

## Relevance of Signal Transduction Pathway Mutations

1. *TLX3*-rearranged T-ALL is associated with strongly activating *NOTCH1* mutations whereas the incidence of NOTCH-activating mutations is reduced in *TAL1*- or *LMO2*-rearranged pediatric T-ALL patients. (*this thesis*)
2. Some seemingly wild-type *NOTCH1* or *FBXW7* T-ALL patients express an activated NOTCH1-driven expression signature, implying that not all NOTCH1-activating mutational mechanisms have been revealed yet. (*this thesis*)
3. *PTEN/AKT* mutations and NOTCH1-activating mutations are nearly mutual exclusive genetic events in pediatric T-ALL that are each associated with specific T-ALL genetic subtypes. (*this thesis*)
4. Micro-deletions represent a novel, RAG-mediated PTEN inactivational event in pediatric T-cell leukemic cells and thymocytes of healthy individuals. (*this thesis*)
5. The immature T-ALL subtype as identified by unsupervised gene expression profiling analysis, is strongly predicted by the early T-cell precursor ALL gene signature and thus both reflect a single disease entity. (*this thesis*)
6. A uniform prognostic factor for pediatric T-ALL has not been identified yet.
7. RAG activity is associated with a developing immune system in children and thus may explain the increased incidence of lymphatic leukemia in children compared to adults.
8. Biologists typically concentrate on fold change, statisticians on *p*-value.
9. Great discoveries and improvements invariably involve the cooperation of many minds. (*Alexander Graham Bell*)
10. Don't let perfection get in the way of progress. (*Tony Bombardino*)
11. Er zijn meer volwassenen bang in het licht, dan kinderen in het donker. (*van Kooten en de Bie*)