Grade	Key tissue features observed
0: surface and	Matrix: normal architecture
cartilage morphology intacted	Cells: intact and appropriate orientation
1: surface intacted	Matrix: superficial zone intact, oedema and/or superficial
	fibrillation, focal superficial matrix condensation.
	Cells: death, proliferation (clusters), hypertrophy, superficial
	zone – deeper than only the superficial zone
2: surface discontinuity	Matrix discontinuity at superficial zone (deep fibrillation);
	Cationic stain matrix depletion (Safranin O or Toluidine Blue)
	upper third of cartilage; Focal perichondronal increased stain
	(mid zone); Disorientation of chondron columns
	Cells: death, proliferation (clusters), hypertrophy
	Matrix vertical fissures into mid zone, branched fissures:
3: vertical fissures (clefts)	Cationic stain depletion (Safranin O or Toluidine Blue) into
	lower two-thirds of cartilage (deep zone); New collagen
	formation (polarized light
	microscopy, Picro Sirius Red stain)
	Cells: death, regeneration (clusters), hypertrophy, cartilage
	domains adjacent to fissures
4: erosion	Cartilage matrix loss: delamination of superficial layer, mid
	layer cyst formation
	Excavation: matrix loss superficial layer and mid zone
	Surface: sclerotic bone or reparative tissue including
5: denudation	fibrocartilage within denuded surface. Microfracture with

6: deformation

repair limited to bone surface Bone remodeling (more than osteophyte formation only). Includes: microfracture with fibrocartilaginous and osseous

repair extending above the previous surface

Table S1. Summary of the OARSI scoring table for rabbit articular cartilage used to evaluate defects (adapted from Pritzker et al, 2006 and Laverty et al, 2010) [30,31].

Parameters	Key features and score	
Filling of defect	111%-125%	1
	91%-110%	0
relative to surface of	76%–90%	1
normal adjacent	51%-75%	2
cartilage	26%-50%	3
	< 25%	4
Integration of repair	Normal continuity and integration	0
tissue with	Decreased cellularity	1
surrounding articular	Gap or lack of continuity on one side	2
cartilage	Gap or lack of continuity on two sides	3
Matrix staining with	Normal	0
	Slightly reduced	1
Safranin O-fast green	Moderately reduced	2
Jananin O-last green	Substantially reduced	3
	None	4
	(a) Normal	0
	(b) Mostly round cells with the morphology of	
	chondrocytes	
Cellular morphology	>75% of tissue with columns in radial zone	0
	25%–75% of tissue with columns in radial zone	1
(choose first between	< 25% of tissue with columns in radial zone (disorganized)	2
a-b-c-d)	(c) 50% round cells with the morphology of chondrocytes	
	>75% of tissue with columns in radial zone	2
	25%–75% of tissue with columns in radial zone	3
	< 25% of tissue with columns in radial zone (disorganized)	4
	(d) Mostly spindle-shape (fibroblast-like) cells	5
	Normal	0
Architecture within entire defect (excluding margins)	1–3 small voids	1
	1–3 large voids	2
	> 3 large voids	3
	Clefts or fibrillation	4
Architecture of surface	Normal	0
	Slight fibrillation or irregularity	1
	Moderate fibrillation or irregularity	2
	Severe fibrillation or disruption	3
Percentage of new subchondral bone	90%-100%	0
	75%-89%	1
	50%-74%	2
	25%-49%	3
	< 25%	4
Formation of tidemark	Completed	0
	75%–99%	1
	50%-74%	2
	25%-49%	3
	< 25%	4

Table S2. Summary of the Sellers score used to evaluate cartilage regeneration in treated defects. (according to Sellers et al, 1997) [36].