	Examin	ation Meas	ures an	d Freq	uency								
Measurement		Enrollment Visit											
			12 -mo	<b>18- mo</b> <sup>1</sup>	24 - mo	30- mo <sup>1</sup>	36 - mo	48 - mo	60- mo	72- mo	84- mo	96- mo	
Blood collection, fasting <sup>2, 12</sup>													
- Blood draw for serum		X	X	X	X	X	X	X		X			
- Blood draw for plasma and buffy coat		X	X	X	X	X	X	X		X			
- Blood draw for RNA			X					X					
- Blood draw for lymphocyte extraction and cryopreservation								X		X <sup>14</sup>			
Urine collection													
- Fasting second AM void		X	X	X	X	X	X	X		X			
- Pregnancy test for premenopausal women	X	X	X	X	X	X	X	X		X		X	
Height, standing	X				X			X				X	
Weight	X		X		X		X	X		X		X	
Knee size screen for MRI knee coil	X		$X^3$	3	3	3	3	3		$X^3$		$X^3$	
Body size screen for MRI bore	X		X	X	X	X	X						
Abdominal circumference <sup>4</sup>		X	2.2		X			X				X	
Hand examination (DIP bony enlargements) 4	X												
Hallux valgus												X	
Knee examination <sup>4</sup>													
- Alignment (by goniometer)		X	X		X		X						
- Anserine bursa tenderness		X	X				X	X					
- Effusion		X			X			X					

Examination Measures and Frequency													
Measurement	Screening Visit	Enrollment Visit	Follow-up Visit										
			12 -mo	18- mo <sup>1</sup>	24 - mo	30- mo <sup>1</sup>	36 - mo	48 - mo	60- mo	72- mo	84- mo	96- mo	
- Flexion contracture and hyperextension		X											
- Tibiofemoral joint line tenderness		X	X		X		X	X					
- Knee flexion pain/tenderness		X											
- Patellar tenderness		X	X		X		X	X					
- Patellar quadriceps tenderness/tendinitis		X											
- Patello-femoral crepitus		X	X		X		X	X					
- Medial-lateral laxity					X		X						
- Knee pain location (knee pain map)					X							X	
Thrust, walk <sup>4</sup>			X		X		X						
Blood pressure, seated			X		X		X	X		X		X	
Resting heart rate <sup>4</sup>		X			X			X		X		X	
Performance Measures / Physical Activity													
- 20-meter timed walk	X	X	X		X		X	X		X		X	
- 400-meter timed walk <sup>4</sup>		X			X			X				X	
- Chair stands, timed		X	X		X		X	X		X		X	
- Isometric quadriceps and hamstring strength <sup>4</sup>		X	$X^5$		X		$X^5$	X					
- Accelerometry <sup>13</sup>								$X^{13}$		$X^{17}$			

	Examin	ation Meas	ures an	d Freq	uency	7								
Measurement	Screening Visit	Enrollment Visit												
			12 -mo	18- mo <sup>1</sup>	24 - mo	30- mo <sup>1</sup>	36 - mo	48 - mo	60- mo	72- mo	84- mo	96- mo		
MRI														
- Knee, right and left			X	X <sup>6</sup> (Unilat)	X	X <sup>6</sup> (Unilat)	$X^7$	X		X		X		
- Thigh, right and left <sup>4</sup>		X	$X^5$		X		$X^5$	X				X		
X-ray	X													
- Knee: bilateral PA fixed flexion view	X		X		X		X	X		X		X		
- Knee: unilateral fluoroscopic-guided view (one or both knees) <sup>8</sup>		8	8		X <sup>8</sup>									
- Knee: unilateral lateral view (both knees) 9		<sup>9</sup> X	5				$X^9$	5						
- Hip: AP pelvis view	X		$X^5$				X	X				X		
- Hand: dominant PA hand		X X	$X^5$					X						
- Bilateral full limb for mechanical alignment 10	X		$X^{10}$		$X^5$		$X^5$	5						
- Spine	X						X							
DXA														
- Knee (bilateral)						$X^{11}$	11	11		$X^{15}$		$X^{18}$		
- Hip						$X^{11}$	11 2	11		$X^{15}$		$X^{16}$		
- Body composition						X		(		$X^{16}$				

<sup>&</sup>lt;sup>1</sup>This visit obtained in a subset of progression cohort participants at either the 18-month or 30-month follow-up visit

<sup>&</sup>lt;sup>2</sup> Most participants will have AM blood draws after an overnight fast; a small percent will have PM blood draws after a minimum 2 hour fast. AM vs PM blood draws will be consistent for the same participant across visits.

<sup>&</sup>lt;sup>3</sup> Optional

<sup>&</sup>lt;sup>4</sup>The majority of the Reference (Non-exposed) controls had a streamlined clinic visit which did not include the following exams: abdominal circumference, hand examination, knee examination, walk thrust, resting heart rate, 400-meter timed walk, isometric strength, and thigh MRI.

<sup>&</sup>lt;sup>5</sup> To be obtained if participant eligible and a valid measurement was not obtained at the previous visit.

<sup>&</sup>lt;sup>6</sup> Obtained in the knee that had the extended set of sequences at baseline, usually the right knee. 30-month visit may include trabecular sequences.

<sup>&</sup>lt;sup>7</sup> Shorter MRI protocol is an option. Shorter protocol is defined as a scan of one knee (usually right) that had the extended set of sequences at previous visit. If both knees scanned, MRI may include trabecular sequences.

<sup>&</sup>lt;sup>8</sup> Obtained in a subset of Progression subcohort participants at 2 clinical centers.

<sup>&</sup>lt;sup>9</sup>Obtained in Reference (Non-exposed) controls.

<sup>&</sup>lt;sup>10</sup> Obtained in the Progression and Incidence subcohort participants.

<sup>&</sup>lt;sup>11</sup>BMD of bilateral knees and unilateral hip obtained in a subset of Progression subcohort participants at 1) either the 30-month or 36-month follow-up visit and 2) a follow-up measure at the 48-month follow-up visit.

<sup>&</sup>lt;sup>12</sup> Selected assays for retention purposes only at clinic's discretion.

<sup>&</sup>lt;sup>13</sup> Funded by D. Dunlop/P.Semanik ancillary study.

<sup>&</sup>lt;sup>14</sup> To be obtained if not obtained at the 48-month follow-up visit.

<sup>&</sup>lt;sup>15</sup> BMD of bilateral knees and unilateral hip obtained in the incidence and non-exposed control cohort, and those progression cohort participants enrolled in the McAlindon bone ancillary study who did not have an MRI with trabecular sequences at the 48-month follow-up visit.

<sup>&</sup>lt;sup>16</sup>Body composition and hip DXA only if clinic wants to for retention purposes (not funded)

<sup>&</sup>lt;sup>17</sup> Participants who completed the accelerometry measurement at the 48-month follow-up visit will be asked to do the accelerometry measurement again at the 72-month follow-up visit

<sup>&</sup>lt;sup>18</sup> To be obtained in the incidence cohort participants who had knee DXA at the 72-month follow-up visit