## Precision medicine estimates

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## **Notes**

- 1. In Tables 1-4, the models estimated in columns 4 and 8 are the same as those estimated in 3 and 7, respectively, the key difference being (4) and (8) are estimated with standard errors clustered at the level of most common ICD-9 chapter.
- 2. Columns listed as All include all Phase I-III trials since 1995 while Recent include only trials since 2005.
- 3. Marginal effects cannot be reported for interaction terms but are reported for models including interactions.
- 4. Start year fixed effects are estimated in Table 5-6 Columns 1-3, but are not reported because of mechanical relationship with the outcome variable (trial duration). Column 4 reports the linear trend in duration by trial start year, but this is also mechanically linked to duration in that all observed trials have an upper bound on their observed actual end date (the date data was collected). Therefore recent trials with observed durations will be shorter on average than trials that began earlier.
  - Some trials have estimated dates after the date of data collection, but the meaning of these dates is somewhat unclear. Table 5 includes trials with actual or estimated end dates while Table 6 subsets to only actual end dates.
  - Removing estimated-date trials does not seem to radically change the pattern of results, but R-squared does increase substantially indicating much of the unexplained variation in duration may be due to measurement error.

Table 1: Logit Coefficients Dependent variable: Generous PPM

	(1) All b/se	(2) All b/se	(3) All b/se	(4) All (clustered SE) b/se	(5) Recent b/se	(6) Recent b/se	(7) Recent b/se	(8) Recent (clustered SE) b/se
Trial start year	0.068***	0.068***	0.067***	0.069***	0.102***	0.102***	0.102***	0.104***
	(0.003)	(0.003)	(0.003)	(0.004)	(0.005)	(0.005)	(0.005)	(0.006)
Phase 2 Clinical (includes Phase 2/Phase 3 trials)	0.279***		0.268***	0.275**	0.328***		0.314***	0.322***
Di a di i i	(0.033)	0. 400 distrib	(0.033)	(0.089)	(0.036)	0. 40 <b>5</b> de la	(0.036)	(0.090)
Phase 3 Clinical	0.405***	0.403***	0.383***	0.397*	0.440***	0.437***	0.417***	0.430*
	(0.047)	(0.047)	(0.047)	(0.182)	(0.051)	(0.051)	(0.051)	(0.182)
Trial site in US=1	0.139***	0.136***	0.711***	0.727***	0.149***	0.147***	0.761***	0.776***
	(0.031)	(0.032)	(0.066)	(0.114)	(0.033)	(0.033)	(0.069)	(0.127)
Drug indication for neoplasm	2.745***	2.739***	3.101***	3.120***	2.794***	2.787***	3.170***	3.188***
	(0.038)	(0.039)	(0.055)	(0.119)	(0.040)	(0.041)	(0.058)	(0.124)
Received NIH funding	-0.151*	-0.151*	-0.104	-0.104***	-0.192**	-0.193**	-0.136*	-0.134***
	(0.063)	(0.063)	(0.063)	(0.015)	(0.070)	(0.070)	(0.069)	(0.019)
Biomarker type: genomic	4.713***	4.713***	4.728***	4.727***	4.735***	4.734***	4.754***	4.757***
	(0.068)	(0.068)	(0.068)	(0.279)	(0.075)	(0.075)	(0.075)	(0.317)
Phase 2 only		0.285***				0.335***		
		(0.034)				(0.036)		
Phase 2/3 Clinical		0.143				0.162		
		(0.105)				(0.111)		
Trial site in US=1				. = . =			. =	0.004111
× Drug indication for neoplasm=1			-0.732*** (0.074)	-0.745*** (0.113)			-0.789*** (0.078)	-0.801*** (0.129)
Constant	-143.869***	-143.910***	-143.462***	-145.976***	-212.276***	-212.390***	-213.810***	-217.810***
	(7.030)	(7.030)	(7.021)	(7.894)	(10.107)	(10.108)	(10.104)	(12.564)
N	113410	113410	113410	106626	96254	96254	96254	90650
p<0.05, ** p<0.01, *** p<0.001								

Table 2: Logit Marginal Effects Dependent variable: Generous PPM

	(1)	(2)	(3)	(4)	(5)	(6)	(7)	(8)
	All	All	All	All (clustered SE)	Recent	Recent	Recent	Recent (clustered SE)
	b/se	b/se	b/se	b/se	b/se	b/se	b/se	b/se
Trial start year	0.0027***	0.0027***	0.0027***	0.0029***	0.0042***	0.0042***	0.0042***	0.0045***
Trial start year	(0.0027)	(0.0027)	(0.0027)	(0.0029	(0.0042)	(0.0042)	(0.0042)	
Phase 2 Clinical	(0.0001)	(0.0001)	(0.0001)	(0.0002)	(0.0002)	(0.0002)	(0.0002)	(0.0003)
(includes Phase 2/Phase 3 trials)	0.0111***		0.0107***	0.0115**	0.0135***		0.0129***	0.0138***
	(0.0013)		(0.0013)	(0.0035)	(0.0015)		(0.0015)	(0.0036)
Phase 3 Clinical	0.0162***	0.0161***	0.0153***	0.0166*	0.0181***	0.0180***	0.0171***	0.0185*
	(0.0019)	(0.0019)	(0.0019)	(0.0073)	(0.0021)	(0.0021)	(0.0021)	(0.0074)
Trial site in US=1	0.0056***	0.0055***	0.0054***	0.0056***	0.0062***	0.0060***	0.0061***	0.0062***
	(0.0013)	(0.0013)	(0.0013)	(0.0016)	(0.0014)	(0.0014)	(0.0014)	(0.0018)
Drug indication for neoplasm	0.1096***	0.1093***	0.1173***	0.1207***	0.1149***	0.1146***	0.1259***	0.1294***
	(0.0011)	(0.0011)	(0.0013)	(0.0030)	(0.0012)	(0.0012)	0.1259***	0.1294***
Received NIH funding	-0.0060*	-0.0060*	-0.0042	-0.0043***	-0.0079**	-0.0079**	-0.0056*	-0.0058***
	(0.0025)	(0.0025)	(0.0025)	(0.0006)	(0.0029)	(0.0029)	(0.0028)	(0.0008)
Biomarker type: genomic	0.1881***	0.1881***	0.1884***	0.1976***	0.1947***	0.1946***	0.1951***	0.2043***
	(0.0025)	(0.0025)	(0.0025)	(0.0102)	(0.0028)	(0.0028)	(0.0028)	(0.0117)
Phase 2 only		0.0114***				0.0138***		
		(0.0013)				(0.0015)		
Phase 2/3 Clinical		0.0057				0.0067		
		(0.0042)				(0.0046)		
N	113410	113410	113410	106626	96254	96254	96254	90650
p<0.05, ** p<0.01, *** p<0.001								

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Table 3: Logit Coefficients Dependent variable: Restrictive PPM

	(1) All b/se	(2) All b/se	(3) All b/se	(4) All (clustered SE) b/se	(5) Recent b/se	(6) Recent b/se	(7) Recent b/se	(8) Recent (clustered SE) b/se
Trial start year	0.062***	0.062***	0.062***	0.064***	0.093***	0.093***	0.093***	0.095***
	(0.004)	(0.004)	(0.004)	(0.004)	(0.005)	(0.005)	(0.005)	(0.005)
Phase 2 Clinical (includes Phase 2/Phase 3 trials)	0.351***		0.343***	0.355***	0.397***		0.387***	0.401***
Phase 3 Clinical	(0.035) 0.589***	0.586***	(0.035) 0.573***	(0.058) 0.595***	(0.037) 0.621***	0.617***	(0.037) 0.605***	(0.058) 0.626***
Phase 5 Chinical	(0.050)	(0.050)	(0.050)	(0.110)	(0.054)	(0.054)	(0.054)	(0.114)
Trial site in US=1	0.076*	0.073*	0.600***	0.610***	0.073*	0.070*	0.638***	0.642***
111ai site iii 05–1	(0.033)	(0.033)	(0.081)	(0.114)	(0.035)	(0.035)	(0.086)	(0.130)
Drug indication for neoplasm	3.137***	3.129***	3.429***	3.453***	3.195***	3.185***	3.502***	3.521***
Brag maleation for neoplasm	(0.045)	(0.046)	(0.064)	(0.096)	(0.048)	(0.048)	(0.068)	(0.099)
Received NIH funding	-0.168**	-0.169**	-0.138*	-0.132***	-0.181*	-0.182*	-0.145*	-0.141***
10001/00 1 (111 funding	(0.065)	(0.065)	(0.065)	(0.018)	(0.071)	(0.071)	(0.071)	(0.017)
Biomarker type: genomic	4.716***	4.715***	4.725***	4.725***	4.750***	4.749***	4.762***	4.766***
Phase 2 only	(0.073)	(0.073) 0.358*** (0.035)	(0.073)	(0.247)	(0.081)	(0.081) 0.405*** (0.037)	(0.081)	(0.271)
Phase 2/3 Clinical		0.151 (0.117)				0.163 (0.124)		
Trial site in US=1 ×Drug indication for neoplasm=1		(0117)	-0.621***	-0.628***		(0.12.)	-0.672***	-0.673***
Constant	-133.586*** (7.274)	-133.665*** (7.275)	(0.088) -133.397*** (7.268)	(0.111) -136.362*** (8.352)	-195.002*** (10.493)	-195.187*** (10.495)	(0.093) -196.035*** (10.491)	(0.130) -199.952*** (11.274)
N p<0.05, ** p<0.01, *** p<0.001	113410	113410	113410	106626	96254	96254	96254	90650

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Table 4: Logit Marginal Effects Dependent variable: Generous PPM

	(1)	(2)	(3)	(4)	(5)	(6)	(7)	(8)
	All	All	All	All (clustered SE)	Recent	Recent	Recent	Recent (clustered SE)
	b/se	b/se	b/se	b/se	b/se	b/se	b/se	b/se
Total start seem	0.0022***	0.0022***	0.0022***	0.0025***	0.0025***	0.0025***	0.0025***	0.0020***
Trial start year	0.0023***	0.0023***	0.0023***	0.0025***	0.0035***	0.0035***	0.0035***	0.0038***
Dhasa 2 Clinical	(0.0001)	(0.0001)	(0.0001)	(0.0002)	(0.0002)	(0.0002)	(0.0002)	(0.0002)
Phase 2 Clinical (includes Phase 2/Phase 3 trials)	0.0129***		0.0126***	0.0137***	0.0150***		0.0146***	0.0159***
,	(0.0013)		(0.0013)	(0.0021)	(0.0014)		(0.0014)	(0.0021)
Phase 3 Clinical	0.0216***	0.0215***	0.0211***	0.0230***	0.0234***	0.0233***	0.0228***	0.0248***
	(0.0018)	(0.0018)	(0.0018)	(0.0040)	(0.0020)	(0.0020)	(0.0020)	(0.0042)
Trial site in US=1	0.0028*	0.0027*	0.0027*	0.0028**	0.0028*	0.0026*	0.0028*	0.0028*
	(0.0012)	(0.0012)	(0.0012)	(0.0009)	(0.0013)	(0.0013)	(0.0013)	(0.0011)
Drug indication for neoplasm	0.1153***	0.1150***	0.1167***	0.1202***	0.1207***	0.1203***	0.1252***	0.1288***
	(0.0013)	(0.0013)	(0.0013)	(0.0016)	(0.0013)	(0.0013)	(0.0014)	(0.0018)
Received NIH funding	-0.0062**	-0.0062**	-0.0051*	-0.0051***	-0.0068*	-0.0069*	-0.0055*	-0.0056***
_	(0.0024)	(0.0024)	(0.0024)	(0.0007)	(0.0027)	(0.0027)	(0.0027)	(0.0007)
Biomarker type: genomic	0.1734***	0.1733***	0.1736***	0.1825***	0.1794***	0.1794***	0.1797***	0.1889***
71 0	(0.0025)	(0.0025)	(0.0025)	(0.0076)	(0.0028)	(0.0028)	(0.0028)	(0.0086)
Phase 2 only		0.0132***				0.0153***		
•		(0.0013)				(0.0014)		
Phase 2/3 Clinical		0.0055				0.0062		
		(0.0043)				(0.0047)		
N	113410	113410	113410	106626	96254	96254	96254	90650
p<0.05, ** p<0.01, *** p<0.001								

Table 5: OLS regression
Dependent variable: Duration in months (winsorized to remove extreme outliers)
Trials ending after 2005 with actual or estimated end dates

	(1)	(2)	(3)	(4)
	All trials b/se	US trials b/se	US cand b/se	b/se
Start year control	Fixed Effects	Fixed Effects	Fixed Effects	Linear
Phase 2 Clinical (includes Phase 2/Phase 3 trials)	7.473***	6.195***	4.870***	4.225***
,	(0.208)	(0.322)	(0.533)	(0.582)
Phase 3 Clinical	11.283***	7.989***	14.850***	16.634***
	(0.245)	(0.404)	(0.947)	(1.025)
Trial site in US	5.398***			
Days indication for magnicum	(0.184) 22.541***	20.567***		
Drug indication for neoplasm	(0.209)	(0.326)		
Received NIH funding	12.733***	12.745***	8.527***	7.352***
Treceived Till Tallaling	(0.529)	(0.595)	(0.882)	(0.959)
Generous PPM	8.751***	9.805***	9.203***	7.804***
	(0.357)	(0.516)	(0.626)	(0.938)
Biomarker role (detailed): diagnosis				2.035
				(1.320)
Biomarker role (detailed): differential diagnosis				4.076***
Diamontan note (detailed), mustisting during registernes				(1.052) -2.948*
Biomarker role (detailed): predicting drug resistance				-2.948** (1.149)
Biomarker role (detailed): predicting treatment efficacy				-0.452
Biomarker role (detailed): predicting treatment emeters				(1.351)
Biomarker role (detailed): predicting treatment toxicity				0.952
				(1.735)
Biomarker role (detailed): screening				1.312
P'and and a (144'lal) and a'm Condition				(1.466)
Biomarker role (detailed): selection for therapy				-4.031*** (1.148)
Biomarker role (detailed): disease profiling				2.656*
promiser rote (detailed), discuss promise				(1.055)
Biomarker role (detailed): monitoring disease progression				-0.286
				(1.336)
Biomarker role (detailed): monitoring treatment efficacy				2.237
Diamandan mala (datailad), manufaning turaturant taninita				(1.176)
Biomarker role (detailed): monitoring treatment toxicity				1.193 (2.226)
Biomarker role (detailed): not determined				3.795
Biomarker fore (detailed). Not determined				(2.091)
Biomarker role (detailed): prognosis				1.198
				(1.241)
Biomarker role (detailed): prognosis - risk stratification				2.960*
D' 1 1 (1 ( '1 1) 1 1 C (				(1.408)
Biomarker role (detailed): risk factor				0.523 (1.084)
Biomarker role (detailed): staging				-4.272***
Diomarker fore (detailed). staging				(1.175)
Biomarker role (detailed): toxicity profiling				-3.957
, , , ,				(11.956)
Constant	114.556***	119.542***	134.880***	6319.361***
N.	(3.781)	(4.591)	(5.587)	(123.376)
N 2	80334	38668	14663	13208
r2 p<0.05, ** p<0.01, *** p<0.001	0.352	0.327	0.245	0.207
p \ 0.00, p \ 0.01, p \ 0.001				

Table 6: OLS regression Dependent variable: Duration in months (winsorized to remove extreme outliers) Trials ending after 2005 with *actual* end dates

	(1) All trials	(2) US trials		(4) cer trials
	b/se	b/se	b/se	b/se
Start year control Phase 2 Clinical (includes Phase 2/Phase 3 trials)	Fixed Effects 5.800***	Fixed Effects 4.469***	Fixed Effects 2.182***	Linear 1.274*
Thase 2 chinear (merades 1 hase 2/1 hase 3 trials)	(0.213)	(0.315)	(0.553)	(0.603)
Phase 3 Clinical	8.189***	5.663***	8.528***	9.224***
	(0.248)	(0.386)	(1.025)	(1.113)
Trial site in US	3.467***			
Drug indication for neoplasm	(0.186) 16.740***	15.473***		
Drug indication for heophasin	(0.226)	(0.327)		
Received NIH funding	11.672***	11.555***	8.388***	7.017***
Ç	(0.520)	(0.579)	(0.885)	(0.959)
Generous PPM	7.512***	7.719***	7.890***	5.066***
D' 1 (1 (1 (1 (1 (1 (1 (1 (1 (1 (1 (1 (1 (	(0.418)	(0.573)	(0.716)	(1.033)
Biomarker role (detailed): diagnosis				2.654 (1.410)
Biomarker role (detailed): differential diagnosis				3.968***
Biomarker rote (detailed). differential diagnosis				(1.178)
Biomarker role (detailed): predicting drug resistance				-1.745
				(1.281)
Biomarker role (detailed): predicting treatment efficacy				3.832**
Discussion and a (detailed), and distinct the standard to sixty				(1.429)
Biomarker role (detailed): predicting treatment toxicity				0.493 (1.948)
Biomarker role (detailed): screening				1.826
<i>5</i>				(1.641)
Biomarker role (detailed): selection for therapy				-0.348
				(1.299)
Biomarker role (detailed): disease profiling				1.117
Biomarker role (detailed): monitoring disease progression				(1.146) -1.813
biomarker fore (detailed). Monitoring disease progression				(1.482)
Biomarker role (detailed): monitoring treatment efficacy				1.802
				(1.242)
Biomarker role (detailed): monitoring treatment toxicity				-2.811
P'1 (14.71-1)1 (14.71-1)				(2.515)
Biomarker role (detailed): not determined				-1.689 (2.463)
Biomarker role (detailed): prognosis				0.472
1 18				(1.329)
Biomarker role (detailed): prognosis - risk stratification				0.808
				(1.687)
Biomarker role (detailed): risk factor				-0.551
Biomarker role (detailed): staging				(1.176) -3.929**
Diomarker role (detailed). Suiging				(1.320)
Biomarker role (detailed): toxicity profiling				7.927
				(10.477)
Constant	122.058***	125.346***	137.731***	11021.899***
N	(3.195) 49591	(3.843) 25898	(4.846) 8867	(149.589) 8020
r2	0.507	0.502	0.461	0.432
p<0.05, ** p<0.01, *** p<0.001	0.507	0.502	0.101	0.152
1 , 1 , 1 , 1 , 1 , 1 , 1 , 1 , 1 , 1 ,				