Estimating the effects of Aducanumab on lifetime and ten-year risks of Alzheimer's disease

Inflating treatment effects

Aducanumab has not been shown to have a massive treatment effect on the probability of developing Alzheimer's disease (AD). The relative risk (RR) for 1-year transitions is likely in the range of 0.75-1. However, it may be useful to see how a treatment with a larger effect would impact lifetime risks of AD. To that end, we assume RRs of 0.75, 0.50, and 0.25 to see how they change lifetime and ten-year risks.

Lifetime Risks

Females

We format tables such that each table represents a given starting state in the multistate model. Columns correspond to lifetime risks for each $RR \in (1, 0.75, 0.50, 0.25)$ for each age (rows).

Table 1: Lifetime risks for females in state 1 (Normal)

Age	RR=1	RR = 0.75	RR=0.5	RR=0.25
60	20.14 %	17.84 %	14.56 %	9.62 %
65	18.73~%	16.48~%	13.32~%	8.68~%
70	16.63~%	14.48~%	11.55~%	7.4~%
75	13.82~%	11.87~%	9.3~%	5.83~%
80	10.42~%	8.78 %	6.73~%	4.1~%
85	7.07~%	5.82~%	4.34~%	2.57~%
90	4.14~%	3.33~%	2.41~%	1.38~%
95	2.07~%	1.62 %	1.14~%	0.63 %

Table 2: Lifetime risks for females in state 2 (Amyloid)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	30.97~%	27.81 %	23 %	15 %
65	29.29~%	26.1 %	21.32~%	13.63~%
70	26.86~%	23.66~%	19.01~%	11.85~%
75	23.52~%	20.38~%	16.02~%	9.68~%
80	19.09 %	16.2~%	12.38~%	7.21~%
85	13.77 %	11.38~%	8.42~%	4.72~%
90	8.39~%	6.74~%	4.82~%	2.6~%
95	4.27~%	3.34~%	2.32~%	1.21~%

Table 3: Lifetime risks for females in state 3 (Neurodegeneration)

Age	RR=1	RR = 0.75	RR=0.5	RR=0.25
60	30.27~%	28.08 %	24.85~%	19.69 %
65	27.58~%	25.31 %	22.01~%	16.88~%
70	24.45~%	22.12~%	18.82~%	13.87~%
75	20.8~%	18.46~%	15.27~%	10.72~%
80	16.54~%	14.32~%	11.44~%	7.58~%
85	11.87~%	9.98~%	7.67~%	4.79~%
90	7.31~%	5.96~%	4.4~%	2.59~%
95	3.82~%	3.02~%	2.15~%	1.21~%

Table 4: Lifetime risks for females in state 4 (A + N) $\,$

Age	RR=1	RR = 0.75	RR=0.5	RR=0.25
60	41.87 %	38.47~%	32.93~%	22.69~%
65	40.8~%	37.21~%	31.46~%	21.18~%
70	38.9~%	35.08~%	29.13~%	19.04~%
75	35.85~%	31.81~%	25.78~%	16.22~%
80	31.19~%	27.04~%	21.23~%	12.79 %
85	24.69 %	20.78 %	15.71~%	9.02~%
90	16.87~%	13.73 %	9.99~%	5.48~%
95	9.91~%	7.81 %	5.48~%	2.89 %

Table 5: Lifetime risks for females in state 5 (A + N + MCI)

Age	RR=1	RR = 0.75	RR=0.5	RR=0.25
60	95.63 %	93.65 %	88.86 %	72.95 %
65	93.57~%	90.7~%	84.23~%	65.84~%
70	90.06~%	85.96~%	77.56~%	57.26~%
75	84.71~%	79.07~%	68.7~%	47.55~%
80	76.19~%	68.96~%	57.21~%	36.95~%
85	63.81~%	55.51~%	43.66~%	26.27~%
90	46.68~%	38.83~%	28.94~%	16.32~%
95	29.89 %	23.87 %	16.99 %	9.09 %

Males

Table 6: Lifetime risks for males in state 1 (Normal)

Age	RR=1	RR = 0.75	RR=0.5	RR=0.25
60	13.87~%	12.19~%	9.87~%	6.48~%
65	12.89~%	11.25~%	9.01~%	5.84~%
70	11.33~%	9.78~%	7.73~%	4.92~%
75	9.25~%	7.87~%	6.11~%	3.81~%
80	6.75~%	5.64~%	4.28~%	2.6~%
85	4.38~%	3.57~%	2.64~%	1.56~%
90	2.38~%	1.9 %	1.36~%	0.78~%
95	1.03~%	0.8~%	0.56~%	0.31~%

Table 7: Lifetime risks for males in state 2 (Amyloid)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	23.1 %	20.54 %	16.76 %	10.71 %
65	21.85~%	19.26~%	15.51~%	9.71~%
70	19.89~%	17.32~%	13.7~%	8.37~%
75	17.19~%	14.71~%	11.38~%	6.74~%
80	13.63~%	11.42~%	8.59~%	4.91~%
85	9.5~%	7.75~%	5.65~%	3.11~%
90	5.43~%	4.31 %	3.04~%	1.62~%
95	2.43~%	1.88~%	1.29~%	0.67~%

Table 8: Lifetime risks for males in state 3 (Neurodegeneration)

Age	RR=1	RR = 0.75	RR=0.5	RR=0.25
60	23.1 %	21.41 %	18.98 %	15.22 %
65	20.83~%	19.07~%	16.58~%	12.85~%
70	18.18~%	16.38 %	13.9~%	10.33~%
75	15.16~%	13.38 %	11.01~%	7.77~%
80	11.72~%	10.08~%	8 %	5.31~%
85	8.12~%	6.77~%	5.16~%	3.23~%
90	4.7~%	3.79 %	2.78~%	1.64~%
95	2.16~%	1.7~%	1.2~%	0.68~%

Table 9: Lifetime risks for males in state 4 (A + N) $\,$

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	33.62~%	30.61~%	25.86~%	17.45 %
65	32.88~%	29.69 %	24.73~%	16.28~%
70	31.27~%	27.87~%	22.77~%	14.53~%
75	28.6~%	25.04~%	19.94~%	12.25~%
80	24.46~%	20.91~%	16.13~%	9.5~%
85	18.88~%	15.67~%	11.65~%	6.55~%
90	12.38~%	9.94~%	7.12~%	3.84~%
95	6.56~%	5.11 %	3.54~%	1.84~%

Table 10: Lifetime risks for males in state 5 (A + N + MCI)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	92.93~%	90.1 %	83.93 %	66.37~%
65	90.37~%	86.56~%	78.74~%	59.24~%
70	85.99~%	80.9 %	71.31~%	50.68~%
75	79.54~%	73.01~%	61.89~%	41.31~%
80	69.91~%	62.11~%	50.26~%	31.4~%
85	56.66~%	48.4 %	37.23~%	21.79 %
90	40.15~%	32.86~%	24.03~%	13.26~%
95	23.87~%	18.79 %	13.17~%	6.93~%

Ten-year risks

Females

Table 11: Ten-year risks for females in state 1 (Normal)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	0.2 %	0.18 %	0.16 %	0.13 %
65	0.47~%	0.41~%	0.35~%	0.26~%
70	1.07~%	0.92~%	0.74~%	0.53~%
75	2.2~%	1.84~%	1.43~%	0.95~%
80	3.68~%	3.04~%	2.3~%	1.44~%
85	4.68~%	3.81~%	2.82~%	1.68~%
90	3.84~%	3.08~%	2.22~%	1.27~%
95	2.06~%	1.62~%	1.14~%	0.63~%

Table 12: Ten-year risks for females in state 2 (Amyloid)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	1.26 %	1.04 %	0.77 %	0.43 %
65	2.5~%	2.06~%	1.52~%	0.85~%
70	4.65~%	3.84~%	2.84~%	1.58~%
75	7.83~%	6.47~%	4.78~%	2.66~%
80	11.05~%	9.11~%	6.71~%	3.73~%
85	11.5~%	9.39~%	6.85~%	3.76~%
90	8.17~%	6.55~%	4.68~%	2.51~%
95	4.27~%	3.34~%	2.32~%	1.21 %

Table 13: Ten-year risks for females in state 3 (Neurodegeneration)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	3.6~%	3.51~%	3.39 %	3.25 %
65	4.32~%	4.11~%	3.85~%	3.54~%
70	5.52~%	5.09~%	4.55~%	3.88~%
75	7.33~%	6.5~%	5.48~%	4.21~%
80	9.34~%	8.01~%	6.38~%	4.38~%
85	9.67~%	8.08~%	6.17~%	3.88~%
90	7.07~%	5.76 %	4.24~%	2.5~%
95	3.82~%	3.02~%	2.15~%	1.21~%

Table 14: Ten-year risks for females in state 4 (A + N) $\,$

Age	RR=1	RR = 0.75	RR=0.5	RR=0.25
60	7.07 %	6.08 %	4.7 %	2.76 %
65	10.71~%	9.2~%	7.11~%	4.18~%
70	15.5~%	13.3~%	10.27~%	6.02~%
75	20.79 %	17.79 %	13.69~%	7.99~%
80	24.43~%	20.76~%	15.85~%	9.17~%
85	23.1 %	19.31~%	14.47~%	8.2~%
90	16.76 %	13.63 %	9.9~%	5.42~%
95	9.9~%	7.81 %	5.48~%	2.89~%

Table 15: Ten-year risks for females in state 5 (A + N + MCI)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	93.47 %	87.93 %	75.88 %	50.63 %
65	91.72 %	85.9 %	73.74 %	48.92 %
70	88.64~%	82.41 %	70.13~%	46.08~%
75	83.83~%	76.95~%	64.5~%	41.67~%
80	75.84~%	68.14~%	55.66~%	34.92~%
85	63.74~%	55.36 %	43.4~%	25.94~%
90	46.68~%	38.83 %	28.93~%	16.3~%
95	29.89~%	23.87~%	16.99~%	9.09~%

Males

Table 16: Ten-year risks for females in state 1 (Normal)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	0.2 %	0.18 %	0.16 %	0.13 %
65	0.47~%	0.41~%	0.35~%	0.26~%
70	1.07~%	0.92~%	0.74~%	0.53~%
75	2.2~%	1.84~%	1.43~%	0.95~%
80	3.68~%	3.04~%	2.3~%	1.44~%
85	4.68~%	3.81~%	2.82~%	1.68~%
90	3.84~%	3.08~%	2.22~%	1.27~%
95	2.06~%	1.62~%	1.14~%	0.63~%

Table 17: Ten-year risks for females in state 2 (Amyloid)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	1.26 %	1.04 %	0.77 %	0.43 %
65	2.5~%	2.06~%	1.52~%	0.85~%
70	4.65~%	3.84~%	2.84~%	1.58~%
75	7.83~%	6.47~%	4.78~%	2.66~%
80	11.05~%	9.11~%	6.71~%	3.73~%
85	11.5~%	9.39~%	6.85~%	3.76~%
90	8.17~%	6.55~%	4.68~%	2.51~%
95	4.27~%	3.34~%	2.32~%	1.21~%

Table 18: Ten-year risks for females in state 3 (Neurodegeneration)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	3.6 %	3.51 %	3.39 %	3.25 %
65	4.32~%	4.11~%	3.85~%	3.54~%
70	5.52~%	5.09~%	4.55~%	3.88~%
75	7.33 %	6.5~%	5.48~%	4.21~%
80	9.34~%	8.01~%	6.38~%	4.38~%
85	9.67~%	8.08~%	6.17~%	3.88~%
90	7.07~%	5.76 %	4.24~%	2.5~%
95	3.82~%	3.02~%	2.15~%	1.21 %

Table 19: Ten-year risks for females in state 4 (A + N)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	7.07 %	6.08 %	4.7 %	2.76 %
65	10.71~%	9.2~%	7.11~%	4.18~%
70	15.5~%	13.3 %	10.27~%	6.02~%
75	20.79 %	17.79 %	13.69~%	7.99~%
80	24.43~%	20.76~%	15.85~%	9.17~%
85	23.1~%	19.31 %	14.47~%	8.2~%
90	16.76~%	13.63 %	9.9~%	5.42~%
95	9.9~%	7.81~%	5.48~%	2.89~%

Table 20: Ten-year risks for females in state 5 (A + N + MCI)

Age	RR=1	RR=0.75	RR=0.5	RR=0.25
60	93.47~%	87.93 %	75.88 %	50.63 %
65	91.72~%	85.9 %	73.74~%	48.92~%
70	88.64~%	82.41~%	70.13~%	46.08~%
75	83.83~%	76.95~%	64.5~%	41.67~%
80	75.84~%	68.14~%	55.66~%	34.92~%
85	63.74~%	55.36~%	43.4~%	25.94~%
90	46.68~%	38.83~%	28.93~%	16.3~%
95	29.89~%	23.87~%	16.99~%	9.09~%