SUPPORTING INFORMATION

Table SI1 Alanine transaminase (ALT) and aspartate transaminase (AST) of plasma collected from rats treated with acoustic pulses involving various inter-pulse periods (IPP).

| Enzyme Activity (IU/L) at Indicated Inter-Pulse Period | | | | | | | |
|--|--------------|---------------|--------------|--------------|--------------|-------------|--|
| | 72 ms | 400 ms | 800 ms | 2000 ms | 4000 ms | Control | |
| ALT | 90 ± 70 | 118 ± 120 | 159 ± 97 | 127 ± 40 | 180 ± 82 | 45 ± 8 | |
| AST | 205 ± 30 | 225 ± 102 | 239 ± 75 | 217 ± 69 | 236 ± 58 | 87 ± 21 | |

Table SI2 Alanine transaminase (ALT) and aspartate transaminase (AST) of plasma collected from rats treated with various pulse-train US exposure regimens. ('m,n': US pulsing 'on' time, 'off' time; C+:continuous US with PFR 13.9 Hz, C+':continuous US with PFR 55.6 Hz; details shown in Table 2)

| Enzyme Activity (IU/L) at Indicated pulse-train US exposure regimens | | | | | | | | | | |
|--|--------------|--------------|--------------|--------------|-------------|---------------|--------------|--------------|--------------|------------|
| | C+ | '1 s, 2 s' | '2 s, 2 s' | '2 s, 3 s' | '3 s, 2 s' | C+' | '1 s, 2 s' | '2 s, 2 s' | '2 s, 3 s' | '3 s, 2 s' |
| ALT | 65 ± 22 | 60 ± 15 | 63 ± 23 | 84 ± 29 | 72 ± 35 | 117 ± 35 | 73 ± 16 | 80 ± 24 | 80 ± 16 | 74 ± 16 |
| AST | 215 ± 61 | 206 ± 43 | 220 ± 68 | 228 ± 61 | 242 ± 68 | 344 ± 116 | 240 ± 41 | 235 ± 48 | 227 ± 37 | 245 ± 32 |

Table SI3 The extents of hepatic necrosis damage on day 1 from variable treatments (150 µg plasmid).

| MB concentration | US PNP (MPa) | Necrotic area / total area |
|------------------|------------------|----------------------------|
| 0 | 0 | 0 |
| 10% | 0 | 0 |
| 0 | 2.7 | 1.5% |
| 0.5% | 2.7 | 6.0% |
| 1.5% | 2.7 | 6.8% |
| 10% | 1.0 | 3.7% |
| 10% | 2.0 | 9.1% |
| 10% | 2.7 | 10.4% |
| 10% | 3.3 | 14.6% |
| 10% | 2.7(pulse-train) | 5.1% |

Table SI4 Summary of various animal experiments.

| Experiment | Number of groups | Animal numbers per group |
|---|------------------|--------------------------|
| Comparison of gene transfer enhancement with focused and unfocused transducer | 3 | 8 |
| Gene delivery mediated by US with MBs | 4 | 6 |
| Dependence of transgene expression on MBs concentration | 8 | 6 |
| Gene transfer efficiency upon various US peak negative pressures (PNP) | 6 | 7-9 |
| Effect of inter-pulse period of US on gene transfer efficiency | 5 | 6 |
| Effect of pulse train US exposure on gene transfer efficiency | 10 | 6-7 |
| Histological assay for liver damage | 2 | 15 |