^{V600E} mutations ($\approx 60\%$)• TERT promoter mutations (in a subset)• Few fusion events	<ul style="list-style-type: none">• Classical and tall-cell PTC histology• More frequent lymph node metastases• Higher recurrence risk
RAS-like	<ul style="list-style-type: none">• High thyroid differentiation score; retention of follicular gene programs• Lower MAPK output, more PI3K/AKT signaling	<ul style="list-style-type: none">• NRAS/HRAS/KRAS mutations ($\approx 13\%$)• PAX8-PPARG and other rare fusions• Rare TERT mutations	<ul style="list-style-type: none">• Follicular variant PTC and follicular carcinoma• Vascular invasion more common than nodal spread• Generally favorable prognosis
Kinase-fusion	<ul style="list-style-type: none">• Intermediate differentiation score; MAPK activation via receptor tyrosine kinases• Distinct fusion-driven expression	<ul style="list-style-type: none">• RET/PTC1, RET/PTC3, NTRK1/3, ALK and other RTK fusions ($\approx 5\text{--}10\%$)	<ul style="list-style-type: none">• Often in younger or radiation-exposed patients• Variable nodal spread• Targetable by specific kinase inhibitors