Gene	$\log_2 FC$	Gene	$\log_2 FC$	Gene	$\log_2 FC$
WIF1	-1,814	CCND2	+3,376	FZD2	+1,536
FGF4	-1,664	CXXC4	+2,976	APC	+1,366
PITX2	-1,174	SFRP4	+2,826	KREMEN1	+1,236
WNT6	-1,054	WNT2B	+2,096	FZD8	+1,186
DVL1	-1,014	SFRP1	+1,926	FZD4	+1,086
		DKK3	+1,766	PRICKLE1	+1,066

Table S1. Stiffness-dependent mRNA expression profile of members of the Wnt/beta-catenin signaling pathway in ARPE-19 cells (RT² Profiler PCR Array). Declaration of differences in expression between soft ($E = 30 \,\mathrm{kPa}$) and stiff ($E = 80 \,\mathrm{kPa}$) substrate as $\log_2 FC$. Cutoff of genes shown based on $|\log_2 FC| \ge 1$. Resulting selection of 17 genes with a change in expression out of 84 investigated genes. Order based on $\log_2 FC$. Normalization on *ACTB*, *B2M*, *GAPDH*, *HPRT1* and *RPLP0* as endogenous controls. Expression data from n = 1 experiment.

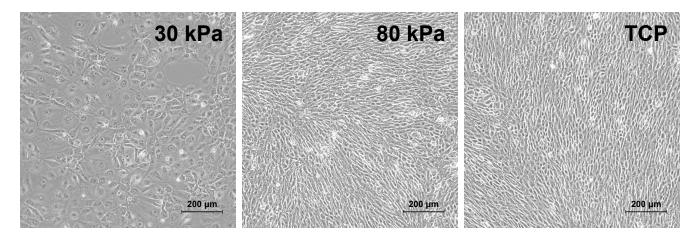


Figure S1. Stiffness-dependent morphology of ARPE-19 cells after three weeks of culture on substrates of different Young's modulus (phase contrast microscopy). Morphology of cells on soft ($E = 30 \,\mathrm{kPa}$) and stiff ($E = 80 \,\mathrm{kPa}$) substrate as well as on standard tissue culture plastic (TCP) is shown.

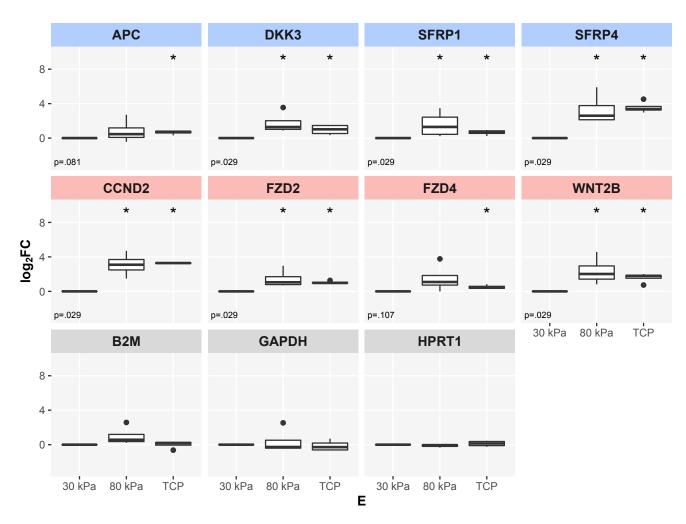


Figure S2. Stiffness-dependent transcription of members of the Wnt/beta-catenin signaling pathway in ARPE-19 cells for validation of qPCR array data (qPCR). Expression data from n=4 independent experiments, visualization as $\log_2 FC$ with $E=30\,\mathrm{kPa}$ as reference. Coloring of antagonistic (blue) and agonistic (red) impact on the activity of the Wnt/beta-catenin signaling pathway. Normalization on B2M, GAPDH and HPRT1 as endogenous controls (gray). Statistical significance between each two interventions was assessed by Wilcoxon-Mann-Whitney test with $E=30\,\mathrm{kPa}$ as reference. $p\le0.05$ (*), $p\le0.01$ (***) and $p\le0.001$ (***). Global p-values were determined using Kruskal-Wallis test. Benjamini-Hochberg procedure was applied for multiple testing correction.

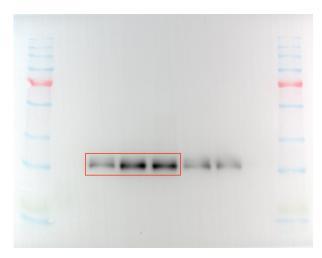


Figure S3. Stiffness-dependent expression of SFRP1 ($\approx 35 \, \text{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \, \text{kPa}$, $E = 80 \, \text{kPa}$ and TCP (red rectangle).

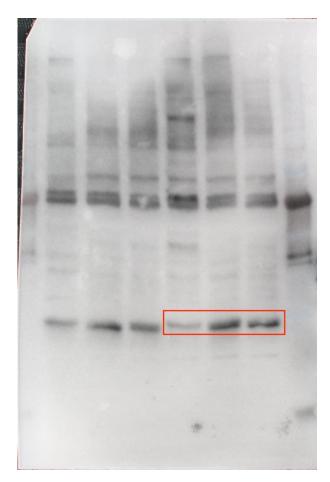


Figure S5. Stiffness-dependent expression of Wnt-2b ($\approx 41 \, \text{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \, \text{kPa}$, $E = 80 \, \text{kPa}$ and TCP (red rectangle).

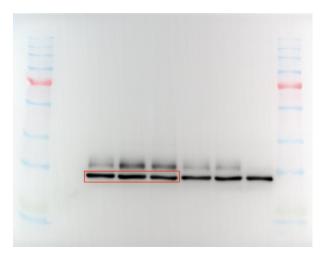


Figure S4. Loading control GAPDH ($\approx 38 \, \text{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \, \text{kPa}$, $E = 80 \, \text{kPa}$ and TCP (red rectangle).



Figure S6. Loading control GAPDH ($\approx 38\,\mathrm{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E=30\,\mathrm{kPa}$, $E=80\,\mathrm{kPa}$ and TCP (red rectangle).

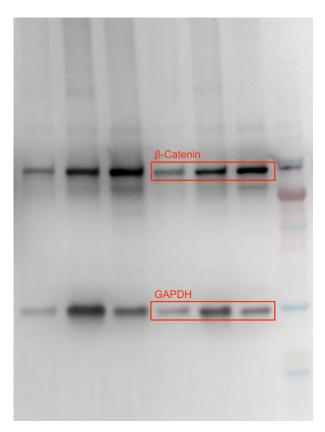


Figure S7. Stiffness-dependent expression of beta-catenin (\approx 92 kDa) and loading control GAPDH (\approx 38 kDa) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E=30\,\mathrm{kPa}$, $E=80\,\mathrm{kPa}$ and TCP (red rectangles).

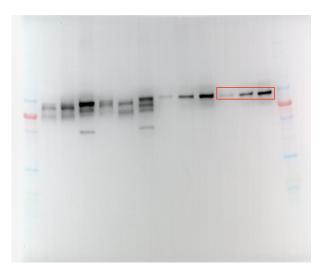


Figure S8. Stiffness-dependent expression of Non-phospho (active) beta-catenin (Ser45) (\approx 92 kDa) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E=30\,\mathrm{kPa}$, $E=80\,\mathrm{kPa}$ and TCP (red rectangle).



Figure S9. Loading control GAPDH ($\approx 38 \,\mathrm{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \,\mathrm{kPa}$, $E = 80 \,\mathrm{kPa}$ and TCP (red rectangle).

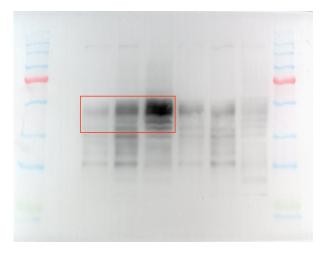


Figure S10. Stiffness-dependent expression of MITF ($\approx 50 \, \text{kDa}$ to $75 \, \text{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \, \text{kPa}$, $E = 80 \, \text{kPa}$ and TCP (red rectangle).



Figure S12. Stiffness-dependent expression of Dicer ($\approx 220 \,\mathrm{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \,\mathrm{kPa}$, $E = 80 \,\mathrm{kPa}$ and TCP (red rectangle).

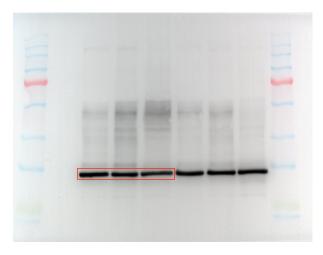


Figure S11. Loading control GAPDH ($\approx 38 \, \text{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \, \text{kPa}$, $E = 80 \, \text{kPa}$ and TCP (red rectangle).

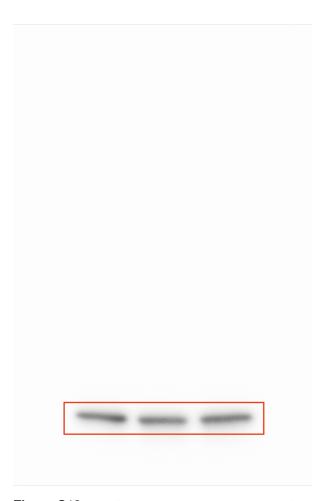


Figure S13. Loading control GAPDH ($\approx 38 \, \text{kDa}$) in ARPE-19 cells (full western blot) with the three lanes of interest from left to right representing $E = 30 \, \text{kPa}$, $E = 80 \, \text{kPa}$ and TCP (red rectangle).