Online supplement

Influence of age on upper-arm cuff blood pressure measurement

Dean S. Picone¹, Martin G. Schultz¹, Petr Otahal¹, J. Andrew Black^{1,2}, Willem J. Bos^{3,4}, Chen-Huan Chen⁵, Hao-Min Cheng⁵, Antoine Cremer⁶, Nathan Dwyer^{1,3}, Ricardo Fonseca¹, Alun D. Hughes⁷, Hack-Lyoung Kim⁸, Peter S. Lacy⁹, Esben Laugesen¹⁰, Nobuyuki Ohte¹¹, Stefano Omboni^{12,13}, Christian Ott¹⁴, Telmo Pereira¹⁵, Giacomo Pucci¹⁶, Philip Roberts-Thomson^{1,2}, Niklas B. Rossen¹⁰, Roland E. Schmieder¹⁴, Daisuke Sueta¹⁷, Kenji Takazawa¹⁸, Jiguang Wang¹⁹, Thomas Weber²⁰, Berend E. Westerhof²¹, Bryan Williams⁹, Hirotsugu Yamada²², Eiichiro Yamamoto¹⁷, James E. Sharman¹ for the InvaSive blood PressurE ConsorTium

¹Menzies Institute for Medical Research, University of Tasmania, Hobart, Australia ²Royal Hobart Hospital, Hobart, Tasmania

³St Antonius Hospital, Department of Internal Medicine, Nieuwegein, The Netherlands ⁴Department of Internal Medicine, Leiden University Medical Center, Leiden, The Netherlands

⁵Department of Medicine, National Yang-Ming University School of Medicine, Department of Medical Education, Taipei Veterans General Hospital, Taipei, Taiwan ⁶Department of Cardiology/Hypertension, University Hospital of Bordeaux, Bordeaux,

France

⁷Institute of Cardiovascular Sciences, University College London, London, United Kingdom
 ⁸Division of Cardiology, Seoul National University Boramae Hospital, Seoul, South Korea
 ⁹Institute of Cardiovascular Sciences University College London (UCL) and National
 Institute for Health Research (NIHR) UCL/UCL Hospitals Biomedical Research Centre,
 London, United Kingdom

1

¹⁰Department of Endocrinology and Internal Medicine, Aarhus University Hospital, Aarhus, Denmark

¹¹Department of Cardio-Renal Medicine and Hypertension, Nagoya City University Graduate School of Medical Sciences, Nagoya, Japan

¹²Clinical Research Unit, Italian Institute of Telemedicine, Varese, Italy

¹³Scientific Research Department of Cardiology, Science and Technology Park for Biomedicine, Sechenov First Moscow State Medical University, Moscow, Russian Federation

¹⁴Department of Nephrology and Hypertension, University Hospital Erlangen, Friedrich-Alexander University Erlangen-Nürnberg, Erlangen, Germany

¹⁵Polytechnic Institute of Coimbra, ESTES, Department of Physiology, General Humberto Delgado Street 102, Lousã, Portugal

¹⁶Unit of Internal Medicine at Terni University Hospital, Department of Medicine, University of Perugia, Perugia, Italy

¹⁷Department of Cardiovascular Medicine, Graduate School of Medical Sciences, Kumamoto University, Kumamoto, Japan

¹⁸Center for Health Surveillance and Preventive Medicine, Tokyo Medical University Hospital, Tokyo, Japan

¹⁹Centre for Epidemiological Studies and Clinical Trials, Shanghai Key Laboratory of Hypertension, The Shanghai Institute of Hypertension, Department of Hypertension, Ruijin Hospital, Shanghai Jiao Tong University School of Medicine, Shanghai, China ²⁰Cardiology Department, Klinikum Wels-Grieskirchen, Wels, Austria ²¹Department of Pulmonary Diseases, VU University Medical Center, Amsterdam, The

Netherlands

²²Department of Community Medicine for Cardiology, Tokushima Graduate School of

Biomedical Sciences, Tokushima, Japan

Address for correspondence:

Professor James Sharman Menzies Institute for Medical Research University of Tasmania Private Bag 23 Hobart, 7000 Australia Telephone: +61 3 6226 4709 Fax: +61 3 6226 7704 Email: James.Sharman@utas.edu.au

Table of Contents

Expanded Methods	5
Online supplement Table S1.	6
Online supplement Table S2.	9
Online supplement Table S3.	11
Online supplement Table S4.	12
Online supplement Table S5.	13
Online supplement Table S6.	14
Online supplement Table S7.	15
Online supplement Table S8.	16
Online supplement Table S9.	17
Online supplement Table S10.	18
Online supplement Figure S1	19
Online supplement Figure S2	20
Online supplement Figure S3	21
Online supplement Figure S4	22
Online supplement Figure S5	23
Online supplement Figure S6	24
Online supplement Figure S7	25
Online supplement Figure S8	26
Online supplement Figure S9	27
Online supplement Figure S10	28
Online supplement references	29

Expanded Methods.

Description of the order of the cuff BP measurements.

Cuff and invasive BP were measured simultaneously in seventeen studies (n=1078). Cuff BP was measured just prior to the invasive BP in seven studies (n=313), whereas invasive BP was measured just prior to cuff BP in five studies (n=204). In one study (n=27), duplicate cuff BP measurements were recorded, then invasive BP, then duplicate cuff BP again with the average of the four cuff values used for analysis. In another study (n=52), an invasive BP was recorded, then cuff BP and then another invasive BP with the average of the duplicate invasive BP recordings used for analysis.

Description of the study quality score used. One point was awarded for each of the five attributes when the highest standard of study quality was met.

1. Type of catheter

- a) micromanometer tip: 1 point OR
- **b**) fluid filled catheter manometer system description of frequency and damping characteristics: **1 point** OR
- c) Fluid filled catheter manometer system insufficient detail for b): 0 points

2. Sequence of aortic and brachial BP measurements

- a) Simultaneous: 1 point \square OR
- b) sequential, describing the time between measurements and that no major hemodynamic changes occurred: 1 point □ OR
- c) sequential, insufficient detail for b): **0 points**

3. Position of catheter in aorta

- a) described with sufficient detail to ascertain position (aortic BP was required to be measured in the proximal aorta or aortic arch): **1 point** OR
- **b**) general description: **0** points

4. Pressure wave capture length

- a) > 1 beat of continuously captured data, with a description that the recording was of good quality (i.e period of capture was stable): **1 point** OR
- **b**) 1 beat: **0 points** OR
- c) or no description: **0 points**

5. Participant characteristics

- a) description of patient inclusion/exclusion criteria (with reference to conditions that may cause hemodynamic instability / difficulty to obtain accurate measurements): **1 point** OR
- **b**) detailed description of the patient clinical characteristics (with reference to conditions that may cause hemodynamic instability / difficulty to obtain accurate measurements): **1 point** OR
- c) no, or poor, description of the patient inclusion/exclusion criteria (with reference to conditions that may cause hemodynamic instability / difficulty to obtain accurate measurements): **0 points** OR
- d) no or poor description of patient clinical characteristics (with reference to conditions that may cause hemodynamic instability / difficulty to obtain accurate measurements): **0 points**

Online supplement Table S1. Studies included in the cull versus invasive aortic BP analyses.	
Citation	Cuff blood pressure device
1. Aakhus et al, ¹ Noninvasive Estimates of Aortic Root Pressures - External Subclavian Arterial	UA 751, Takeda Medical Inc
Pulse Tracing Calibrated by Oscillometrically Determined Brachial Arterial Pressures. Clin	
Physiol. 1993;13(6):573-586.	
2. Bhatt SD, Hinderliter AL, Stouffer GA. ² Influence of Sex on the Accuracy of Oscillometric-	Omron device
Derived Blood Pressures. J Clin Hypertens. 2011;13(2):112-119.	
3. Borow KM, Newburger JW. ³ Noninvasive estimation of central aortic pressure using the	Dinamap 845
oscillometric method for analyzing systemic artery pulsatile blood flow: comparative study of	
indirect systolic, diastolic, and mean brachial artery pressure with simultaneous direct ascending	
aortic pressure measurements. Am Heart J. 1982;103(5):879-886.	
4. Bos et al, ⁴ Pseudohypertension and the measurement of blood pressure. <i>Hypertension</i> .	Mercury sphygmomanometer
1992;20(1):26-31.	
5. Broyd et al, unpublished*	PulseCor R6.5 (POEM2 oscillometric
	BP [Welch Allyn])
6. Cheng HM, Wang KL, Chen YH, et al. ⁵ Estimation of central systolic blood pressure using an	VP-2000, Colin corporation
oscillometric blood pressure monitor. Hypertens Res. 2010;33(6):592-599.	
7. Cheng et al, unpublished*	VP-2000, Colin corporation
8. Costello BT, Schultz MG, Black JA, Sharman JE. ⁶ Evaluation of a brachial cuff and	Pulsecor R7.0 (Welch Allyn brachial
suprasystolic waveform algorithm method to noninvasively derive central blood pressure. Am J	BP)
Hypertens. 2015;28(4):480-486.	
9. Cremer A, Butlin M, Codjo L, et al. ⁷ Determination of central blood pressure by a noninvasive	Diasys Integra II
method (brachial BP and QKD interval). J Hypertens. 2012;30(8):1533-1539.	
10. Ding FH, Li Y, Zhang RY, Zhang Q, Wang JG. ⁸ Comparison of the SphygmoCor and Omron	Omron HEM 9000AI
devices in the estimation of pressure amplification against the invasive catheter measurement. J	
Hypertens. 2013;31(1):86-93.	
11. Jeon WK, Kim MA, Kim HL, et al. ⁹ Association between aortic knob width and invasively	Mennen Medical oscillometric monitor

Online supplement Table S1. Studies included in the cuff versus invasive aortic BP analyses.

measured aortic pulse pressure. Blood Press Monit. 2018; 23(3): 121-6.

12. Laugesen E, Rossen NB, Peters CD, et al. ¹⁰ Assessment of central blood pressure in patients	Riester Champion N automatic blood
with type 2 diabetes: a comparison between SphygmoCor and invasively measured values. Am J	pressure monitor
Hypertens. 2014;27(2):169-176.	
13. Lin MM, Cheng HM, Sung SH, et al. ¹¹ Estimation of central aortic systolic pressure from the	WatchBP Office, Microlife
second systolic peak of the peripheral upper limb pulse depends on central aortic pressure	
waveform morphology. J Hypertens. 2012;30(3):581-586.	
14. Nagle et al, ¹² Comparisons of direct and indirect blood pressure with pressure-flow dynamics	Mercury sphygmomanometer
during exercise. J Appl Physiol. 1966;21(1):317-320.	
15. Nakagomi A, Okada S, Shoji T, Kobayashi Y. ¹³ Aortic pulsatility assessed by an oscillometric	Mobil-o-graph, IEM
method is associated with coronary atherosclerosis in elderly people. <i>Blood Press.</i> 2016:1-8.	
16. Ohte N, Saeki T, Miyabe H, et al. ¹⁴ Relationship between blood pressure obtained from the	BP-8800, Omron Colin
upper arm with a cuff-type sphygmomanometer and central blood pressure measured with a	
catheter-tipped micromanometer. Heart Vessels. 2007;22(6):410-415.	
17. Ott C, Haetinger S, Schneider MP, Pauschinger M, Schmieder RE. ¹⁵ Comparison of two	Dinamap Pro 100 V2
noninvasive devices for measurement of central systolic blood pressure with invasive measurement	
during cardiac catheterization. J Clin Hypertens (Greenwich). 2012;14(9):575-579.	
18. Park CM, Korolkova O, Davies JE, et al. ¹⁶ Arterial pressure: agreement between a brachial	PulseCor R6.5 (POEM2 oscillometric
cuff-based device and radial tonometry. J Hypertens. 2014;32(4):865-872.	BP [Welch Allyn])
19. Park et al, unpublished*	PulseCor R6.5 (POEM2 oscillometric
	BP [Welch Allyn])
20. Pereira T, Maldonado J, Coutinho R, et al. ¹⁷ Invasive validation of the Complior Analyse in the	Colson MAM BP 3AA1-2
assessment of central artery pressure curves: a methodological study. Blood Press Monit.	
2014;19(5):280-287.	
21. Picone et al, unpublished*	Sphygmocor Xcel (oscillometric BP
	Suntech Advantage NIBP)
22. Pucci G, Cheriyan J, Hubsch A, et al. ¹⁸ Evaluation of the Vicorder, a novel cuff-based device	Vicorder
for the noninvasive estimation of central blood pressure. J Hypertens. 2013;31(1):77-85.	

 23. Pucci et al, unpublished* 24. Rossen NB, Laugesen E, Peters CD, et al.¹⁹ Invasive validation of arteriograph estimates of central blood pressure in patients with type 2 diabetes. <i>Am J Hypertens</i>. 2014;27(5):674-679. 	Omron HEM 9000AI Arteriograph
25. Sueta D, Yamamoto E, Tanaka T, et al. ²⁰ The accuracy of central blood pressure waveform by novel mathematical transformation of non-invasive measurement. <i>Int J Cardiol.</i> 2015;189:244-246.	Pasesa AVE-1500 (Shisei Datum)
 26. Smulyan H, Siddiqui DS, Carlson RJ, London GM, Safar ME.²¹ Clinical utility of aortic pulses and pressures calculated from applanated radial-artery pulses. <i>Hypertension</i>. 2003;42(2):150-155. 	Colin Medical Instruments (oscillometric)
27. Smulyan H, Sheehe PR, Safar ME. ²² A preliminary evaluation of the mean arterial pressure as measured by cuff oscillometry. <i>Am J Hypertens</i> . 2008;21(2):166-171.	Colin Medical Instruments (oscillometric)
28. Smulyan H, Mukherjee R, Sheehe PR, Safar ME. ²³ Cuff and aortic pressure differences during dobutamine infusion: a study of the effects of systolic blood pressure amplification. <i>Am Heart J</i> .	Colin Medical Instruments (oscillometric)
 2010;159(3):399-405. 29. Takazawa K, Kobayashi H, Shindo N, Tanaka N, Yamashina A.²⁴ Relationship between radial 	TM2740; Colin Medical Technology
and central arterial pulse wave and evaluation of central aortic pressure using the radial arterial pulse wave. <i>Hypertens Res.</i> 2007;30(3):219-228.	Co, Komaki, Japan)
30. Takazawa K, Kobayashi H, Kojima I, et al. ²⁵ Estimation of central aortic systolic pressure using late systolic inflection of radial artery pulse and its application to vasodilator therapy. <i>J</i>	Omron HEM-9000AI
<i>Hypertens</i> . 2012;30(5):908-916. 31. Weber T, Wassertheurer S, Rammer M, et al. ²⁶ Validation of a brachial cuff-based method for estimating central systolic blood pressure. <i>Hypertension</i> . 2011;58(5):825-832.	Mobil-o-graph, IEM

estimating central systolic blood pressure. *Hypertension*. 2011;58(5):825-832. *Unpublished study details are reported in Online Appendix 1 of Picone et al, J Am Coll Cardiol, 2017; 70(5):572-86.

Citation	Cuff blood pressure device
1. Berliner K, Yildiz M, Garnier B, Lee DH, Fujiy H. ²⁷ Blood Pressure	Auscultation (Baumanometer)
Measurements in Obese Persons - Comparison of Intra-Arterial and	
Auscultatory Measurements. Am J Cardiol. 1961;8(1):10-	
2. Bos et al, ⁴ Pseudohypertension and the measurement of blood pressure.	Mercury sphygmomanometer (Groups B, C). Hawksley
Hypertension. 1992;20(1):26-31.	Random Zero sphygmomanometer (Group D)
3. Cheng HM, Wang KL, Chen YH, et al. ⁵ Estimation of central systolic blood	VP-2000 Colin Corporation
pressure using an oscillometric blood pressure monitor. Hypertens Res.	
2010;33(6):592-599.	
4. Cheng et al, unpublished (methods identical to study 3.)	VP-2000 Colin Corporation
5. Ding FH, Li Y, Zhang RY, Zhang Q, Wang JG. ⁸ Comparison of the	Omron HEM 9000AI
SphygmoCor and Omron devices in the estimation of pressure amplification	
against the invasive catheter measurement. J Hypertens. 2013;31(1):86-93.	
6. Lin MM, Cheng HM, Sung SH, et al. ¹¹ Estimation of central aortic systolic	WatchBP Office, Microlife
pressure from the second systolic peak of the peripheral upper limb pulse	
depends on central aortic pressure waveform morphology. J Hypertens.	
2012;30(3):581-586.	
7. Melamed R, Johnson K, Pothen B, Sprenkle MD, Johnson PJ. ²⁸ Invasive	Oscillometric device
blood pressure monitoring systems in the ICU: influence of the blood-	
conserving device on the dynamic response characteristics and agreement with	
noninvasive measurements. Blood Press Monit. 2012;17(5):179-183.	
8. Muecke S, Bersten A, Plummer J. ²⁹ The mean machine; accurate non-	Oscar2
invasive blood pressure measurement in the critically ill patient. J Clin Monit	
Comput. 2009;23(5):283-297.	
9. Omboni S, Parati G, Groppelli A, Ulian L, Mancia G. ³⁰ Performance of the	
AM-5600 blood pressure monitor: comparison with ambulatory intra-arterial	

Online supplement Table S2. Studies included in the cuff versus invasive brachial BP sensitivity analyses.

pressure. J Appl Physiol (1985). 1997;82(2):698-703.

10. Picone DS, Schultz MG, Peng X, et al.³¹ Discovery of New Blood
Pressure Phenotypes and Relation to Accuracy of Cuff Devices Used in Daily
Clinical Practice. Hypertension. 2018;71(6):1239-47.
11. Pucci et al, unpublished*
12. Raftery EB, Ward AP.³² The indirect method of recording blood pressure.
Cardiovasc Res. 1968;2(2):210-218.
Sphygmocor Xcel (oscillometric BP Suntech Advantage NIBP)
London School of Hygiene mercury sphygmomanometer

*Unpublished study details are reported in Online Appendix 1 of Picone et al, J Am Coll Cardiol, 2017; 70(5):572-86.

	Difference	95% confidence interval	p-value
Cuff – invasive aortic			-
systolic blood pressure			
40 to 49 years	3.4	0.5 to 6.3	< 0.0001
50 to 59 years	1.2	-1.3 to 3.7	
60 to 69 years	-0.6	-3.1 to 1.8	
70 to 79 years	-2.7	-5.2 to -0.2	
80 to 89 years	-3.7	-7.0 to -0.4	
Cuff – invasive aortic			
diastolic blood pressure			
40 to 49 years	3.8	1.6 to 6.1	< 0.0001
50 to 59 years	5.4	3.4 to 7.4	
60 to 69 years	6.1	4.1 to 8.0	
70 to 79 years	7.9	5.9 to 10.0	
80 to 89 years	9.9	7.4 to 12.5	
Cuff – invasive aortic pulse			
pressure			
40 to 49 years	-0.4	-3.6 to 2.7	< 0.0001
50 to 59 years	-4.2	-7.0 to-1.4	
60 to 69 years	-6.7	-9.4 to-4.0	
70 to 79 years	-10.7	-13.4 to-7.9	
80 to 89 years	-13.6	-17.2 to-10.1	

Online supplement Table S3. Cuff blood pressure compared with invasive aortic systolic, diastolic and pulse pressure differences across decades of age.

The 'difference' column is the marginal effects of the mean calculated for cuff minus invasive aortic BP at each age decade. The number of subjects in each age decade is as follows: 40 to 49 years, n=168; 50 to 59 years, n=403; 60 to 69 years, n=550; 70 to 79 years, n=447; 80 to 89 years, n=106. Units for all BP data are mm Hg.

	Estimate	95% confidence interval	p-value
Cuff – invasive aortic systolic blood pressure			
• •	-1.7	-2.3 to -1.2	< 0.0001
Age category	-1./	-2.5 10 -1.2	<0.0001
Sex	4.8	3.4 to 6.1	< 0.0001
Cuff – invasive aortic			
diastolic blood pressure			
Age category	1.5	1.05 to 1.9	< 0.0001
Sex	1.9	0.9 to 2.9	0.00020
Cuff – invasive aortic pulse			
pressure			
Age category	-3.2	-3.8 to -2.6	< 0.0001
Sex	2.9	1.5 to 4.2	< 0.0001

Online supplement Table S4. Cuff blood pressure compared with invasive aortic systolic, diastolic and pulse pressure differences across decades of age and adjusted for sex.

Linear mixed modelling was used to account for participant clustering within individual studies. The age category refers to subjects grouped according to the following age decades: 40 to 49 years, n=166; 50 to 59 years, n=398; 60 to 69 years, n=537; 70 to 79 years, n=440; 80 to 89 years, n=106. There was no interaction between the age category and sex.

	Estimate	95% confidence interval	p-value
Cuff – invasive aortic			
systolic blood pressure			
Age category	-1.4	-2.0 to -0.8	< 0.0001
Invasive mean arterial pressure	-0.40	-0.44 to -0.36	< 0.0001
Heart rate	0.15	0.10 to 0.20	< 0.0001
Body mass index	0.18	0.055 to 0.31	0.0051
Cuff – invasive aortic			
diastolic blood pressure			
Age category	1.6	1.1 to 2.1	< 0.0001
Invasive mean arterial pressure	-0.13	-0.16 to -0.092	< 0.0001
Heart rate	-0.029	-0.071 to 0.013	0.18
Body mass index	-0.097	-0.20 to 0.0095	0.075
Cuff – invasive aortic puls	se		
pressure			
Age category	-3.0	-3.6 to -2.3	< 0.0001
Invasive mean arterial pressure	-0.28	-0.32 to -0.23	< 0.0001
Heart rate	0.18	0.13 to 0.24	< 0.0001
Body mass index	0.28	0.14 to 0.42	< 0.0001

Online supplement Table S5. Cuff blood pressure compared with invasive aortic systolic, diastolic and pulse pressure differences across decades of age, adjusted for mean arterial pressure, heart rate and body mass index.

Linear mixed modelling was used to account for participant clustering within individual studies. The age category refers to subjects grouped according to the following age decades: 40 to 49 years, n=150; 50 to 59 years, n=350; 60 to 69 years, n=443; 70 to 79 years, n=355; 80 to 89 years, n=84.

••	40 to 49 years	50 to 59 years	60 to 69 years	70 to 79 years	80 to 89 years
	40 to 49 years (n=71)	50 to 59 years (n=141)	60 to 69 years (n=159)	70 to 79 years (n=118)	80 to 89 years (n=31)
Subject characteristics	(m -7 x)	(m - x ix)	(II-IU))	(m=110)	(II - CI)
Age, years	45.4±3	54.8±3	64.0 [61.0 to 66.0]	74.6±3	81 [80.0 to 82.5]
Female sex, %*	21 (27)	36 (32)	47 (37)	29 (26)	1 (4)
Height, cm^{\dagger}	166.5±9.6	165.2±9.1	165.6±10.5	164.7±9.8	164.4±6.7
Weight, kg [‡]	82.3±20.6	81.0±21.5	79.4±17.4	72.6±14.8	67.3±11.5
Body mass index, kg/m ^{2§}	29.7±7.3	29.7±7.6	28.9±6.0	26.7±5.3	24.9±3.6
Heart rate, beats/min	69±13	67±11	68±12	66±12	63±9
Blood pressure					
Cuff systolic blood pressure	143±38	138±26	147±30	149±28	150±27
Cuff diastolic blood	85±21	82±15	78±16	78±13	75±13
pressure					
Cuff pulse pressure	58±20	56±17	69±22	71±22	75±25
Invasive brachial systolic	145±32	143±24	153±30	156±27	157±27
blood pressure					
Invasive brachial diastolic	80±18	76±14	73±15	71±13	66±11
blood pressure					
Invasive brachial pulse	65±18	67±17	80±22	84±21	91±25
pressure					

Online supplement Table S6. Sample characteristics across each decade of age for the cuff and invasive brachial blood pressure measurements.

Data are mean±standard deviation or median [interquartile range]. All blood pressure units are mm Hg. *n=433; $^{\dagger}n=477$; $^{\dagger}n=481$; $^{\$}n=477$; $^{\ddagger}n=364$.

Decade of age	n	Difference	95% confidence interval	p-value (trend)
Cuff – invasive brachial systolic blood pressure				
40 to 49 years	71	-4.2	-8.8 to 0.3	0.025
50 to 59 years	141	-7.1	-11.3 to -2.9	
60 to 69 years	159	-7.5	-11.7 to -3.4	
70 to 79 years	118	-8.1	-12.4 to -3.8	
80 to 89 years	31	-9.5	-15.1 to -3.9	
Cuff – invasive brachial diastolic blood pressure				
40 to 49 years	71	4.4	1.3 to 7.4	0.015
50 to 59 years	141	5.0	2.3 to 7.8	
60 to 69 years	159	4.6	1.9 to 7.3	
70 to 79 years	118	6.2	3.4 to 9.0	
80 to 89 years	31	9.3	5.6 to 13.1	
Cuff – invasive brachial pulse pressure				
40 to 49 years	71	-8.6	-12.9 to -4.2	< 0.0001
50 to 59 years	141	-12.1	-16.1 to -8.1	
60 to 69 years	159	-12.2	-16.1 to -8.2	
70 to 79 years	118	-14.3	-18.4 to -10.2	
80 to 89 years	31	-18.8	-24.2 to -13.5	

Online supplement Table S7. Unadjusted data for cuff blood pressure compared with invasive brachial systolic, diastolic and pulse pressure differences across 10-year age groups.

The 'difference' column is the marginal effects of the mean calculated for cuff minus invasive brachial BP at each age group. Units for all BP data are mm Hg.

SCA.			
	Estimate	95% confidence interval	p-value
Cuff – invasive aortic			
systolic blood pressure			
Age category	-0.58	-1.5 to 0.3	0.20
Sex	0.41	-1.7 to 2.6	0.71
Cuff – invasive aortic			
diastolic blood pressure			
Age category	1.1	0.45 to 1.7	0.00068
Sex	1.3	-0.22 to 2.8	0.096
Cuff – invasive aortic pulse			
pressure			
Age category	-1.6	-2.5 to -0.67	0.0006
Sex	-0.87	-3.1 to 1.3	0.44

Online supplement Table S8. Cuff blood pressure compared with invasive brachial systolic, diastolic and pulse pressure differences across decades of age and adjusted for sex.

Linear mixed modelling was used to account for participant clustering within individual studies. The age category refers to subjects grouped according to the following age decades: 40 to 49 years, n=56; 50 to 59 years, n=111; 60 to 69 years, n=128; 70 to 79 years, n=110; 80 to 89 years, n=28. There was no interaction between the age category and sex.

	Estimate	95% confidence interval	p-value
Cuff – invasive brachial			
systolic blood pressure			
Age category	-0.48	-1.5 to 0.53	0.35
Invasive mean arterial pressure	-0.33	-0.42 to -0.24	< 0.0001
Heart rate	0.022	-0.069 to 0.11	0.63
Body mass index	0.077	-0.17 to 0.33	0.55
Cuff – invasive brachial			
diastolic blood pressure			
Age category	1.2	0.47 to 1.96	0.0016
Invasive mean arterial pressure	-0.030	-0.097 to 0.034	0.37
Heart rate	0.036	-0.032 to 0.010	0.30
Body mass index	-0.29	-0.47 to -0.095	0.0030
Cuff – invasive brachial			
pulse pressure			
Age category	-1.7	-2.7 to -0.69	0.0011
Invasive mean arterial pressure	-0.30	-0.38 to -0.21	< 0.0001
Heart rate	-0.013	-0.10 to 0.078	0.78
Body mass index	0.35	0.096 to 0.60	0.0064

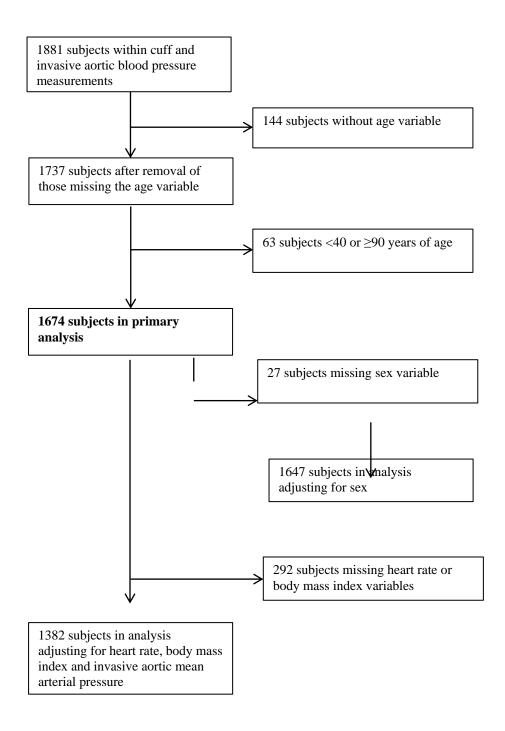
Online supplement Table S9. Cuff blood pressure compared with invasive brachial systolic, diastolic and pulse pressure differences across decades of age, adjusted for mean arterial pressure, heart rate and body mass index.

Linear mixed modelling was used to account for participant clustering within individual studies. The age category refers to subjects grouped according to the following age decades: 40 to 49 years, n=40; 50 to 59 years, n=86; 60 to 69 years, n=106; 70 to 79 years, n=79; 80 to 89 years, n=21.

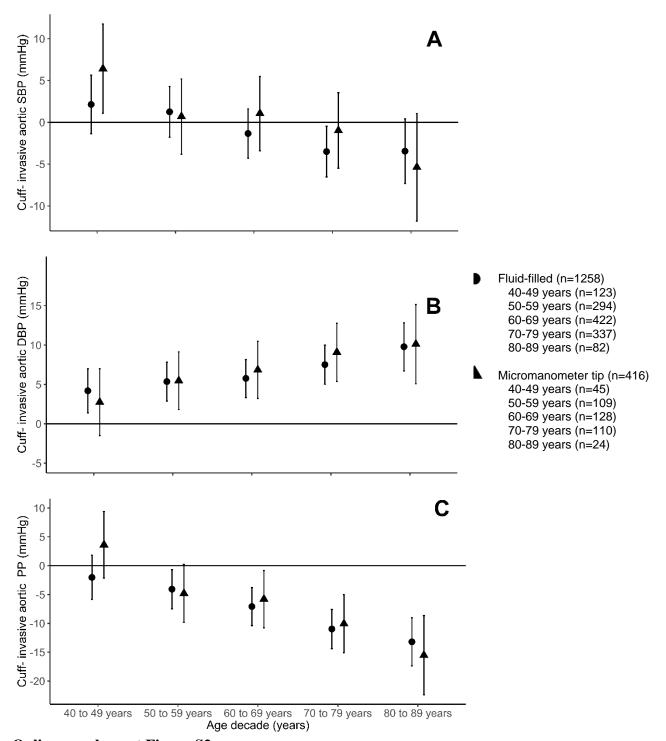
Online supplement Table S10. Influence of age on cuff blood pressure (BP) compared with invasive aortic BP when adjusted for the difference in invasive brachial and invasive aortic systolic, diastolic or pulse pressure (BP amplification).

	Cuff-aortic SBP	Cuff-aortic DBP	Cuff-aortic PP
Age	-1.1 [-2.1 to -0.2]	1.3 [0.56 to 2.0]	-2.5 [-3.4 to -1.5]
	P=0.018	P=0.00043	P<0.0001
Invasive brachial – invasive aortic blood pressure	0.47 [0.35 to 0.59]	0.68 [0.49 to 0.88]	0.42 [0.29 to 0.54]
	P<0.0001	P<0.0001	P<0.0001

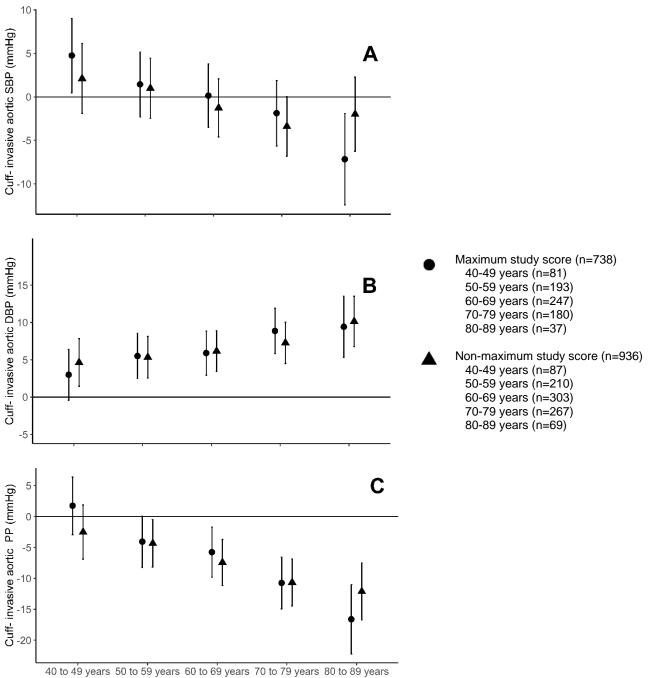
Data are estimate [95% confidence interval] and calculated using linear mixed modelling. SBP, systolic blood pressure; DBP, diastolic blood pressure; PP, pulse pressure.



Online supplement Figure S1. Flow of subjects according to the study inclusion criteria from the database of cuff and invasive aortic blood pressure measurements.

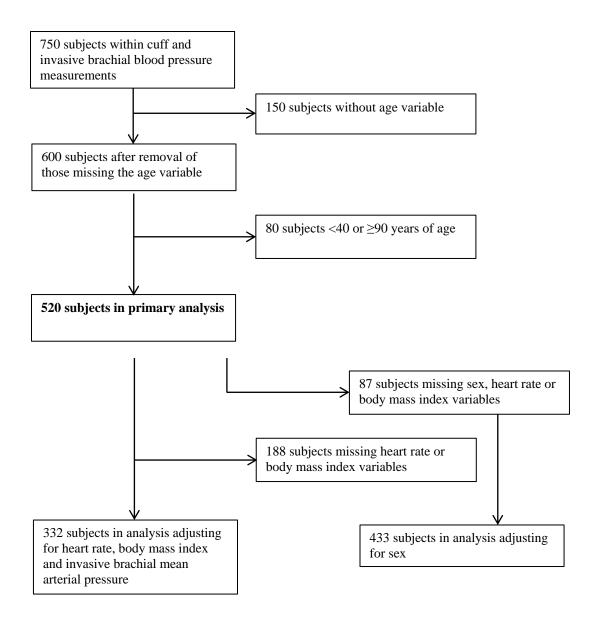


Online supplement Figure S2. Cuff blood pressure (BP) compared with invasive aortic systolic (SBP; A), diastolic (DBP; B) and pulse pressure (PP; C) measurements (y-axis) stratified by the type of catheter (fluid-filled or micromanometer tip) across decades of age (x-axis). Data are mean difference and 95% confidence interval (error bars). The number of subjects in each age and catheter type grouping is shown in the table on the middle right of the figure. Data above the solid horizontal zero line indicates cuff BP is higher than invasive aortic BP and vice versa below the zero line.

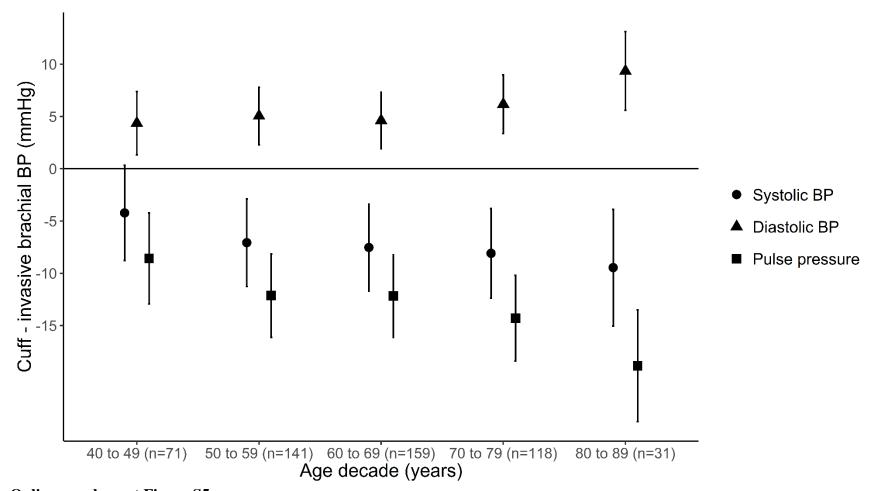


Age decade (years)

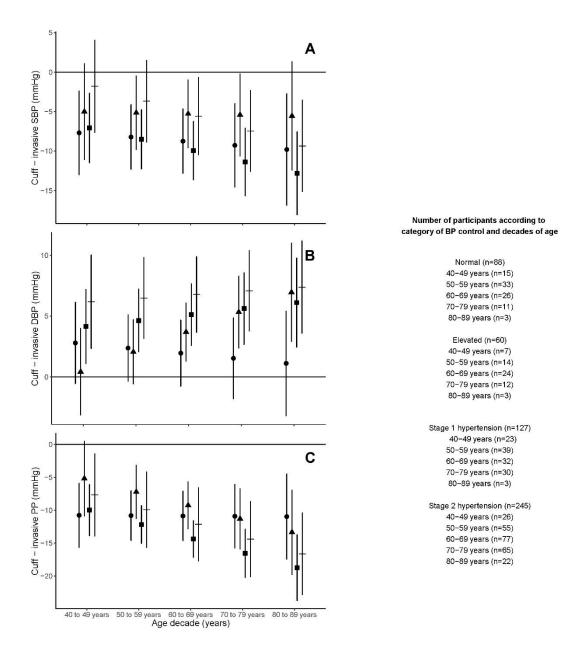
Online supplement Figure S3. Cuff blood pressure (BP) compared with invasive aortic systolic (SBP; A), diastolic (DBP; B) and pulse pressure (PP; C) measurements (y-axis) stratified by the study quality score (maximum or non-maximum score) across ten-year age groups (x-axis). Data are mean difference and 95% confidence interval (error bars). The number of subjects in each age and study quality score grouping is shown in the table on the middle right of the figure. Data above the solid horizontal zero line indicates cuff BP is higher than invasive aortic BP and vice versa below the zero line.



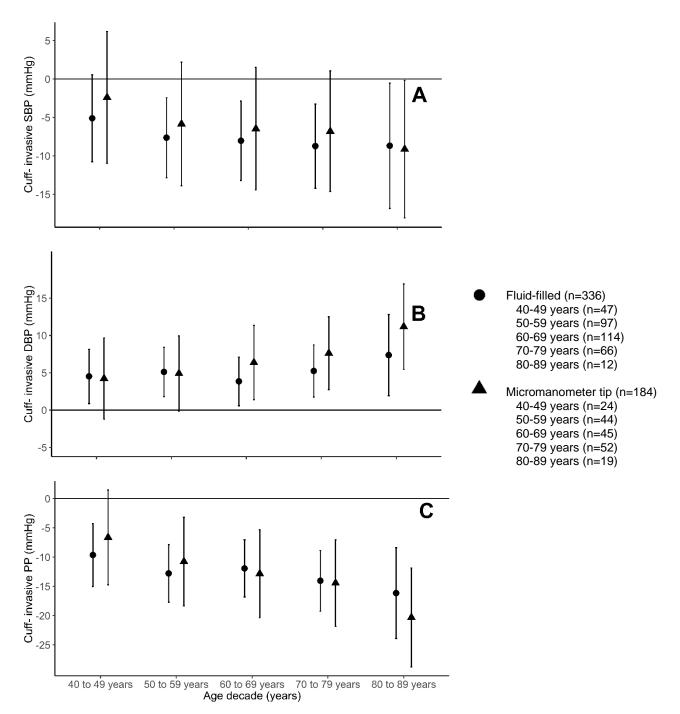
Online supplement Figure S4. Flow of subjects according to the study inclusion criteria from the database of cuff and invasive brachial blood pressure measurements.



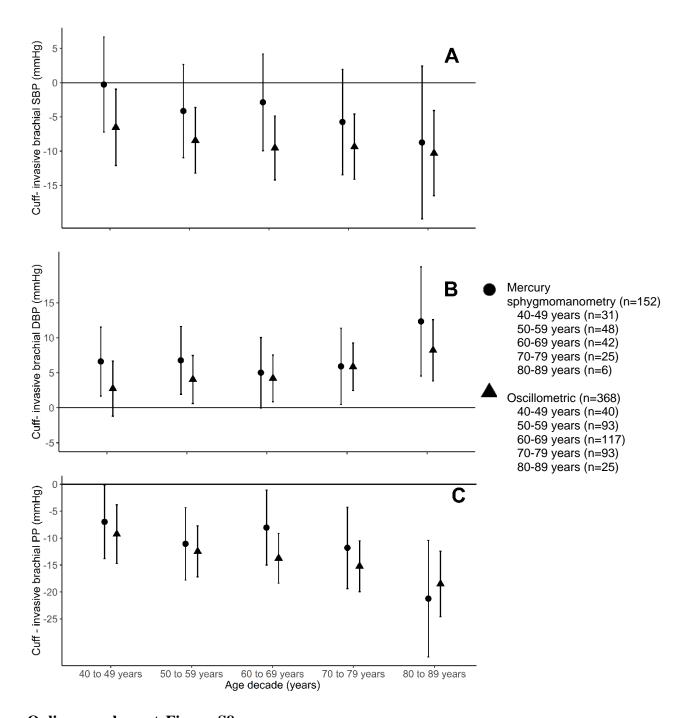
Online supplement Figure S5. Cuff blood pressure (BP) compared with invasive brachial systolic BP (red), diastolic BP (green) and pulse pressure (blue) measurements across age decades. Data are mean difference and 95% confidence interval (error bars). Data above the solid horizontal zero line indicates cuff BP is higher than invasive brachial BP and vice versa below the zero line. The trend for age related difference in cuff BP compared with invasive aortic BP was significant for each of BP variable, p=0.025 systolic BP, p=0.015 diastolic BP and p<0.0001 pulse pressure.



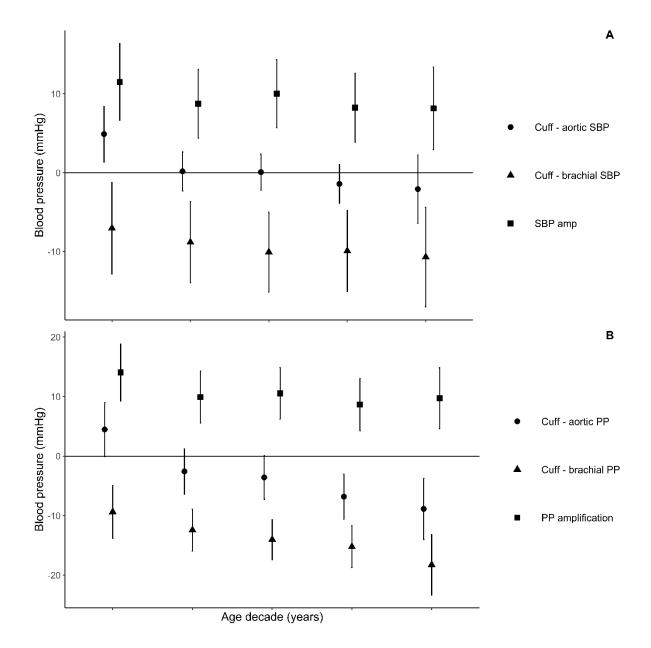
Online supplement Figure S6. Cuff blood pressure (BP) compared with invasive brachial systolic BP (SBP; A), diastolic BP (DBP; B) and pulse pressure (PP; C) measurements across decades of age and stratified according to the category of cuff BP control (according to the 2017 American Heart Association/American College of Cardiology arterial hypertension guidelines). Data are mean difference and 95% confidence interval (error bars). Data above the solid horizontal zero line indicates cuff BP is higher than invasive brachial BP and vice versa below the zero line. The number of subjects in each age and BP guideline grouping is shown in the table on the middle right of the figure. Within each BP category, there were only significant trends for the influence of age on cuff BP compared with invasive brachial BP for SBP in stage 2 hypertension, DBP in the elevated category and PP in stage 1 and 2 hypertension. Circles, normal BP; triangles, elevated BP; squares, stage 1 hypertension; crosses; stage 2 hypertension.



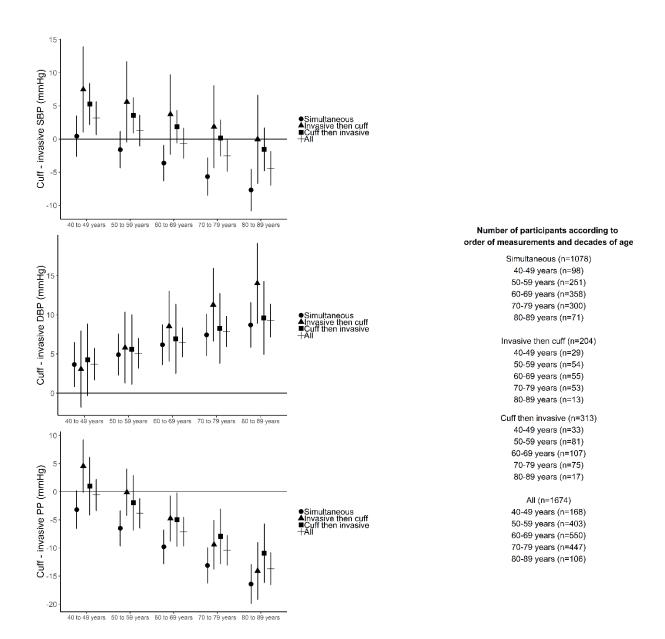
Online supplement Figure S7. Cuff blood pressure (BP) compared with invasive brachial systolic (SBP; A), diastolic (DBP; B) and pulse pressure (PP; C) measurements (y-axis) stratified by the type of catheter (fluid-filled or micromanometer tip) across tenyear age groups (x-axis). Data are mean difference and 95% confidence interval (error bars). The number of subjects in each age and catheter type grouping is shown in the table on the middle right of the figure. Data above the solid horizontal zero line indicates cuff BP is higher than invasive aortic BP and vice versa below the zero line.



Online supplement Figure S8. Cuff blood pressure (BP) compared with invasive brachial systolic (SBP; A), diastolic (DBP; B) and pulse pressure (PP; C) measurements (y-axis) stratified by the type of sphygmomanometer used (oscillometric or mercury) across decades of age (x-axis). Data are mean difference and 95% confidence interval (error bars). The number of subjects in each age and sphygmomanometer grouping is shown in the table on the middle right of the figure. Data above the solid horizontal zero line indicates cuff BP is higher than invasive brachial BP and vice versa below the zero line.



Online supplement Figure S9. The difference between cuff systolic blood pressure (SBP) and invasive aortic (circles) or invasive brachial (triangles) SBP is plotted alongside SBP amplification (SBP amp; invasive brachial SBP– invasive aortic SBP; squares) in panel A. The difference between cuff pulse pressure (PP) and invasive aortic (circles) or invasive brachial (triangles) PP isplotted with PP amplification (invasive brachial PP – invasive aortic PP; squares) in panel B. Data are mean and 95% confidence interval (error bars). Data above the solid horizontal zero line indicates cuff BP is higher than invasive BP and vice versa below the zero line. Data is from a subset of 372 subjects.



Online supplement Figure S10. Cuff blood pressure (BP) compared with invasive aortic systolic BP (SBP; A), diastolic BP (DBP; B) and pulse pressure (PP; C) measurements across decades of age and stratified according to the order of BP measurements. Data are mean difference and 95% confidence interval (error bars). Circles, cuff and invasive BP measured simultaneously; triangles, invasive BP measured just prior to cuff BP; squares, cuff BP measured just prior to invasive BP; crosses; all data together.

Online supplement references

- 1. Aakhus S, Torp H, Haugland T, Hatle L. Noninvasive estimates of aortic root pressures external subclavian arterial pulse tracing calibrated by oscillometrically determined brachial arterial pressures. *Clin. Physiol.* 1993;13:573-586
- 2. Bhatt SD, Hinderliter AL, Stouffer GA. Influence of sex on the accuracy of oscillometric-derived blood pressures. *J. Clin. Hypertens.* 2011;13:112-119
- 3. Borow KM, Newburger JW. Noninvasive estimation of central aortic pressure using the oscillometric method for analyzing systemic artery pulsatile blood flow: Comparative study of indirect systolic, diastolic, and mean brachial artery pressure with simultaneous direct ascending aortic pressure measurements. *Am. Heart J.* 1982;103:879-886
- 4. Bos WJW, Van Goudoever J, Wesseling KH, Rongen GAPJM, Hoedemaker G, Lenders JWM, Van Montfrans GA. Pseudohypertension and the measurement of blood pressure. *Hypertension*. 1992;20:26-31
- 5. Cheng HM, Wang KL, Chen YH, Lin SJ, Chen LC, Sung SH, Ding PY, Yu WC, Chen JW, Chen CH. Estimation of central systolic blood pressure using an oscillometric blood pressure monitor. *Hypertens. Res.* 2010;33:592-599
- 6. Costello BT, Schultz MG, Black JA, Sharman JE. Evaluation of a brachial cuff and suprasystolic waveform algorithm method to noninvasively derive central blood pressure. *Am. J. Hypertens.* 2015;28:480-486
- 7. Cremer A, Butlin M, Codjo L, Coulon P, Ranouil X, Joret C, Coste P, Asmar R, Avolio A, Gosse P. Determination of central blood pressure by a noninvasive method (brachial bp and qkd interval). *J. Hypertens*. 2012;30:1533-1539
- 8. Ding FH, Li Y, Zhang RY, Zhang Q, Wang JG. Comparison of the sphygmocor and omron devices in the estimation of pressure amplification against the invasive catheter measurement. *J. Hypertens.* 2013;31:86-93
- 9. Jeon WK, Kim MA, Kim HL, Ki YJ, Rhee TM, Lim WH, Seo JB, Kim SH, Zo JH. Association between aortic knob width and invasively measured aortic pulse pressure. *Blood Press. Monit.* 2018
- Laugesen E, Rossen NB, Peters CD, Maeng M, Ebbehoj E, Knudsen ST, Hansen KW, Botker HE, Poulsen PL. Assessment of central blood pressure in patients with type 2 diabetes: A comparison between sphygmocor and invasively measured values. *Am. J. Hypertens.* 2014;27:169-176
- 11. Lin MM, Cheng HM, Sung SH, Liao CF, Chen YH, Huang PH, Chen CH. Estimation of central aortic systolic pressure from the second systolic peak of the peripheral upper limb pulse depends on central aortic pressure waveform morphology. *J. Hypertens.* 2012;30:581-586
- 12. Nagle FJ, Naughton J, Balke B. Comparisons of direct and indirect blood pressure with pressure-flow dynamics during exercise. *J. Appl. Physiol.* 1966;21:317-320
- 13. Nakagomi A, Okada S, Shoji T, Kobayashi Y. Aortic pulsatility assessed by an oscillometric method is associated with coronary atherosclerosis in elderly people. *Blood Press.* 2016:1-8
- 14. Ohte N, Saeki T, Miyabe H, Sakata S, Mukai S, Hayano J, Niki K, Sugawara M, Kimura G. Relationship between blood pressure obtained from the upper arm with a cuff-type sphygmomanometer and central blood pressure measured with a catheter-tipped micromanometer. *Heart Vessels*. 2007;22:410-415
- 15. Ott C, Haetinger S, Schneider MP, Pauschinger M, Schmieder RE. Comparison of two noninvasive devices for measurement of central systolic blood pressure with invasive measurement during cardiac catheterization. *J. Clin. Hypertens.* (*Greenwich*). 2012;14:575-579

- 16. Park CM, Korolkova O, Davies JE, Parker KH, Siggers JH, March K, Tillin T, Chaturvedi N, Hughes AD. Arterial pressure: Agreement between a brachial cuffbased device and radial tonometry. *J. Hypertens.* 2014;32:865-872
- 17. Pereira T, Maldonado J, Coutinho R, Cardoso E, Laranjeiro M, Andrade I, Conde J. Invasive validation of the complior analyse in the assessment of central artery pressure curves: A methodological study. *Blood Press. Monit.* 2014;19:280-287
- Pucci G, Cheriyan J, Hubsch A, Hickson SS, Gajendragadkar PR, Watson T, O'Sullivan M, Woodcock-Smith J, Schillaci G, Wilkinson IB, McEniery CM. Evaluation of the vicorder, a novel cuff-based device for the noninvasive estimation of central blood pressure. J. Hypertens. 2013;31:77-85
- 19. Rossen NB, Laugesen E, Peters CD, Ebbehoj E, Knudsen ST, Poulsen PL, Botker HE, Hansen KW. Invasive validation of arteriograph estimates of central blood pressure in patients with type 2 diabetes. *Am. J. Hypertens.* 2014;27:674-679
- 20. Sueta D, Yamamoto E, Tanaka T, Hirata Y, Sakamoto K, Tsujita K, Kojima S, Nishiyama K, Kaikita K, Hokimoto S, Jinnouchi H, Ogawa H. The accuracy of central blood pressure waveform by novel mathematical transformation of non-invasive measurement. *Int. J. Cardiol.* 2015;189:244-246
- 21. Smulyan H, Siddiqui DS, Carlson RJ, London GM, Safar ME. Clinical utility of aortic pulses and pressures calculated from applanated radial-artery pulses. *Hypertension*. 2003;42:150-155
- 22. Smulyan H, Sheehe PR, Safar ME. A preliminary evaluation of the mean arterial pressure as measured by cuff oscillometry. *Am. J. Hypertens.* 2008;21:166-171
- 23. Smulyan H, Mukherjee R, Sheehe PR, Safar ME. Cuff and aortic pressure differences during dobutamine infusion: A study of the effects of systolic blood pressure amplification. *Am. Heart J.* 2010;159:399-405
- 24. Takazawa K, Kobayashi H, Shindo N, Tanaka N, Yamashina A. Relationship between radial and central arterial pulse wave and evaluation of central aortic pressure using the radial arterial pulse wave. *Hypertens. Res.* 2007;30:219-228
- 25. Takazawa K, Kobayashi H, Kojima I, Aizawa A, Kinoh M, Sugo Y, Shimizu M, Miyawaki Y, Tanaka N, Yamashina A, Avolio A. Estimation of central aortic systolic pressure using late systolic inflection of radial artery pulse and its application to vasodilator therapy. *J. Hypertens.* 2012;30:908-916
- 26. Weber T, Wassertheurer S, Rammer M, Maurer E, Hametner B, Mayer CC, Kropf J, Eber B. Validation of a brachial cuff-based method for estimating central systolic blood pressure. *Hypertension*. 2011;58:825-832
- 27. Berliner K, Yildiz M, Garnier B, Lee DH, Fujiy H. Blood pressure measurements in obese persons comparison of intra-arterial and auscultatory measurements. *Am. J. Cardiol.* 1961;8:10-&
- 28. Melamed R, Johnson K, Pothen B, Sprenkle MD, Johnson PJ. Invasive blood pressure monitoring systems in the icu: Influence of the blood-conserving device on the dynamic response characteristics and agreement with noninvasive measurements. *Blood Press. Monit.* 2012;17:179-183
- 29. Muecke S, Bersten A, Plummer J. The mean machine; accurate non-invasive blood pressure measurement in the critically ill patient. *J. Clin. Monit. Comput.* 2009;23:283-297
- 30. Omboni S, Parati G, Groppelli A, Ulian L, Mancia G. Performance of the am-5600 blood pressure monitor: Comparison with ambulatory intra-arterial pressure. *J Appl Physiol* (1985). 1997;82:698-703
- 31. Picone DS, Schultz MG, Peng X, Black JA, Dwyer N, Roberts-Thomson P, Chen CH, Cheng HM, Pucci G, Wang JG, Sharman JE. Discovery of new blood pressure

phenotypes and relation to accuracy of cuff devices used in daily clinical practice. *Hypertension*. 2018;71:1239-1247 Raftery EB, Ward AP. The indirect method of recording blood pressure. *Cardiovasc*.

32. Res. 1968;2:210-218