

Supplementary Results

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Table S1: Risk group level cost and effects (discounted at 5%) of 25 screening scenarios and a scenario without screening, per 1,000 simulated 40-year-olds, assuming perfect adherence

a. Effects of screening for the *very low* risk group

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^{ab}
Test	Start Age	Interval									
No Screening			0	36	0.03	36	12	17,910	17,899	531,056	
FIT	60	3	4,087	294	0.11	28	8	17,917	17,908	841,686	Dominated
FIT	60	2	6,110	385	0.13	26	7	17,919	17,909	881,028	Dominated
FIT	54	3	5,661	370	0.14	27	7	17,920	17,911	915,641	31,995
FIT	50	3	7,061	428	0.15	26	7	17,921	17,912	989,979	Dominated
FIT	54	2	8,219	482	0.16	25	6	17,921	17,913	981,909	33,639
FIT	60	1	10,181	559	0.17	24	6	17,920	17,911	983,128	Dominated
FIT	46	3	7,835	460	0.15	26	7	17,922	17,913	1,075,629	Dominated
FIT	50	2	9,538	538	0.17	25	6	17,923	17,914	1,075,858	63,911
FIT	46	2	10,778	588	0.18	25	6	17,923	17,915	1,192,966	Dominated
FIT	54	1	13,384	701	0.20	22	6	17,923	17,915	1,149,747	86,929
FIT	40	3	9,205	511	0.16	27	7	17,922	17,913	1,243,293	Dominated
FIT	50	1	15,233	778	0.22	22	6	17,925	17,917	1,300,489	96,014
FIT	40	2	12,485	649	0.19	25	7	17,924	17,915	1,417,130	Dominated
FIT	46	1	16,858	841	0.24	23	6	17,925	17,917	1,484,265	226,884
COL	60	10	0	1,978	0.34	18	5	17,922	17,913	1,527,701	Dominated
FIT	40	1	18,889	909	0.24	24	7	17,925	17,917	1,833,001	Dominated
COL	60	5	0	2,868	0.52	16	4	17,923	17,913	1,923,484	Dominated
COL	54	10	0	2,808	0.52	17	4	17,925	17,916	2,090,952	Dominated
COL	50	10	0	2,991	0.55	16	4	17,927	17,917	2,480,460	Dominated
COL	46	10	0	3,069	0.66	17	4	17,928	17,918	2,939,036	Dominated
COL	54	5	0	4,556	0.80	13	3	17,927	17,917	2,958,508	Dominated
COL	50	5	0	4,828	0.84	13	3	17,929	17,919	3,582,891	1,793,697
COL	40	10	0	4,003	0.70	15	4	17,929	17,918	4,102,124	Dominated
COL	46	5	0	5,681	0.97	13	3	17,930	17,919	4,577,339	3,977,790
COL	40	5	0	6,824	1.14	12	3	17,932	17,918	6,435,949	Dominated

b. Effects of screening for the *low risk group*

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^{ab}
Test	Start Age	Interval									
No Screening			0	57	0.05	57	20	17,893	17,876	840,912	
FIT	60	3	4,042	372	0.17	45	13	17,905	17,890	1,124,633	Dominated
FIT	60	2	6,037	471	0.19	42	11	17,907	17,892	1,153,974	19,124
FIT	54	3	5,625	455	0.20	44	12	17,909	17,894	1,195,225	Dominated
FIT	60	1	10,097	658	0.25	38	10	17,909	17,895	1,242,543	Dominated
FIT	54	2	8,175	577	0.23	40	11	17,911	17,897	1,249,685	19,258
FIT	50	3	7,035	515	0.22	43	12	17,911	17,897	1,266,575	Dominated
FIT	50	2	9,519	636	0.24	40	11	17,913	17,900	1,343,028	Dominated
FIT	46	3	7,821	549	0.21	43	12	17,911	17,898	1,353,677	Dominated
FIT	54	1	13,410	815	0.29	36	9	17,914	17,901	1,404,854	39,685
FIT	46	2	10,784	687	0.26	40	11	17,914	17,901	1,461,318	Dominated
FIT	40	3	9,203	597	0.22	44	12	17,912	17,898	1,524,008	Dominated
FIT	50	1	15,338	897	0.32	36	9	17,916	17,904	1,556,566	63,213
FIT	40	2	12,517	746	0.26	41	11	17,915	17,902	1,689,545	Dominated
FIT	46	1	17,029	960	0.33	37	10	17,917	17,905	1,746,359	182,494
COL	60	10	0	2,087	0.46	29	8	17,912	17,899	1,751,467	Dominated
FIT	40	1	19,117	1,022	0.33	39	11	17,917	17,904	2,104,788	Dominated
COL	60	5	0	2,961	0.66	26	7	17,914	17,900	2,116,045	Dominated
COL	54	10	0	2,858	0.68	26	7	17,917	17,905	2,284,883	Dominated
COL	50	10	0	3,113	0.71	25	6	17,920	17,907	2,688,824	Dominated
COL	54	5	0	4,549	0.96	21	5	17,920	17,907	3,086,369	Dominated
COL	46	10	0	3,197	0.82	27	7	17,921	17,908	3,155,140	Dominated
COL	50	5	0	4,923	1.02	20	5	17,923	17,910	3,726,554	370,130
COL	40	10	0	4,129	0.88	24	6	17,924	17,909	4,306,323	Dominated
COL	46	5	0	5,688	1.14	20	5	17,926	17,912	4,686,894	695,898
COL	40	5	0	6,920	1.33	18	4	17,929	17,912	6,555,452	2,053,361

c. Effects of screening for the *average risk group*

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^{ab}
Test	Start Age	Interval									
No Screening			0	83	0.07	83	28	17,873	17,848	1,211,915	
FIT	60	3	3,988	464	0.24	65	18	17,889	17,868	1,462,876	Dominated
FIT	60	2	5,948	573	0.27	61	16	17,892	17,871	1,480,195	11,509
FIT	54	3	5,580	556	0.27	64	18	17,894	17,874	1,527,861	Dominated
FIT	60	1	9,987	776	0.34	56	14	17,896	17,876	1,552,752	Dominated
FIT	54	2	8,119	690	0.31	59	16	17,898	17,879	1,568,124	12,402
FIT	50	3	7,001	619	0.30	62	17	17,897	17,877	1,595,055	Dominated
FIT	50	2	9,492	753	0.33	59	15	17,901	17,882	1,661,451	Dominated
FIT	46	3	7,800	654	0.29	63	17	17,899	17,879	1,685,377	Dominated
FIT	54	1	13,427	950	0.40	53	14	17,902	17,884	1,707,549	24,290
FIT	46	2	10,786	805	0.35	59	16	17,902	17,884	1,781,135	Dominated
FIT	40	3	9,197	701	0.29	64	18	17,899	17,880	1,858,394	Dominated
FIT	50	1	15,446	1,037	0.42	53	14	17,905	17,888	1,862,042	45,573
FIT	40	2	12,549	861	0.35	60	16	17,903	17,884	2,013,880	Dominated
COL	60	10	0	2,210	0.61	42	11	17,900	17,881	2,017,744	Dominated
FIT	46	1	17,217	1,103	0.44	54	14	17,907	17,889	2,057,988	128,069
COL	60	5	0	3,065	0.82	37	9	17,902	17,884	2,345,635	Dominated
FIT	40	1	19,373	1,158	0.43	57	16	17,906	17,888	2,427,946	Dominated
COL	54	10	0	2,920	0.86	38	10	17,907	17,890	2,518,294	Dominated
COL	50	10	0	3,253	0.91	37	9	17,911	17,894	2,936,137	Dominated
COL	54	5	0	4,540	1.16	30	7	17,912	17,895	3,241,023	Dominated
COL	46	10	0	3,344	1.00	39	10	17,913	17,895	3,410,330	Dominated
COL	50	5	0	5,029	1.24	29	7	17,916	17,899	3,896,161	180,213
COL	40	10	0	4,275	1.08	35	9	17,916	17,898	4,548,108	Dominated
COL	46	5	0	5,697	1.35	29	7	17,919	17,902	4,819,181	349,629
COL	40	5	0	7,028	1.56	26	6	17,924	17,905	6,696,070	708,260

d. Effects of screening for the *high* risk group

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^{ab}
Test	Start Age	Interval									
No Screening			0	135	0.11	135	46	17,832	17,792	1,975,690	
FIT	60	2	5,749	777	0.42	99	27	17,864	17,830	2,139,982	4,300
FIT	60	3	3,870	649	0.38	106	30	17,859	17,824	2,148,370	Dominated
FIT	60	1	9,707	1,007	0.52	90	23	17,870	17,837	2,178,183	5,141
FIT	54	3	5,474	762	0.43	103	29	17,867	17,834	2,200,583	Dominated
FIT	54	2	7,976	919	0.48	96	25	17,873	17,842	2,211,588	7,751
FIT	50	3	6,913	833	0.46	101	27	17,872	17,839	2,261,428	Dominated
FIT	50	2	9,403	991	0.51	95	25	17,878	17,847	2,303,291	Dominated
FIT	54	1	13,369	1,217	0.61	86	22	17,881	17,851	2,315,132	10,752
FIT	46	3	7,739	870	0.45	102	28	17,874	17,842	2,356,947	Dominated
FIT	46	2	10,752	1,046	0.53	96	25	17,880	17,850	2,427,374	Dominated
FIT	50	1	15,562	1,321	0.65	86	22	17,885	17,857	2,472,317	28,069
FIT	40	3	9,167	914	0.45	103	29	17,875	17,843	2,534,984	Dominated
COL	60	10	0	2,425	0.88	68	18	17,877	17,847	2,551,250	Dominated
FIT	40	2	12,580	1,098	0.53	98	26	17,881	17,851	2,670,407	Dominated
FIT	46	1	17,495	1,391	0.66	87	23	17,887	17,859	2,682,285	Dominated
COL	60	5	0	3,231	1.13	59	15	17,880	17,852	2,806,337	Dominated
COL	54	10	0	3,059	1.22	62	16	17,888	17,861	2,996,730	Dominated
FIT	40	1	19,807	1,436	0.65	92	26	17,886	17,858	3,076,509	Dominated
COL	50	10	0	3,507	1.30	59	15	17,894	17,868	3,430,659	Dominated
COL	54	5	0	4,524	1.53	49	12	17,895	17,870	3,565,696	Dominated
COL	46	10	0	3,623	1.37	63	17	17,897	17,871	3,922,409	Dominated
COL	50	5	0	5,199	1.66	46	11	17,902	17,878	4,235,107	81,838
COL	40	10	0	4,544	1.49	57	14	17,903	17,877	5,033,423	Dominated
COL	46	5	0	5,719	1.76	48	12	17,907	17,883	5,097,948	172,568
COL	40	5	0	7,201	2.03	42	10	17,915	17,890	6,973,676	282,915

e. Effects of screening for the *very high* risk group

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^{ab}
Test	Start Age	Interval									
No Screening			0	222	0.18	222	76	17,756	17,689	3,296,374	Dominated
FIT	60	1	9,016	1,358	0.81	144	37	17,822	17,768	3,209,564	Dominates
FIT	60	2	5,345	1,101	0.68	159	42	17,813	17,756	3,231,453	Dominated
FIT	54	2	7,610	1,292	0.78	152	40	17,829	17,777	3,268,440	Dominated
FIT	60	3	3,638	950	0.62	170	48	17,805	17,746	3,283,787	Dominated
FIT	54	1	12,890	1,638	0.95	136	34	17,841	17,793	3,300,695	3,687
FIT	54	3	5,237	1,107	0.70	165	45	17,819	17,764	3,309,921	Dominated
FIT	50	2	9,101	1,386	0.83	151	39	17,837	17,786	3,355,258	Dominated
FIT	50	3	6,679	1,195	0.75	161	43	17,827	17,774	3,358,728	Dominated
COL	60	10	0	2,657	1.25	112	29	17,832	17,784	3,432,092	Dominated
FIT	50	1	15,306	1,775	1.02	136	35	17,850	17,804	3,455,984	14,182
FIT	46	3	7,548	1,240	0.73	163	44	17,831	17,778	3,461,465	Dominated
FIT	46	2	10,529	1,453	0.86	152	40	17,842	17,793	3,485,059	Dominated
COL	60	5	0	3,352	1.53	99	25	17,837	17,791	3,580,870	Dominated
FIT	40	3	9,027	1,283	0.74	165	45	17,833	17,781	3,655,055	Dominated
FIT	46	1	17,489	1,867	1.05	138	36	17,854	17,809	3,680,416	Dominated
FIT	40	2	12,465	1,506	0.86	156	42	17,844	17,795	3,745,337	Dominated
COL	54	10	0	3,314	1.75	102	26	17,852	17,809	3,810,691	Dominated
FIT	40	1	20,104	1,909	1.03	146	40	17,853	17,808	4,123,967	Dominated
COL	54	5	0	4,510	2.06	84	20	17,863	17,823	4,162,832	38,064
COL	50	10	0	3,835	1.88	97	24	17,862	17,821	4,233,917	Dominated
COL	46	10	0	4,025	1.95	102	26	17,868	17,828	4,751,928	Dominated
COL	50	5	0	5,329	2.28	77	18	17,875	17,839	4,793,549	39,568
COL	46	5	0	5,780	2.40	80	20	17,884	17,848	5,602,818	82,326
COL	40	10	0	4,909	2.13	94	23	17,877	17,838	5,831,509	Dominated
COL	40	5	0	7,334	2.75	70	16	17,897	17,862	7,423,645	135,377

Abbreviations: COL = colonoscopy; CRC = colorectal cancer, FIT = faecal immunological test; QALYs = quality-adjusted life years

Note: details of the risk groups can be found in Table 2 of the manuscript
 Grey shading highlights strategies on the efficient frontier.

- a. Results are discounted at an annual rate of 5%
- b. Costs are presented in Australian Dollars (\$AUD)

Table S2: Costs and effects of all a) uniform screening scenarios and b) personalised screening scenarios per 1,000 simulated 40-year-olds, assuming perfect adherence when costs are discounted at 5% and QALYs are discounted and weighted by age

a. Effects of uniform screening scenarios assuming perfect adherence

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{bc}	ICER ^c
Test	Start Age	Interval									
No Screening			0	84	0.07	84	29	15,176	15,156	1,234,089	
FIT	60	3	3,981	467	0.24	66	19	15,190	15,172	1,240,498	413
FIT	60	2	5,935	576	0.27	62	16	15,192	15,175	1,256,805	5,643
FIT	54	3	5,571	561	0.28	64	18	15,194	15,177	1,304,332	Dominated
FIT	60	1	9,954	777	0.34	56	15	15,195	15,178	1,327,965	Dominated
FIT	54	2	8,101	695	0.32	59	16	15,197	15,181	1,343,651	14,720
FIT	50	3	6,990	625	0.30	63	17	15,196	15,180	1,371,842	Dominated
FIT	50	2	9,473	759	0.34	59	15	15,199	15,183	1,436,505	Dominated
FIT	46	3	7,789	660	0.29	63	17	15,197	15,181	1,462,077	Dominated
FIT	54	1	13,381	953	0.40	53	14	15,200	15,185	1,480,562	29,828
FIT	46	2	10,767	811	0.35	59	16	15,200	15,185	1,556,681	Dominated
FIT	50	1	15,397	1,042	0.43	53	14	15,202	15,188	1,634,262	52,279
FIT	40	3	9,187	707	0.30	64	18	15,198	15,182	1,635,282	Dominated
FIT	40	2	12,532	868	0.36	60	16	15,201	15,186	1,789,931	Dominated
COL	60	10	0	2,198	0.60	42	11	15,198	15,182	1,789,986	Dominated
FIT	46	1	17,171	1,109	0.44	54	14	15,204	15,190	1,830,442	139,134
COL	60	5	0	3,048	0.82	37	10	15,200	15,184	2,117,448	Dominated
FIT	40	1	19,338	1,165	0.44	57	16	15,204	15,189	2,201,540	Dominated
COL	54	10	0	2,928	0.86	39	10	15,204	15,189	2,294,199	Dominated
COL	50	10	0	3,245	0.91	37	9	15,207	15,192	2,706,770	Dominated
COL	54	5	0	4,540	1.16	31	7	15,207	15,193	3,016,912	Dominated
COL	46	10	0	3,341	1.01	39	10	15,208	15,194	3,181,465	Dominated
COL	50	5	0	5,012	1.24	29	7	15,211	15,197	3,664,480	251,583
COL	40	10	0	4,269	1.09	36	9	15,211	15,196	4,319,443	Dominated
COL	46	5	0	5,700	1.35	30	7	15,214	15,199	4,593,188	427,976
COL	40	5	0	7,011	1.57	26	6	15,217	15,201	6,462,019	869,224

b. Effects of personalised screening scenarios assuming perfect adherence^d

Screening Strategy	FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{b,c}	ICER ^c
No Screening	0	84	0.07	84	29	15,176	15,156	1,234,089	
PS01	360	130	0.09	81	27	15,178	15,159	1,467,668	Dominated
PS02	515	141	0.10	81	27	15,179	15,160	1,471,312	Dominated
PS03	1,722	276	0.16	73	23	15,184	15,166	1,505,808	Dominated
PS04	2,554	324	0.18	71	22	15,185	15,167	1,513,829	Dominated
PS05	2,190	305	0.18	72	23	15,186	15,168	1,520,843	Dominated
PS06	3,322	368	0.20	70	22	15,187	15,169	1,542,584	Dominated
PS07	5,226	525	0.27	63	18	15,192	15,175	1,628,434	Dominated
PS08	5,920	562	0.28	63	18	15,193	15,177	1,656,571	Dominated
PS09	6,017	568	0.28	63	18	15,193	15,177	1,662,780	20,620
PS10	7,898	687	0.33	59	16	15,197	15,181	1,756,804	24,788
PS12	10,057	792	0.36	57	15	15,198	15,183	1,834,425	32,204
PS11	10,800	825	0.37	56	15	15,198	15,183	1,837,112	Dominated
PS13	11,189	859	0.38	55	14	15,200	15,185	1,911,354	41,373
PS14	11,700	881	0.39	55	14	15,200	15,185	1,924,610	43,491
PS15	11,788	885	0.39	55	14	15,200	15,186	1,933,583	49,167
PS16	11,089	1,023	0.44	52	13	15,201	15,186	1,978,086	Dominated
PS17	11,398	1,037	0.44	52	13	15,202	15,187	1,999,556	50,356
PS18	12,293	1,078	0.45	51	13	15,202	15,187	2,013,777	51,739
PS19	12,939	1,106	0.46	51	13	15,202	15,188	2,063,215	55,162
PS20	13,382	1,125	0.47	51	13	15,203	15,189	2,098,112	76,583
PS21	13,646	1,136	0.47	51	13	15,203	15,189	2,116,904	78,423
PS22	13,646	1,154	0.47	51	13	15,203	15,189	2,149,259	98,736
PS23	14,052	1,169	0.48	52	14	15,204	15,190	2,193,345	98,758
PS24	10,378	1,969	0.69	43	11	15,206	15,193	2,519,388	108,054
PS25	11,518	2,017	0.70	43	11	15,206	15,193	2,564,321	118,157
PS26	12,085	2,037	0.70	43	11	15,207	15,193	2,627,025	144,858
PS27	12,085	2,100	0.71	43	11	15,207	15,194	2,699,821	168,462
PS28	12,085	2,209	0.74	43	11	15,208	15,195	2,880,990	Dominated
PS29	12,473	2,223	0.74	43	11	15,208	15,195	2,924,646	208,087
PS30	6,964	3,480	0.99	35	9	15,211	15,197	3,512,866	244,752
PS31	7,289	3,492	1.00	35	9	15,211	15,197	3,549,627	264,513
PS32	7,289	3,804	1.05	34	8	15,212	15,199	3,943,469	358,974
PS33	7,289	4,017	1.09	34	8	15,213	15,199	4,238,838	429,957
PS34	3,372	4,929	1.25	30	7	15,214	15,200	4,694,318	522,899
PS35	3,372	5,105	1.28	30	7	15,214	15,200	4,915,213	882,218
PS36	3,372	5,531	1.35	29	7	15,216	15,201	5,515,822	951,617
PS37	3,372	5,814	1.39	29	7	15,216	15,201	5,945,624	3,457,466
PS38	0	6,612	1.51	27	6	15,217	15,201	6,365,416	3,689,986
PS39	0	6,782	1.54	27	6	15,217	15,201	6,564,338	6,542,166

Abbreviations: COL = colonoscopy; CRC = colorectal cancer, FIT = faecal immunological test;

ICER = incremental cost-effectiveness ratio; QALYs = quality-adjusted life years

Grey shading highlights screening scenarios on the efficient frontier.

a. Life years and QALYs are discounted at 5% and weighted by age

b. Costs are discounted at 5%

c. Costs are presented in Australian Dollars (\$AUD)

d. The personalised screening scenarios are described in Table 3

Table S3: Specifics of the personalised screening scenarios when costs are discounted at 5% and QALYs are discounted at 5% and weighted by age^a

Screening Strategy	Risk Groups				
	Very Low	Low	Average	High	Very High
PS1	NoScr	NoScr	NoScr	NoScr	FIT_60_1
PS2	NoScr	NoScr	NoScr	NoScr	FIT_54_1
PS3	NoScr	NoScr	NoScr	FIT_60_2	FIT_54_1
PS4	NoScr	NoScr	NoScr	FIT_60_1	FIT_54_1
PS5	NoScr	NoScr	NoScr	FIT_54_2	FIT_54_1
PS6	NoScr	NoScr	NoScr	FIT_54_1	FIT_54_1
PS7	NoScr	NoScr	FIT_60_2	FIT_54_1	FIT_54_1
PS8	NoScr	NoScr	FIT_54_2	FIT_54_1	FIT_54_1
PS9	NoScr	NoScr	FIT_54_2	FIT_54_1	FIT_50_1
PS10	NoScr	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1
PS11	NoScr	FIT_54_2	FIT_54_1	FIT_54_1	FIT_50_1
PS12	NoScr	FIT_54_2	FIT_54_1	FIT_50_1	FIT_50_1
PS13	FIT_54_3	FIT_54_2	FIT_54_1	FIT_50_1	FIT_50_1
PS14	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1	FIT_50_1
PS15	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1	FIT_46_1
PS16	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1	COL_50_5
PS17	FIT_54_2	FIT_50_2	FIT_54_1	FIT_50_1	COL_50_5
PS18	FIT_54_2	FIT_54_1	FIT_54_1	FIT_50_1	COL_50_5
PS19	FIT_54_2	FIT_54_1	FIT_50_1	FIT_50_1	COL_50_5
PS20	FIT_54_2	FIT_50_1	FIT_50_1	FIT_50_1	COL_50_5
PS21	FIT_50_2	FIT_50_1	FIT_50_1	FIT_50_1	COL_50_5
PS22	FIT_50_2	FIT_50_1	FIT_50_1	FIT_50_1	COL_46_5
PS23	FIT_50_2	FIT_50_1	FIT_50_1	FIT_46_1	COL_46_5
PS24	FIT_50_2	FIT_50_1	FIT_50_1	COL_50_5	COL_46_5
PS25	FIT_50_1	FIT_50_1	FIT_50_1	COL_50_5	COL_46_5
PS26	FIT_50_1	FIT_50_1	FIT_46_1	COL_50_5	COL_46_5
PS27	FIT_50_1	FIT_50_1	FIT_46_1	COL_50_5	COL_40_5
PS28	FIT_50_1	FIT_50_1	FIT_46_1	COL_46_5	COL_40_5
PS29	FIT_50_1	FIT_46_1	FIT_46_1	COL_46_5	COL_40_5
PS30	FIT_50_1	FIT_46_1	COL_50_5	COL_46_5	COL_40_5
PS31	FIT_46_1	FIT_46_1	COL_50_5	COL_46_5	COL_40_5
PS32	FIT_46_1	FIT_46_1	COL_50_5	COL_40_5	COL_40_5
PS33	FIT_46_1	FIT_46_1	COL_46_5	COL_40_5	COL_40_5
PS34	FIT_46_1	COL_50_5	COL_46_5	COL_40_5	COL_40_5
PS35	FIT_46_1	COL_46_5	COL_46_5	COL_40_5	COL_40_5
PS36	FIT_46_1	COL_46_5	COL_40_5	COL_40_5	COL_40_5
PS37	FIT_46_1	COL_40_5	COL_40_5	COL_40_5	COL_40_5
PS38	COL_50_5	COL_40_5	COL_40_5	COL_40_5	COL_40_5
PS39	COL_46_5	COL_40_5	COL_40_5	COL_40_5	COL_40_5

Abbreviations: COL = Colonoscopy; FIT = faecal immunochemical test; NoScr = no screening
Screening strategies: screening test, screening start age, screening interval
a. All screening ends at or before the age of 74 years

Table S4: Costs and effects of all a) uniform screening scenarios and b) personalised screening scenarios per 1,000 simulated 40-year-olds, assuming perfect adherence when costs and QALYs are discounted at 3%

a. Effects of uniform screening scenarios assuming perfect adherence

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^b
Test	Start Age	Interval									
No Screening		0	84	84	0.07	84	29	24,044	23,998	2,131,337	Dominated
FIT	60	3	3,981	467	0.24	66	19	24,081	24,042	2,039,884	Dominates
FIT	60	2	5,935	576	0.27	62	16	24,088	24,050	2,047,061	861
FIT	54	3	5,571	561	0.28	64	18	24,091	24,054	2,108,708	Dominated
FIT	60	1	9,954	777	0.34	56	15	24,095	24,060	2,142,041	Dominated
FIT	54	2	8,101	695	0.32	59	16	24,099	24,064	2,144,612	7,275
FIT	50	3	6,990	625	0.30	63	17	24,096	24,060	2,178,311	Dominated
FIT	50	2	9,473	759	0.34	59	15	24,103	24,069	2,250,511	Dominated
FIT	46	3	7,789	660	0.29	63	17	24,098	24,062	2,279,045	Dominated
FIT	54	1	13,381	953	0.40	53	14	24,108	24,076	2,323,627	15,069
FIT	46	2	10,767	811	0.35	59	16	24,105	24,072	2,381,682	Dominated
FIT	40	3	9,187	707	0.30	64	18	24,098	24,063	2,452,550	Dominated
FIT	50	1	15,397	1,042	0.43	53	14	24,112	24,081	2,501,392	32,204
FIT	40	2	12,532	868	0.36	60	16	24,106	24,072	2,616,871	Dominated
FIT	46	1	17,171	1,109	0.44	54	14	24,113	24,083	2,716,181	Dominated
COL	60	10	0	2,198	0.60	42	11	24,105	24,072	2,815,371	Dominated
FIT	40	1	19,338	1,165	0.44	57	16	24,111	24,079	3,085,943	Dominated
COL	60	5	0	3,048	0.82	37	10	24,110	24,078	3,338,895	Dominated
COL	54	10	0	2,928	0.86	39	10	24,118	24,088	3,455,297	Dominated
COL	50	10	0	3,245	0.91	37	9	24,124	24,095	3,893,169	98,778
COL	46	10	0	3,341	1.01	39	10	24,126	24,098	4,338,784	Dominated
COL	54	5	0	4,540	1.16	31	7	24,127	24,099	4,560,398	Dominated
COL	50	5	0	5,012	1.24	29	7	24,134	24,108	5,259,267	110,258
COL	40	10	0	4,269	1.09	36	9	24,132	24,104	5,491,543	Dominated
COL	46	5	0	5,700	1.35	30	7	24,139	24,112	6,281,315	234,415
COL	40	5	0	7,011	1.57	26	6	24,147	24,119	8,165,431	285,040

b. Effects of personalised screening scenarios assuming perfect adherence^c

Screening Strategy	FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^b
No Screening	0	84	0.07	84	29	24,044	23,998	2,131,337	
PS01	1,568	264	0.16	73	23	24,064	24,022	2,325,151	Dominated
PS02	1,722	276	0.16	73	23	24,066	24,024	2,328,301	Dominated
PS03	2,554	324	0.18	71	22	24,068	24,028	2,334,210	Dominated
PS04	4,457	481	0.25	64	19	24,082	24,044	2,388,587	5,634
PS05	5,226	525	0.27	63	18	24,086	24,049	2,421,113	6,202
PS06	5,920	562	0.28	63	18	24,089	24,053	2,452,809	7,685
PS07	7,309	658	0.32	59	16	24,096	24,061	2,517,012	7,840
PS08	7,406	663	0.32	59	16	24,097	24,062	2,524,438	10,732
PS09	7,898	687	0.33	59	16	24,098	24,064	2,550,682	12,607
PS10	8,715	739	0.34	57	15	24,102	24,068	2,612,964	Dominated
PS11	10,414	822	0.37	55	14	24,104	24,072	2,671,669	15,773
PS12	10,728	837	0.37	55	14	24,105	24,073	2,689,632	16,721
PS13	10,117	947	0.42	53	14	24,107	24,075	2,725,078	19,526
PS14	10,577	968	0.42	53	14	24,108	24,076	2,763,970	21,599
PS15	11,089	991	0.43	52	14	24,109	24,077	2,782,499	22,238
PS16	11,089	1,023	0.44	52	13	24,110	24,078	2,811,605	26,284
PS17	12,293	1,078	0.45	51	13	24,111	24,080	2,860,808	26,934
PS18	12,939	1,106	0.46	51	13	24,112	24,082	2,918,096	34,370
PS19	13,203	1,117	0.46	51	13	24,113	24,082	2,939,743	44,551
PS20	13,646	1,136	0.47	51	13	24,113	24,083	2,979,627	46,083
PS21	10,378	1,595	0.61	46	12	24,117	24,089	3,226,411	46,622
PS22	10,378	1,951	0.68	43	11	24,121	24,093	3,465,361	52,016
PS23	11,148	1,983	0.69	42	11	24,121	24,094	3,491,503	56,483
PS24	11,148	2,001	0.69	43	11	24,122	24,094	3,526,082	60,225
PS25	11,518	2,017	0.70	43	11	24,122	24,095	3,560,357	68,807
PS26	11,518	2,079	0.71	42	11	24,123	24,096	3,633,921	76,512
PS27	6,575	2,788	0.87	37	9	24,127	24,100	4,079,776	96,653
PS28	6,575	3,356	0.97	34	8	24,130	24,104	4,518,470	109,516
PS29	6,575	3,465	0.99	35	9	24,132	24,106	4,711,626	121,339
PS30	6,575	3,776	1.05	34	8	24,134	24,109	5,113,837	149,883
PS31	3,047	4,286	1.14	31	7	24,136	24,110	5,459,786	176,486
PS32	3,372	4,299	1.14	31	7	24,136	24,111	5,498,219	190,456
PS33	3,372	4,715	1.21	30	7	24,138	24,112	5,839,990	199,604
PS34	3,372	4,929	1.25	30	7	24,140	24,114	6,163,155	230,476
PS35	3,372	5,355	1.32	29	7	24,142	24,116	6,770,919	305,621
PS36	3,372	5,531	1.35	29	7	24,143	24,116	7,016,964	410,008
PS37	0	6,328	1.47	27	6	24,144	24,118	7,613,850	511,280
PS38	0	6,612	1.51	27	6	24,146	24,118	8,044,907	557,935
PS39	0	6,782	1.54	27	6	24,146	24,119	8,270,010	1,157,855
PS40	0	7,011	1.57	26	6	24,147	24,119	8,639,531	1,798,841

Abbreviations: COL = colonoscopy; CRC = colorectal cancer, FIT = faecal immunological test;

ICER = incremental cost-effectiveness ratio; QALYs = quality-adjusted life years

Grey shading highlights screening scenarios on the efficient frontier.

a. Results are discounted at 3%

b. Costs are presented in Australian Dollars (\$AUD)

c. The personalised screening scenarios are described in Table 5

Table S5: Specifics of the personalised screening scenarios by willingness-to-pay threshold when cost and QALYs are discounted by 3%^a

Screening Strategy	Risk Groups				
	Very Low	Low	Average	High	Very High
PS1	NoScr	NoScr	NoScr	FIT_60_2	FIT_60_1
PS2	NoScr	NoScr	NoScr	FIT_60_2	FIT_54_1
PS3	NoScr	NoScr	NoScr	FIT_60_1	FIT_54_1
PS4	NoScr	NoScr	FIT_60_2	FIT_60_1	FIT_54_1
PS5	NoScr	NoScr	FIT_60_2	FIT_54_1	FIT_54_1
PS6	NoScr	NoScr	FIT_54_2	FIT_54_1	FIT_54_1
PS7	NoScr	FIT_60_2	FIT_54_2	FIT_54_1	FIT_54_1
PS8	NoScr	FIT_60_2	FIT_54_2	FIT_54_1	FIT_50_1
PS9	NoScr	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1
PS10	FIT_60_74_3	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1
PS11	FIT_60_74_3	FIT_54_2	FIT_54_1	FIT_54_1	FIT_50_1
PS12	FIT_54_3	FIT_54_2	FIT_54_1	FIT_54_1	FIT_50_1
PS13	FIT_54_3	FIT_54_2	FIT_54_1	FIT_54_1	COL_54_5
PS14	FIT_54_3	FIT_54_2	FIT_54_1	FIT_50_1	COL_54_5
PS15	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1	COL_54_5
PS16	FIT_54_2	FIT_54_2	FIT_54_1	FIT_50_1	COL_50_5
PS17	FIT_54_2	FIT_54_1	FIT_54_1	FIT_50_1	COL_50_5
PS18	FIT_54_2	FIT_54_1	FIT_50_1	FIT_50_1	COL_50_5
PS19	FIT_50_2	FIT_54_1	FIT_50_1	FIT_50_1	COL_50_5
PS20	FIT_50_2	FIT_50_1	FIT_50_1	FIT_50_1	COL_50_5
PS21	FIT_50_2	FIT_50_1	FIT_50_1	COL_50_10	COL_50_5
PS22	FIT_50_2	FIT_50_1	FIT_50_1	COL_50_5	COL_50_5
PS23	FIT_54_1	FIT_50_1	FIT_50_1	COL_50_5	COL_50_5
PS24	FIT_54_1	FIT_50_1	FIT_50_1	COL_50_5	COL_46_5
PS25	FIT_50_1	FIT_50_1	FIT_50_1	COL_50_5	COL_46_5
PS26	FIT_50_1	FIT_50_1	FIT_50_1	COL_50_5	COL_40_5
PS27	FIT_50_1	FIT_50_1	COL_50_10	COL_50_5	COL_40_5
PS28	FIT_50_1	FIT_50_1	COL_50_5	COL_50_5	COL_40_5
PS29	FIT_50_1	FIT_50_1	COL_50_5	COL_46_5	COL_40_5
PS30	FIT_50_1	FIT_50_1	COL_50_5	COL_40_5	COL_40_5
PS31	FIT_50_1	COL_50_10	COL_50_5	COL_40_5	COL_40_5
PS32	FIT_46_1	COL_50_10	COL_50_5	COL_40_5	COL_40_5
PS33	FIT_46_1	COL_50_5	COL_50_5	COL_40_5	COL_40_5
PS34	FIT_46_1	COL_50_5	COL_46_5	COL_40_5	COL_40_5
PS35	FIT_46_1	COL_50_5	COL_40_5	COL_40_5	COL_40_5
PS36	FIT_46_1	COL_46_5	COL_40_5	COL_40_5	COL_40_5
PS37	COL_50_5	COL_46_5	COL_40_5	COL_40_5	COL_40_5
PS38	COL_50_5	COL_40_5	COL_40_5	COL_40_5	COL_40_5
PS39	COL_46_5	COL_40_5	COL_40_5	COL_40_5	COL_40_5
PS40	COL_40_5	COL_40_5	COL_40_5	COL_40_5	COL_40_5

Abbreviations: COL = Colonoscopy; FIT = faecal immunochemical test; NoScr = no screening
Screening strategies: screening test, screening start age, screening interval

a. All screening ends at or before the age of 74 years

Table S6: Costs and effects of all a) uniform screening scenarios and b) personalised screening scenarios per 1,000 simulated 40-year-olds, assuming perfect adherence when costs and QALYs are undiscounted

a. Effects of uniform screening scenarios assuming perfect adherence

Screening Strategy			FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^a	Total QALYs ^a	Total Costs ^{ab}	ICER ^b
Test	Start Age	Interval									
No Screening			0	84	0.07	84	29	43,001	42,869	5,460,172	Dominated
FIT	60	2	5,935	576	0.27	62	16	43,149	43,049	4,739,664	Dominates
FIT	60	3	3,981	467	0.24	66	19	43,126	43,020	4,817,049	Dominated
FIT	54	2	8,101	695	0.32	59	16	43,176	43,084	4,850,988	3,179
FIT	60	1	9,954	777	0.34	56	15	43,173	43,081	4,852,104	Dominated
FIT	54	3	5,571	561	0.28	64	18	43,150	43,050	4,886,396	Dominated
FIT	50	3	6,990	625	0.30	63	17	43,164	43,067	4,935,602	Dominated
FIT	50	2	9,473	759	0.34	59	15	43,186	43,096	4,987,686	Dominated
FIT	46	3	7,789	660	0.29	63	17	43,165	43,069	5,077,689	Dominated
FIT	54	1	13,381	953	0.40	53	14	43,204	43,121	5,087,843	6,415
FIT	46	2	10,767	811	0.35	59	16	43,189	43,100	5,150,353	Dominated
FIT	40	3	9,187	707	0.30	64	18	43,164	43,067	5,270,930	Dominated
FIT	50	1	15,397	1,042	0.43	53	14	43,212	43,131	5,322,533	22,161
FIT	40	2	12,532	868	0.36	60	16	43,185	43,095	5,416,504	Dominated
FIT	46	1	17,171	1,109	0.44	54	14	43,211	43,130	5,590,545	Dominated
COL	60	10	0	2,198	0.60	42	11	43,205	43,125	5,984,373	Dominated
FIT	40	1	19,338	1,165	0.44	57	16	43,197	43,113	6,003,156	Dominated
COL	54	10	0	2,928	0.86	39	10	43,238	43,166	6,946,819	Dominated
COL	60	5	0	3,048	0.82	37	10	43,222	43,147	7,040,041	Dominated
COL	50	10	0	3,245	0.91	37	9	43,253	43,185	7,349,473	38,029
COL	46	10	0	3,341	1.01	39	10	43,254	43,185	7,669,729	Dominated
COL	40	10	0	4,269	1.09	36	9	43,271	43,205	8,919,959	Dominated
COL	54	5	0	4,540	1.16	31	7	43,268	43,203	9,072,841	Dominated
COL	50	5	0	5,012	1.24	29	7	43,287	43,226	9,720,205	56,975
COL	46	5	0	5,700	1.35	30	7	43,295	43,234	10,917,234	Dominated
COL	40	5	0	7,011	1.57	26	6	43,313	43,255	12,815,034	106,718

b. Effects of personalised screening scenarios assuming perfect adherence^c

Screening Strategy	FITs	Colonoscopies	Complications	CRC Incidence	CRC Mortality	Life Years ^b	Total QALYs ^b	Total Costs ^{bc}	ICER ^b
No Screening	0	84	0.07	84	29	43,001	42,869	5,460,172	Dominated
PS03	6,148	668	0.31	58	15	43,161	43,065	4,919,457	Dominated
PS01	4,871	594	0.29	60	17	43,146	43,047	4,922,589	Dominated
PS02	5,330	617	0.30	60	16	43,149	43,052	4,926,793	Dominated
PS04	6,916	713	0.33	57	15	43,171	43,079	4,959,849	3,026
PS05	7,611	750	0.34	56	15	43,179	43,089	4,995,978	3,359
PS06	7,611	776	0.36	56	15	43,183	43,094	5,015,775	4,277
PS07	8,103	801	0.37	56	15	43,187	43,099	5,049,437	6,193
PS08	9,802	884	0.40	54	14	43,196	43,111	5,128,736	6,850
PS09	10,206	902	0.40	53	14	43,198	43,113	5,147,483	7,753
PS10	10,206	950	0.41	53	14	43,201	43,117	5,177,225	7,789
PS11	10,628	969	0.42	52	14	43,203	43,120	5,210,998	10,596
PS12	10,628	1,002	0.43	52	13	43,205	43,123	5,246,635	12,905
PS13	11,832	1,056	0.44	51	13	43,210	43,129	5,323,387	13,478
PS14	9,025	1,443	0.57	46	12	43,221	43,145	5,562,545	14,690
PS15	9,025	1,537	0.59	46	12	43,226	43,151	5,674,015	17,799
PS16	9,671	1,565	0.60	46	12	43,229	43,155	5,749,938	24,226
PS17	9,935	1,576	0.60	46	12	43,230	43,156	5,776,707	24,681
PS18	9,935	1,932	0.68	43	11	43,241	43,170	6,180,918	27,246
PS19	10,704	1,964	0.68	42	11	43,243	43,172	6,235,349	28,675
PS20	11,148	1,983	0.69	42	11	43,244	43,174	6,284,712	29,170
PS21	11,148	2,063	0.71	42	11	43,247	43,178	6,397,904	32,093
PS22	6,205	2,772	0.86	37	9	43,260	43,195	7,046,014	37,125
PS23	6,575	2,788	0.87	37	9	43,261	43,196	7,086,658	41,179
PS24	6,575	3,356	0.97	34	8	43,272	43,209	7,849,972	56,864
PS25	6,575	3,776	1.05	34	8	43,281	43,220	8,480,538	61,177
PS26	3,047	4,286	1.14	31	7	43,287	43,228	9,020,556	68,106
PS27	3,047	4,703	1.21	30	7	43,292	43,234	9,620,268	98,538
PS28	3,047	5,342	1.32	29	7	43,301	43,243	10,611,955	109,959
PS29	0	5,785	1.38	28	6	43,304	43,246	11,134,728	149,697
PS30	0	5,801	1.40	28	6	43,304	43,247	11,171,639	150,421
PS31	0	6,260	1.47	27	6	43,308	43,251	11,894,768	172,055
PS32	0	6,612	1.51	27	6	43,311	43,253	12,415,828	208,145
PS33	0	7,011	1.57	26	6	43,313	43,255	13,052,084	327,048

Abbreviations: COL = colonoscopy; CRC = colorectal cancer, FIT = faecal immunological test; ICER = incremental cost-effectiveness ratio; QALYs = quality-adjusted life years
 Grey shading highlights screening scenarios on the efficient frontier.

- Results are undiscounted
- Costs are presented in Australian Dollars (\$AUD)
- The personalised screening scenarios are described in Table 7

Table S7: Specifics of the personalised screening scenarios by willingness-to-pay threshold, when costs and QALYs are undiscounted^a

Screening Strategy	Risk Groups				
	Very Low	Low	Average	High	Very High
PS1	NoScr	FIT_60_3	FIT_60_2	FIT_60_1	COL_60_10
PS2	NoScr	FIT_60_2	FIT_60_2	FIT_60_1	COL_60_10
PS3	FIT_60_3	FIT_60_2	FIT_60_2	FIT_60_1	COL_60_10
PS4	FIT_60_3	FIT_60_2	FIT_60_2	FIT_54_1	COL_60_10
PS5	FIT_60_3	FIT_60_2	FIT_54_2	FIT_54_1	COL_60_10
PS6	FIT_60_3	FIT_60_2	FIT_54_2	FIT_54_1	COL_54_10
PS7	FIT_60_3	FIT_54_2	FIT_54_2	FIT_54_1	COL_54_10
PS8	FIT_60_3	FIT_54_2	FIT_54_1	FIT_54_1	COL_54_10
PS9	FIT_60_2	FIT_54_2	FIT_54_1	FIT_54_1	COL_54_10
PS10	FIT_60_2	FIT_54_2	FIT_54_1	FIT_54_1	COL_54_5
PS11	FIT_54_2	FIT_54_2	FIT_54_1	FIT_54_1	COL_54_5
PS12	FIT_54_2	FIT_54_2	FIT_54_1	FIT_54_1	COL_50_5
PS13	FIT_54_2	FIT_54_1	FIT_54_1	FIT_54_1	COL_50_5
PS14	FIT_54_2	FIT_54_1	FIT_54_1	COL_54_10	COL_50_5
PS15	FIT_54_2	FIT_54_1	FIT_54_1	COL_50_10	COL_50_5
PS16	FIT_54_2	FIT_54_1	FIT_50_1	COL_50_10	COL_50_5
PS17	FIT_50_2	FIT_54_1	FIT_50_1	COL_50_10	COL_50_5
PS18	FIT_50_2	FIT_54_1	FIT_50_1	COL_50_5	COL_50_5
PS19	FIT_54_1	FIT_54_1	FIT_50_1	COL_50_5	COL_50_5
PS20	FIT_54_1	FIT_50_1	FIT_50_1	COL_50_5	COL_50_5
PS21	FIT_54_1	FIT_50_1	FIT_50_1	COL_50_5	COL_40_5
PS22	FIT_54_1	FIT_50_1	COL_50_10	COL_50_5	COL_40_5
PS23	FIT_50_1	FIT_50_1	COL_50_10	COL_50_5	COL_40_5
PS24	FIT_50_1	FIT_50_1	COL_50_5	COL_50_5	COL_40_5
PS25	FIT_50_1	FIT_50_1	COL_50_5	COL_40_5	COL_40_5
PS26	FIT_50_1	COL_50_10	COL_50_5	COL_40_5	COL_40_5
PS27	FIT_50_1	COL_50_5	COL_50_5	COL_40_5	COL_40_5
PS28	FIT_50_1	COL_50_5	COL_40_5	COL_40_5	COL_40_5
PS29	COL_50_10	COL_50_5	COL_40_5	COL_40_5	COL_40_5
PS30	COL_46_10	COL_50_5	COL_40_5	COL_40_5	COL_40_5
PS31	COL_46_10	COL_40_5	COL_40_5	COL_40_5	COL_40_5
PS32	COL_50_5	COL_40_5	COL_40_5	COL_40_5	COL_40_5
PS33	COL_40_5	COL_40_5	COL_40_5	COL_40_5	COL_40_5

Abbreviations: COL = Colonoscopy; FIT = faecal immunochemical test; NoScr = no screening
Screening strategies: screening test, screening start age, screening interval
a. All screening ends at or before the age of 74 years

Table S8: Threshold analysis: estimated cost of determining polygenic risk when personalised screening would be equally cost-effective to uniform annual screening from 50-74 years.

Scenario ^a	Cost ^b	Incremental cost compared to uniform screening	QALYs	Incremental QALYs compared to uniform screening	Incremental costs required to be cost-effective compared to uniform screening ^c	Difference between actual costs and required costs to be cost-effective ^d	Estimated cost to determine risk per individual
FIT_1_50_74	\$1,634,262		17887.34			-	-
PS14	\$1,437,254	-\$197,008	17883.63	-3.71	-\$185,413	\$11,595	\$11.59
PS15	\$1,450,510	-\$183,752	17884.03	-3.31	-\$165,690	\$18,062	\$18.06
PS16	\$1,478,770	-\$155,493	17884.77	-2.57	-\$128,564	\$26,929	\$26.93
PS17	\$1,503,986	-\$130,277	17885.41	-1.93	-\$96,695	\$33,581	\$33.58
PS18	\$1,539,677	-\$94,585	17886.31	-1.03	-\$51,734	\$42,851	\$42.85
PS19	\$1,589,115	-\$45,147	17887.39	0.05	\$2,377	\$47,524	\$47.52
PS20	\$1,624,012	-\$10,250	17887.94	0.60	\$29,979	\$40,229	\$40.23
PS21	\$1,642,804	\$8,542	17888.23	0.89	\$44,647	\$36,104	\$36.10

Abbreviations: QALYs = Quality Adjusted Life Years; FIT = faecal immunochemical test

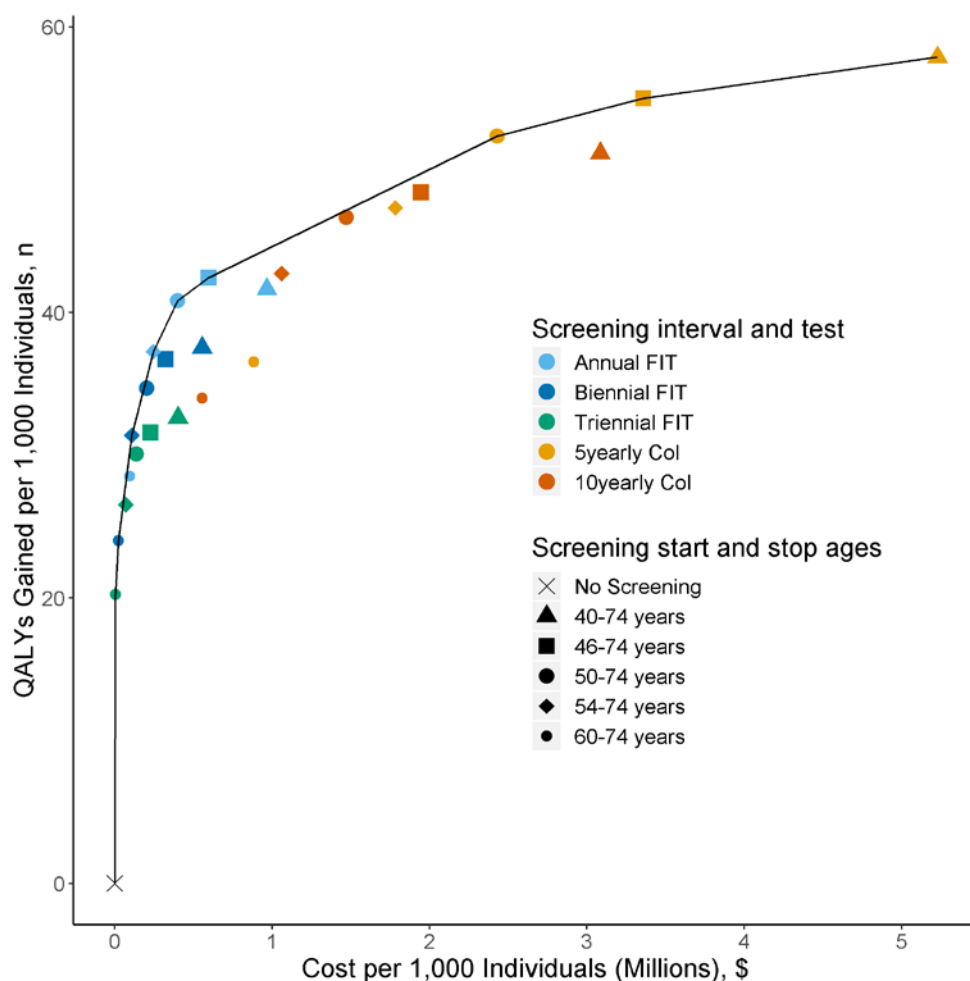
a. Only scenarios with positive threshold costs are included

b. Costs for determining risk are excluded

c. Calculated by multiplying the incremental number of QALYs by the willingness-to-pay threshold of \$50,000

d. Calculated by subtracting the incremental cost compared to uniform screening from the incremental costs required to be cost-effective compared to uniform screening

Figure S1: Costs and quality-adjusted life years (discounted at 5%) per 1,000 40-year-olds assuming perfect adherence for 25 uniform colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies^a

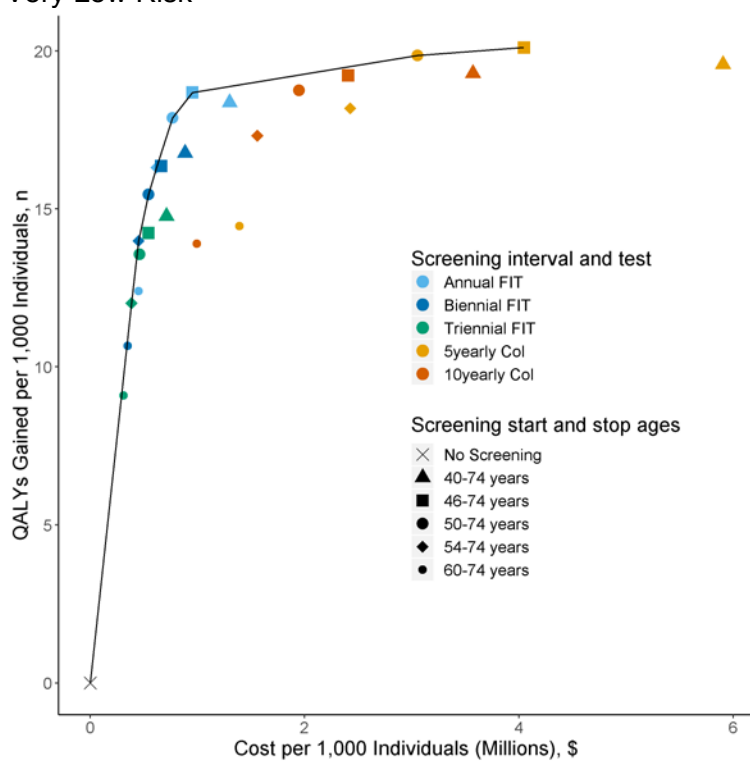


Abbreviations: Col = colonoscopy; FIT = faecal immunochemical test; QALYs = quality-adjusted life years

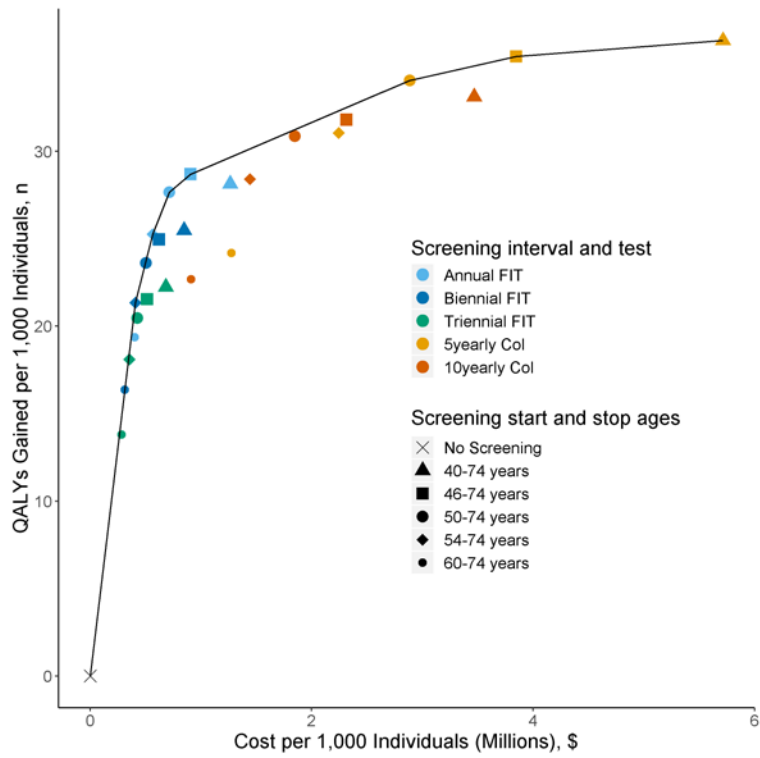
- a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.

Figure S2: Risk group level costs and quality-adjusted life years (discounted at 5%) per 1,000 40-year-olds assuming perfect adherence for 25 colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies^a

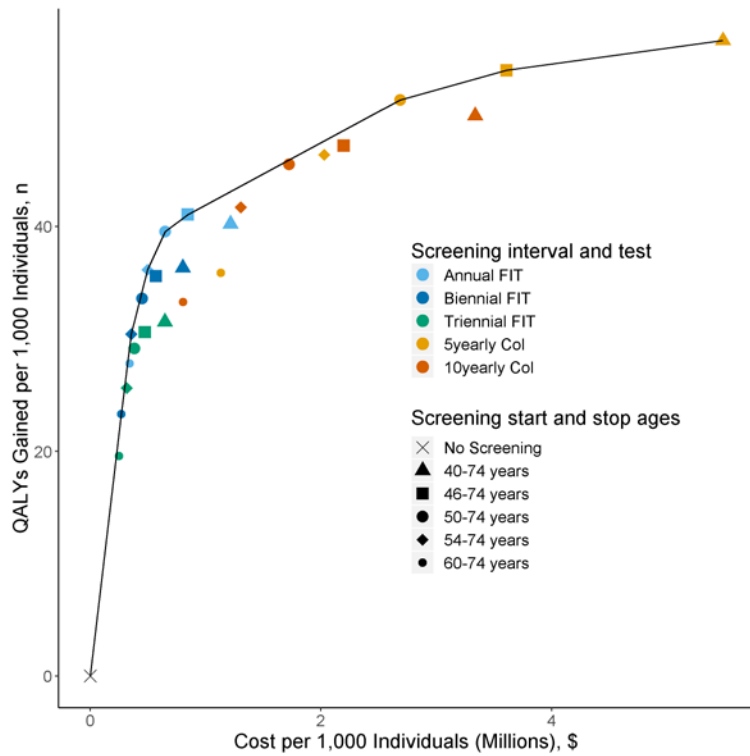
a) Very Low Risk



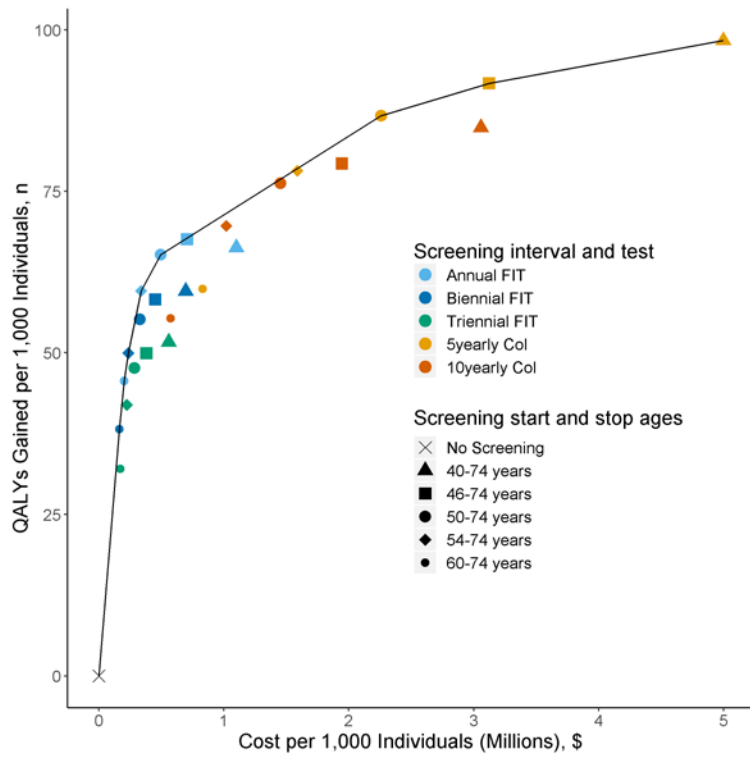
b) Low Risk



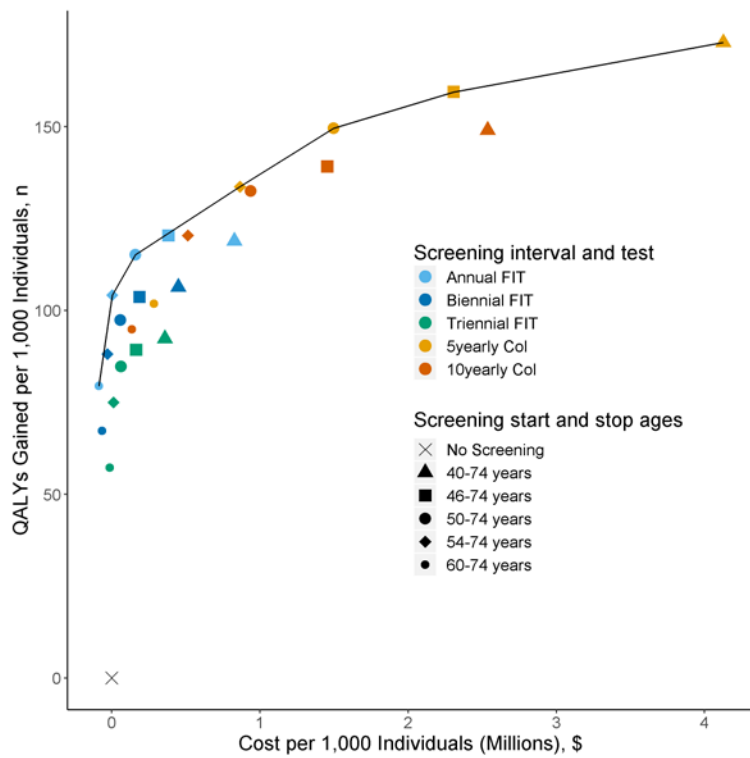
c) Average Risk



d) High Risk



e) Very High Risk

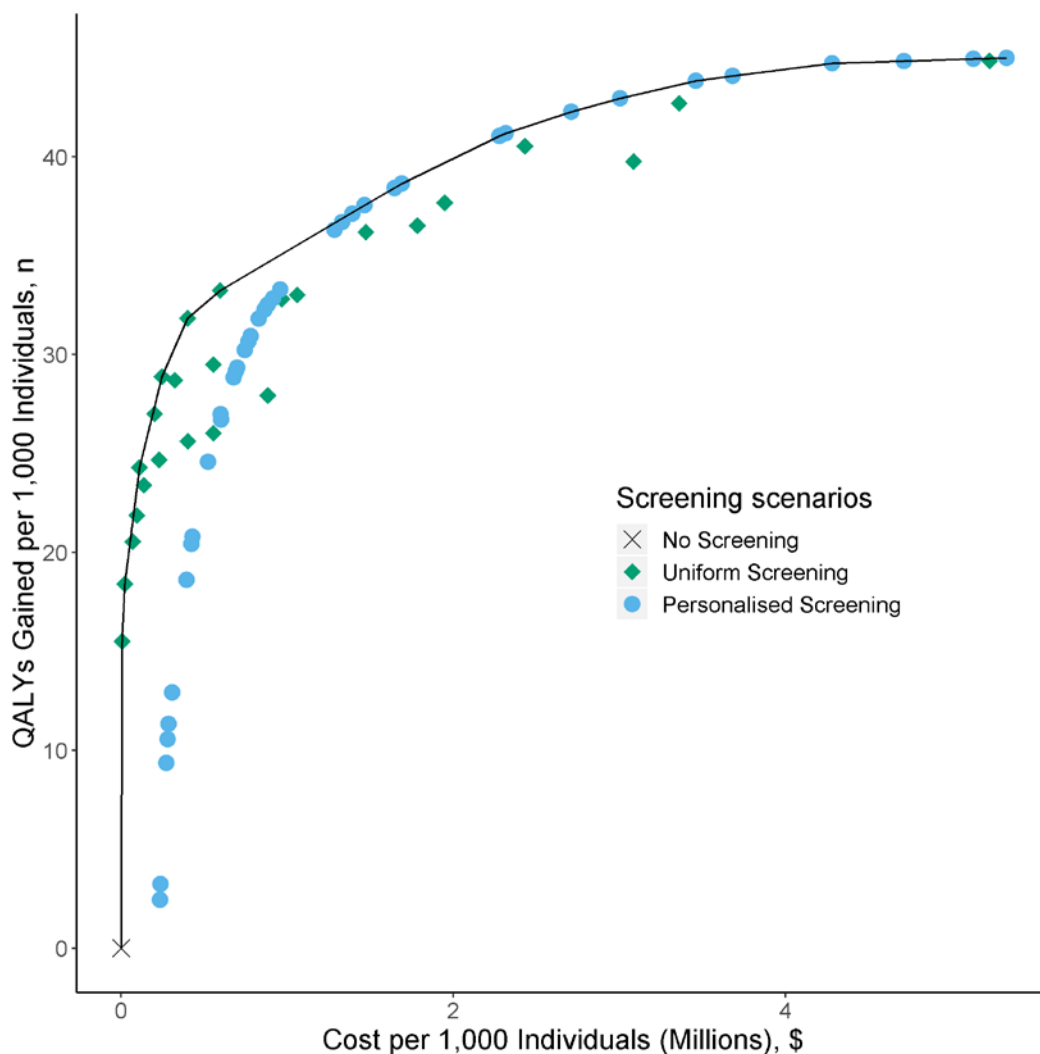


Abbreviations: Col = colonoscopy; FIT = faecal immunochemical test; QALY = quality-adjusted life years

a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life

years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.

Figure S3: Costs (discounted at 5%) and quality-adjusted life years (discounted at 5% and weighted by age) per 1,000 40-year-olds assuming perfect adherence for all uniform and personalised colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies^a

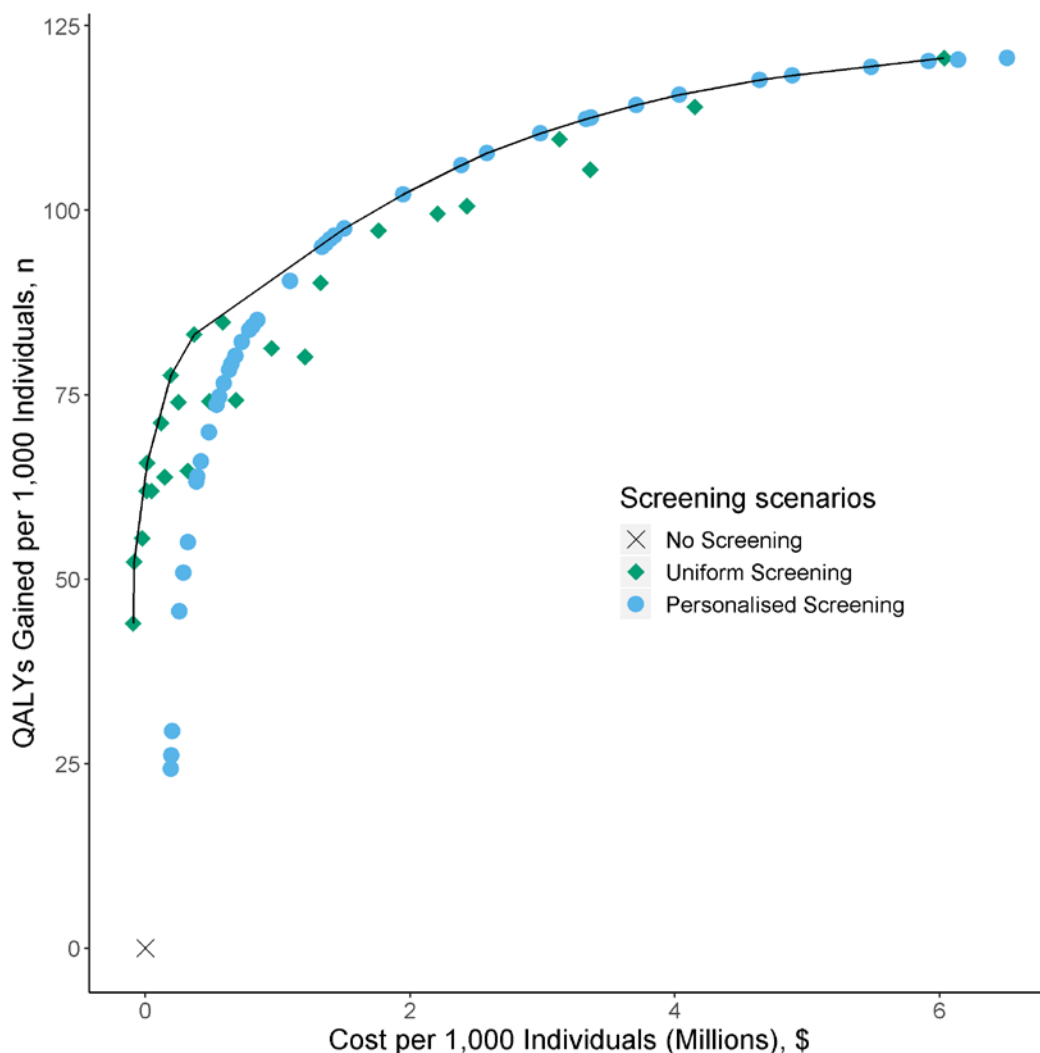


Abbreviations: QALY = quality-adjusted life years

Note: A description of the personalised screening scenarios can be found in Table 3.

- a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.

Figure S4: Costs and quality-adjusted life years (discounted at 3%) per 1,000 40-year-olds assuming perfect adherence for all uniform and personalised colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies^a

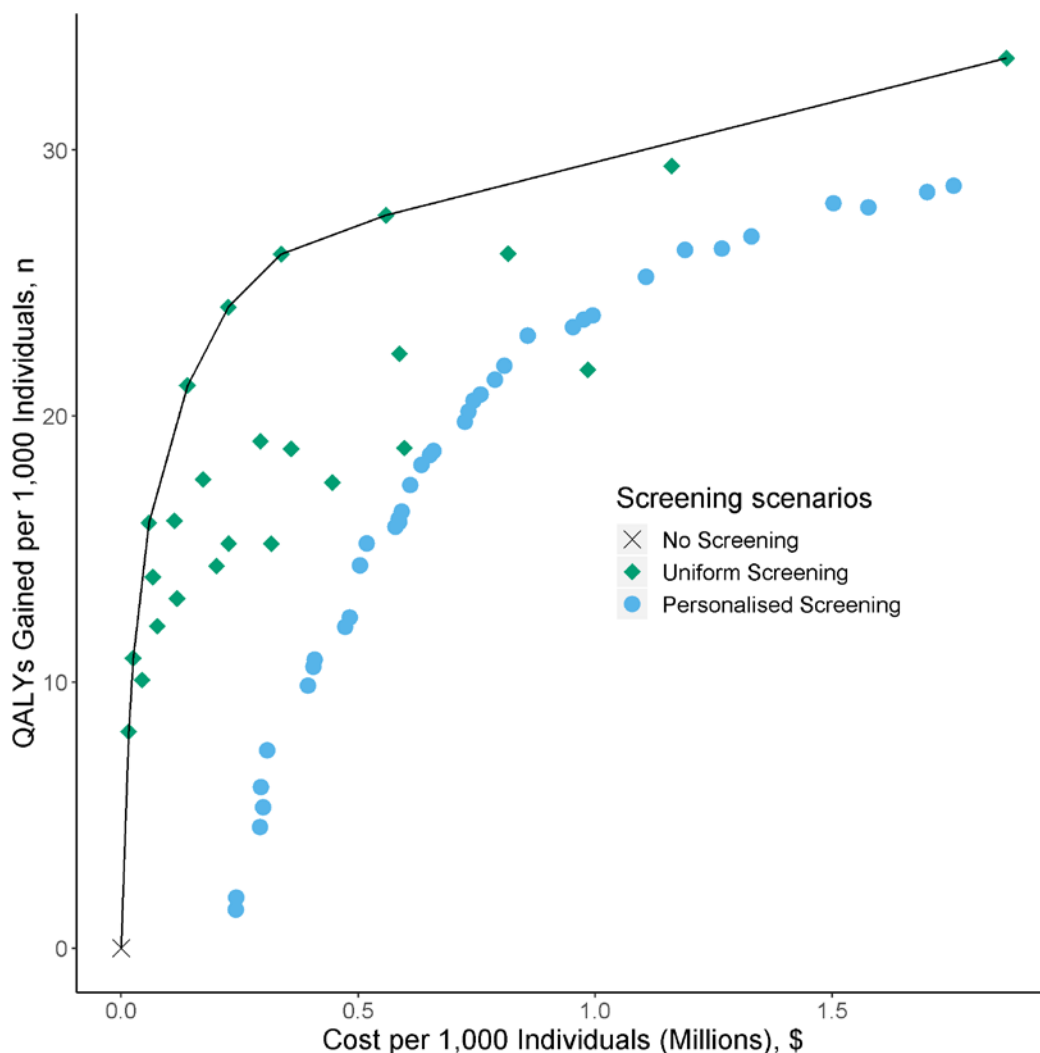


Abbreviations: QALY = quality-adjusted life years

Note: A description of the personalised screening scenarios can be found in Table 5.

- a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.

Figure S5: Costs and quality-adjusted life years (discounted at 5%) per 1,000 40-year-olds assuming realistic participation for uniform screening and lowered adherence for personalised colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies^a

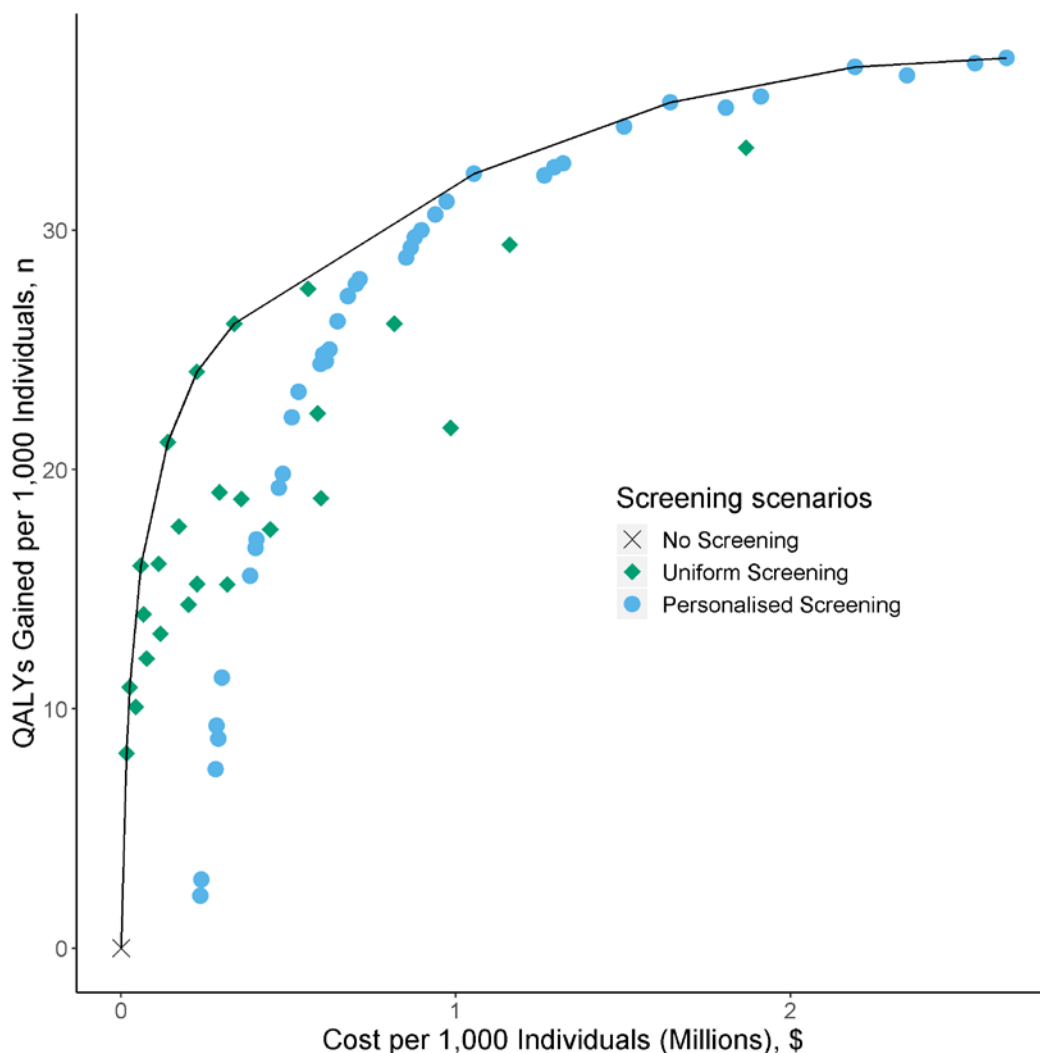


Abbreviations: QALY = quality-adjusted life years

Note: A description of the personalised screening scenarios can be found in Table 4 of the manuscript.

- a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.

Figure S6: Costs and quality-adjusted life years (discounted at 5%) per 1,000 40-year-olds assuming realistic participation for uniform screening and higher adherence for personalised colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies^a

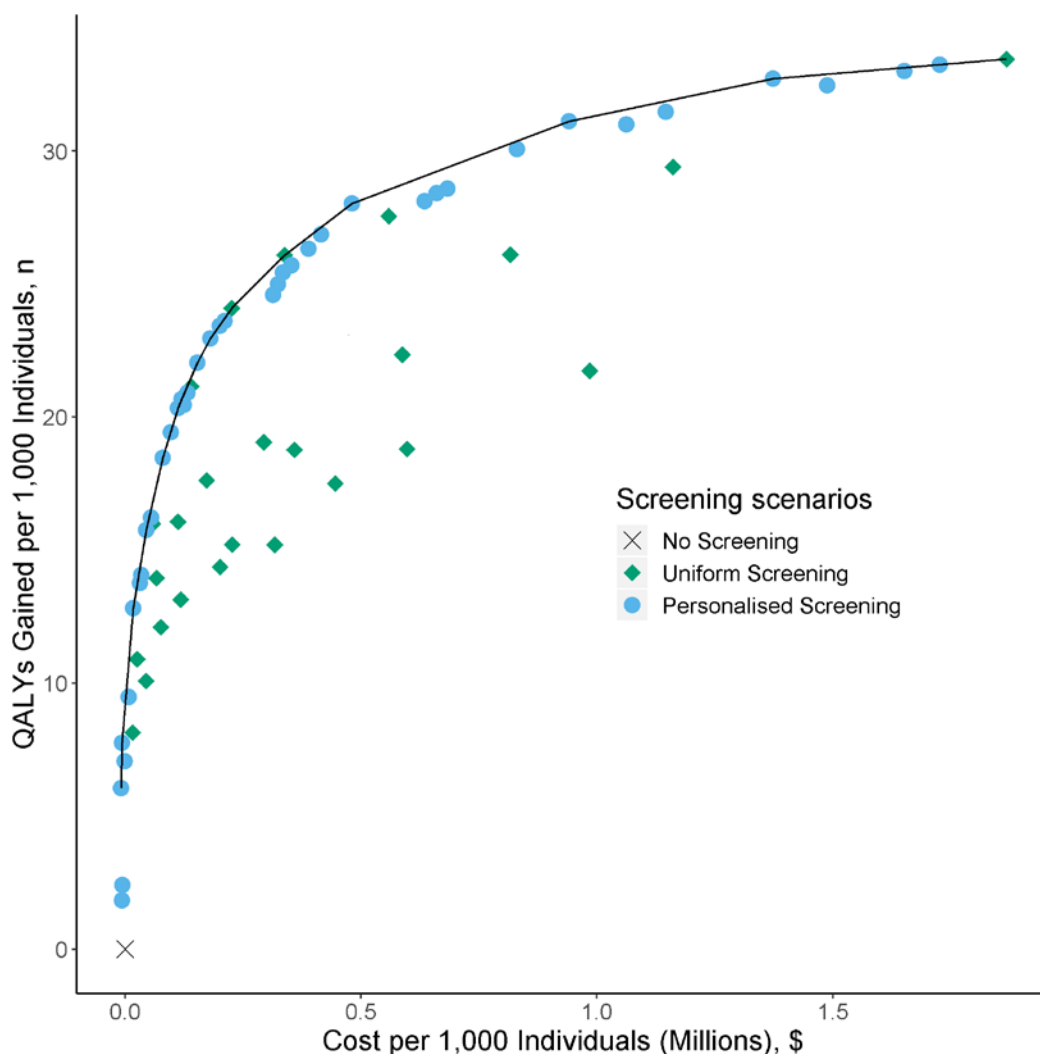


Abbreviations: QALY = quality-adjusted life years

Note: A description of the personalised screening scenarios can be found in Table 4 of the manuscript.

- a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.

Figure S7: Costs and quality-adjusted life years (discounted at 5%) per 1,000 40-year-olds assuming realistic adherence for all uniform and personalised colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies.^a Costs for determining polygenic risk are excluded.

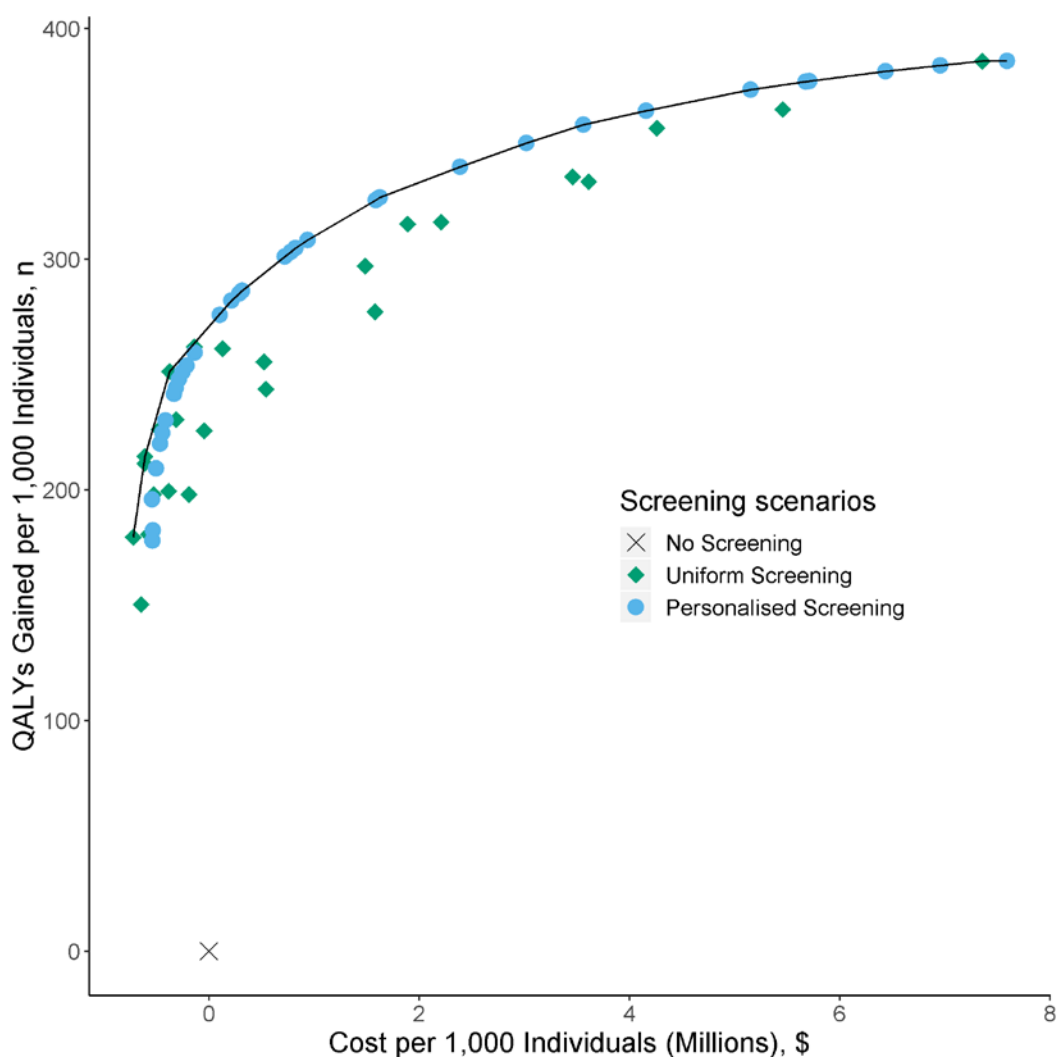


Abbreviations: QALY = quality-adjusted life years

Note: A description of the personalised screening scenarios can be found in Table 4 of the manuscript.

- a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.

Figure S8: Costs and quality-adjusted life years (undiscounted) per 1,000 40-year-olds assuming perfect adherence for all uniform and personalised colorectal cancer screening scenarios and a scenario without screening, with the efficient frontier connecting the economically efficient strategies^a



Abbreviations: QALY = quality-adjusted life years;

Note: A description of the personalised screening scenarios can be found in Table 7.

- a. Discounted costs and life years gained reflect total costs and life years gained of a screening program, accounting for time preference for present over future outcomes. Quality-adjusted life years gained are plotted on the y-axis, and total costs are plotted on the x-axis. Each possible screening strategy is represented by a point. Strategies that form the solid line connecting the points lying left and upward are the economically rational subset of choices. This line is called the efficient frontier. The inverse slope of the line represents the incremental cost-effectiveness ratio of the connected strategies. Points lying to the right and beneath the line represent the dominated strategies.