#### Supplementary Table 1: Likelihood ratios and post-test probabilities for EBV (Model 1 and

#### Model 3)

Model 2	L		Model 3		
Score	LR+	Post-test probability % *	Score	LR+	Post-test probability % *
		(95% CI)			(95% CI)
>= 0	1.00	8.0 (4.6 to 11.4)	>= 0	1.00	8.0 (4.7 to 11.3)
>= 4	1.00	8.0 (4.6 to 11.5)	>= 3	1.00	8.0 (4.7 to 11.4)
>= 6	1.05	8.4 (4.8 to 12)	>= 5	1.01	8.1 (4.7 to 11.4)
>= 7	1.10	8.7 (5 to 12.4)	>= 6	1.08	8.5 (5.1 to 12.0)
>= 9	1.16	9.1 (5.2 to 13)	>= 7	1.13	8.9 (5.4 to 12.5)
>= 11	1.18	9.3 (5.3 to 13.3)	>= 8	1.17	9.3 (5.6 to 12.9)
>= 12	1.19	9.4 (5.4 to 13.4)	>= 12	1.20	9.4 (5.7 to 13.2)
>= 14	1.21	9.5 (5.4 to 13.6)	>= 13	1.21	9.5 (5.8 to 13.3)
>= 15	1.18	9.3 (5.2 to 13.3)	>= 14	1.22	9.6 (5.9 to 13.4)
>= 16	1.19	9.4 (5.3 to 13.5)	>= 15	1.24	9.8 (6.0 to 13.6)
>= 17	1.21	9.5 (5.3 to 13.7)	>= 16	1.22	9.6 (5.8 to 13.3)
>= 18	1.22	9.6 (5.4 to 13.8)	>= 17	1.24	9.7 (5.9 to 13.5)
>= 20	1.51	11.6 (6.3 to 16.9)	>= 18	1.48	11.4 (7.3 to 15.5)
>= 21	1.65	12.6 (6.9 to 18.3)	>= 19	1.78	13.4 (8.9 to 17.9)
>= 22	2.09	15.4 (8.3 to 22.4)	>= 20	1.89	14.1 (9.5 to 18.8)
>= 23	2.11	15.5 (8.4 to 22.6)	>= 21	2.07	15.2 (10.3 to 20.1)
>= 24	2.40	17.3 (9.3 to 25.3)	>= 22	2.15	15.7 (10.7 to 20.7)
>= 25	2.47	17.7 (9.5 to 25.9)	>= 23	2.09	15.4 (10.3 to 20.4)
>= 26	2.41	17.3 (9.1 to 25.6)	>= 24	2.17	15.9 (10.6 to 21.2)
>= 27	2.44	17.5 (9.2 to 25.9)	>= 25	2.20	16.1 (10.5 to 21.6)
>= 28	2.74	19.2 (10.2 to 28.3)	>= 26	2.54	18.1 (12.3 to 23.9)
>= 29	3.48	23.2 (12.2 to 34.2)	>= 27	2.70	19.0 (13.0 to 25.0)

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Model 1	L		Model 3			
>= 31	4.04	26.0 (14.0 to 38.1)	>= 28	3.30	22.3 (15.9 to 28.7)	
>= 32	4.27	27.1 (14.6 to 39.5)	>= 29	3.57	23.7 (17.1 to 30.3)	
>= 33	4.84	29.6 (15.8 to 43.4)	>= 31	4.40	27.7 (20.5 to 34.9)	
>= 34	5.19	31.1 (16.8 to 45.4)	>= 33	4.57	28.4 (20.9 to 36.0)	
>= 35	5.39	31.9 (17.4 to 46.5)	>= 34	5.73	33.2 (24.3 to 42.1)	
>= 37	6.10	34.6 (19.2 to 50.1)	>= 35	6.61	36.5 (23.7 to 49.2)	
>= 38	5.34	31.7 (15.5 to 47.9)	>= 36	6.29	35.4 (22.3 to 48.4)	
>= 40	5.71	33.2 (15.7 to 50.7)	>= 38	6.78	37.1 (23.3 to 50.9)	
>= 41	5.50	32.3 (14.3 to 50.4)	>= 39	7.34	39.0 (24.6 to 53.3)	
>= 42	6.68	36.7 (17.1 to 56.4)	>= 40	8.01	41.0 (26.2 to 55.9)	
>= 44	6.01	34.3 (14.5 to 54.1)	>= 41	7.21	38.5 (23.3 to 53.7)	
>= 45	5.34	31.7 (11.9 to 51.6)	>= 43	7.05	38.0 (21.8 to 54.2)	
>= 46	5.95	34.1 (11.7 to 56.4)	>= 44	10.57	47.9 (28.7 to 67.1)	
>= 49	11.22	49.4 (19.2 to 79.5)	>= 45	8.81	43.4 (18.6 to 68.2)	
>= 51	15.58	57.5 (22.6 to 92.5)	>= 53	8.81	43.4 (0.01 to 99.9)	
>= 53	18.70	61.9 (22.3 to 99.9)				
>= 56	14.02	54.9 (10.4 to 99.4)				
>= 57	18.70	61.9 (5.8 to 99.9)				

 $^{\ast}$  Based on pre-test probability of 8%

### Supplementary Table 2: Likelihood ratios and post-test probabilities for EBV (Model 2 and

#### Model 4)

Model 2			Model 4			
Score	LR+	Post-test probability % * (95% CI)	Score	LR+	Post-test probability % * (95% CI)	
≥0	1	8.0 (4.8 to 11.2)	≥0	1	8.0 (4.8- to 11.2)	
≥2	3.27	22.1 (12.1 to 32.2)	≥1	2.14	15.7 (9.2 to 22.2)	
≥5	3.49	23.3 (11.6 to 35.0)	≥2	2.70	19.0 (10.0 to 28.0)	
≥7	12.56	52.2 ( 24.5 to 79.9)	≥3	3.48	23.2 (8.5 to 37.9)	

\*Based on pre-test probability of 8%

Supplementary Table 3:Likelihood ratios and post-test probabilities for positive EBV in

#### validation cohort

Score	Positive	Post-test Prob	Post-test Prob (95%	Post-test Prob (95%
	Likelihood	(95% Cl) 8% prior	CI) 12% prior	CI) 16% prior
	Ratio (LR+)			

≥0	1	8.0 (4.8 to 11.2)	12.0 (10.7 to 13.3)	16.0 (14.5 to 17.5)
≥1	2.14	15.7 (9.2 to 22.2)	22.6 (15.1 to 30.1)	28.9 (20.8 to 37.1)
≥2	2.70	19.0 (10.0 to 28.0)	26.9 (16.8 to 37.1)	34.0 (23.1 to 44.9)
≥3	3.48	23.2 (8.5 to 37.9)	32.2 (15.9 to 48.4)	39.8 (22.8 to 56.9)

### Supplementary Table 4: Post-test probabilities incorporating atypical lymphocytosis as a

### diagnostic test in series (Model 1 and Model 3)

Model 1	Pre-test	Post-test probability	Model 3	Pre-test probability	Post-test probability %
Score	probability %	%	Score	%	(95% CI)
	(95% CI)	(95% CI)		(95% CI)	
>= 0	8.0 (4.6 to 11.4)	83.9 (79.3 to 88.6)	>= 0	8.0 (4.7 to 11.3)	83.9 (79.4 to 88.4)
>= 4	8.0 (4.6 to 11.5)	84.0 (79.3 to 88.7)	>= 3	8.0 (4.7 to 11.4)	84.0 (79.5 to 88.5)
>= 6	8.4 (4.8 to 12)	84.6 (79.9 to 89.3)	>= 5	8.1 (4.7 to 11.4)	84.0 (79.6 to 88.5)
>= 7	8.7 (5 to 12.4)	85.1 (80.4 to 89.8)	>= 6	8.5 (5.1 to 12.0)	84.9 (80.4 to 89.3)
>= 9	9.1 (5.2 to 13)	85.8 (81.0 to 90.5)	>= 7	8.9 (5.4 to 12.5)	85.5 (81.1 to 89.9)
>= 11	9.3 (5.3 to 13.3)	86.0 (81.3 to 90.8)	>= 8	9.3 (5.6 to 12.9)	86.0 (81.6 to 90.4)
>= 12	9.4 (5.4 to 13.4)	86.2 (81.4 to 90.9)	>= 12	9.4 (5.7 to 13.2)	86.2 (81.8 to 90.6)
>= 14	9.5 (5.4 to 13.6)	86.3 (81.5 to 91.1)	>= 13	9.5 (5.8 to 13.3)	86.3 (82.0 to 90.7)
>= 15	9.3 (5.2 to 13.3)	86.0 (81.1 to 90.9)	>= 14	9.6 (5.9 to 13.4)	86.5 (82.1 to 90.8)
>= 16	9.4 (5.3 to 13.5)	86.1 (81.2 to 91.0)	>= 15	9.8 (6.0 to 13.6)	86.7 (82.3 to 91.0)
>= 17	9.5 (5.3 to 13.7)	86.3 (81.4 to 91.2)	>= 16	9.6 (5.8 to 13.3)	86.4 (82.0 to 90.8)
>= 18	9.6 (5.4 to 13.8)	86.4 (81.5 to 91.3)	>= 17	9.7 (5.9 to 13.5)	86.6 (82.2 to 91.0)
>= 20	11.6 (6.3 to 16.9)	88.7 (83.5 to 93.9)	>= 18	11.4 (7.3 to 15.5)	88.5 (84.4 to 92.7)
>= 21	12.6 (6.9 to 18.3)	89.6 (84.4 to 94.9)	>= 19	13.4 (8.9 to 17.9)	90.3 (86.3 to 94.2)
>= 22	15.4 (8.3 to 22.4)	91.6 (86.2 to 97.0)	>= 20	14.1 (9.5 to 18.8)	90.8 (87.0 to 94.7)
>= 23	15.5 (8.4 to 22.6)	91.7 (86.3 to 97.1)	>= 21	15.2 (10.3 to 20.1)	91.5 (87.7 to 95.3)
>= 24	17.3 (9.3 to 25.3)	92.6 (87.1 to 98.2)	>= 22	15.7 (10.7 to 20.7)	91.8 (88.0 to 95.6)
>= 25	17.7 (9.5 to 25.9)	92.8 (87.3 to 98.4)	>= 23	15.4 (10.3 to 20.4)	91.6 (87.7 to 95.5)
>= 26	17.3 (9.1 to 25.6)	92.6 (86.9 to 98.3)	>= 24	15.9 (10.6 to 21.2)	91.9 (88.0 to 95.8)
>= 27	17.5 (9.2 to 25.9)	92.7 (87.0 to 98.4)	>= 25	16.1 (10.5 to 21.6)	92.0 (87.9 to 96.1)
>= 28	19.2 (10.2 to 28.3)	93.5 (87.8 to 99.1)	>= 26	18.1 (12.3 to 23.9)	93.0 (89.1 to 96.8)
>= 29	23.2 (12.2 to 34.2)	94.8 (89.0 to 100)	>= 27	19.0 (13.0 to 25.0)	93.4 (89.6 to 97.2)

>= 31	26.0 (14.0 to 38.1)	95.5 (89.8 to 100)	>= 28	22.3 (15.9 to 28.7)	94.5 (91.0 to 98.0)
>= 32	27.1 (14.6 to 39.5)	95.7 (90.0 to 100)	>= 29	23.7 (17.1 to 30.3)	94.9 (91.5 to 98.3)
>= 33	29.6 (15.8 to 43.4)	96.2 (90.4 to 100)	>= 31	27.7 (20.5 to 34.9)	95.8 (92.6 to 99.1)
>= 34	31.1 (16.8 to 45.4)	96.4 (90.7 to 100)	>= 33	28.4 (20.9 to 36.0)	96.0 (92.7 to 99.3)
>= 35	31.9 (17.4 to 46.5)	96.6 (90.9 to 100)	>= 34	33.2 (24.3 to 42.1)	96.8 (93.4 to 100)
>= 37	34.6 (19.2 to 50.1)	97.0 (91.4 to 100)	>= 35	36.5 (23.7 to 49.2)	97.2 (92.8 to 100)
>= 38	31.7 (15.5 to 47.9)	96.5 (90.2 to 100)	>= 36	35.4 (22.3 to 48.4)	97.0 (92.4 to 100)
>= 40	33.2 (15.7 to 50.7)	96.8 (90.2 to 100)	>= 38	37.1 (23.3 to 50.9)	97.3 (92.6 to 100)
>= 41	32.3 (14.3 to 50.4)	96.6 (89.7 to 100)	>= 39	39.0 (24.6 to 53.3)	97.5 (92.8 to 100)
>= 42	36.7 (17.1 to 56.4)	97.2 (90.5 to 100)	>= 40	41.0 (26.2 to 55.9)	97.7 (93.1 to 100)
>= 44	34.3 (14.5 to 54.1)	96.9 (89.7 to 100)	>= 41	38.5 (23.3 to 53.7)	97.4 (92.4 to 100)
>= 45	31.7 (11.9 to 51.6)	96.5 (88.7 to 100)	>= 43	38.0 (21.8 to 54.2)	97.4 (92.0 to 100)
>= 46	34.1 (11.7 to 56.4)	96.9 (88.7 to 100)	>= 44	47.9 (28.7 to 67.1)	98.2 (93.1 to 100)
>= 49	49.4 (19.2 to 79.5)	98.3 (90.6 to 100)	>= 45	43.4 (18.6 to 68.2)	97.9 (90.7 to 100)
>= 51	57.5 (22.6 to 92.5)	98.8 (91.0 to 100)	>= 53	43.4 (0.01 to 99.9)	97.9 (77.5 to 100)
>= 53	61.9 (22.3 to 99.9)	99.0 (90.8 to 100)			
>= 56	54.9 (10.4 to 99.4)	98.7 (88.3 to 100)			
>= 57	61.9 (5.8 to 99.9)	99.0 (87.4 to 100)			

Pre-test probability is based on the results of application of the CPR

Post-test probability is calculated using the pre-test probability and a LR+ of 60.03 for atypical lymphocytosis

# Supplementary Table 5: Post-test probabilities incorporating atypical lymphocytosis for

## positive EBV (Model 2 and Model 4)

Model 2			Model 4					
Score	Pre-test probability	Post-test probability	Score	Pre-test probability	Post-test probability %			
	% (95% CI)	% (95% CI)		% (95% CI)	(95% CI)			
≥0	8.0 (4.8 to 11.2)	83.9 (79.6 to 88.2)	≥0	8.0 (4.8 to 11.2)	83.9 (80.0 to 87.9)			
≥2	22.1 (12.1 to 32.2)	94.5 (88.9 to 100)	≥1	15.7 (9.2 to 22.2)	91.8 (86.8 to 96.7)			
≥5	23.3 (11.6 to 35.0)	94.8 (88.6 to 100)	≥2	19.0 (10.0 to 28.0)	93.4 (87.7 to 99.1)			
≥7	52.2 ( 24.5 to 79.9)	98.5 (91.7 to 100)	≥3	23.2 (8.5 to 37.9)	94.8 (87.0 to 100)			

Pre-test probability is based on the results of application of the CPR

Post-test probability is calculated using the pre-test probability and a LR+ of 60.03 for atypical lymphocytosis

# Supplementary Table 6: The number of patients with and without EBV at each level of each point score

	Model 1		Model 2		Model 3		Model 4		Model 4	
							(derivatio	n)	(validation	)
Total	EBV	No EBV	EBV	No EBV	EBV	No EBV	EBV	No EBV	EBV	No EBV
Score/	n (%)	n (%)	n (%)	n (%)	n (%)	n (%)	n (%)	n (%)	n (%)	n (%)
Points										
0	0	1 (100)	13 (5.8)	211		1 (100)	13	194	147	1410
				(94.2)			(6.28)	(93.72)	(9.44)	(90.56)
1	0	0	0	0	0	0	7	37	42	234
							(15.91)	(84.09)	(15.22)	(84.78)
2	0	0	4 (25)	12 (75)	0	0	10	32	106	242
							(23.81)	(76.19)	(30.46)	(69.54)
3	0	0	0	0	0	1 (100)	11	22	84	77
							(33.33)	(66.67)	(52.17)	(47.83)
4	0	10 (100)	0	0	0	0	0	0	0	0
5	0	0	8	31	0	14 (100)	0	0	0	0
			(20.51)	(79.49)						
6	0	8 (100)	0	0	0	10 (100)	0	0	0	0
7	0	10 (100)	8	5 (38.46)	0	8	0	0	0	0
,	0	10 (100)	(61.54)	5 (58.40)	Ū	0	U	U	U	0
8	0	0	01.34)	0	0	4 (100)	0	0	0	0
9	0	4 (100)	0	0	0	0	0	0	0	0
11	0	2 (100)	0	0	0	0	0	0	0	0
12	0	1 (100)	0	0	0	2 (100)	0	0	0	0
13	0	0	0	0	0	2 (100)	0	0	0	0
14	0	1 (100)	0	0	0	3 (100)	0	0	0	0
15	0	2 (100)	0	0	1 (25)	3 (75)	0	0	0	0
16	0	1 (100)	0	0	0	3 (100)	0	0	0	0
17	0	1 (100)	0	0	3 (6)	47 (94)	0	0	0	0
10	2 (4 26)	45	-		4 (2 57)	27	-	0	-	-
18	2 (4.26)	45	0	0	1 (3.57)	27	0	0	0	0
		(95.74)	-		4 (0.00)	(96.43)	-	-	-	~
19	0	0	0	0	1 (8.33)	11	0	0	0	0
20	0	11 (100)	0	0	1 (7.60)	(91.67)	0	0	0	0
20	0	11 (100)	0	0	1 (7.69)	12 (92.31)	0	0	0	0
21	1 (3.33)	29	0	0	0	(92.31)	0	0	0	0
21	1 (3.33)	29 (96.67)	U	U	U	2 (100)	U	U	U	U
		(10.07)								

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22	0	1 (100)	0	0	1	2	0	0	0	0
					(33.33)	(66.67)				
23	1 (6.67)	14	0	0	0	3 (100)	0	0	0	0
		(93.33)								
24	0	1 (100)	0	0	0	1 (100)	0	0	0	0
25	1 (25)	3 (75)	0	0	1 (7.14)	13	0	0	0	0
						(92.86)				
26	1 (25)	3 (75)	0	0	2	10	0	0	0	0
					(16.67)	(83.33)				
27	1 (100)	0	0	0	0	9 (100)	0	0	0	0
28	0	4 (100)	0	0	0	3 (100)	0	0	0	0
29	1 (6.67)	14	0	0	1 (10)	9 (90)	0	0	0	0
		(93.33)								
31	1	2	0	0	0	1 (100)	0	0	0	0
	(33.33)	(66.67)								
32	1	6	0	0	0	0	0	0	0	0
	(14.29)	(85.71)								
33	0	1 (100)	0	0	1 (12.5)	7 (87.5)	0	0	0	0
34	0	2 (100)	0	0	1 (20)	4 (80)	0	0	0	0
35	0	3 (100)	0	0	2 (50)	2 (50)	0	0	0	0
36	0	0	0	0	0	1 (100)	0	0	0	0
37	3 (50)	3 (50)	0	0	0	0	0	0	0	0
38	1 (25)	3 (75)	0	0	0	1 (100)	0	0	0	0
39	0	0	0	0	0	1 (100)	0	0	0	0
40	0	8 (100)	0	0	1 (100)	0	0	0	0	0
41	0	1 (100)	0	0	1 (50)	1 (50)	0	0	0	0
42	1 (12.5)	7 (87.5)	0	0	0	0	0	0	0	0
43	0	0	0	0	2	5	0	0	0	0
					(28.57)	(71.43)				
44	0	2 (100)	0	0	2	1	0	0	0	0
					(66.67)	(33.33)				
45	0	1 (100)	0	0	3 (50)	3 (50)	0	0	0	0
46	1	2	0	0	0	0	0	0	0	0
	(33.33)	(66.67)								
49	1 (50)	1 (50)	0	0	0	0	0	0	0	0
51	1 (50)	1 (50)	0	0	0	0	0	0	0	0
53	1 (100)	0	0	0	0	1 (100)	0	0	0	0
56	1 (50)	1 (50)	0	0	0	0	0	0	0	0
57	1	5	0	0	0	0	0	0	0	0
	(16.67)	(83.33)								
64	1 (100)	0	0	0	0	0	0	0	0	0

71	0	0	0	0	1 (2.38)	0	0	0	0	0
74	1 (100)	0	0	0	0	0	0	0	0	0