

Supplementary data

Table 2. Baseline characteristics of our study population versus the baseline characteristics of the excluded patients/non-responders

| | Current study population n = 868 | Excluded patients/ non-responders n = 134 | p-value |
|--------------------------------------|-------------------------------------|---|---------|
| Age | 56 (5.2) | 56 (5.3) | 0.6 |
| Sex: | | | |
| Male | 179 (21) | 31 (23) | |
| Female | 689 (79) | 103 (77) | 0.5 |
| BMI | 26 (4.1) | 26 (3.6) | 0.3 |
| Comorbidities | 1.9 (1.5) | 1.9 (1.8) | 0.7 |
| Number of painful joints: | | | |
| 1 | 262 (30) | 46 (34) | |
| 2 | 370 (43) | 55 (41) | |
| 3 | 125 (14) | 19 (14) | |
| 4 | 111 (13) | 14 (10) | 0.8 |
| Maximal Kellgren and Lawrence score: | | | |
| 0 | 302 (35) | 10 (8) | |
| 1 | 566 (65) | 10 (8) | 0.2 |
| WOMAC subscales | | | |
| Pain | 25 (17) | 29 (19) | 0.01 |
| Function | 23 (17) | 27 (20) | 0.01 |
| Stiffness | 33 (21) | 36 (23) | 0.1 |

WOMAC; Western Ontario and McMaster Universities Osteoarthritis Index. Continuous variables are shown as mean (SD), categorical variables are shown as number (percentage)

Table 5. Sensitivity analyses

| | HR (95% CI) adjusted for maximal K&L at baseline | HR (95% CI) corrected for maximal K&L during follow-up |
|---|---|---|
| Joint Model with slope included for pain | | |
| Pain | 1.08 (1.06–1.10) | 1.07 (1.05–1.09) |
| Pain slope | 225 (0–276×10 ⁶) | 2.46 (0–15×10 ⁶) |
| Sex ^a | 0.60 (0.34–1.05) | 0.68 (0.39–1.20) |
| Age (years) | 1.06 (1.01–1.10) | 1.05 (1.01–1.10) |
| BMI | 0.94 (0.89–0.99) | 0.95 (0.90–1.00) |
| Maximal K&L score | 2.93 (1.53–5.61) | 1.85 (1.38–2.47) |
| Number of painful joints | 0.73 (0.57–0.93) | 0.76 (0.60–0.98) |
| Comorbidities | 0.78 (0.67–0.92) | 0.79 (0.67–0.93) |
| Joint Model with slope included for function | | |
| Function | 1.06 (1.05–1.08) | 1.06 (1.04–1.07) |
| Function slope | 1,801 (0–683×10 ⁶) | 33.5 (0–52×10 ⁶) |
| Sex ^a | 0.75 (0.43–1.30) | 0.81 (0.46–1.40) |
| Age (years) | 1.05 (1.01: 1.10) | 1.05 (1.00: 1.10) |
| BMI | 0.95 (0.90–1.00) | 0.95 (0.90–1.00) |
| Maximal K&L score | 2.95 (1.54–5.64) | 1.85 (1.39–2.46) |
| Number of painful joints | 0.73 (0.57–0.94) | 0.78 (0.61–1.00) |
| Comorbidities | 0.81 (0.69–0.94) | 0.82 (0.71–0.96) |

HR: hazard ratio.

K&L: Kellgren and Lawrence

^a Men as reference category

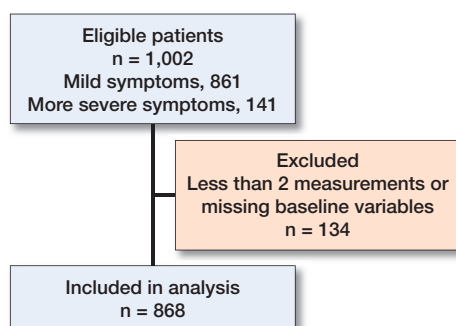


Figure 1. Flow chart.

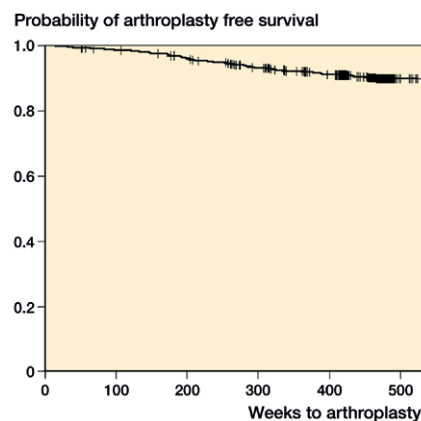


Figure 2. Kaplan–Meier curve for arthroplasty free survival. Censoring is symbolized by “+”.

SUPPLEMENTARY FILE**R-code mixed effects model and joint model.****#Mixed model pain:**

```
fitlmeconv <- lme(womacpain~timetofollowup+gender+age+bmi+max_kl+painfulljoints+comorbidities, random=~timetofollowup |patientid,
data=d2, na.action=na.omit)
```

#Mixed model function:

```
fitlmeconv2 <- lme(womacfunction~timetofollowup +gender+age+bmi+max_kl+painfulljoints+comorbidities, random=~timetofollowup |patientid,
data=d2, na.action=na.omit)
```

#Cox regression:

```
coxFit <- coxph(Surv(timetoevent, arthroplasy) ~ gender + age +bmi+max_kl+painfulljoints+comorbidities, data = ds2, x = TRUE)
```

#Joint Model pain:

```
fit.JM <- jointModel(fitlmeconv, coxFit, timeVar="timetofollowup", method="piecewise-PH-aGH", verbose=TRUE)
summary(fit.JM)
confint(fit.JM, parm = "Event")
round(exp(confint(fit.JM, parm = "Event")), 2)
```

#Joint Model pain + slope:

```
dForm <- list(fixed = ~ 1,
             indFixed = 2,
             random = ~ 1,
             indRandom = 2)
fit.JM.slope <- jointModel(fitlmeconv, coxFit,
                          timeVar="timetofollowup",
                          method="piecewise-PH-aGH",
                          parameterization = "slope",
                          derivForm = dForm,
                          verbose=TRUE)
summary(fit.JM.slope)
```

```
fit.JM.both <- jointModel(fitlmeconv, coxFit,
                          timeVar="timetofollowup",
                          method="piecewise-PH-aGH",
                          parameterization = "both",
                          derivForm = dForm,
                          verbose=TRUE,
                          control = list(GHk = 9))
summary(fit.JM.both)
confint(fit.JM.both, parm = "Event")
round(exp(confint(fit.JM.both, parm = "Event")), 2)
```

#Joint Model function:

```
fit.JM2 <- jointModel(fitlmeconv2, coxFit, timeVar="timetofollowup", method="piecewise-PH-aGH", verbose=TRUE)
summary(fit.JM2)
confint(fit.JM2, parm = "Event")
round(exp(confint(fit.JM2, parm = "Event")), 2)
```

#Joint Model function + slope:

```
fit.JM.slope2 <- jointModel(fitlmeconv2, coxFit,
                           timeVar="timetofollowup",
                           method="piecewise-PH-aGH",
                           parameterization = "slope",
                           derivForm = dForm,
                           verbose=TRUE)
summary(fit.JM.slope2)
```

```
fit.JM.both2 <- jointModel(fitlmeconv2, coxFit,
                           timeVar="timetofollowup",
                           method="piecewise-PH-aGH",
                           parameterization = "both",
                           derivForm = dForm,
                           verbose=TRUE,
                           control = list(GHk = 9))
summary(fit.JM.both2)
confint(fit.JM.both2, parm = "Event")
round(exp(confint(fit.JM.both2, parm = "Event")), 2)
```