

eTable 3. Predictive performance of different definitions of progression independent of relapse activity against the risk of reaching EDSS score = 6.0 in 15978 patients with age < 37 years and baseline EDSS \leq 5.5

Definition	Sensitivity	Specificity	NPV	PPV	Accuracy
B _{fix} -24w-90d	56.6 (54.2-59.1)	41.1 (39.8-42.4)	76.2 (74.6-77.8)	22.2 (20.9-23.4)	44.6
B _{fix} -24w-180d	43.1 (40.7-45.6)	51.6 (50.2-52.9)	75.4 (74.0-76.8)	20.9 (19.5-22.2)	49.6
B _{fix} -24w-ABS	26.6 (24.4-28.8)	72.2 (71.0-73.4)	76.9 (75.7-78.0)	22.1 (20.2-24.0)	61.8
B _{fix} -48w-90d	56.6 (54.2-59.1)	41.7 (40.4-43.0)	76.1 (74.5-77.7)	22.7 (21.4-24.0)	45.2
B _{fix} -48w-180d	43.1 (40.7-45.6)	51.4 (50.0-52.7)	74.9 (73.5-76.3)	21.1 (19.7-22.5)	49.4
B _{fix} -48w-ABS	26.6 (24.4-28.8)	72.6 (71.4-73.8)	76.6 (75.4-77.8)	22.7 (20.8-24.6)	61.9
B _{fix} -Sust-90d	59.3 (56.6-62.0)	32.5 (30.7-34.3)	63.1 (60.5-65.6)	29.1 (27.3-30.9)	41.0
B _{fix} -Sust-180d	46.4 (43.6-49.1)	42.9 (41.0-44.8)	63.1 (60.9-65.3)	27.5 (25.6-29.4)	44.0
B _{fix} -Sust-ABS	28.1 (25.6-30.6)	69.4 (67.7-71.2)	67.4 (65.6-69.1)	30.1 (27.5-32.7)	56.3
B _{rov} -24w-90d	56.6 (54.2-59.1)	38.3 (37.2-39.5)	8.3 (79.0-81.6)	16.6 (15.6-17.6)	41.6
B _{rov} -24w-180d	43.1 (40.7-45.6)	47.6 (46.4-48.7)	79.4 (78.2-80.6)	15.1 (14.1-16.2)	46.8
B _{rov} -24w-ABS	26.6 (24.4-28.8)	67.7 (66.6-68.8)	81.0 (80.0-81.9)	15.2 (13.8-16.5)	60.4
B _{rov} -48w-90d	56.6 (54.2-59.1)	39.6 (38.4-40.7)	80.5 (79.2-81.9)	17.1 (16.1-18.1)	42.7
B _{rov} -48w-180d	43.1 (40.7-45.6)	47.4 (46.3-48.6)	79.1 (77.9-80.3)	15.3 (14.3-16.4)	46.7
B _{rov} -48w-ABS	26.6 (24.4-28.8)	68.7 (67.6-69.7)	80.9 (79.9-81.9)	15.8 (14.4-17.2)	61.6
B _{rov} -Sust-90d	60.4 (57.6-63.1)	29.5 (28.0-31.0)	68.6 (66.3-70.9)	22.6 (21.1-24.0)	37.3
B _{rov} -Sust-180d	44.7 (44.9-50.5)	38.3 (36.7-39.9)	68.3 (66.3-70.3)	20.8 (19.3-22.3)	40.7
B _{rov} -Sust-ABS	29.0 (26.5-31.5)	64.5 (62.9-66.0)	72.8 (71.2-74.3)	21.7 (19.8-23.7)	55.5

Legend to eTable 3

EDSS: Expanded Disability Status Scale, NPV: Negative Predictive Value; PPV: Positive Predictive Value; B_{fix} : fixed baseline; B_{rov} : roving baseline; 24w: 24-week confirmation; 48w: 48-week confirmation; Sust: sustained disability accrual; 90d: no relapses ≤ 90 days before and ≤ 30 days after the event score; 180d: no relapses ≤ 180 days before and ≤ 30 days after the event score; ABS: no relapses from baseline to the confirmation score