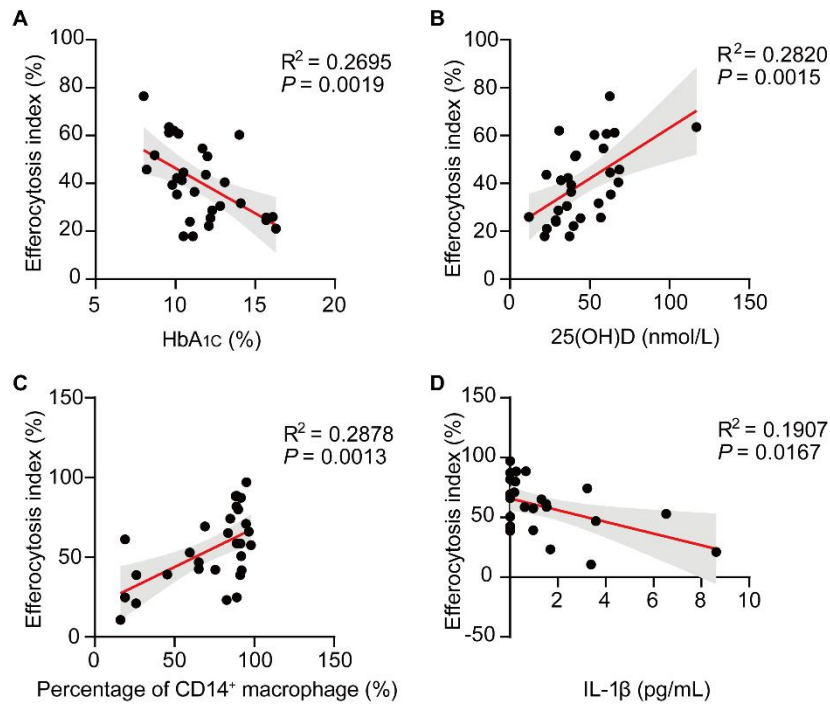


**Supplementary Figure 1** Gating strategy for identification of monocyte/macrophage efferocytosis.

Monocytes/macrophages were incubated with CFSE-labeled apoptotic cells for 2 hours and were then stained with anti-CD14 or anti-CD11b antibodies respectively. The efferocytosis index was calculated as follows: (number of CFSE+CD14+ monocytes/number of total CD14+ monocytes)  $\times$  100 (%), or (number of CFSE+CD11b+ macrophages/number of total CD11b+ macrophages)  $\times$  100 (%). A blank control as well as a isotype control as controls for gating, with two replicate wells per sample to increase the stability and reliability of the experiment.



**Supplementary Figure 2** Parameters correlated with the efferocytosis index of monocytes and monocyte-derived macrophages in patients with type 2 diabetes. Univariate regression analysis of the relationships between each variable and efferocytosis index of monocyte (A-B) and monocyte-derived macrophage (C-D) in diabetes patients. The regression line (black line) and corresponding 95%CI (gray area) are shown for each univariate regression analysis.

**Supplementary Table 1-Monocyte classification in subjects with or without type 2 diabetes**

|                                   |                           | <b>Control</b> | <b>T2D</b>    | <b><i>P</i> value</b> |
|-----------------------------------|---------------------------|----------------|---------------|-----------------------|
| Monocyte count<br>and frequency   | Monocyte numbers          | 0.45 ± 0.23    | 0.41 ± 0.14   | 0.394                 |
|                                   | Monocytes (%)             | 7.58 ± 2.99    | 6.59 ± 2.18   | <b>0.027</b>          |
| Monocyte<br>subset<br>frequencies | Classical monocyte (%)    | 71.79 (16.92)  | 70.29 (17.58) | <b>0.036</b>          |
|                                   | Intermediate monocyte (%) | 3.98 (3.08)    | 3.17 (3.03)   | 0.513                 |
|                                   | Nonclassical monocyte (%) | 2.87 (1.51)    | 2.98 (3.33)   | 0.248                 |

Data are mean ± SD, n (%), or median (IQR). *P* values indicate differences between T2D patients (n=30) and healthy control (n=30).

**Supplementary Table 2** Multivariate linear regression analysis for variables associated with the monocyte efferocytosis index in the total sample population

| <b>Variables</b>                         | <b>Unstandardized Coefficients</b> |           | <b>Standardized Coefficients</b> | <b>t</b> | <b><i>P</i> value</b> | <b>95.0% Confidence Interval for <b>B</b></b> |                    |
|--|------------------------------------|-----------|----------------------------------|----------|-----------------------|---|--------------------|
|  | <b>β</b>                           | <b>SE</b> | <b>stβ</b>                       |          |                       | <b>Lower Bound</b>                            | <b>Upper Bound</b> |
| <b>(Constant)</b>                        | 60.587                             | 11.301    |                                  | 5.361    | < 0.001               | 37.878  | 83.297             |
| <b>HbA<sub>1C</sub></b>                  | -3.792                             | 0.505     | -0.697                           | -7.505   | < 0.001               | -4.807  | -2.777             |
| <b>Percentage of classical monocytes</b> | 0.367                              | 0.144     | 0.236                            | 2.542    | 0.014                 | .077  | .657               |

Age, waist-to-hip ratio, HbA<sub>1C</sub>, FPG, percentage of classical monocytes and serum 25(OH)D concentrations were introduced as independent variables in the stepwise linear regression analysis.

**Supplementary Table 3** Multivariate linear regression analysis for variables associated with the macrophage efferocytosis index in the total sample population

| Variables                                   | Unstandardized Coefficients |       | Standardized Coefficients | t     | P value | 95.0% Confidence Interval for B |             |
|---|-----------------------------|-------|---------------------------|-------|---------|---------------------------------|-------------|
|   | $\beta$                     | SE    | st $\beta$                |       |         | Lower Bound                     | Upper Bound |
| (Constant)                                  | 17.460                      | 9.051 |                           | 1.929 | .061    | -.806                           | 35.726      |
| Percentage of CD14 <sup>+</sup> macrophages | 0.575                       | 0.106 | 0.641                     | 5.419 | < 0.001 | .361                            | .789        |

Age, HbA<sub>1C</sub>, FPG, percentage of CD14<sup>+</sup> macrophages and serum IL- $\beta$  concentrations, were introduced as independent variables in the stepwise linear regression analysis.

**Supplementary Table 4** Multivariate linear regression analysis for variables associated with monocyte efferocytosis index in patients with type 2 diabetes

| Variables         | Unstandardized Coefficients |        | Standardized Coefficients | t      | P value | 95.0% Confidence Interval for B |             |
|-------------------|-----------------------------|--------|---------------------------|--------|---------|---------------------------------|-------------|
|                   | $\beta$                     | SE     | st $\beta$                |        |         | Lower Bound                     | Upper Bound |
| (Constant)        | 57.581                      | 15.476 |                           | 3.268  | <0.001  | 25.827                          | 89.334      |
| 25(OH)D           | 0.314                       | 0.118  | 0.482                     | 3.013  | 0.013   | .071                            | .556        |
| HbA <sub>1C</sub> | -2.729                      | 1.073  | 0.344                     | -2.149 | 0.017   | -4.932                          | -.526       |

Age, HbA<sub>1C</sub> and 25(OH)D levels were introduced as independent variables in stepwise linear regression analysis.

**Supplementary Table 5** Multivariate linear regression analysis for variables associated with monocyte efferocytosis index in patients with type 2 diabetes

| Variables                                   | Unstandardized Coefficients |        | Standardized Coefficients | t     | P value | 95.0% Confidence Interval for B |             |
|---|-----------------------------|--------|---------------------------|-------|---------|---------------------------------|-------------|
|   | $\beta$                     | SE     | st $\beta$                |       |         | Lower Bound                     | Upper Bound |
| (Constant)                                  | 19.662                      | 12.149 |                           | 1.189 | 0.249   | -2.624                          | 41.947      |
| Percentage of CD14 <sup>+</sup> macrophages | 0.534                       | 0.138  | 0.627                     | 3.862 | 0.001   | .248                            | .819        |

Age, percentage of CD14<sup>+</sup> macrophage, IL-1 $\beta$  levels were introduced as independent variables in stepwise linear regression analysis.

**Supplementary Table 6** Participant characteristics of the healthy controls (CON) and diabetes patients with HbA1c lower than 11.0% (D1) or higher than 11.0% (D2)

| Parameter                | CON           | D1             | D2                            |
|--------------------------|---------------|----------------|-------------------------------|
| N (females)              | 30 (8)        | 14 (5)         | 16 (4)                        |
| Age (years)              | 50 ± 10       | 50 ± 10        | 51±11                         |
| BMI (kg/m <sup>2</sup> ) | 22.83 ± 2.37  | 26.66 ± 4.47** | 24.23 ± 3.36                  |
| Waist to hip ratio       | 0.90 ± 0.09   | 0.98 ± 0.04**  | 0.98 ± 0.08 <sup>†</sup>      |
| FPG (mmol/L)             | 5.07 ± 0.42   | 9.00 ± 1.81**  | 11.08 ± 3.81 <sup>*, †</sup>  |
| HbA <sub>1C</sub> (%)    | 5.1 ± 0.4     | 9.8 ± 0.9***   | 13.3 ± 1.8 <sup>***, ††</sup> |
| hsCRP (mg/dl)            | 0.50 (1.50)   | 2.00 (5.75) ** | 2.00 (4.00) *                 |
| TNF- $\alpha$ (pg/ml)    | 1.00 (1.08)   | 1.09 (0.70)    | 0.64 (0.54)                   |
| IL-1 $\beta$ (pg/ml)     | 0.17 (0.95)   | 0.82 (1.37)    | 0.00 (1.31)                   |
| IL-6 (pg/ml)             | 2.25 (3.27)   | 2.38 (2.73)    | 3.18 (1.18)                   |
| IL-8 (pg/ml)             | 5.37 (5.05)   | 5.55 (7.85)    | 11.09 (9.33) *                |
| IL-10 (pg/ml)            | 1.14 (1.38)   | 0.94 (1.03)    | 1.31 (1.42)                   |
| IL-12 (pg/ml)            | 0.22 (0.97)   | 0.10 (0.53)    | 0.32 (0.92)                   |
| 25(OH)D (nmol/L)         | 50.59 ± 20.70 | 52.10 ± 23.85  | 40.28 ± 14.71                 |

Data are mean ± SD, n (%), or median (IQR). \*  $P < 0.05$ , \*\*  $P < 0.01$ , \*\*\*  $P < 0.001$  vs CON. <sup>†</sup>  $P < 0.05$ , <sup>††</sup>  $P < 0.01$  vs D1 from one-way ANOVA or Kruskal-Wallis rank test for normally and nonnormally distributed continuous variables, respectively. CON-control subjects, D1-diabetes patients with HbA<sub>1C</sub> < 11%, D2-diabetes patients with HbA<sub>1C</sub> ≥ 11%.