# **Supplementary Online Content**

Golledge J, Pinchbeck J, Tomee SM, et al. Efficacy of telmisartan to slow growth of small abdominal aortic aneurysms: a randomized clinical trial. *JAMA Cardiol*. Published online August 26, 2020. doi:10.1001/jamacardio.2020.3524

eTable 1. Adherence to Allocated Medication

**eTable 2.** Effect of Telmisartan on Health-Related Quality of Life at the End of the Treatment Period

eTable 3. Effect of Telmisartan on Safety Outcomes

eFigure 1. Effect of Telmisartan on C-reactive Protein

**eFigure 2.** Lack of Interaction Between Initial AAA Diameter Grouping and Effect f Telmisartan on AAA Growth Measured With Ultrasound

eFigure 3. Effect of Telmisartan on Risk of AAA Repair or Death From AAA Rupture

eFigure 4. Effect of Telmisartan on Ankle Brachial Pressure Index

eFigure 5. Effect of Telmisartan on Short Form-36 Physical Component Score

eFigure 6. Effect of Telmisartan on Short Form-36 Mental Component Score

eAppendix. TEDY Team, Collaborating Sites, and Investigators

This supplementary material has been provided by the authors to give readers additional information about their work.

	Telmisartan		Placebo	
Time since randomization	Number assessed (%)	Taking ≥80% tablets (%)	Number assessed (%)	Taking ≥80% tablets (%)
3 Months	83 (78)	77 / 83 (93)	78 (77)	72 / 78 (92)
6 Months	79 (75)	72 / 79 (91)	79 (78)	77 / 79 (97)
One Year	72 (68)	66 / 72 (92)	73 (72)	69 / 73 (95)
18 Months	67 (63)	61 / 67 (91)	63 (62)	58 / 63 (92)
Two Years	76 (72)	68 / 76 (89)	68 (67)	62 / 68 (91)

Treatment adherence assessed by pill counting. Table shows the number of participants who had capsules counted, and the proportion who were taking  $\geq 80\%$  of the study medications.

SF-36 outcome	Telmisartan (n=106)	Placebo (n=101)	Mean difference (95% CI)	<i>p</i> -value*
Physical functioning	66.1 (2.7)	64.8 (2.8)	-1.3 (-1.3 to -1.3)	0.731
Role physical	74.1 (3.0)	75.6 (2.7)	1.5 (1.5 to 1.5)	0.708
Bodily pain	72.8 (2.7)	74.6 (2.6)	1.8 (1.8 to 1.8)	0.630
General health	61.5 (2.1)	64.5 (2.1)	2.9 (2.9 to 2.9)	0.329
Vitality	60.3 (2.0)	62.7 (2.0)	2.4 (2.4 to 2.4)	0.392
Social functioning	84.4 (2.1)	84.5 (2.2)	0.1 (0.1 to 0.1)	0.975
Role emotional	75.6 (2.6)	74.4 (2.6)	-1.1 (-1.1 to -1.1)	0.758
Mental health	80.4 (1.4)	81.0 (1.6)	0.6 (0.6 to 0.6)	0.782
Physical component score <sup>†</sup>	43.9 (1.0)	44.2 (1.0)	0.4 (0.4 to 0.4)	0.802
Mental component score†	51.9 (0.8)	52.3 (0.9)	0.4 (0.4 to 0.4)	0.732

eTable 2. Effect of telmisartan on health-related quality of life at the end of the treatment period

Shown are mean (SD) SF-36 health-related quality of life scores at the end of the trial by allocated treatment group. Missing data were imputed using the last observation carried forward method.

\*Calculated using t-test comparing means at 24 months.

<sup>†</sup>The physical- and mental- component summary scores are based on the weighted T-scores of the four physical and mental SF-36 scales, respectively. The raw SF-36 component summary scores were transformed to a norm-based T-score with a mean ( $\pm$ SD) score of 50 $\pm$ 10 in the general population (higher scores indicate a better quality of life).

## eTable 3. Effect of telmisartan on safety outcomes

Assessment	Telmisartan (n=106)	Placebo (n=101)	<i>p</i> -value <sup>a</sup>
Any adverse event	65 (61)	58 (57)	0.575
Hospitalization	27 (25)	33 (33)	0.285
Death	2 (2)	6 (6)	0.162
Syncope or hypotensive symptoms	28 (26)	13 (13)	0.015
Cough	1 (1)	5 (5)	0.112
Gastrointestinal symptoms	6 (6)	4 (4)	0.749
Angioedema	0 (0)	0 (0)	
End-stage renal disease	0 (0)	0 (0)	•

Values shown are frequency (%). Only the first adverse event for each participant was counted. Percentages may not total 100 because of rounding.

<sup>a</sup>Calculated using Fisher's exact test to account for low frequencies.

eFigure 1. Effect of telmisartan on C-reactive protein



eFigure 2: Lack of interaction between initial AAA diameter grouping and effect of telmisartan on AAA growth measured with ultrasound



Effects of telmisartan vs placebo on AAA growth by baseline ultrasound diameter. The mean difference for each group is the average difference in annual AAA growth rates between randomly allocated groups (intention-to-treat). Box areas are inversely proportional to the variance within each group. AAA, abdominal aortic aneurysm; SE, standard error; MD, mean difference; CI, confidence interval.





Kaplan-Meier illustration of the risk of a first AAA-related event (composite of AAA repair or AAA-related mortality) among participants allocated to telmisartan or placebo. Follow-up was continued up until two years. The number of patients at risk at the start of each six months of follow-up are shown below the graph.

Placebool

Telmis**lant**an



#### eFigure 4. Effect of telmisartan on ankle brachial pressure index

Ankle brachial pressure index (ABI) calculated as the highest of the dorsalis pedis or posterior tibial artery Doppler pressure divided by the brachial artery Doppler pressure. Representative ABI taken as the lowest ABI of both lower limbs. Incompressible ABIs (>1.4) not included. I bars represent 95% CI. No significant difference in rate of change of ABI (p = 0.959).



## eFigure 5. Effect of telmisartan on Short Form-36 physical component score

Shown is the mean SF-36 summary score for each randomized group at baseline and at subsequent study visits (numbers shown below graph). Missing data were imputed using the last value carried forward method.



## eFigure 6. Effect of telmisartan on Short Form-36 mental component score

Shown is the mean SF-36 summary score for each randomized group at baseline and at subsequent study visits (numbers shown below graph). Missing data were imputed using the last value carried forward method.

#### eAppendix. TEDY Team, Collaborating Sites and Investigators

**Independent trial randomisation** Chris Reid, Monash and Curtin University, Australia Data and safety monitoring board John Walsh, South Australia Andrew Tonkin, Victoria Cindy Sealey, Queensland Trial executive committee Jonathan Golledge, Queensland Rene Jaeggi, Queensland Paul Norman, Western Australia Sophie Rowbotham, Queensland Jenna Pinchbeck, Queensland Data quality oversight and checking Cindy Sealey, Queensland Rene Jaeggi, Queensland Tejas Singh, Queensland Joseph Moxon, Queensland Jonathan Golledge, Queensland <u>Data analysis</u> Dylan Morris, Queensland **Core Imaging Laboratory** Jenna Pinchbeck, Queensland Georgina Anderson, Queensland **Recruitment sites** Australia **Gosford Vascular Services, New South Wales** Principal Investigator Dr Bernie Bourke Trial Coordinator Dr Michael Bourke Queen Elizabeth Hospital, Adelaide, South Australia Principal Investigator Professor Robert Fitridge Trial Coordinators Ms Ruth Battersby and Dr. Prue Cowled Royal Brisbane and Women's Hospital, Queensland Principal Investigator Professor Jason Jenkins Trial Coordinator Dr Sophie Rowbotham and Dr Brad Stefanovic The Townsville University Hospital and the Mater Hospital Pilmico, Queensland Principal Investigators Professor Jonathan Golledge and Dr Frank Quigley Trial Coordinators Ms Jenna Pinchbeck and Ms Barbara Bradshaw The Netherlands Trial Coordinator Ms Stephanie Tomee Leiden University Medical Centre, Leiden Principal Investigators Associate Professor Jan Lindeman and Prof. Jaap F. Hamming Jeroen Bosch Hospital, Hertogenbosch Principal Investigator Dr. Jan Willem Hinnen **Deventer ziekenhuis** Principal Investigator Dr Robert B van Tongeren Haga Ziekenhuis, Den Haag Principal Investigator Dr. Jan J Wever Haaglanden Medisch Centrum, Den Haag Principal Investigator Dr. Daniël Eefting Sint Franciscus Gasthuis en Vlietland, Rotterdam Principal Investigator Dr. Jerome P van Brussel, Antonius Ziekenhuis Nieuwegein Principal Investigator Dr. Jean-Paul de Vries (current address: University Medical Center Groningen) Elisabeth Tweesteden Ziekenhuis, Tilburg Principal Investigator Dr. Patrick W. Vriens The United States of America Veterans Administration Hospital and Stanford University, Palo Alto, California Principal Investigators Professor Ron Dalman and Dr. Oliver A. Aalami Trial Coordinator Ms Lori McDonnell