Decision Models to Compare Treatments in Older Patients with AML

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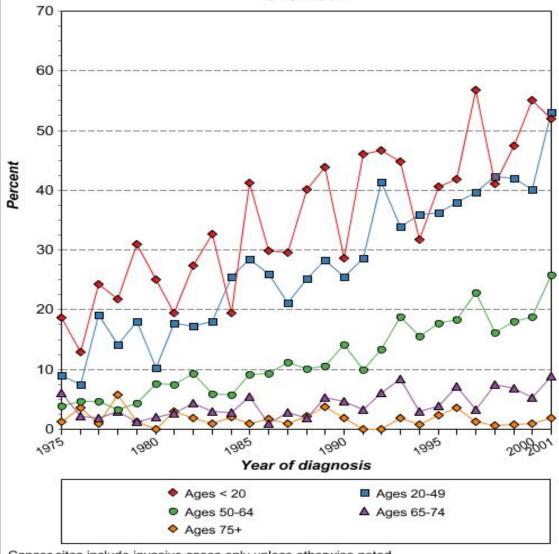
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Jeff Lancet

Statistician: Jongphil Kim



5-Year Relative Survival Rates By Year Dx By Age At Diagnosis/Death Acute Myeloid Leukemia, All Races, Both Sexes 1975-2001



Cancer sites include invasive cases only unless otherwise noted.

Survival source: SEER 9 areas (San Francisco, Connecticut, Detroit, Hawaii, Iowa, New Mexico, Seattle, Utah, and Atlanta).

Survival rates are relative rates expressed as percents. The 5-year survival estimates are calculated using monthly intervals.



Our hypothesis

- Although part of the lack of progress in the elderly is due to changes in AML biology and unavailability of transplant, a significant part is due to suboptimal use of currently available treatments
- Progress could be made if the available treatments were better applied. Data suggest a potential to triple the 2-year survival.

Swedish data

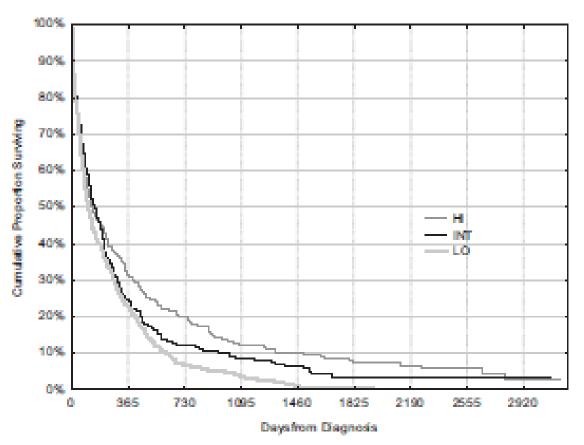
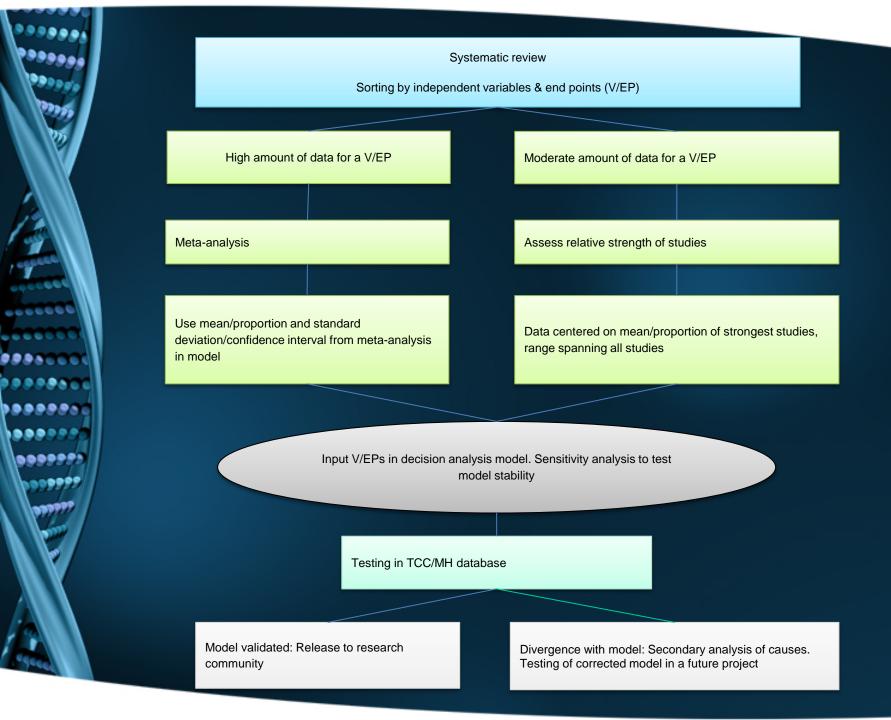


Figure 7. Overall survival of all patients, treated and untreated, 70 to 79 years of age according to geographic region, with different proportions of patients given intensive therapy (Table 4).



The questions

- Although "old" is defined in the AML world as age >60, what data do we have on treating AML patients age 70 and above?
- What choices should be made in patients with poor PS or high comorbidity?
- How can we provide objective prognostic estimates to help physician and patient decisionmaking?





Step I: Systematic review (Ben Djulb.)

Systematic review

Sorting by independent variables & end points (V/EP)

High amount of data for a V/EP

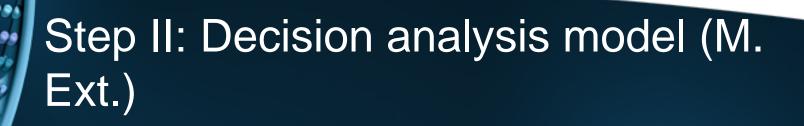
Meta-analysis

Use mean/proportion and standard deviation/confidence interval from meta-analysis in model

Moderate amount of data for a V/EP

Assess relative strength of studies

Data centered on mean/proportion of strongest studies, range spanning all studies



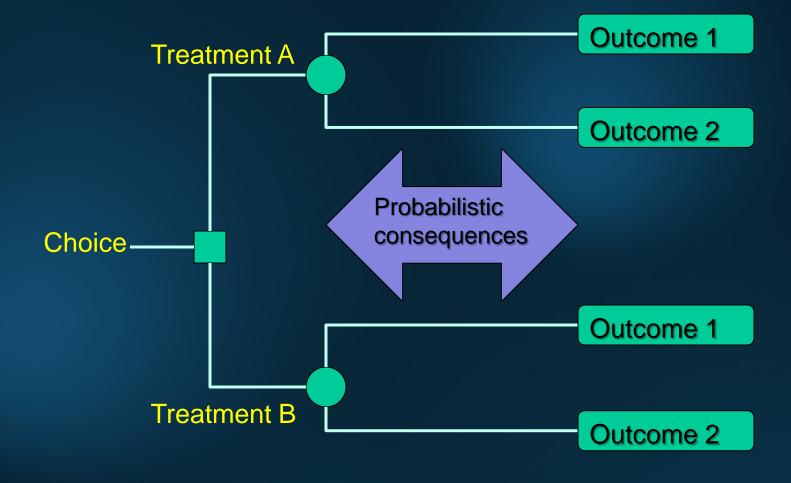
Input V/EPs in decision analysis model. Sensitivity analysis to test model stability



Variables included

- Cytogenetics (3 categories)
- Age
- Comorbidity (Charlson /Sorror)
- Functional status (ECOG/ Karnofsky PS)
- 4 treatment options:
 - Intensive chemotherapy
 - Low-dose chemotherapy
 - Hypomethylating agents
 - Supportive care

Decision model



General structure of a microsimulation model

Decision model outline Initial active disease T0 CR1 Resistant Dead (1st disease month) CR1 Relapse Resistant Dead T1 disease 000000 CR1 CR2 Resistant Relapse Dead disease T2 CR1 CR2 Resistant Dead Relapse disease



Decision models

- Primary: 1-year survival
- 2-year survival
- 30d mortality vs 1y and 2y survival
- Regret model



Combined Therapy:

Adjuvant! Online Decision making tools for health care professionals Adjuvant! for Breast Cancer (Version 8.0)

Patient Infor	mation		
Age:	72	No additional therapy:	
Comorbidity:	Major Prob. (+10)		
ER Status:	Positive -	18.1 alive in 10 years.	
	2 12	29.9 die of cance	er.
Tumor Grade: Grade 3		52.0 die of other causes.	
Tumor Size:	2.1 - 3.0 cm	With hormonal the	rapy: Benefit = 4.4 alive.
Positive Nodes:	1-3		Sa Partie
Calculate For:	Mortality -	With chemotherapy	: Benefit = 5.7 alive.
10 Year Risk:	45 Prognostic		
Adjuvant The	rapy Effectiveness	With combined the	rapy: Benefit = 8.9 alive.
Horm: Arome	atase Inhibitor for 5 yrs		
Chemo: 3rd G	eneration Regimens		
Hormonal Therap	ру: 32	Print Results PDF	Access Help and Clinical
Chemotherapy:	40	In	nages for Consultations



Step III: Test in Transmed database (Jeff Lancet)

Testing in TCC/MH database

Model validated: Release to research community

Divergence with model: Secondary analysis of causes. Testing of corrected model in a future project



Model testing

- A critical step that is too rarely done with prognostic/predictive models
- We will use the TCC database: estimated 900 Moffitt patients aged 70+ at time of analysis



The goal

