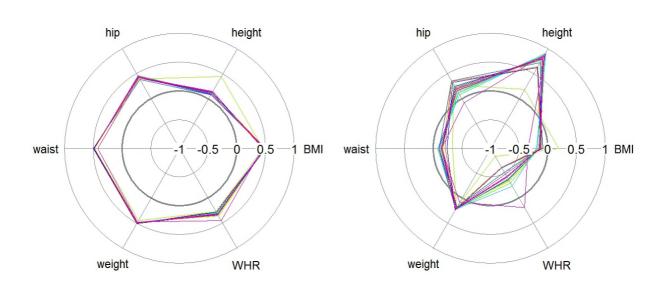
Supplementary Figures

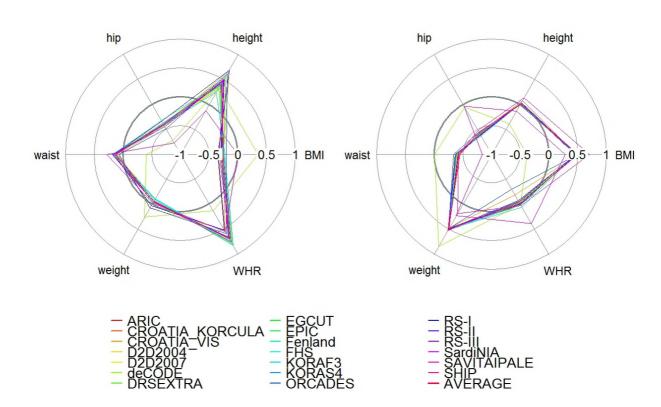
Supplementary Figure 1: Results of the pilot analysis.

(1) Study specific loadings for the PCs in the pilot analysis.

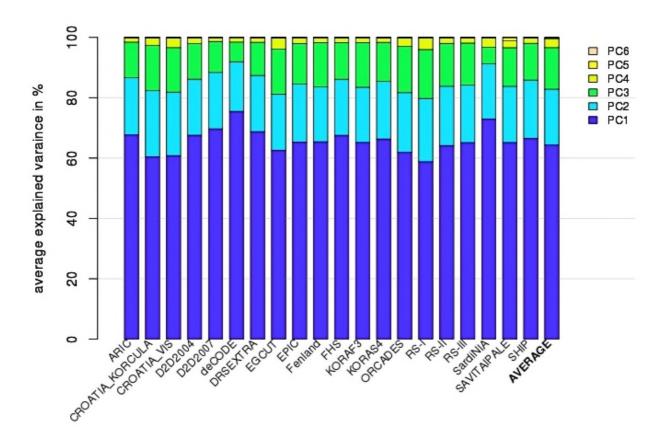
avPC1 avPC2



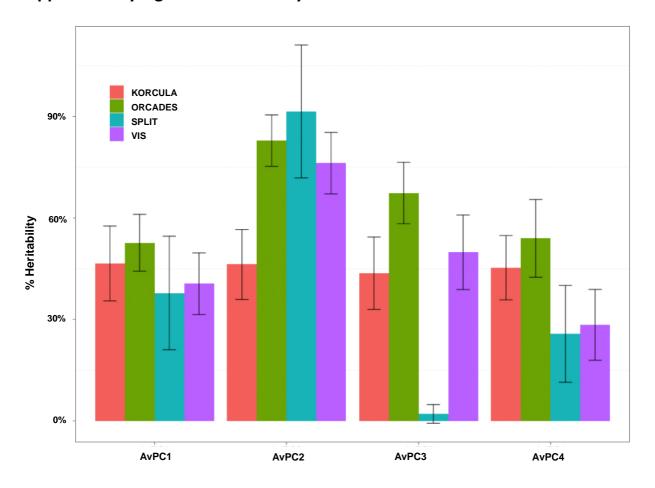
avPC3 avPC4



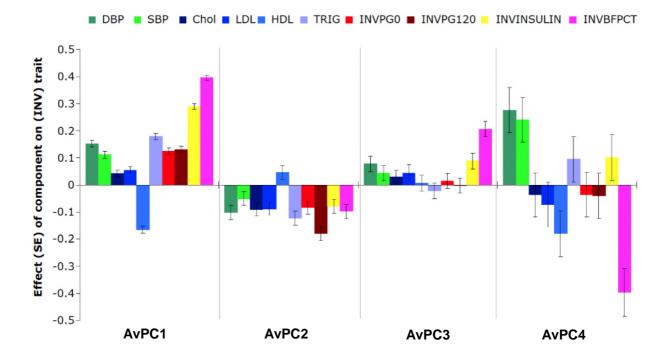
(2) Explained variance of PCs in each study of the pilot analysis.



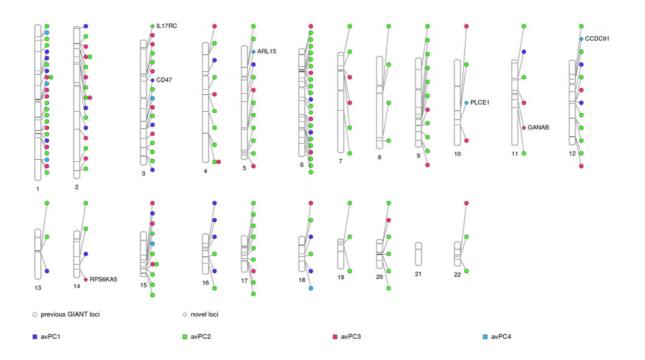
Supplementary Figure 2: Heritability of AvPCs.



Supplementary Figure 3: Correlation of avPCs with clinical traits. Association results for n=1,402 persons from the from FENLAND study.

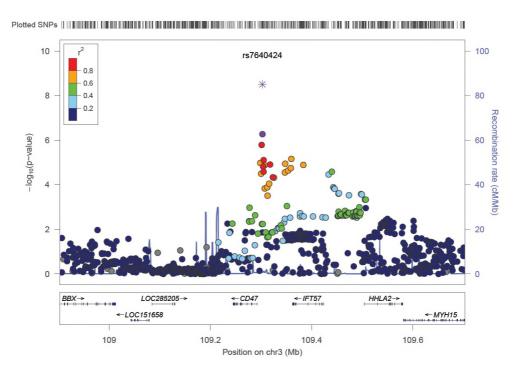


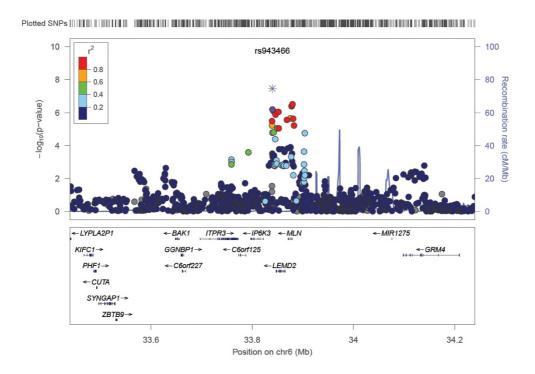
Supplementary Figure 4: Phenograms of all 189 genome-wide significant loci for avPCs. All genome wide significantly associated loci (promising p-value in the first stage meta analysis ($<5x10^{-6}$) and genome-wide significant in first and second stage combined analysis ($<5x10^{-8}$)) with one of the avPCs. Different colours are used for the four avPCs. Novel loci are highlighted by a diamond shape and the name of the nearest gene. (PhenoGram was generated with the online tool: http://visualization.ritchielab.psu.edu/phenograms/plot.)



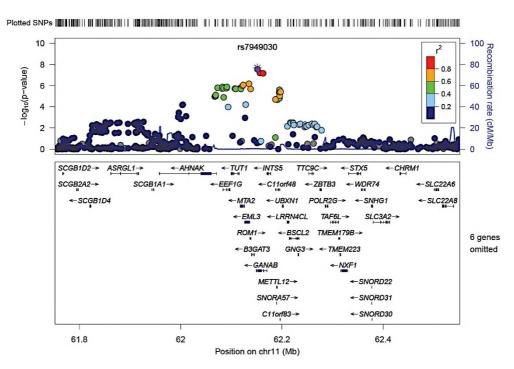
Supplementary Figure 5: Local association plots for novel loci. Local association plots of the six loci that were significantly associated (promising p-value in the first stage meta analysis (<5x10⁻⁶) and genome-wide significant in first and second stage combined analysis (<5x10⁻⁸)) with an avPC of body shape and independent of findings on BMI, WHR or height of previous GIANT analyses. In the plots the p-values of the first stage meta analyses are presented.

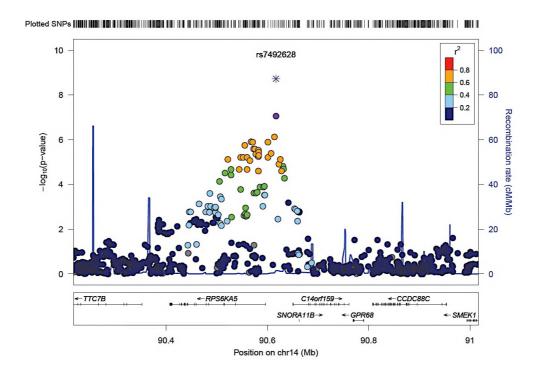
(1) avPC1



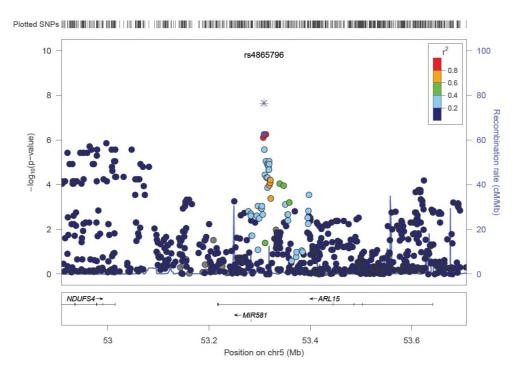


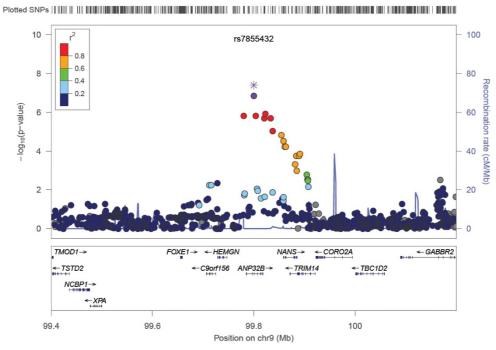
(2) avPC3



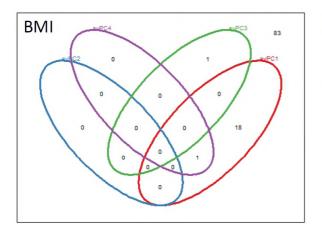


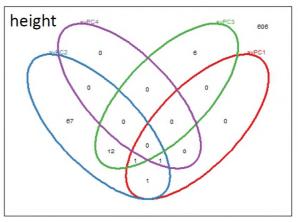
(4) avPC4

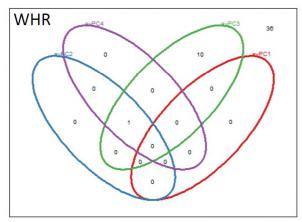


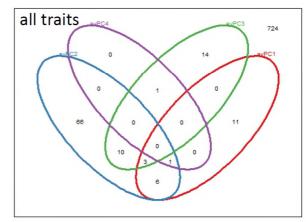


Supplementary Figure 6: Venn Diagram of known anthropometric loci in body shape results. The Venn Diagrams report the overlap of loci that were reported in the latest GIANT meta analyses on one trait of BMI, height and WHR or for any of those traits (all traits) and identified for an avPC. In the upper right corner of each diagram the number of loci associated with the trait is given that were not identified for any avPC. A locus is regarded to the same if the lead SNPs are in LD > 0.8 and less than 500kb away from each other.



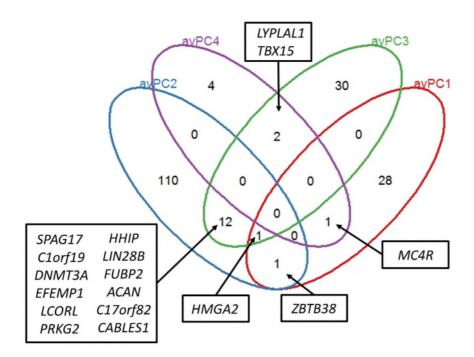






Supplementary Figure 7: Loci that were genome-wide significant for more than one avPC.

This Venn diagram shows which loci were significantly associated with more than one avPC. Loci of two avPCs are considered to overlap if the best genomewide significant SNPs per locus are in strong LD ($R^2 > 0.8$).



Supplement Tables:

Supplementary Table 1: Studies analysed in the pilot study. All studies are population based and are unrelated.

Study	N			
ARIC	9,713			
CROATIA_KORCULA	530			
CROATIA_VIS	518			
D2D2004	2,429			
D2D2007	2,711			
deCODE	4,779			
DRSEXTRA	1,408			
EGCUT	35,125			
EPIC	2,390			
Fenland	1,402			
FHS	1,659			
KORAF3	1,605			
KORAS4	1,813			
METSIM	0			
ORCADES	312			
RS-I	5,974			
RS-II	1,911			
RS-III	1,927			
SardiNIA	887			
SAVITAIPALE	1,194			
SHIP	4,068			
total	82,355			

Supplementary Table 2: Loadings and explained variance of the avPCs. The average loadings that are used for calculation of the avPVs are given in this table. For each avPC the average of explained variance is given in this table.

	avPC1	avPC2	avPC3	avPC4	avPC5	avPC6
BMI	-0.473	-0.128	-0.284	0.504	0.635	-0.148
Height	-0.131	0.803	0.513	0.018	0.265	-0.062
Hip	-0.444	0.197	-0.399	-0.583	0.110	0.503
Waist	-0.488	-0.159	0.138	-0.424	-0.163	-0.715
Weight	-0.486	0.186	-0.056	0.475	-0.690	0.155
WHR	-0.297	-0.490	0.689	-0.019	0.105	0.432
Explained	64.37%	18.46%	13.79%	2.97%	0.26%	0.15%
variance						
Explained		99.5	9%			
variance avPC1-						
avPC4						

Supplementary Table 3: Number of studies and individuals per analysis stage.

For each stage the maximal number of individuals is given that are used for analysis on body shape avPCs. In the second stage studies with genome wide data are analysed as well as studies with Metabochip data (N).

	Number of studies	Number of individuals
1st stage	43	133,376
2nd stage	22 (metabochip =12)	39,904 (metabochip= 32,170)
1st + 2nd combined	65	173,278

Supplementary Table 4: Summary of genome-wide significant, promising and novel loci.

	Promising Loci	Genome-Wide Significant	Novel
avPC1	56	31	2
avPC2	205	124	0
avPC3	89	45	2
avPC4	35	7	2
	385	207	6

Supplementary Table 5: Loci that were genome wide significant for more than one avPC.

Supplementary Table 5a: Loci that were genome wide significant for two avPCs. Loci of two avPCs are considered to be the same if the best genome wide significant SNPs per locus are in LD > 0.8. (Genome wide significant as defined in the text: promising p-value in the first stage meta analysis (<5x10-6) and genome wide significant in first and second stage combined analysis (<5x10-8).)

avPCs	snp1	snp2		pos. snp1	pos. snp2	distance	LD	next gene
avPC1,avPC2	rs1582874	rs724016	3	142,597,909	142,588,260	9,649	1.00	ZBTB38
avPC1,avPC4	rs6567160	rs476828	18	55,980,115	56,003,567	23,452	1.00	MC4R
avPC3,avPC4	rs10923724	rs10923712	1	119,348,365	119,306,957	41,408	0.86	TBX15
avPC3,avPC4	rs2791550	rs2605100	1	217,721,992	217,710,847	11,145	0.94	LYPLAL1
avPC2,avPC3	rs7536458	rs7536458	1	118,666,125	118,666,125	0	1.00	SPAG17
avPC2,avPC3	rs1046934	rs2274432	1	182,290,152	182,287,568	2,584	1.00	C1orf19
avPC2,avPC3	rs2289195	rs2289195	2	25,316,987	25,316,987	0		
avPC2,avPC3	rs3791675	rs3791675	2	55,964,813	55,964,813	0	1.00	EFEMP1
avPC2,avPC3	rs6853216	rs4057984	4	17,579,753	17,566,298	13,455	1.00	LCORL
avPC2,avPC3	rs1975474	rs1115919	4	82,397,961	82,392,421	5,540	0.86	PRKG2
avPC2,avPC3	rs7689420	rs7689420	4	145,787,802	145,787,802	0	1.00	HHIP
avPC2,avPC3	rs314263	rs7759938	6	105,499,438	105,485,647	13,791	0.95	LIN28B
avPC2,avPC3	rs7466269	rs7021911	9	132,453,905	132,490,839	36,934	0.91	FUBP3

avPC2,avPC3	rs2280470	rs2280470	15	87,196,630	87,196,630	0 1.0	00 ACAN
avPC2,avPC3	rs2079795	rs9892365	17	56,851,431	56,846,166	5,265 1.0	00 <i>C17orf82</i>
avPC2,avPC3	rs4239437	rs4239436	18	18,986,222	18,985,928	294 1.0	00 CABLES1

Supplementary Table 5b: Loci that were genome wide significant for three avPCs. Loci of two avPCs are considered to be the same if the best genome wide significant SNPs per locus are in LD > 0.9. (Genome wide significant as defined in the text: promising p-value in the first stage meta analysis (<5x10-6) and genome wide significant in first and second stage combined analysis (<5x10-8).)

avPCs	snp1	snp2	snp3	chr	pos.	distance (snp1-	LD (snp1-2/snp1-	next
					SNP1/SNP2/SNP3	2/snp1-3/snp2-3)	3/snp2-3)	gene
avPC1,avPC2,	rs7970350	rs1351394	rs8756	12	64,646,431/	8338/412/7926	0.91/0.89/0.97	HMGA2
avPC3					64,638,093/			
					64,646,019			

Supplement Notes:

AUTHOR CONTRIBUTIONS

Ruth J.F. Loos, Martina Müller-Nurasyid lead the steering committee and oversaw the consortium. The writing group consisted of Janina S Ried, Janina Jeff, Audrey Y Chu, Jennifer L Bragg-Gresham, Jenny van Dongen, Jennifer E Huffman, Martina Müller-Nurasyid, and Ruth JF Loos. The method development for the PCA approach was conducted by Janina S Ried, Martina Müller-Nurasyid. Data cleaning and preparation was performed by Janina S Ried, Jenny van Dongen, and Jennifer E Huffman. The GWAS and Metabochip Meta-analyses and follow-up analyses were carriedout by Janina S Ried, Janina Jeff, Audrey Y Chu, Jennifer L Bragg-Gresham, Jenny van Dongen, and Jennifer E Huffman.

The data analysis was a collective effort by Tarunveer Singh Ahluwalia, Eva Albrecht, Traci M Bartz, John Blangero, Jennifer L Bragg-Gresham, Gemma Cadby, Daniel I. Chasman, Charleston W.K. Chiang, L Adrienne Cupples, Niina Eklund, Joel Eriksson, Tõnu Esko, Teresa Ferreira, Krista Fischer, Anuj Goel, Mathias Gorski, Mariaelisa Graff, Caroline Hayward, Nancy L Heard-Costa, Frank Hu, Jennifer E Huffman, David Hunter, Aaron Isaacs, Anne U Jackson,

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Genotyping of contributing studies was performed by Cristina Barlassina, John Beilby, Claire Bellis, John Blangero, Lori L Bonnycastle, Marcel Bruinenberg, Harry Campbell, John C Chambers, Daniel I. Chasman, Yii-Der Ida Chen, Peter S Chines, Francesca D'Avila, Maria Dimitriou, Tõnu Esko, Nele Friedrich, Anette Prior Gjesing, Harald Grallert, Niels Grarup, Marie Neergaard Harder, Catharina A Hartman, Andrew Tym Hattersley, Caroline Hayward, Andrew A Hicks, Georg Homuth, Frank Hu, Jennie Hui, David Hunter, Pirro G Hysi, Shapour Jalilzadeh, Marjo-Riitta Jarvelin, Johanne M Justesen, Leena Kinnunen, Jaspal S Kooner, Peter Kovacs, Diana Kuh, Theodosios Kyriakou, Claudia Langenberg, Terho Lehtimäki, Mattias Lorentzon, Wendy L McArdle, Mark I McCarthy, Carolina Medina-Gomez, Lili Milani, Narisu Narisu, Kari E North, Jeffrey R O'Connell, Claes Ohlsson, Ken K Ong, Ben A. Oostra, Aarno Palotie, Lavinia Paternoster, Lu Qi, Nigel W Rayner, Rasmus Ribel-Madsen, Marcus Richards, Fernando Rivadeneira, Lynda M. Rose, Sylvain Sebert, Lorraine Southam, Kathleen Stirrups, Michael Stumvoll, Morris A Swertz, Amy J Swift, Ioanna Tachmazido, Alexander Teumer, Anke Tönjes, André G. Uitterlinden, Pim van der Harst, Cornelia M. van Duijn, Cristina Venturini, Veronique Vitart, Uwe Völker, Nicholas J Wareham, Elisabeth Widén, James F Wilson, and Andrew Wong.

Phenotype coordination of contributing studies was a collective effort by the following: Alexessander Couto Alves, Stephan JL Bakker, Matthias Blüher, John C Chambers, Daniel I. Chasman, Daniele Cusi, George Dedoussis, Joel Eriksson, Aliki-Eleni Farmaki, Nita G Forouhi, Caroline S Fox, Ron T Ganzevoort, Nicola Glorioso, Jürgen Gräßler, Jagvir Grewal, Catharina A Hartman, Maija Hassinen, Andrew Tym Hattersley, Aki S Havulinna, Iris M Heid, Hans Hillege, Oddgeir Holmen, Frank Hu, David Hunter, Lise Lotte Husemoen, Pirro G Hysi, Till Ittermann, Alan L James, Eero Jokinen, Torben Jorgensen, Mika Kähönen, Maria Karaleftheri, Ivana Kolcic, Ishminder K Kooner, Jaspal S Kooner, Peter Kovacs, Diana Kuh, Jari Lahti, Tomi Laitinen, Alexandra M. Lewin, Peter Lichtner, Jaana Lindström, Allan Linneberg, Roberto Lorbeer, Mattias Lorentzon, Reedik Mägi, Massimo Mangino, Satu Männistö, Paolo Manunta, Carolina Medina-Gomez, Andres Metspalu, Rebecca Mills, Gabriele Müller, Arthur W Musk, Kari E North, Claes Ohlsson, Lyle J Palmer, Annette Peters, Irene Pichler, Maria G Pilia, Ozren Polašek, Inga Prokopenko, Bruce M. Psaty, Lu Qi, Olli T Raitakari, Nigel W Rayner, Marcus Richards, Fernando Rivadeneira, Lynda M. Rose, Igor Rudan, Veikko Salomaa, Salome Scholtens, Alan R Shudiner, Thorkild I. A. Sorensen, Ronald P Stolk, David P Strachan, Heather M Stringham, Morris A Swertz, Anke Tönjes, Angelo Tremblay, Emmanouil Tsafantakis, André G. Uitterlinden, Peter J Van der Most, Liesbeth Vandenput,

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