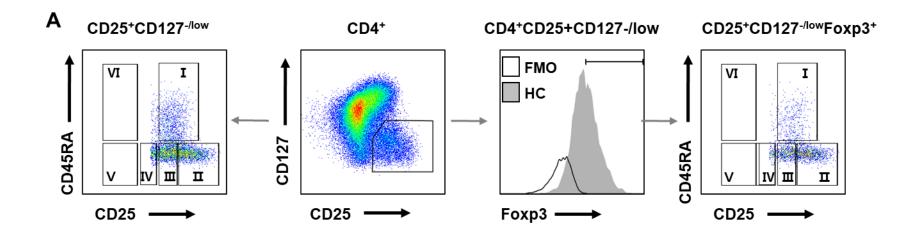
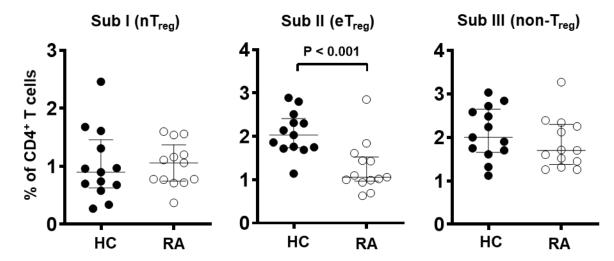
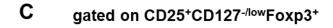
Figure S1.









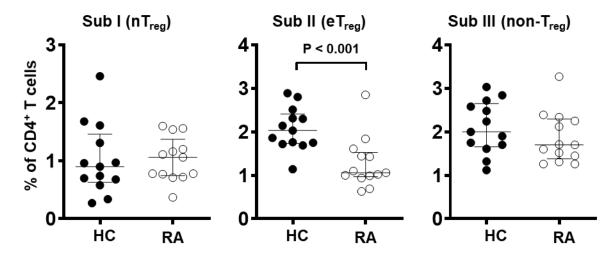
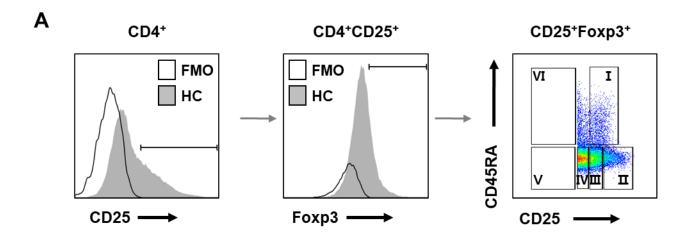


Figure S1. Frequency of effector T_{reg} cells among total T_{reg} cells defined as CD25+CD127-/low or CD25+CD127-/lowFoxp3+ is decreased in PBMCs of RA patients. Blood samples were collected from healthy donors (HC, n = 13) and RA patients (RA, n = 13) and T_{reg} cell subpopulations were analyzed by flow cytometry. (A) Representative plots show the gating strategy used to identify subgroup I (CD25int CD45RA+ cells), II (CD25hiCD45RA- cells), and III (CD25intCD45RA- cells) in CD25+CD127-/low or CD25+CD127-/lowFoxp3+ T_{reg} cells by flow cytometry. (B) Frequency of subgroup I, II, and III in T_{reg} cells defined by CD25+CD127-/low Foxp3+. Statistical differences were calculated by Mann-Whitney test.

Figure S2.



B gated on CD25*Foxp3*

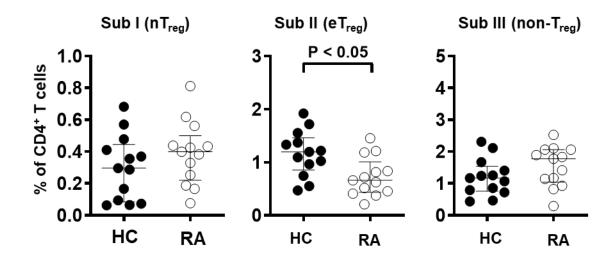


Figure S2. Frequency of effector T_{reg} cells among CD25⁺Foxp3⁺ T_{reg} cells is decreased in PBMCs of RA patients. (A) Representative plots show the gating strategy used to identify subgroup I (CD25^{int}CD45RA⁺ cells), II (CD25^{int}CD45RA⁻ cells), and III (CD25^{int}CD45RA⁻ cells) in CD25⁺Foxp3⁺ Treg cells by flow cytometry. FMO (fluorescence minus one control); HC (healthy control). (B) Frequency of subgroup I, II, and III in CD25⁺Foxp3⁺ T_{reg} cells of PBMCs from healthy controls (HC, n = 13) and RA patients (RA, n = 13). Statistical differences were calculated by Mann-Whitney test.

Figure S3.

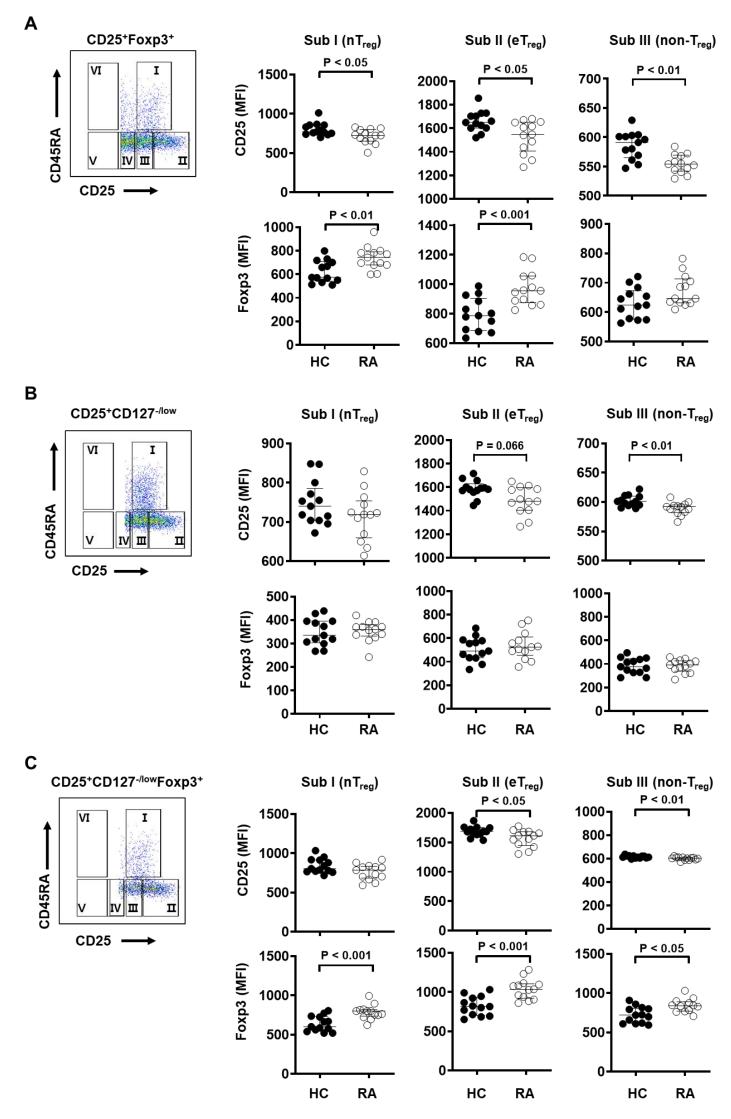


Figure S3. Naïve and effector T_{reg} cells from RA patients show decreased CD25 expression. Expression of CD25 and Foxp3 was measured by flow cytometry in T_{reg} cell subgroups I (CD25^{int}CD45RA+ cells), II (CD25^{int}CD45RA- cells), and III (CD25^{int}CD45RA- cells) in PBMCs from healthy controls (HC, n = 13) and RA patients (RA, n = 13). T_{reg} cells were defined as (A) CD25+Foxp3+, (B) CD25+CD127-/low, or (C) CD25+CD127-/lowFoxp3+. Statistical differences were calculated by Mann-Whitney test.

Table S1. Characteristics of Controls and RA subjects

	Controls	RA subjects	
Variables	(N = 13)	(N = 13)	Unit
Gender	9 (69.2 %)	9 (69.2 %)	Female n (%)
Age	43 (26-70)	52 (22-78)	Median yr (range ¹)
Body-mass index ²	22.4 (19.4-28.1)	20.2 (17.7-31.6)	Median (range ¹)
Disease activity (DAS28 score)) na³	2.7 (0.5-4.5)	Median (range ¹)
Remission (< 2.6))	6 (46.2)	N (%)
Mild (2.6-3.2))	2 (15.4)	N (%)
Moderate (3.2-5.1))	5 (38.4)	N (%)
Severe (>5.1))	0 (0)	N (%)

¹ Minimum to maximum
² Body-mass index is the weight in kilograms divided by the square of the height in meters.
³ not applicable